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As filed with the Securities and Exchange Commission on February 1, 2021

Registration No. 333-252210

**UNITED STATES
SECURITIES AND EXCHANGE COMMISSION**
Washington, D.C. 20549

AMENDMENT NO.1 TO

FORM F-1

REGISTRATION STATEMENT
UNDER THE SECURITIES ACT OF 1933

Adagene Inc.

(Exact name of Registrant as specified in Its charter)

Not Applicable

(Translation of Registrant's name into English)

Cayman Islands
(State or other jurisdiction of
incorporation or
organization)

2834
(Primary Standard Industrial
Classification Code Number)

Not Applicable
(I.R.S. Employer
Identification Number)

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**Approximate date of commencement of proposed sale to the public:
As soon as practicable after the effective date of this registration statement.**

If any of the securities being registered on this Form are to be offered on a delayed or continuous basis pursuant to Rule 415 under the Securities Act of 1933, check the following box.

If this Form is filed to register additional securities for an offering pursuant to Rule 462(b) under the Securities Act, check the following box and list the Securities Act registration statement number of the earlier effective registration statement for the same offering.

If this Form is a post-effective amendment filed pursuant to Rule 462(c) under the Securities Act, check the following box and list the Securities Act registration statement number of the earlier effective registration statement for the same offering.

If this Form is a post-effective amendment filed pursuant to Rule 462(d) under the Securities Act, check the following box and list the Securities Act registration statement number of the earlier effective registration statement for the same offering.

Indicate by check mark whether the registrant is an emerging growth company as defined in Rule 405 of the Securities Act of 1933.

Emerging growth company

If an emerging growth company that prepares its financial statements in accordance with U.S. GAAP, indicate by check mark if the registrant has elected not to use the extended transition period for complying with any new or revised financial accounting standards† provided pursuant to Section 7(a)(2)(B) of the Securities Act.

CALCULATION OF REGISTRATION FEE

Title of each class of securities to be registered	Proposed maximum aggregate offering price ⁽¹⁾	Amount of registration fee ⁽⁴⁾
Ordinary shares, par value US\$0.0001 per share ⁽²⁾⁽³⁾	US\$125,000,000	US\$13,637.50

- (1) Estimated solely for the purpose of determining the amount of registration fee in accordance with Rule 457(o) under the Securities Act of 1933.
- (2) Includes ordinary shares initially offered and sold outside the United States that may be resold from time to time in the United States either as part of their distribution or within 40 days after the later of the effective date of this registration statement and the date the shares are first bona fide offered to the public, and also includes ordinary shares that may be purchased by the underwriters pursuant to an over-allotment option. These ordinary shares are not being registered for the purpose of sales outside the United States.
- (3) American depositary shares issuable upon deposit of the ordinary shares registered hereby will be registered under a separate registration statement on Form F-6 (Registration No.333-252543). Each American depositary share represents one and one quarter (1.25) ordinary shares.
- (4) Previously paid.

The Registrant hereby amends this registration statement on such date or dates as may be necessary to delay its effective date until the Registrant shall file a further amendment which specifically states that this registration statement shall thereafter become effective in accordance with Section 8(a) of the Securities Act of 1933, as amended, or until the registration statement shall become effective on such date as the Commission, acting pursuant to such Section 8(a), may determine.

† The term "new or revised financial accounting standard" refers to any update issued by the Financial Accounting Standards Board to its Accounting Standards Codification after April 5, 2012.

The information in this prospectus is not complete and may be changed. We may not sell these securities until the registration statement filed with the Securities and Exchange Commission is effective. This prospectus is not an offer to sell these securities and it is not soliciting an offer to buy these securities in any state where the offer or sale is not permitted.

Subject to completion
Preliminary Prospectus dated _____, 2021

American Depositary Shares
ADAGENE
Adagene Inc.

Representing _____ Ordinary Shares

This is an initial public offering of American depositary shares, or ADSs, representing ordinary shares of Adagene Inc.

We are offering _____ ADSs. Each ADS represents _____ of our ordinary shares, par value US\$0.0001 per share.

Prior to this offering, there has been no public market for the ADSs. It is currently estimated that the initial public offering price per share will be between US\$ _____ and US\$ _____.

We have applied for listing the ADSs on the Nasdaq Global Market under the symbol "ADAG."

We are an "emerging growth company" under applicable U.S. federal securities laws and are eligible for reduced public company reporting requirements.

See "Risk Factors" beginning on page 20 for factors you should consider before buying the ADSs.

Neither the United States Securities and Exchange Commission nor any other regulatory body has approved or disapproved of these securities or passed upon the accuracy or adequacy of this prospectus. Any representation to the contrary is a criminal offense.

	Per ADS	Total
Public offering price	US\$	US\$
Underwriting discounts and commissions ⁽¹⁾	US\$	US\$
Proceeds, before expenses, to us	US\$	US\$

(1) See "Underwriting" for additional disclosure regarding compensation payable by us to the underwriters.

The underwriters have a 30-day option to purchase up to an additional _____ ADSs from us at the initial public offering price less the underwriting discount.

The underwriters expect to deliver the ADSs against payment in U.S. dollars in New York, New York on _____, 2021.

**Goldman Sachs
(Asia) L.L.C.**

Morgan Stanley

Jefferies

China Renaissance

The date of this prospectus is _____, 2021.

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We are responsible for the information contained in this prospectus. We have not authorized anyone to provide you with different information, and we take no responsibility for any other information others may give you. We are not, and the underwriters are not, making an offer to sell these securities in any jurisdiction where the offer or sale is not permitted. You should not assume that the information contained in this prospectus is accurate as of any date other than its date.

We have not authorized anyone to provide you with information different from that contained in this prospectus. We are offering to sell, and seeking offers to buy, the ADSs only in jurisdictions where offers and sales are permitted. The information contained in this prospectus is current only as of the date of this prospectus, regardless of the time of delivery of this prospectus or of any sale of the ADSs.

We have not taken any action to permit a public offering of the ADSs outside the United States or to permit the possession or distribution of this prospectus outside the United States. Persons outside the United States who come into possession of this prospectus must inform themselves about and observe any restrictions relating to the offering of the ADSs and the distribution of the prospectus outside the United States.

Until _____, 2021 (the 25th day after the date of this prospectus), all dealers that buy, sell or trade ADSs, whether or not participating in this offering, may be required to deliver a prospectus. This is in addition to the obligation of dealers to deliver a prospectus when acting as underwriters and with respect to their unsold allotments or subscriptions.

PROSPECTUS SUMMARY

The following summary is qualified in its entirety by, and should be read in conjunction with, the more detailed information and financial statements and the related notes appearing elsewhere in this prospectus. In addition to this summary, we urge you to read the entire prospectus carefully, especially the risks of investing in the ADSs discussed under "Risk Factors," "Business," and information contained in "Management's Discussion and Analysis of Financial Condition and Results of Operations" before deciding whether to buy the ADSs. In particular, we call your attention to the risk that we could be delisted from The Nasdaq Global Market pursuant to the Holding Foreign Companies Accountable Act enacted on December 18, 2020 if the Public Company Accounting Oversight Board continues to be unable to inspect our independent registered public accounting firm for three consecutive years.

OVERVIEW

We are a platform-driven, clinical-stage biopharmaceutical company committed to transforming the discovery and development of novel antibody-based cancer immunotherapies. Our platform is designed to generate therapeutic antibody candidates with unique functional epitopes and species cross-reactivity as highlighted by our immunotherapy pipeline. These features enable our novel drug discovery strategy to advance from lead identification through vigorous preclinical modeling to biomarker-guided mono- and combination immunotherapy development in clinical settings. We have pioneered a dynamic interface design to harness the conformational diversity of antibodies, which enlarges epitope sampling of a given drug target for differentiated therapeutic antibody development. We aim to push the boundaries of antibody discovery and engineering through the precise design, construction, and selection of antibody product candidates intractable to traditional antibody technology.

Life is motion. The motion of proteins and their dynamic interactions trigger a cascade of complex biological and pharmacological effects. Our core technology is built upon our fundamental understanding of the role that protein folding and the motion of molecules play in giving rise to dynamic conformational diversity, where an amino acid sequence can adopt multiple structures and functions. Our approach recognizes that a protein's native state is not accurately represented by a single static structure but rather by a variety of structures in dynamic equilibrium, resulting in a high level of functional diversity, in contrast to the conventional static antibody drug discovery paradigm of "one sequence, one structure and one function."

We have developed our proprietary AI-Powered DPL platform to explore the dynamic conformational diversity of protein sequences, and the flexible binding sites of antibody sequences in particular, as a new paradigm for antibody drug discovery. Our DPL platform combines artificial intelligence, or AI, algorithms and ever-increasing big data in antibody sequence, structure, binding epitope and affinity from public and our own proprietary databases to design, construct and screen high-quality proprietary antibody libraries with well-defined sequence, scaffold and biophysical attributes for antibody drug discovery. Powered by computational physics in combination with AI and big data, our DPL platform samples a potentially infinite number of dynamic binding interface structures arising from the conformational diversity of a finite number of antibody amino acid sequences, allowing us to exponentially expand the universe of candidate antibody binding sites far beyond conventional natural or synthetic antibody repertoires. By exploiting conformational diversity, we have designed and precisely constructed approximately one trillion (10^{12}) antibody sequences in our DPL. These antibodies feature broad epitope (the portion of an antigen that are recognized by an antibody) coverage and robust chemistry, manufacturing, and control, or CMC, attributes.

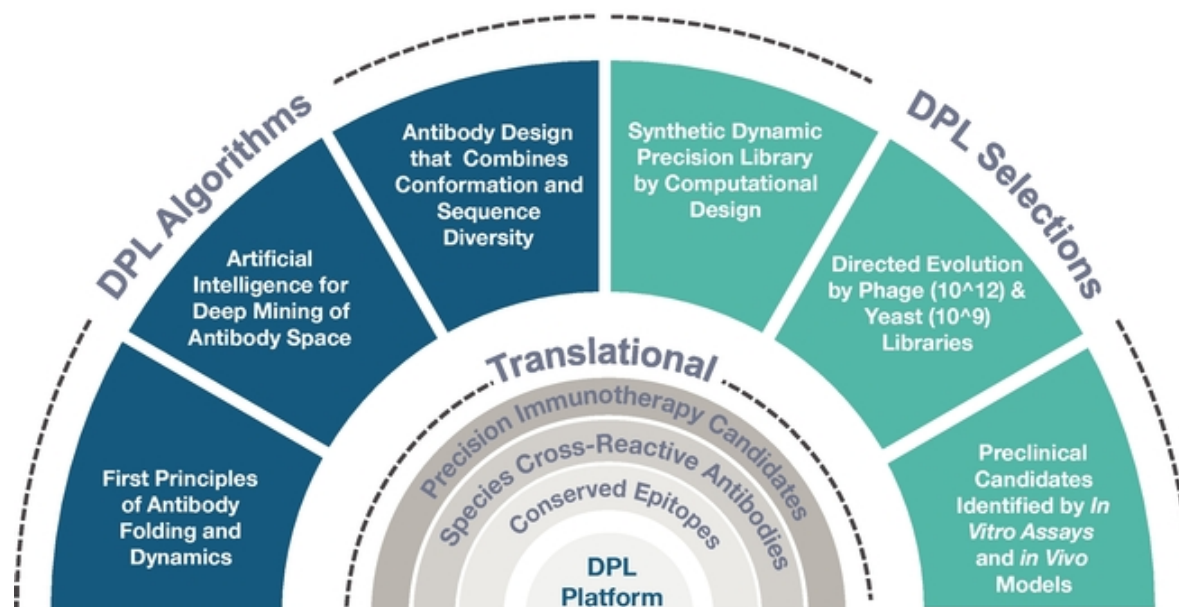
We believe our AI-powered antibody discovery and engineering DPL platform significantly increase R&D productivity for antibody drug discovery, as illustrated by our clinical anti-CD137 and anti-CTLA-4 programs. DPL library screening against CD137 or CTLA-4 antigens has yielded a large number of high affinity primary hits. The abundant discovery hits with diversified binding epitopes show

the power of our AI-powered DPL platform not only in creating novel antibodies in targeting different epitopes of a given antigen, but also in targeting the conserved epitope across different species of a given antigen with broad species cross-reactivity from human, monkey to mouse, which enables us to study their efficacy and safety in extensive immuno-competent or syngeneic animal models, to explore their pharmacodynamics and predictive biomarkers in responding vs nonresponding tumor models *in vivo*, and to understand their deep target biology and novel MOA before testing them in human clinical trials to look for their clinical signals consistent with their MOA. Our DPL platform empowers us to engage the dynamic epitope of the conformationally dynamic target which might be challenging using conventional antibody discovery approaches. We believe that the high-affinity and cross-reactive primary hits from our AI-Powered DPL library screening save time and cost from discovery to early clinical proof of concept. The broad species cross-reactivity of the primary hits also streamlines lead identification and potentially enables high fidelity translation from preclinical to clinical studies. Furthermore, we believe our AI-powered dynamic precision library expands the diversity at the start of discovery to maximize the chance that suitable leads are found on the first pass.

Translational fidelity from preclinical modeling to informed clinical development is one of the top challenges to developing cancer immunotherapies. Most traditionally developed antibodies do not cross react between their human and murine targets due to their limited species cross-reactivity, making it very difficult to reliably evaluate the same antibody in both the preclinical and clinical settings. Some of the most contentious issues related to preclinical and clinical studies of CD137 and CTLA-4, the targets of our lead product candidates, immunotherapies are traceable to the differences between the antibodies used for preclinical and clinical studies. For example, according to Frost & Sullivan, two of the leading clinical anti-CD137 agonist antibodies bind to different epitopes of CD137 and exhibit dramatic differences in their respective clinical safety and efficacy results, underscoring the importance of finding suitable species cross-reactive antibodies like those we have utilized for comprehensive preclinical evaluation before entering clinical trials.

We believe that it is essential to model the interactions between tumors and an intact host immune system *in vivo* to evaluate the therapeutic potential of antibodies in preclinical studies. The flexibility of antibody binding interface is fundamental to the NEObody technology of our DPL Platform and allows us to generate species cross-reactive antibodies to assess the safety and efficacy potential of mono- and combination therapy candidates in syngeneic animal models before launching clinical trials. We use syngenic mouse models which are known for their intact *in vivo* immune systems to provide the original proof of concept for cancer immunotherapies by blocking immune check points with monoclonal antibodies, or mAbs. We believe that the use of species cross-reactive antibodies, rather than surrogate antibodies used in traditional syngeneic animal models, should facilitate the translational relevance and clinical utility of these well-established preclinical models for determining optimal dose, schedule, sequencing, combination synergy, risk and benefit features. The results from the assessment of new species cross-reactive antibodies in rigorous preclinical models may allow us to control the scope and cost of clinical trials, enable the identification of potential clinical biomarkers useful to monitor clinical pharmacological and safety signals, and help preselect patients for precision mono- and combination therapies.

The figure below illustrates how our DPL platform integrates our computational algorithm-enabled high-throughput screening and functional antibody evaluation for preclinical candidates suitable for clinical development as explained above.



Our DPL platform is further composed of three proprietary enabling technologies tailored to three key attributes of antibody-based therapeutic modalities:

- NEObody technology is a fully synthetic phage display and yeast display-based antibody discovery technology, which we believe is differentiated from other synthetic antibody technologies through its innovative designs and precise constructions. NEObody technology enables the generation of antibodies designed with dynamic binding sites that adapt kinetically to unique epitopes, triggering a novel MOA. The species cross-reactive antibodies generated by NEObody technology not only have the potential to reveal new biological functions of the targets, but also facilitate preclinical studies using various immune system intact animal models, resulting in high fidelity translation from preclinical to clinical studies. We refer to antibodies generated by our NEObody technology as NEObodies.
- SAFEbody technology is designed to mask an antibody binding interface with a masking motif, which then prevents an antibody from binding to its target in healthy tissues. The masking motif is designed to activate, or unmask, the antibody to allow binding in the tumor microenvironment, or TME, where certain activation conditions such as a protease is upregulated as compared to healthy tissues, allowing the antibody to bind to its target for tumor killing. Our SAFEbody enabled therapeutic candidates are therefore designed to be activated predominantly in the TME while remaining largely in an inactive state in healthy tissues. Our SAFEbody technology can be applied to mask the binding sites of any antibodies including but not limited to NEObodies. We refer to such masked antibodies including NEObodies as SAFEbodies.
- POWERbody technology, which enables the creation of new versions of bispecific T-cell engagers, or TCEs, with enhanced safety profiles, antibody-drug conjugates, or ADCs, or antibodies that are designed to reach beyond the therapeutic potency of traditional monospecific antibodies.
- SAFEbody technology can be applied to our NEObodies, such as what we did with ADG116 to convert into ADG126. POWERbody is created by applying the SAFEbody technology to a bispecific TCE in either CD3 or both in CD3 and antigen binding arms to enhance the safety

profile. Our POWERbody technology is designed to improve antitumor activity while maintaining the enhanced safety profile.

We believe that comprehensive *in vivo* preclinical evaluations are the key to assess the efficacy and safety potential of tailor-made antibody candidates before progressing them into lengthy and costly clinical trials. NEObody, SAFEbody and POWERbody technologies are all designed to facilitate favorable druggability, manageable CMC attributes, and reduced immunogenicity. As highlighted by our lead product candidates, such as ADG106, ADG126 and ADG116, our NEObody technology allows us to engineer and select species cross-reactive NEObodies designed to dynamically adapt to unique and evolutionally conserved epitopes. ADG126 has been further engineered using our SAFEbody technology to address the safety concerns associated with existing CTLA-4 therapeutics.

The figure below shows how our NEObody, SAFEbody, and POWERbody technologies and DPL platform are inter-connected and utilized for the building of our product pipeline of mono- and combination immunotherapies.



Notes:

- (1) Antibodies generated by NEObody technology, which are designed with dynamic binding sites that adapt kinetically to unique epitopes, triggering a novel MOA.
- (2) NEObodies or traditional antibodies are masked in their binding sites by SAFEbody technology as shown by the blocking bar in the antibody binding sites in the figure, which are designed to be selectively activated in the TME, potentially limiting on-target off-tumor toxicity in normal tissues.
- (3) Multiple potent modalities, including bispecific T-cell engagers, ADC, and Fc engineering for enhanced ADCC, etc., are masked in their binding sites by SAFEbody to create a POWERbody.

Our most advanced NEObody product candidate, ADG106, is a fully human ligand-blocking agonistic anti-CD137 mAb currently being evaluated in Phase Ib clinical trials in the United States and China. ADG106 is designed to target a unique epitope of CD137 that is different from other anti-CD137 antibodies currently under clinical development. Epitope mapping and X-ray structural analysis of ADG106 with CD137 have shown in preclinical studies that ADG106 is capable of binding to CD137 in a fashion similar to its natural ligand, CD137L. Our first SAFEbody product candidate, ADG126, is a fully human anti-CTLA-4 masked antibody. It is designed to enhance the safety features by masking the antibody binding site of ADG126, which would be unmasked in the TME, where the activated ADG126 would block CTLA-4 and deplete regulatory T-cells by means of enhanced antibody-dependent cellular cytotoxicity, or ADCC. In preclinical studies, ADG126 was tolerated at doses of up to 200 mg/kg in nonhuman primate models. As ADG126 is also species cross-reactive in humans,

cynomolgus monkeys and mice, we believe that preclinical studies of ADG126 will support the rational design of clinical trials to expedite its development. Our third product candidate, ADG116, is a fully human anti-CTLA-4 NEObody. Epitope mapping and X-ray structural analysis have shown that in preclinical studies, ADG116 is capable of binding to a novel epitope of CTLA-4 different from ipilimumab, the only CTLA-4 mAb approved globally. The dynamic interface of ADG116 enabled not only its species cross-reactivity with human, cynomolgus monkey, and mouse CTLA-4 for preclinical studies, but also its dynamic engagement on a unique epitope of CTLA-4 to trigger a novel MOA, distinct from ipilimumab, by softer ligand blocking and stronger regulatory T-cell depletion via strong ADCC.

Our species cross-reactive ADG106, ADG126 and ADG116 have enabled a deep understanding of the interaction between tumor and host immune system *in vivo* in syngeneic animal models. This understanding has been utilized to design and guide the clinical development of rational, mechanism-based mono- and combination therapies using ADG106, ADG126 and ADG116. Because there is limited clinical safety and efficacy data available for anti-CD137 agonists we have followed our preclinical and mechanistic study for the clinical development of ADG106. We observed three patients with Grade 3 liver enzyme increase; among them, one patient with abnormal liver enzyme baseline showed a Grade 3 AST increase. ADG106 showed preliminary clinical antitumor activity in patients who have progressed after several lines of treatment in our completed Phase Ia dose escalation and ongoing Phase Ib dose expansion trials in the United States and China. It is very encouraging to observe the clinical response in connection with the changes in PD biomarkers due to target engagement in a dose dependent manner. We have also observed a correlation with a potential predictive biomarker for patient selection in three patients with more than 30% tumor shrinkage across different indications in retrospective analysis in our ongoing Phase I trials. This predictive biomarker is related to the CD137 pathway. We intend to further explore this predictive biomarker in order to guide our development of precision mono- and combination immunotherapies based on our preclinical and preliminary clinical data.

Our Pipeline

By leveraging our proprietary DPL platform, we have developed a robust pipeline of innovative product candidates in various stages of development, ranging from research and discovery to preclinical and clinical development. Our highly differentiated clinical-stage pipeline consists of ADG106, ADG126 and ADG116. We also have a robust preclinical pipeline in various stages of development. In addition, we have out-licensed the Greater China rights of ADG104, a PD-L1 mAb in Phase Ib and Phase II trials concurrently in China, to our partner, Sanjin and its affiliates. We have the right to apply for the patents derived from our core and key technologies pertaining to ADG104 in the rest of the world and we retain a majority of the economic benefits derived from ADG104 in the rest of the world.

The following chart provides an overview of the status of each of our programs at clinical or IND-enabling stages, for which we have global rights:

Drug Target	Trial	Design	Preclinical	IND Enabling	Phase I	Phase II	Phase III	Anticipated Milestones
ADG106 Anti-CD137 NEObody	1001† (US)	Mono	Solid tumors and NHL					End of Phase I meeting in Q1 2021
	1002† (China)	Mono	Solid tumors and NHL					End of Phase I meeting in Q1 2021
	1003 (Australia)	Combo	All tumor types					Phase Ib read out in H2, 2021
	1008 (China)	Combo	All tumor types					Phase Ib read out in H1, 2021
	2001 (Global)	Mono/Combo (Biomarker*)	All tumor types					Phase II read out in H2, 2021
ADG116 Anti-CTLA-4 NEObody	1001 (US)	Mono/Combo	Solid tumors					Expansion pending on ADG116-1003 trial
	1002 (China)	Mono/Combo	Solid tumors					Expansion in selected indications following ADG116-1003 trial
	1003 (Australia)	Mono/Combo	Solid tumors					Safety and efficacy read out in selected indications in H2 2021
ADG126 Anti-CTLA-4 SAFEbody	1001 (Global)	Mono/Combo	Solid tumors					Safety and efficacy read out in selected indications in H2 2021, lead SAFEbody
	1002 (China)	Mono/Combo	Solid tumors					Expansion in selected indications following ADG126-1001 trial

Notes:
† ADG106-1001 and ADG106-1002 trials are currently in Phase Ib dose expansion phase
* denotes biomarker driven patient enrollment
We intend to use clinical data collected from the Australia trials to facilitate clinical development in China and the United States
NHL = non-Hodgkin's lymphoma

ADG106: Novel agonistic anti-CD137 NEObody candidate

ADG106, is a fully human ligand-blocking, agonistic anti-CD137 immunoglobulin G4, or IgG4, mAb generated using our NEObody technology. ADG106 is being developed for the treatment of advanced solid tumors and non-Hodgkin's lymphoma, or NHL. CD137 stimulates the immune system to attack cancer cells and is a key driver for T-cell and natural killer cell, or NK cell, proliferation. ADG106 is designed to target a unique conserved epitope of CD137 with a novel MOA for CD137 agonism by its natural ligand-like binding and cross-linking by Fcγ receptors. The broad species cross-reactivity of ADG106 observed in preclinical studies has enabled us to explore robust translational studies concerning the biology of CD137 in mouse, rat, nonhuman primate, and human, especially for anti-CD137 as a single agent and in combination therapies in immuno-competent syngeneic tumor models. In preclinical studies to date, we observed that ADG106 had encouraging antitumor activity and was well tolerated as a monotherapy and in combination with the existing standard-of-care, or SOC, and other immuno-oncology therapies, which was confirmed in the Phase I clinical trials in cancer patients as a monotherapy. These data indicate that ADG106 has the potential to address the limitations of other existing anti-CD137 therapies.

As of the November 30, 2020, the Data Cut-off Date, we have completed the Phase Ia dose escalation in each of our Phase I studies of ADG106 in both the United States and China as a monotherapy in patients with advanced or metastatic solid tumors and/or NHL. We are currently in the Phase Ib dose expansion phase for both trials in the United States and China. ADG106 was dose escalated up to 10 mg/kg and is well tolerated at dose expansion at 3 mg/kg, 5 mg/kg and at 300mg and 400 mg flat doses. A total number of 92 patients have been dosed. Both clinical trials, namely ADG106-1001 and ADG106-1002, have limited Treatment Emergent Adverse Effects, or TEAEs, liver toxicity or hematologic abnormalities. We observed three patients with ³Grade 3 liver enzyme increase; among them, one patient with abnormal liver enzyme baseline showed a Grade 3 AST increase. Two patients developed hemolytic anemia in ADG106-1002 which were controlled; one of the two patients received prior therapies which were known to cause autoimmune hemolytic anemia. One patient with a solid tumor who previously failed chemotherapies, radiotherapy, and an anti-PD-L1 related antibody treatment showed a partial response to ADG106 treatment with a 40% tumor size

reduction after two treatment cycles. In addition, two NHL patients showed more than a 30% tumor size reduction after one and two ADG106 treatment cycles, respectively. Furthermore, biomarker studies showed target engagement with specific PD biomarkers indicative of immune system activation, and clinical response correlated with changes in CD137 target engagement. These data are encouraging given that the enrolled population was not preselected and was heavily pretreated. We have identified a potential predictive biomarker which correlates with patient response to ADG106 treatment from a retrospective analysis of the ongoing Phase I clinical trial. Based on this finding, we are in the process of preparing an additional Phase II trial which we expect to initiate in 2021 and for which we intend to stratify and preselect patients using this predictive biomarker to potentially enhance clinical response of patients to ADG106 treatment. We also plan to pursue potential registrational trials evaluating ADG106 in biomarker enriched patient populations.

We have also evaluated ADG106 in combination with other therapies including chemotherapies, immune modulators, and immuno-oncology therapies in preclinical studies. Data from combination studies in tumor bearing mice showed that the combination of ADG106 with immune checkpoint inhibitors, including an anti-PD-1/L1 mAb or anti-CTLA-4 mAb, enhanced *in vivo* antitumor activity. We plan to explore the combination of ADG106 with other targeted antibody therapies for the treatment of hematologic malignancies and solid tumors. We also have identified tumor-specific biomarkers that we believe may correlate with ADG106 antitumor activity in multiple mouse tumor models. These preclinical study findings are consistent with the interim results from our ongoing Phase Ib clinical trials. Going forward, we are focusing on evaluating combinations of ADG106 with various therapies in clinical trials. We have received NMPA approval for the ADG106-1008 study in China, which is a Phase Ib/II combination trial of ADG106 with anti-PD-1 in advanced solid tumors and hematological malignancies. We are also in the preparation for a submission of a Clinical Trial Notification, or CTN, for our ADG106-1003 clinical trial in Australia, which will combine ADG106 with anti-PD-1, anti-VEGF, or anti-CD20 in advanced solid tumors and hematological malignancies.

We have observed responses to ADG106 in all three T cell lymphoma patients enrolled in the ADG106-1002 Phase I trial. We are enrolling additional patients in our ADG106-1002 trial and will conduct an investigator initiated trial in the United States to further explore ADG106 as a monotherapy and in combination with other therapies in T cell lymphoma. We expect to submit an Orphan Drug Designation application to the FDA for ADG106 requesting for expedited regulatory approval for this indication.

ADG126: Novel anti-CTLA-4 SAFEbody candidate

Our most advanced SAFEbody program, ADG126, is a masked fully-human anti-CTLA-4 mAb engineered to address the safety concerns associated with existing CTLA-4 therapeutics, while maintaining potency when locally activated in the TME. The FDA approval of ipilimumab validated anti-CTLA-4 for cancer treatment. However, due to its on-target off-tumor toxicity, the approved indications for ipilimumab are limited, which we believe has caused sales of ipilimumab to trail other immuno-oncology therapies such as anti-PD-1/L1 antibodies.

ADG126 is designed to address the toxicity and efficacy issues related to the MOA of existing approved CTLA-4 immuno-oncology therapies and to expand the potential of CTLA-4 as a target for the treatment of cancer. In preclinical studies, ADG126 was tolerated at doses of up to 200 mg/kg in nonhuman primate models. We believe the encouraging preclinical tolerability of ADG126 suggests its potential in combination with other immunotherapies such as an anti-PD-1/PD-L1 antibody or an anti-CD137 antibody, including ADG106.

To better address the unmet clinical need for a safe and potent anti-CTLA-4 antibody for chemotherapy-free mono- and combination immunotherapy, we have submitted a CTN and received HREC approval for an ADG126 Phase I dose escalation trial in Australia and are expecting to commence patient enrollment by February 2021. We also have received approval from the FDA to initiate the Phase I clinical trial of ADG126 in the United States in 2021. These two Phase I dose escalation trials will test five doses at 0.1, £0.3, £1, £3, and £10 mg/kg, based on our preclinical primate toxicity study in comparison with ipilimumab. We anticipate that these clinical trials will allow us to determine the ADG126 effective dose in late 2021. Meanwhile, we are preparing IND submissions to initiate ADG126 clinical trials in China.

ADG116: Novel anti-CTLA-4 NEObody candidate

ADG116 is a fully-human ligand-blocking anti-CTLA-4 mAb generated using our NEObody technology. ADG116 is designed to target a unique conserved epitope of CTLA-4. In preclinical studies, ADG116 was observed to have softer CTLA-4 ligand blocking and stronger ADCC for depleting regulatory T-cells than ipilimumab. In a head-to-head *in vivo* efficacy study, ADG116 was observed to have at least a five-fold greater anti-tumor activities in comparison with ipilimumab. In addition, ADG116 was observed to induce antitumor responses concomitant with reduced immunosuppressive regulatory T-cell and enhanced cytotoxic T lymphocyte (CD8⁺ T-cells) activities in the TME. We believe that these preclinical results support the further clinical evaluation of ADG116 both as monotherapies and combination therapies for a wide range of tumor types.

As of January 15, 2021, we have successfully completed dose escalation at the first three doses in our ADG116-1003 clinical trial in Australia, including a dose level at 0.03mg/kg, the dose level used for the first patient who suffered a fatal incident in the ADG116 Phase I trial in the United States. As of January 15, 2021, no dose limiting toxicity or treatment related serious adverse events have been observed in any of the patients treated. There was no treatment induced liver toxicity and all three patients treated at 0.03mg/kg showed normal liver enzyme level post ADG116 treatment. The Safety Review Committee approved dose escalation for ADG116-1003 clinical trial to 0.1mg/kg. The ADG116-1001 Phase I trial in the United States was placed on clinical hold on September 30, 2019 by the FDA, after we reported to the FDA the death of the only patient dosed in the trial. The FDA removed the clinical hold on December 5, 2019 after we submitted an amendment to the study protocol.

Our Global Partnership and Collaborations

We have a successful track record of collaboration and partnerships with global biopharmaceutical companies and academic institutions. So far, we have established multiple collaboration programs and will continue to seek to continue to seek partnership opportunities where we can leverage our proprietary technology platform to develop novel antibodies to address unmet medical needs. Over the past two years, we have established partnerships and collaborations with multiple biopharmaceutical companies.

- We entered into a material transfer and collaboration and license agreement with ADC Therapeutics SA, or ADC Therapeutics, under which ADC Therapeutics intends to use our SAFEbody technology to generate a masked antibody that could be combined with the pyrrolbenzodiazepine cytotoxic payload technology used in ADC Therapeutics' ADCs for the development of a novel ADC against a solid tumor target. Under the agreement, we could be eligible to receive royalty payments and could have an exclusive option to negotiate a license to develop and commercialize co-developed assets in certain territories; and
- We are also collaborating with Guilin Sanjin Pharmaceutical Co., Ltd., or Sanjin, and its affiliates to develop two different monoclonal antibodies with the first being ADG104, a

monospecific antibody that targets PD-L1 and is in Phase Ib and Phase II clinical trials concurrently in China, and the second being an undisclosed monoclonal antibody, which Sanjin and its affiliates have recently filed an IND application that has been accepted by the NMPA;

- We have recently extended the term of our research collaboration agreement, or the NIH RCA, with the National Heart Lung and Blood Institute, part of the National Institutes of Health, or NIH, to 2023. We first entered into the NIH RCA in March 2018 to collaborate with Dr. Richard Childs, Chief of the Laboratory of Transplantation Immunotherapy at NHLBI, who has used antibodies discovered by us to develop a CAR-T cell therapy candidate targeting a human endogenous retrovirus expressed in the majority of clear cell kidney tumors. The extension of the NIH RCA will allow us to use our proprietary antibody technology to further collaborate with Dr. Richard Childs and his team to develop other antibodies; and
- We entered into a collaboration and license agreement with Exelixis, Inc., or Exelixis, under which Exelixis will utilize our SAFEbody technology platform to generate masked versions of mAbs from Exelixis' growing preclinical pipeline for the development of ADCs or other innovative biologics against Exelixis-nominated targets. Under the agreement, we could be eligible to receive upfront payment as well as milestone and royalty payments on net sales of any potential products developed around each of these targets.

We are also working with global biopharmaceutical companies to potentially develop additional strategic partnerships.

- We had recently worked with Celgene (now Bristol-Myers Squibb) to discover antibodies targeting novel antigens using our proprietary DPL platform;
- Under a material transfer agreement, we are developing SAFEbody drug conjugates against a tumor target selected by Tanabe Research Laboratories, Inc., or TRL, with potential for negotiating a future license agreement with TRL if our pilot work proves successful;
- We recently entered into an evaluation agreement with a global biotechnology company, or the Evaluation Agreement, under which we are developing bispecific antibodies leveraging our SAFEbody and bispecific technologies against tumor targets provided by the counterparty. The Evaluation Agreement could facilitate the parties' interest in a further collaboration concerning the development of a new antibody therapeutic; and
- We had previously collaborated with GlaxoSmithKline (China), or GSK China, where we were engaged to generate high affinity antibodies against multiple epitopes of multi-transmembrane targets; and we previously worked with Jiangsu Hengrui Medicine Company Limited, or Jiangsu Hengrui, where we were able to discover cross-reactive agonistic antibody for immuno-oncology.

Our Team and Investors

We were founded in 2011 by Dr. Peter Luo and is led by an experienced management team. Dr. Luo, who previously founded the biopharmaceutical company Abmaxis which was subsequently acquired by Merck, has a proven track record of more than two decades in antibody discovery and engineering using a multidisciplinary approach that combines computational and experimental technology based on physical, chemical, and biological sciences. Our management team is composed of industry veterans with extensive experience in therapeutic antibody research and development and collectively has decades of experience in molecular biology, immunotherapy, immunology, antibody discovery, protein engineering, and clinical development. Our management team brings a strong history of leadership, innovation, and research and development experience at leading companies, including Merck/Abmaxis, Affomix/Illumina, Amgen, AnaptysBio, Bristol-Myers Squibb, Celgene, Corixa, Forty Seven Inc., Xencor, Novartis, Pfizer, Prometheus, Quantice, and Teva Biopharmaceuticals. Our company is further supported by a strong group of investors that share our commitment to developing

next-generation immuno-oncology therapies for the treatment of cancers. Our investors include leading institutional investors such as F-Prime, Eight Roads, General Atlantic, GP Healthcare Capital, Sequoia China and Wuxi AppTec.

OUR STRATEGIES

We are utilizing our proprietary DPL platform to design, construct and develop novel immunotherapies and precision antibodies to address unmet patient needs globally. Our strategy encompasses the following key elements:

- Advance clinical development of our lead product candidates, ADG106, ADG126 and ADG116, as monotherapies and in combination with other therapies.
- Develop a deep and broad pipeline of promising antibodies using NEObody™, SAFEbody™ and POWERbody™ technologies and advance into preclinical and clinical development.
- Leverage our technology to develop our pipeline and strengthen our DPL platform.
- Continue to collaborate with leading biopharmaceutical companies and academic institutions to discover and develop novel candidates based upon our DPL platform.
- Maximize value creation by advancing our product candidates to potential commercialization in key markets alone or with strategic partners.
- Build global operations for global markets, while leveraging a global supply chain and China cost effectiveness.

RISK FACTORS

Our business is subject to a number of risks and uncertainties, including, among others, the following:

- We have a limited operating history, which may make it difficult to evaluate our current business and predict our future performance.
- We have incurred net losses historically and we may continue to incur net losses in the future.
- We may need to obtain substantial additional financing to fund our growth and operations, which may not be available on acceptable terms, if at all.
- We may not be able to identify or discover new product candidates, and may allocate our limited resources to pursue a particular product candidate and fail to capitalize on product candidates that may later prove to be more profitable, or for which there is a greater likelihood of success.
- We may not be successful in our efforts to use and expand our proprietary platforms to build a pipeline of product candidates.
- We depend substantially on the success of our product candidates, particularly ADG106, ADG126 and ADG116, which are in clinical development, and our ability to identify additional product candidates. If we are unable to successfully identify new product candidates, complete clinical development, obtain regulatory approval and commercialize our product candidates, or experience significant delays in doing so, our business will be materially harmed.
- If we are ultimately unable to obtain regulatory approval for our product candidates, our business will be substantially harmed. Even if we receive regulatory approval for our product candidates, we will be subject to ongoing regulatory obligations and continued regulatory review, which may result in significant additional expense and we may be subject to penalties if we fail

to comply with regulatory requirements or experience unanticipated problems with our product candidates.

- We face intense competition and the possibility that our competitors may develop therapies that are similar, more advanced, or more effective than ours, which may adversely affect our financial condition and our ability to successfully commercialize our product candidates.
- We may not be able to enter into additional collaboration agreements beyond our existing strategic partnerships or collaborations with NIH, Exelixis, ADC Therapeutics and Guilin Sanjin. If we are unable to maintain existing and future strategic partnerships, or if these strategic partnerships are not successful, our business could be adversely affected.
- We may be unsuccessful in obtaining or maintaining adequate patent or other intellectual property protection for one or more of our product candidates, due to the failure to obtain issuance from our owned or licensed patent and trademark applications or to maintain the confidentiality and proprietary nature of our trade secrets, and our issued patents covering one or more of our product candidates could be found invalid or unenforceable if challenged in court or before administrative bodies and a third party could misappropriate our trade secrets or independently develop technology that are highly similar to our trade secrets.
- We have certain shareholders who will have board representation rights after the offering and their individual interests may differ from yours.
- There can be no assurance that we will not be a passive foreign investment company ("PFIC") for U.S. federal income tax purposes for any taxable year, which could subject U.S. investors in our ADSs or ordinary shares to significant adverse U.S. federal income tax consequences.
- The audit report included in this prospectus is prepared by an auditor that is not inspected by the Public Company Accounting Oversight Board and, as such, our investors are deprived of the benefits of such inspection. In addition, various legislative and regulatory developments related to U.S.-listed China based companies due to lack of PCAOB inspection and other developments due to political tensions between the United States and China may have a material adverse impact on our listing and trading in the U.S. and the trading prices of our ADSs, and we could be delisted if we are unable to meet the PCAOB inspection requirements in time.
- We could be delisted if we are unable to meet the PCAOB inspection requirements in time.
- The COVID-19 pandemic could adversely impact our business, including our clinical trials.

We also face other challenges, risks and uncertainties that may materially and adversely affect our business, financial condition, results of operations and prospectus. You should consider the risk discussed in "Risk Factors" and elsewhere in this prospectus before investing in the ADSs.

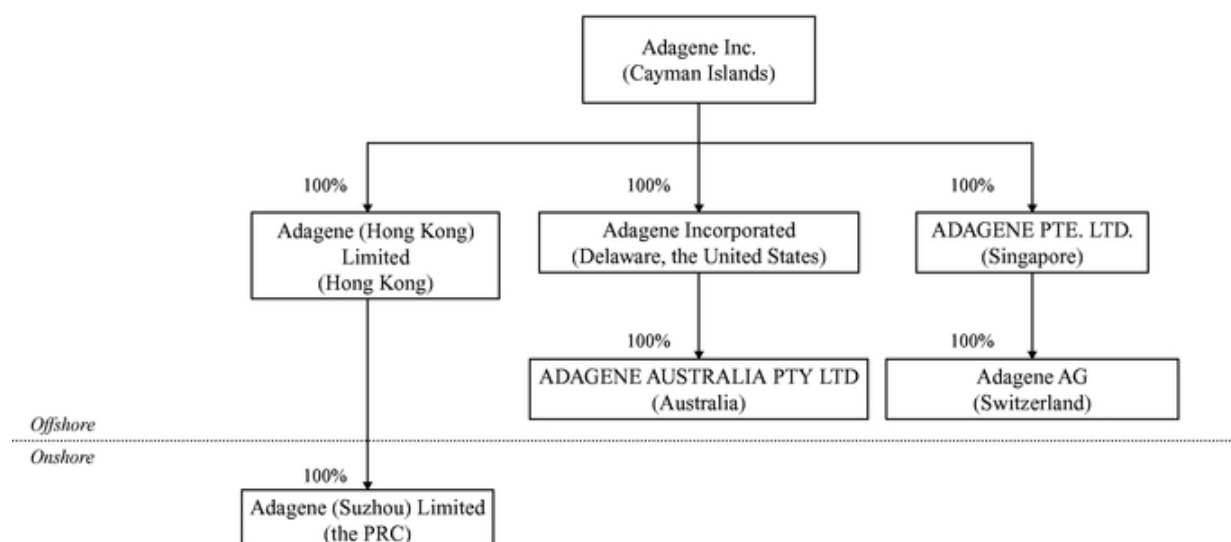
Corporate History and Structure

In February 2011, Adagene Inc. was incorporated under the laws of the Cayman Islands as our offshore holding company.

In December 2011, we established Adagene (Hong Kong) Limited, or Adagene Hong Kong, a wholly-owned subsidiary incorporated under the laws of Hong Kong, as our intermediary holding company. In February 2012, Adagene Hong Kong incorporated Adagene (Suzhou) Limited, or Adagene Suzhou, in China, through which we commenced our research and development activities in China.

In September 2017, we established a wholly-owned subsidiary in the state of Delaware, the United States, Adagene Incorporated, to conduct our research and development activities in the United States to facilitate the discovery and development of product candidates and expand our global presence, we have further incorporated several subsidiaries overseas, such as Australia, Singapore and Switzerland.

The following diagram illustrates our corporate structure as of the date of this prospectus, including our material subsidiaries:



You may refer to "Corporate History and Structure" for more details.

Corporate Information

Our corporate headquarters is located at 4F, Building C14, No. 218, Xinghu Street, Suzhou Industrial Park Suzhou, Jiangsu Province, 215123, People's Republic of China. Our registered office is located at Vistra (Cayman) Limited, P. O. Box 31119 Grand Pavilion, Hibiscus Way, 802 West Bay Road, Grand Cayman, KY1 - 1205 Cayman Islands. Our telephone number is +86-512-8777-3632. Our agent for service of process in the United States is Cogency Global Inc., located at 10E 40th Street, 10th Floor, New York, NY 10016. Our corporate website is www.adagene.com. The information contained on or that can be accessed through our website is not incorporated by reference into this prospectus, and you should not consider information on our website to be part of this prospectus.

IMPLICATIONS OF BEING AN EMERGING GROWTH COMPANY AND A FOREIGN PRIVATE ISSUER

As a company with less than US\$1.07 billion in revenue for the last fiscal year, we qualify as an "emerging growth company" pursuant to the Jumpstart Our Business Startups Act of 2012 (as amended by the Fixing America's Surface Transportation Act of 2015), or the JOBS Act. An emerging growth company may take advantage of specified reduced reporting and other requirements that are otherwise applicable generally to public companies. These provisions include exemption from the auditor attestation requirement under Section 404 of the Sarbanes-Oxley Act of 2002, or Section 404, in the assessment of the emerging growth company's internal control over financial reporting. The JOBS Act also provides that an emerging growth company does not need to comply with any new or revised financial accounting standards until such date that a private company is otherwise required to comply with such new or revised accounting standards. We do not plan to "opt out" of such exemptions afforded to an emerging growth company.

We will remain an emerging growth company until the earliest of (i) the last day of our fiscal year during which we have total annual gross revenues of at least US\$1.07 billion; (ii) the last day of our fiscal year following the fifth anniversary of the completion of this offering; (iii) the date on which we have, during the previous three-year period, issued more than US\$1.0 billion in non-convertible debt;

or (iv) the date on which we are deemed to be a "large accelerated filer" under the Securities Exchange Act of 1934, as amended, or the Exchange Act, which would occur if the market value of our ADSs that are held by non-affiliates exceeds US\$700 million as of the last business day of our most recently completed second fiscal quarter. Once we cease to be an emerging growth company, we will not be entitled to the exemptions provided in the JOBS Act discussed above.

Upon consummation of this offering, we will report under the Securities Exchange Act of 1934, as amended, or the Exchange Act, as a non-U.S. company with foreign private issuer status. As a foreign private issuer, we may take advantage of certain provisions in the Nasdaq listing rules that allow us to follow Cayman Islands law for certain corporate governance matters. Even after we no longer qualify as an emerging growth company, as long as we qualify as a foreign private issuer under the Exchange Act, we will be exempt from certain provisions of the Exchange Act that are applicable to U.S. domestic public companies, including:

- the sections of the Exchange Act regulating the solicitation of proxies, consents or authorizations in respect of a security registered under the Exchange Act;
- the sections of the Exchange Act requiring insiders to file public reports of their stock ownership and trading activities and liability for insiders who profit from trades made in a short period of time;
- the rules under the Exchange Act requiring the filing with the Securities and Exchange Commission, or SEC, of quarterly reports on Form 10-Q containing unaudited financial and other specified information, or current reports on Form 8-K, upon the occurrence of specified significant events; and
- Regulation Fair Disclosure, or Regulation FD, which regulates selective disclosures of material information by issuers.

CONVENTIONS WHICH APPLY TO THIS PROSPECTUS

Unless we indicate otherwise, all information in this prospectus reflects the following:

- no exercise by the underwriters of their over-allotment option to purchase up to additional ADSs representing ordinary shares from us; and

Except where the context otherwise requires and for purposes of this prospectus only:

- "Adagene Suzhou" refers to Adagene (Suzhou) Limited.
- "ADSs" refers to the American depository shares, each representing of our ordinary shares;
- "Antibody binding interface" or "antibody binding sites" refers to the antibody binding surface spots in contact with its recognition antigen;
- "China" or "PRC" refer to the People's Republic of China, excluding, for the purpose of this prospectus only, Taiwan, Hong Kong and Macau;
- "conformational diversity" or "dynamic diversity" refers to the existence of more than one conformation or structure due to dynamic fluctuation of the structures for a given protein sequence, independent of any conformational changes caused by external binding;
- "epitopes" or "epitope of an antigen" refers to the specific binding spots of an antigen in contact with its antibody binding surface;
- "Greater China", for the purpose of this prospectus, refers to the People's Republic of China, Hong Kong, Macau and Taiwan;

- "multi-specificity" refers to a protein exerting a similar function (such as binding) on distinctly different ligands, perhaps while using different active site residues;
- "NEObody" refer to antibody designed with dynamic binding sites that adapt kinetically to unique epitopes through novel MOA, using our NEObody technology;
- "POWERbody" refer to antibody that utilizes our SAFEbody technology to create new bispecific T-cell engagers, antibody-drug conjugates, or antibodies, which are designed to reach beyond the therapeutic potency of traditional monospecific antibodies;
- "RMB" or "Renminbi" refers to the legal currency of the People's Republic of China;
- "SAFEbody" refer to antibody engineered with its binding sites masked, which are designed to be selectively activated in the TME, potentially limiting on-target off-tumor toxicity in normal tissues;
- "species cross-reactivity" refers to reactivity of the same protein that recognizes and binds to similar epitopes of a given class of targets in different species;
- "US\$", "dollars" or "U.S. dollars" refers to the legal currency of the United States; and
- "we," "us," "our company," and "our," refer to Adagene Inc., a Cayman Islands company and its subsidiaries.
- "NEObodies" refer to antibodies designed with dynamic binding sites that adapt kinetically to unique epitopes through novel MOAs, using our NEObody technology;
- "ordinary shares" or "shares" prior to the completion of this offering refers to our ordinary shares of par value US\$0.0001 per share;
- "POWERbodies" refer to antibodies that utilize our SAFEbody technology to create new bispecific T-cell engagers, antibody-drug conjugates, or antibodies, which are designed to reach beyond the therapeutic potency of traditional monospecific antibodies; and
- "SAFEbodies" refer to antibodies engineered with their binding sites masked, which are designed to be selectively activated in the TME, potentially limiting on-target off-tumor toxicity in normal tissues.

This prospectus contains information derived from various public sources and certain information from an industry report dated September 22, 2020 commissioned by us and prepared by Frost & Sullivan, a third-party industry research firm, to provide information regarding our industry and market position. Such information involves a number of assumptions and limitations, and you are cautioned not to give undue weight to these estimates. The industry in which we operate is subject to a high degree of uncertainty and risk due to variety of factors, including those described in the "Risk Factors" section. These and other factors could cause results to differ materially from those expressed in these publications and reports.

THE OFFERING

Offering price	US\$ per ADS.
ADSs offered by us	ADSs (or ADSs if the underwriters exercise their over-allotment option in full).
The ADSs	<p>Each ADS represents ordinary shares, par value US\$0.0001 per share. The depositary will hold the ordinary shares underlying your ADSs. You will have rights as provided in the deposit agreement.</p> <p>We do not expect to pay dividends in the foreseeable future. If, however, we declare dividends on our ordinary shares, the depositary will pay you the cash dividends and other distributions it receives on our ordinary shares, after deducting its fees and expenses in accordance with the terms set forth in the deposit agreement.</p> <p>You may turn in your ADSs to the depositary in exchange for ordinary shares. The depositary will charge you fees for any exchange.</p> <p>We may amend or terminate the deposit agreement without your consent. If you continue to hold your ADSs after an amendment to the deposit agreement, you agree to be bound by the deposit agreement as amended.</p> <p>To better understand the terms of the ADSs, you should carefully read the "Description of American Depositary Shares" section of this prospectus. You should also read the deposit agreement, which is filed as an exhibit to the registration statement that includes this prospectus.</p>
Ordinary shares	<p>We will issue ordinary shares represented by ADSs in this offering.</p> <p>All options, regardless of grant dates, will entitle holders to the equivalent number of ordinary shares once the vesting and exercising conditions on such share-based compensation awards are met.</p> <p>See "Description of Share Capital."</p>
Ordinary shares outstanding immediately after this offering	Immediately upon the completion of this offering, ordinary shares will be outstanding (or ordinary shares if the underwriters exercise their option to purchase additional ADSs in full).
Over-allotment option	We have granted to the underwriters an option, which is exercisable within 30 days from the date of this prospectus, to purchase up to an aggregate of additional ADSs.

Use of proceeds	<p>We expect to receive net proceeds of approximately US\$ million from this offering, after deducting underwriting discounts and commissions and estimated offering expenses payable by us.</p> <p>We plan to use the net proceeds of this offering for (i) approximately 95% for research and development, among which approximately 26% for ADG106 clinical development, approximately 26% for both ADG126 and ADG116 clinical development, and approximately 43% for our technical and platform development and development of our preclinical candidates, and (ii) approximately 5% for working capital and other general corporate purposes. See "Use of Proceeds."</p>
Lockup	<p>We, [our directors, executive officers and existing shareholders] have agreed with the underwriters, subject to certain exceptions, not to sell, transfer or dispose of, directly or indirectly, any of ADSs or ordinary shares or securities convertible into or exercisable or exchangeable for ADSs or ordinary shares for a period of [180] days after the date of this prospectus. See "Shares Eligible for Future Sale" and "Underwriting" for more information.</p>
Nasdaq trading symbol	ADAG.
Payment and settlement	The underwriters expect to deliver the ADSs against payment therefor through the facilities of The Depository Trust Company on , 2020.
Depository	JPMorgan Chase Bank, N.A.
Risk factors	See "Risk Factors" and other information included in this prospectus for discussions of the risks relating to investing in the ADSs. You should carefully consider these risks before deciding to invest in the ADSs.

Unless specifically indicated otherwise or unless the context otherwise requires, all references to our ordinary shares as of the date of this prospectus exclude a total of 2,262,500 ordinary shares, issued but deemed to be not outstanding as of the date of this prospectus, held by Great Han Fortune LP.

OUR SUMMARY CONSOLIDATED FINANCIAL DATA

The following summary consolidated statements of comprehensive loss data for the years ended December 31, 2018 and 2019, summary consolidated balance sheet data as of December 31, 2018 and 2019 and summary consolidated cash flow data for the years ended December 31, 2018 and 2019 have been derived from our audited consolidated financial statements included elsewhere in this prospectus. Our consolidated financial statements are prepared in accordance with accounting principles generally accepted in the United States of America, or U.S. GAAP. The following summary consolidated statements of comprehensive loss for the nine months ended September 30, 2019 and 2020, summary consolidated balance sheet data as of September 30, 2020 and summary consolidated cash flows data for the nine months ended September 30, 2019 and 2020 have been derived from our unaudited interim condensed consolidated financial statements included elsewhere in this prospectus and have been prepared on the same basis as our audited consolidated financial statements and include all adjustments, consisting only of normal and recurring adjustments, that we consider necessary for a fair statement of our financial position and operating results for the periods presented. Our historical results are not necessarily indicative of results expected for future periods. You should read this Summary Consolidated Financial Data section together with our consolidated financial statements and the related notes and "Management's Discussion and Analysis of Financial Condition and Results of Operations" included elsewhere in this prospectus.

Summary Consolidated Statements of Comprehensive Loss Data

The following table presents our summary consolidated statements of comprehensive loss data for the years ended December 31, 2018 and 2019 and our selected unaudited interim condensed

consolidated statements of comprehensive loss data for the nine months ended September 30, 2019 and 2020.

	For the Year Ended December 31,		For the Nine Months Ended September 30,	
	2018	2019	2019	2020
	US\$	US\$	US\$	US\$
	(in thousands)			
Revenue:				
Licensing revenue	1,511	480	—	310
Expenses:				
Research and development expenses	(16,081)	(16,212)	(11,518)	(23,568)
Administrative expenses	(2,765)	(3,438)	(2,268)	(7,448)
Total operating expenses	(18,846)	(19,650)	(13,786)	(31,016)
Loss from operations	(17,335)	(19,170)	(13,786)	(30,706)
Interest income	711	923	712	613
Interest expense	(91)	(138)	(102)	(114)
Other income	902	723	217	655
Foreign exchange gain (loss), net	13	22	(68)	(77)
Change in fair value of warrant liabilities	534	1,207	1,207	—
Loss before income tax	(15,266)	(16,432)	(11,818)	(29,629)
Income tax expense	—	—	—	—
Net loss attributable to Adagene Inc.'s shareholders	(15,266)	(16,432)	(11,818)	(29,629)
Other comprehensive income (loss):				
Foreign currency translation adjustments, net of nil tax	(11)	66	281	(215)
Total comprehensive loss attributable to Adagene Inc.'s shareholders	(15,277)	(16,367)	(11,537)	(29,844)
Net loss attributable to Adagene Inc.'s shareholders	(15,266)	(16,432)	(11,818)	(29,629)
Deemed contribution from convertible redeemable preferred shareholders	1,186	—	—	—
Accretion of convertible redeemable preferred shares to redemption value	(223)	(246)	(184)	(186)
Net loss attributable to ordinary shareholders	(14,303)	(16,678)	(12,002)	(29,815)
Weighted average number of ordinary shares used in per share calculation:				
—Basic	15,159	15,178	15,173	15,757
—Diluted	15,159	15,178	15,173	15,757
Net loss per ordinary share				
—Basic	(0.94)	(1.10)	(0.79)	(1.89)
—Diluted	(0.94)	(1.10)	(0.79)	(1.89)

Summary Consolidated Balance Sheet Data

The following table presents our summary consolidated balance sheet data as of December 31, 2018 and 2019 and our selected unaudited interim consolidated balance sheet data as of September 30, 2019 and 2020.

	As of December 31,		As of September 30,	
	2018	2019	2020	
	Actual	Actual	Actual	Pro forma ⁽¹⁾
	(in USD thousands)			
Current assets:				
Cash and cash equivalents	16,058	92,533	82,895	82,895
Short-term investments	33,000	8,000	—	—
Total current assets	51,817	103,923	86,606	86,606
Total assets	54,417	105,889	89,077	89,077
Current liabilities:				
Amounts due to related parties	3,674	1,896	2,387	2,387
Accruals and other current liabilities	2,574	2,540	2,789	2,789
Short-term borrowings	2,331	717	2,203	2,203
Total current liabilities	10,346	7,181	10,417	10,417
Long-term borrowings	—	1,516	3,135	3,135
Total liabilities	10,488	8,697	13,610	13,610
Total mezzanine equity	84,955	154,201	154,387	—
Total shareholders' equity (deficit)	(41,027)	(57,009)	(78,921)	75,466

Note:

- (1) All of the preferred shares will automatically convert into ordinary shares on a one-on-one basis immediately prior to the completion of this offering. The unaudited pro forma balance sheet information assumes the automatic conversion of all of the outstanding preferred shares into ordinary shares on a one-to-one basis, as if conversion would have occurred on September 30, 2020.

Summary Consolidated Cash Flow Data

The following table presents our summary consolidated cash flow data for the years ended December 31, 2018 and 2019 and our selected unaudited interim consolidated cash flow data for the nine months ended September 30, 2019 and 2020.

	Year Ended		Nine Months Ended	
	December 31,		September 30,	
	2018	2019	2019	2020
	(in USD thousands)			
Net cash used in operating activities	(14,265)	(18,154)	(11,791)	(19,988)
Net cash generated (used in) from investing activities	(29,510)	24,856	18,942	7,434
Net cash generated from financing activities	51,058	69,694	16,310	3,097
Effect of exchange rate on cash and cash equivalents	39	78	153	(180)
Net increase in cash and cash equivalents	7,322	76,474	23,614	(9,638)
Cash and cash equivalents at the beginning of year/period	8,736	16,058	16,058	92,533
Cash and cash equivalents at the end of year/period	16,058	92,533	39,673	82,895

RISK FACTORS

You should consider carefully all of the information in this prospectus, including the risks and uncertainties described below and our consolidated financial statements and related notes, before making an investment in the ADSs. Any of the following risks and uncertainties could have a material adverse effect on our business, financial condition and results of operations. In particular, as we are a China-based company incorporated in the Cayman Islands, you should pay special attention to subsections headed "Risks Related to Doing Business in the PRC" below. The market price of the ADSs could decline significantly as a result of any of these risks and uncertainties, and you may lose all or part of your investment. When determining whether to invest, you should also refer to the other information contained in this prospectus, including our financial statements and the related notes thereto. You should also carefully review the cautionary statements referred to under "Special Note Regarding Forward-looking Statements." Our actual results could differ materially and adversely from those anticipated in this prospectus.

Risks Related to Our Financial Prospects and Need for Additional Capital

We have a limited operating history, which may make it difficult to evaluate our current business and predict our future performance.

We are a clinical stage biopharmaceutical company with a limited operating history. Since our inception in 2011, we have focused substantially all of our efforts and financial resources on the discovery and development of antibody therapeutics for the treatment of cancer. We have no products approved for commercial sale and therefore we have not generated any revenue from product sales. We have not obtained regulatory approvals for any of our product candidates and there is no assurance that we will obtain approvals in the future. We expect to continue to incur significant expenses and operating losses over the next several years and for the foreseeable future. Our prior losses, combined with expected future losses, have had and may continue to have an adverse effect on our working capital.

Our operations to date have focused on developing our product candidates, building our intellectual property portfolio, conducting preclinical testing and clinical trials, and raising capital. These operations provide a limited basis for you to assess our ability to successfully market and commercialize our product candidates. Consequently, predictions about our future success or viability may not be as accurate as they could be if we had a longer operating history. We will encounter risks and difficulties frequently experienced by early-stage companies in rapidly evolving fields as we seek to shift our focus to late stage development and commercial activities. If we do not address these risks and difficulties successfully, we may not be successful in such a transition.

We have incurred net losses historically and we may continue to incur net losses in the near future.

Since our inception in 2011, we have devoted our resources to the development of innovative antibodies in the therapeutic area. While we have generated revenues from licensing and collaboration deals, we have not generated any revenue from commercial product sales to date, and we have had significant operating losses since our inception. For the years ended December 31, 2018 and 2019 and the nine months ended September 30, 2020, we incurred net losses of US\$15.3 million, US\$16.4 million and US\$29.6 million, respectively. Substantially all of our operating losses have resulted from costs incurred in connection with our research and development programs. To date, we have financed our operations principally through private placements. Our product candidates and programs are in preclinical development or early stage clinical development, and we have not received marketing approval for any of our product candidates. Our product candidates will require substantial investments and significant marketing efforts before we generate any revenues from product sales, if ever. We

expect our net losses will increase as more product candidates enter into clinical trial stage. Our ability to generate product revenue and achieve profitability depends on, among other things:

- completing research and development of our product candidates;
- initiating, enrolling patients in and completing clinical trials of product candidates on a timely basis;
- obtaining regulatory approvals and marketing authorizations for any product candidates for which we complete clinical trials;
- developing and maintaining adequate manufacturing capabilities either by ourselves or in connection with third-party manufacturers;
- launching and commercializing any product candidates for which we obtain regulatory approvals and marketing authorizations;
- establishing a sales, marketing and commercialization team for any future products for which we may obtain regulatory approval;
- seeking to identify additional product candidates;
- addressing any competing technological and market developments; and
- maintaining, protecting and expanding our portfolio of intellectual property rights.

We may never succeed in these activities and, even if we do, may never generate revenue that is significant enough to achieve profitability. Because of the numerous risks and uncertainties associated with the development, delivery and commercialization of complex innovative antibody therapeutic, we are unable to accurately predict the timing or amount of expenses or when, or if, we will be able to achieve profitability. If we are required by regulatory authorities to perform studies in addition to those currently expected, or if there are any delays in the initiation and completion of our clinical trials or the development of any of our product candidates, our expenses could increase and profitability could be further delayed.

If we fail to become profitable or are unable to sustain profitability on a continuing basis, we may be unable to continue our operations at planned levels and be forced to reduce our operations. Even if we do achieve profitability, we may not be able to sustain or increase profitability on a quarterly or annual basis. Our failure to become or remain profitable would decrease the value of our company and could impair our ability to raise capital, expand our business or continue our operations. Failure to become and remain profitable may adversely affect the market price of the ADSs and our ability to raise capital and continue operations. A decline in the value of our company could also cause you to lose all or part of your investment.

We may need to obtain substantial additional financing to fund our growth and operations, which may not be available on acceptable terms, if at all.

The development of biopharmaceutical product candidates is capital-intensive. We have used substantial funds to advance our discovery programs and develop our technology and product candidates, and will require significant funds to conduct further research and development, preclinical testing and clinical trials of our product candidates, to seek regulatory approvals for our product candidates and to manufacture and market products, if any, that are approved for commercial sales. If our product candidates enter and advance through preclinical studies and clinical trials, we will need substantial additional funds to expand our development, regulatory and manufacturing capabilities. In addition, upon the closing of this offering, we expect to incur additional costs associated with operating as a public company.

To date, we have funded our operations primarily through capital contributions from our shareholders via private placements. Our operations have consumed substantial amounts of cash since inception. As of September 30, 2020, we had US\$82.9 million in cash and cash equivalents. The net cash used in our operating activities was US\$14.3 million, US\$18.2 million and US\$20.0 million for the years ended December 31, 2018 and 2019 and the nine months ended September 30, 2020, respectively. Our future funding requirements and the period for which we expect increasing capital need may be different than what we are planning. Our monthly spending levels vary based on new and ongoing research and development activities. Because of the numerous risks and uncertainties associated with our product development, we are unable to accurately predict the timing and amount of our operating expenditures, which will depend largely on:

- the scope, timing, progress, costs and results of discovery, preclinical development, laboratory testing and clinical development activities of our current product candidates;
- the number, scope, progress and results of preclinical and clinical programs we decide to pursue;
- the progress of the development efforts of parties with whom we have entered or may in the future enter into collaborations and research and development agreements;
- our ability to maintain our current licenses, research and development programs, and to establish new collaboration arrangements;
- the costs and timing of preparing, filing and prosecuting patent applications, maintaining and enforcing our intellectual property rights and defending any intellectual property-related claims;
- the cost, timing and outcome of regulatory review of any of our product candidates;
- the costs and timing of future commercialization activities, including manufacturing, marketing, sales, and distribution, for any of our product candidates for which we receive marketing approval;
- the revenue, if any, received from commercial sales of our product candidates for which we receive marketing approval; and
- our efforts to enhance operational systems and hire additional personnel, including personnel to support development of our product candidates and satisfy our obligations as a public company.

We will need to continue to rely on additional financing to achieve our business objectives. Adequate additional financing may not be available to us on acceptable terms, or at all. Any additional capital-raising efforts may divert our management from their day-to-day activities, which may adversely affect our ability to develop and commercialize our current and future product candidates, if approved. If we are unable to raise capital when needed or on attractive terms, we could be forced to delay, reduce or altogether cease our research and development programs or future commercialization efforts.

Raising additional capital may lead to dilution of shareholdings by our existing shareholders and restrict our operations or require us to relinquish rights to our technologies or product candidates.

We may seek additional funding through a combination of equity and debt financings and collaborations. To the extent that we raise additional capital through the sale of equity or convertible debt securities, the beneficial ownership interest of existing shareholders and the holders of ADSs will be diluted, and the terms may include liquidation or other preferences that adversely affect the rights of our existing shareholders and the holders of the ADSs. The incurrence of additional indebtedness or the issuance of certain equity securities could result in increased fixed payment obligations and could also result in certain additional restrictive covenants, such as incurring additional debt, making capital expenditures, or declaring dividends. If we raise additional funds through partnerships, collaborations, strategic alliances, or licensing arrangements with third parties, we may have to relinquish valuable

rights to our technologies, product candidates, or future revenue streams, or grant licenses on terms that are not favorable to us.

We have certain shareholders who will have board representation rights after the offering and their individual interests may differ from yours.

In 2014, we completed private placements of our Series A-1 and A-2 preferred shares, in which we raised gross proceeds of approximately US\$8.5 million. In 2016, we completed a private placement of our Series B preferred shares, in which we raised gross proceeds of approximately US\$28.0 million. In 2018, we completed a private placement of our Series C-1 preferred shares, in which we raised gross proceeds of approximately US\$50.0 million. In 2019, we completed private placements of our Series C-2 and C-3 preferred shares, in which we raised gross proceeds of approximately US\$69.0 million in aggregate. As a result of these private placements, a significant portion of our outstanding equity is currently held by multiple separate institutional investors through several separate funds and our founders. Collectively, these institutional investors that are our principal shareholders beneficially owned approximately 55.5% of our outstanding voting stock as of the date of this prospectus on an as-converted basis.

These institutional investors will continue have a significant level of influence because of their level of ownership, including a greater ability than you and our other shareholders to influence the election of directors and the potential outcome of other matters submitted to a vote of our shareholders, such as mergers, the sale of substantially all of our assets and other extraordinary corporate matters. These investors and our founder, Peter Luo, also have certain rights, such as board representation right and registration right that our other shareholders do not have.

For instance, our post-offering memorandum and articles of association will provide that JSR Limited shall have the right to designate, appoint, remove and replace and reappoint one director so long it holds at least five percent of the shares outstanding on a fully-diluted basis and an as-converted basis, respectively; as long as Wuxi Pharmatech Healthcare Fund I L.P., which is controlled by the ultimate controlling party of our sole supplier, holds at least five percent of the shares outstanding on a fully-diluted basis, it shall have the right to nominate one independent non-executive director and such one director shall be appointed and agreed by the board; as long as Peter Luo holds or beneficially owns any shares or is employed by us or any of our subsidiaries, he will serve as one of our directors and the Chairman of the Board of Directors; in addition, during the period commencing upon the effectiveness of our registration statement on Form F-1, of which this prospectus is a part, and ending on the earlier of (i) the date upon which Peter Luo beneficially owns less than five percent of the shares outstanding on a fully diluted basis, (ii) the death or legal determination of Peter Luo's incapacity or (iii) the termination of Peter Luo as an executive officer or principal scientific advisor to us or any of our subsidiaries or for cause (as determined under his related employment or consulting arrangements and subject to all related cure provisions), Peter Luo shall have the right to designate, appoint, remove and replace and reappoint one additional director; and as long as General Atlantic Singapore AI Pte. Ltd. and its affiliates hold at least five percent of the shares outstanding on a fully-diluted basis, they shall have the right to designate, appoint, remove and replace and reappoint one director.

The interests of these investors could conflict with the interests of our other shareholders, including you, and any future transfer by these investors of their shares of preferred or ordinary share to other investors who have different business objectives could have a material adverse effect on our business, results of operations, financial condition and the market value of our ordinary shares or ADSs.

Risks Related to Clinical Development of Our Product Candidates

We may not be able to identify or discover new product candidates, and may allocate our limited resources to pursue a particular product candidate and fail to capitalize on product candidates that may later prove to be more profitable, or for which there is a greater likelihood of success.

Although we will focus our efforts on continued preclinical and clinical developments, regulatory approval process and commercialization with respect to our existing product candidates, the success of our business depends in part upon our ability to identify, license, discover, develop, or commercialize additional product candidates. Research programs to identify new product candidates require substantial technical, financial, and human resources. Our research programs may initially show promise in identifying potential product candidates, yet fail to yield product candidates for clinical development for a number of reasons, including:

- our research or business development methodology or search criteria and process may be unsuccessful in identifying potential product candidates;
- our potential product candidates may be shown to have harmful side effects or may have other characteristics that may make the products unmarketable or unlikely to obtain marketing approval; and
- potential product candidates may not be effective in treating their targeted diseases.

Because we have limited financial and managerial resources, we focus on research programs and product candidates for specific targets. As a result, we may forgo or delay pursuit of opportunities with other product candidates that later may be proved to have greater commercial potential or a greater likelihood of success. On the other hand, if we do not prioritize the allocation of our resources and conduct research programs that cover a broad range of targets or engage clinical programs that are overly expansive, we may be subject to significant risk of loss as a large part of the research and clinical programs fail. Our resource allocation decisions may cause us to fail to capitalize on viable commercial products or profitable market opportunities.

Accordingly, there can be no assurance that we will ever be able to develop suitable potential product candidates through internal research programs, which could materially adversely affect our future growth and prospects.

We may not be successful in our efforts to use and expand our proprietary platforms to build a pipeline of product candidates.

A key element of our strategy is to leverage our technology platform to expand our pipeline of antibody product candidates and in order to do so, we will continue to invest in our platform and development capabilities. Although our research and development efforts to date have resulted in a pipeline of product candidates, these product candidates may not be safe and effective. In addition, although we expect that our platform will allow us to develop a diverse pipeline of novel and differentiated product candidates, we may not prove to be successful at doing so. Even if we are successful in continuing to build our pipeline, the potential product candidates that we identify may not be suitable for clinical development, including as a result of being shown to have harmful side effects or other characteristics that indicate that they are unlikely to be products that will receive marketing approval or achieve market acceptance. Even after approval, if we cannot successfully develop or commercialize our products, or if serious adverse events are discovered after commercialization, we will not be able to generate any product revenue, which would adversely affect business.

Any failures or setbacks in our platforms or our other proprietary technologies could negatively affect our business and financial condition.

Our product candidates are created with, and dependent upon, our proprietary antibody discovery platforms, such as our proprietary Dynamic Precision Library platform, which includes our NEObody platform, SAFEbody platform and POWERbody platform. These proprietary technology platforms are also the basis of our collaborations with certain other partners. To date, no products based on any of these technologies have been approved for commercial sale in any jurisdiction. Any failures or setbacks with respect to our proprietary technologies, including adverse effects resulting from the use of product candidates derived from these technologies in human clinical trials and/or the imposition of clinical holds on trials of any product candidates using our proprietary technologies, could have a detrimental impact on our clinical pipeline, as well as our ability to maintain and enter into new corporate collaborations regarding our technologies or otherwise, which would negatively affect our business and financial conditions.

Our product candidates, for which we intend to seek approval as biologics products, may face competition sooner than anticipated.

Even if we are successful in achieving a final regulatory approval to commercialize a product candidate ahead of our competitors, our product candidates may face competition from biosimilar products. In the United States, our product candidates are regulated by the FDA as biologic products and we intend to seek approval for these product candidates pursuant to the Biologics License Application (BLA) pathway. The Biologics Price Competition and Innovation Act of 2009 (BPCIA) created an abbreviated pathway for the approval of biosimilar and interchangeable biologic products. The abbreviated regulatory pathway establishes legal authority for the FDA to review and approve biosimilar biologics, including the possible designation of a biosimilar as "interchangeable" based on its similarity to an existing brand product. Under the BPCIA, an application for a biosimilar product cannot be approved by the FDA until 12 years after the original branded product was approved under a BLA. The law is complex and is still being interpreted and implemented by the FDA. As a result, its ultimate impact, implementation, and meaning are subject to uncertainty. While it is uncertain when such processes intended to implement BPCIA may be fully adopted by the FDA, any such processes could have a material adverse effect on the future commercial prospects for our product candidates.

There is a risk that any of our product candidates approved as a biological product under a BLA would not qualify for the 12-year period of exclusivity or that this exclusivity could be shortened due to congressional action or otherwise, or that the FDA will not consider our product candidates to be reference products for competing products, potentially creating the opportunity for interchangeable or generic competition sooner than anticipated. Other aspects of the BPCIA, some of which may impact the BPCIA exclusivity provisions, have also been the subject of litigation. Moreover, the extent to which a biosimilar, once approved, will be substituted for any one of our reference products in a way that is similar to traditional generic substitution for non-biological products is not yet clear, and will depend on a number of marketplace and regulatory factors that are still developing.

Jurisdictions in addition to the United States have established abbreviated pathways for regulatory approval of biological products that are biosimilar to earlier approved reference products. For example, the European Union has had an established regulatory pathway for biosimilars since 2005.

The increased likelihood of biosimilar competition has increased the risk of loss of innovators' market exclusivity. Due to this risk, and uncertainties regarding patent protection, if our clinical candidates are approved for marketing, it is not possible to predict the length of market exclusivity for any particular product with certainty based solely on the expiration of the relevant patent(s) or the current forms of regulatory exclusivity. It is also not possible to predict changes in United States regulatory law that might reduce biological product regulatory exclusivity. The loss of market exclusivity

for a product would likely materially and negatively affect revenues and we may not generate adequate or sufficient revenues from them or be able to reach or sustain profitability.

We depend substantially on the success of our product candidates, particularly ADG106, ADG126, ADG116 and ADG104, which are in clinical development, and our ability to identify additional product candidates. Clinical trials of our product candidates may not be successful. If we are unable to successfully identify new product candidates, complete clinical development, obtain regulatory approval and commercialize our product candidates, or experience significant delays in doing so, our business will be materially harmed.

Our business and the ability to generate revenue related to product sales, if ever, will depend on the successful development, regulatory approval and commercialization of our antibody product candidates for the treatment of patients with cancer, particularly ADG106, ADG126, ADG116 and ADG104, which are still in clinical stage. Other than ADG106, ADG126, ADG116 and ADG104 which are currently in Phase I development, our current product candidates are in relatively early stages of development. We have invested a significant portion of our efforts and financial resources in the development of our existing product candidates and all of our product candidates will require significant further development, financial resources. The success of our product candidates, including ADG106, ADG126, ADG116 and ADG104, will depend on several factors, including:

- successful enrollment in, and completion of, preclinical studies and clinical trials;
- receipt of regulatory approvals from the FDA, NMPA and other comparable regulatory authorities for our product candidates;
- establishing commercial manufacturing capabilities, either by building facilities ourselves or making arrangements with third-party manufacturers;
- relying on third parties to conduct our clinical trials safely and efficiently;
- obtaining and maintaining patent, trade secret and other intellectual property protection and regulatory exclusivity;
- protecting our rights in our intellectual property;
- ensuring we do not infringe, misappropriate or otherwise violate the patent, trade secret or other intellectual property rights of third parties;
- launching commercial sales of our product candidates, if and when approved;
- competition with other product candidates and drugs; and
- continued acceptable safety profile for our product candidates following final regulatory approval, if and when received.

Due to the uncertain, time-consuming and costly clinical development and regulatory approval process, we may not successfully develop any of our product candidates, or we or our partners may choose to discontinue the development of product candidates for a variety of reasons. Our failure to effectively advance our development programs could have a material adverse effect on our business, financial condition, results of operations and future growth prospects, and cause the market price of our ADSs to decline.

Clinical trials are expensive, time consuming and difficult to design and implement and may fail to demonstrate adequate safety and efficacy of our product candidates. The results of our current and previous preclinical studies or clinical trials may not be predictive of future results, and the results of our current and planned clinical trials may not satisfy the requirements of the FDA or comparable foreign regulatory authorities or provide the basis for regulatory approval.

Before obtaining marketing approval from regulatory authorities for the sale of our product candidates, we must conduct preclinical studies and extensive clinical trials to demonstrate their safety and efficacy in humans. Clinical testing is expensive and difficult to design and implement. Clinical testing can take many years to complete, and its ultimate outcome is uncertain. A failure of one or more clinical trials can occur at any stage of the process. We will be required to demonstrate with substantial evidence through well-controlled clinical trials that our product candidates are safe, pure, and potent for use in a diverse patient population before we can seek final regulatory approvals for their commercial sale. Our clinical trials may produce negative or inconclusive results, and we may decide, or regulators may require us, to conduct additional and expansive preclinical or clinical testings.

We cannot assure that the results of later clinical trials will replicate the results of prior clinical trials and preclinical testing. The results are subject to the risk that one or more of the clinical outcomes may materially change as patient enrollment continues and more patient data becomes available. Product candidates in later-stage clinical trials may fail to demonstrate sufficient safety and efficacy to the satisfaction of the FDA, the NMPA and comparable foreign regulatory authorities, despite having progressed through preclinical studies or initial clinical trials. Product candidates that have shown promising results in early clinical trials may still suffer significant setbacks in subsequent clinical trials or registration clinical trials. For example, a number of companies in the biopharmaceutical industry, including those with greater resources and experience than us, have suffered significant setbacks in advanced clinical trials, even after obtaining promising results in earlier clinical trials.

A failure of a clinical trial to meet its predetermined endpoints may cause us to abandon a pipeline product or an indication and may delay development of any other pipeline products. Any delay in, or termination of, our clinical trials will delay the submission for regulatory approval and application, and, ultimately, our ability to commercialize any of our pipeline products and generate revenue.

Moreover, principal investigators for our clinical trials may serve and have served as scientific advisors or consultants to us from time to time and receive compensation in connection with such services. Under certain circumstances, we may be required to report some of these relationships to the FDA or comparable foreign regulatory authorities. The FDA or comparable foreign regulatory authority may conclude that a financial relationship between us and a principal investigator has created a conflict of interest or otherwise affected interpretation of the trial. The FDA or comparable foreign regulatory authority may therefore question the integrity of the data generated at the applicable clinical trial site and the utility of the clinical trial itself may be jeopardized. This could result in a delay in approval, or rejection, of our marketing applications by the FDA or comparable foreign regulatory authority, as the case may be, and may ultimately lead to the denial of marketing approval of our pipeline products.

We may experience delays in our ongoing clinical trials.

We may experience delays in our ongoing clinical trials, and we do not know whether planned clinical trials will begin on time, need to be redesigned, enroll patients on time, or be completed on schedule, if at all. Clinical trials can be delayed, suspended, or terminated for a variety of reasons, including the following:

- delays in or failure to obtain regulatory authorization to commence a trial;

- delays in or failure to obtain institutional review board, or IRB, approval at each site;
- delays in or failure to reach agreement on acceptable terms with prospective contract research organizations, or CROs, and clinical trial sites, the terms of which can be subject to extensive negotiation and may vary significantly among different CROs and trial sites;
- difficulty in recruiting clinical trial investigators of appropriate competencies and experience;
- delays in establishing the appropriate dosage levels in clinical trials;
- delays in or failure to recruit and enroll suitable patients to participate in a trial, particularly considering study inclusion and exclusion criteria and patients' prior lines of therapy and treatment;
- the difficulty in certain countries in identifying the sub-populations that we are trying to treat in a particular trial, which may delay enrollment and reduce the power of a clinical trial to detect statistically significant results;
- lower than anticipated retention rates of patients in clinical trials;
- failure to have patients complete a trial or return for post-treatment follow-up;
- clinical sites deviating from trial protocol or dropping out of a trial;
- delays adding new investigators or clinical trial sites;
- safety or tolerability concerns could cause us or our collaborators or governmental authorities, as applicable, to suspend or terminate a trial if it is found that the participants are being exposed to unacceptable health risks, undesirable side effects or other unfavorable characteristics of the product candidate, or if such undesirable effects or risks are found to be caused by a chemically or mechanistically similar therapeutic or therapeutic candidate;
- our third-party research contractors failing to comply with regulatory requirements or meet their contractual obligations to us in a timely manner, or at all;
- changes in regulatory requirements, policies and guidelines;
- failure to comply with the applicable regulatory requirements through the clinical process;
- manufacturing sufficient quantities of a product candidate for use in clinical trials;
- the quality or stability of a product candidate falling below acceptable standards;
- changes in the treatment landscape for our target indications that may make our product candidates no longer relevant; and
- third-party actions claiming infringement by our product candidates in clinical trials outside the United States and obtaining injunctions interfering with our progress.

In addition, disruptions caused by the COVID-19 pandemic may increase the likelihood that we encounter such difficulties or delays in initiating, enrolling, conducting or completing our planned and ongoing clinical trials. Moreover, while we plan to submit additional investigational new drug applications, or INDs, for other product candidates, we may not be able to file such INDs on the timeline we expect. For example, we may experience manufacturing delays or other delays with IND-enabling preclinical studies. Moreover, we cannot be sure that submission of an IND will result in the FDA allowing clinical trials to begin, or that, once begun, issues will not arise that suspend or terminate clinical trials. Additionally, even if such regulatory authorities agree with the design and implementation of the clinical trials set forth in an IND, we cannot guarantee that such regulatory authorities will not change their requirements in the future. These considerations also apply to new clinical trials we may submit as amendments to existing INDs.

If we are required to conduct additional clinical trials or other studies with respect to any of our product candidates beyond those that we initially contemplated, if we are unable to successfully complete our clinical trials or other studies or if the results of these studies are not positive or are only modestly positive, we may be delayed in obtaining regulatory approval for that product candidate, we may not be able to obtain regulatory approval at all or we may obtain approval for indications that are not as broad as intended. Our drug development costs will also increase if we experience delays in testing or approvals, and we may not have sufficient funding to complete the testing and approval process. Significant clinical trial delays could allow our competitors to bring drugs to market before we do and impair our ability to commercialize our drugs, if and when approved. If any of this occurs, our business will be materially harmed.

Clinical trials must be conducted in accordance with the FDA and other applicable regulatory authorities' legal requirements, regulations or guidelines, and are subject to oversight by these governmental agencies and Ethics Committees or IRBs at the medical institutions where the clinical trials are conducted. We could encounter delays if a clinical trial is suspended or terminated by us, by the IRBs or Ethics Committees of the institutions in which such trials are being conducted, by the Data Review Committee or Data Safety Monitoring Board for such trial or by the FDA, or other regulatory authorities. Such authorities may impose such a suspension or termination due to a number of factors, including failure to conduct the clinical trial or to perform obligations in accordance with regulatory requirements or our clinical protocols, inspection of the clinical trial operations or trial site by the FDA or other regulatory authorities resulting in the imposition of a clinical hold, unforeseen safety issues or adverse side effects, failure to demonstrate a benefit from using a drug, changes in governmental regulations or administrative actions or lack of adequate funding to continue the clinical trial. If we experience delays in the completion of, or termination of, any clinical trial of our product candidates, the commercial prospects of our product candidates will be harmed, and our ability to generate product revenues from any of these product candidates will be delayed. In addition, any delays in completing our clinical trials will increase our costs, slow down our product candidate development and approval process and jeopardize our ability to commence product sales and generate revenues. Significant clinical trial delays could also allow our competitors to bring products to market before we do or shorten any periods during which we have the exclusive right to commercialize our product candidates and impair our ability to commercialize our product candidates and may harm our business and results of operations.

Further, clinical trials must be conducted with supplies of our product candidates produced under current good manufacturing practices, or cGMP, requirements and other regulations. Furthermore, we rely on CROs and clinical trial sites to ensure the proper and timely conduct of our clinical trials and while we have agreements governing their committed activities, we have limited influence over their actual performance. We depend on our collaborators and on medical institutions and CROs to conduct our clinical trials in compliance with good clinical practice, or GCP, requirements. To the extent our collaborators or the CROs fail to enroll participants for our clinical trials, fail to conduct the study in accordance with GCP or are delayed for a significant time in the execution of trials, including achieving full enrollment, we may be affected by increased costs, program delays or both, which may harm our business. In addition, clinical trials that are conducted in countries outside the United States may subject us to further delays and expenses as a result of increased shipment costs, additional regulatory requirements and the engagement of non-U.S. CROs, as well as expose us to risks associated with clinical investigators who are unknown to the FDA, and different standards of diagnosis, screening and medical care.

If we encounter difficulties in enrolling patients in our clinical trials, our clinical development activities could be delayed or otherwise adversely affected.

Successful and timely completion of clinical trials will require that we enroll a sufficient number of patients. Our clinical trials may be subject to delays for a variety of reasons, including as a result of enrollment taking longer than anticipated, subject withdrawal or adverse events. These types of developments could cause us to delay the trial or halt further development.

While we believe our differentiated product candidates address highly unmet medical needs that will facilitate our patient enrollment, clinical trials may compete with other clinical trials that are in the same therapeutic areas as our product candidates, and this competition reduces the number and types of patients available to us, as some patients who might have opted to enroll in our trials may instead opt to enroll in a trial being conducted by one of our competitors. Moreover, enrolling patients in clinical trials for cancer therapies is challenging, as cancer patients will first receive the applicable standard of care. Many patients who respond positively to the standard of care antibody therapy (and thus do not enroll in clinical trials) are believed to have tumor types that would have responded well to our product candidates. Patients who fail to respond positively to the standard of care treatment will be eligible for clinical trials of unapproved product candidates. However, these patients may have either compromised immune function from prior administration of chemotherapy or an enhanced immune response from the prior administration of checkpoint inhibitors. Either of these prior treatment regimens may render our therapies less effective in clinical trials. Additionally, patients who have failed approved therapies will typically have more advanced cancer and a poorer long-term prognosis.

Because the number of qualified clinical investigators and clinical trial sites is limited, we expect to conduct some of our clinical trials at the same clinical trial sites that some of our competitors use, which will reduce the number of patients who are available for our clinical trials at such clinical trial sites. We may experience difficulties in patient enrollment in our clinical trials for a variety of reasons, including:

- severity of the disease under investigation;
- the size and nature of the patient population;
- the patient eligibility criteria defined in the protocol;
- the size of the study population required for analysis of the trial's primary endpoints;
- perceived risks and benefits of our pipeline products;
- our resources to facilitate timely enrollment in clinical trials;
- patient referral practices of physicians;
- the proximity of patients to trial sites;
- the design of the trial;
- our ability to recruit clinical trial investigators with the appropriate competencies and experience;
- our ability to obtain and maintain patients' consent; and
- the risk that patients enrolled in clinical trials will not complete a clinical trial.

These factors may make it difficult for us to enroll enough patients to complete our clinical trials in a timely and cost-effective manner. Delays in the completion of any clinical trial of our product candidates will increase our costs, slow down our product candidate development and approval process and delay or potentially jeopardize our ability to commence product sales and generate revenue. In addition, some of the factors that cause, or lead to, a delay in the commencement or completion of clinical trials may also ultimately lead to the denial of regulatory approval of our product candidates.

Interim, topline or preliminary data from our clinical trials that we announce or publish from time to time may change as more patient data become available and are subject to audit and verification procedures that could result in material changes in the final data.

From time to time, we may publicly disclose preliminary or topline or data from our preclinical studies and clinical trials, which is based on a preliminary analysis of then-available data, and the results and related findings and conclusions are subject to change following a more comprehensive review of the data related to the particular study or trial. We also make assumptions, estimations, calculations and conclusions as part of our analyses of data, and we may not have received or had the opportunity to fully and carefully evaluate all data. As a result, the topline results that we report may differ from future results of the same studies, or different conclusions or considerations may qualify such results, once additional data have been received and fully evaluated. Topline data also remain subject to audit and verification procedures that may result in the final data being materially different from the preliminary data we previously published. As a result, topline data should be viewed with caution until the final data are available.

From time to time, we may also disclose interim data from our preclinical studies and clinical trials. Interim data from clinical trials that we may complete are subject to the risk that one or more of the clinical outcomes may materially change as patient enrollment continues and more patient data become available. Adverse differences between preliminary or interim data and final data could significantly harm our business prospects. Further, disclosure of interim data by us or by our competitors could result in volatility in the price of our common stock.

Further, others, including regulatory agencies, may not accept or agree with our assumptions, estimates, calculations, conclusions, or analyses or may interpret or weigh the importance of data differently, which could impact the value of the particular program, the approvability or commercialization of the particular product candidate or product and our company in general. If the interim, topline, or preliminary data that we report differ from actual results, or if others, including regulatory authorities, disagree with the conclusions reached, our ability to obtain approval for, and commercialize, our product candidates may be harmed, which could harm our business, operating results, prospects, or financial condition.

Risks Related to Obtaining Regulatory Approval of Our Drug Candidates

The regulatory approval processes of the FDA, NMPA and other comparable regulatory authorities are lengthy, time consuming and inherently unpredictable, and if we are ultimately unable to obtain regulatory approvals for our product candidates, our business will be substantially harmed.

In the United States, marketing approval of biologics requires the submission of a BLA to the FDA, and we are not permitted to market any product candidate in the United States until we obtain approval from the FDA of the BLA for that product candidate. A BLA must be supported by extensive clinical and preclinical data, as well as extensive information regarding pharmacology, chemistry, manufacturing and controls. Outside the United States, many comparable foreign regulatory authorities employ similar approval processes.

We have not previously submitted a BLA to the FDA or similar regulatory approval filings to the NMPA or other comparable foreign authorities, for any product candidate, and we cannot be certain that any of our product candidates will receive regulatory approval. Obtaining approval of a BLA can be a lengthy, expensive and uncertain process, and as a company we have no experience with the preparation of a BLA submission or any other application for marketing approval. In addition, the FDA has the authority to require a risk evaluation and mitigation strategies, or REMS, plan as part of a BLA or after approval, which may impose further requirements or restrictions on the distribution or use of an approved biologic, such as limiting prescribing to certain physicians or medical centers that

have undergone specialized training, limiting treatment to patients who meet certain safe-use criteria and requiring treated patients to enroll in a registry.

FDA approval is not guaranteed, and the time required to obtain approval by the FDA, NMPA and other comparable regulatory authorities is unpredictable but typically takes many years following the commencement of preclinical trials and clinical trials and depends upon numerous factors, including the substantial discretion of the regulatory authorities. In addition, due to external issues such as pandemics or other public health emergencies, FDA, NMPA and other comparable regulatory authorities may be delayed in their review of product applications. In addition, approval policies, regulations or the type and amount of clinical data necessary to gain approval may change during the course of a product candidate's clinical development and may vary among jurisdictions. We have not obtained regulatory approval for any product candidate, and it is possible that none of our existing product candidates or any product candidates we may discover, in-license or acquire and seek to develop in the future will ever obtain regulatory approval.

Our product candidates could fail to receive regulatory approval from the FDA, NMPA or a comparable regulatory authority for many reasons, including:

- disagreement with the design or implementation of our clinical trials;
- failure to demonstrate that a product candidate is safe and effective or safe, pure, and potent for its proposed indication;
- failure of clinical trial results to meet the level of statistical significance required for approval;
- failure to demonstrate that a product candidate's clinical and other benefits outweigh its safety risks;
- disagreement with our interpretation of data from preclinical trials or clinical trials;
- the insufficiency of data collected from clinical trials of our product candidates to support the submission and filing of a BLA or other submission or to obtain regulatory approval;
- the FDA, NMPA or comparable regulatory authority's finding of deficiencies related to the manufacturing processes;
- failure of our product candidates to pass current Good Manufacturing Practice, or cGMP, inspections during the regulatory review process or across the production cycle of our product; and
- changes in approval policies or regulations that render our preclinical and clinical data insufficient for approval.

The FDA and other regulatory authorities have substantial discretion in the approval process, and determining when or whether regulatory approval will be obtained for any of our product candidates. For example, regulatory authorities in various jurisdictions have in the past had, and may in the future have, differing requirements for, interpretations of and opinions on our preclinical and clinical data. As a result, we may be required to conduct additional preclinical studies, alter our proposed clinical trial designs or conduct additional clinical trials to satisfy the regulatory authorities in each of the jurisdictions in which we hope to conduct clinical trials and develop and market our products, if approved. Further, even if we believe the data collected from clinical trials of our product candidates are promising, such data may not be sufficient to support approval by the FDA or any other regulatory authority.

The FDA, NMPA or a comparable regulatory authority may require more information, including additional preclinical or clinical data, to support approval, which may delay or prevent approval and our commercialization plans, or we may decide to abandon the development program. If we were to

obtain approval, regulatory authorities may approve any of our product candidates for fewer or more limited indications than we request, may grant approval contingent on the performance of costly post-marketing clinical trials, may approve a product candidate with a label that is not desirable for the successful commercialization of that product candidate, or may be difficult to meet manufacturing requirements. In addition, if our product candidate produces undesirable side effects or safety issues, the FDA may require the establishment of Risk Evaluation Mitigation Strategies, or REMS, or the NMPA or a comparable regulatory authority may require the establishment of a similar strategy, that may, for instance, restrict distribution of our drugs and impose burdensome implementation requirements on us. Any of the foregoing scenarios could materially harm the commercial prospects of our product candidates.

Disruptions at the FDA and other government agencies caused by funding shortages or global health concerns could hinder their ability to hire, retain or deploy key leadership and other personnel, or otherwise prevent new or modified products from being developed, approved or commercialized in a timely manner or at all, which could negatively impact our business.

The ability of the FDA to review and approve new products can be affected by a variety of factors, including government budget and funding levels, statutory, regulatory, and policy changes, the FDA's ability to hire and retain key personnel and accept the payment of user fees, and other events that may otherwise affect the FDA's ability to perform routine functions. Average review times at the FDA have fluctuated in recent years as a result. In addition, government funding of other government agencies that fund research and development activities is subject to the political process, which is inherently fluid and unpredictable. Disruptions at the FDA and other agencies may also slow the time necessary for new biologics or modifications to licensed biologics to be reviewed and/or approved by necessary government agencies, which would adversely affect our business. For example, over the last several years, including for 35 days beginning on December 22, 2018, the U.S. government has shut down several times and certain regulatory agencies, such as the FDA, have had to furlough critical FDA employees and stop critical activities.

Separately, in response to the COVID-19 pandemic, on March 10, 2020, the FDA announced its intention to postpone most inspections of foreign manufacturing facilities, and on March 18, 2020, the FDA temporarily postponed routine surveillance inspections of domestic manufacturing facilities. Subsequently, on July 10, 2020, the FDA announced its intention to resume certain on-site inspections of domestic manufacturing facilities subject to a risk-based prioritization system. The FDA intends to use this risk-based assessment system to identify the categories of regulatory activity that can occur within a given geographic area, ranging from mission critical inspections to resumption of all regulatory activities. Regulatory authorities outside the United States may adopt similar restrictions or other policy measures in response to the COVID-19 pandemic. If a prolonged government shutdown occurs, or if global health concerns continue to prevent the FDA or other regulatory authorities from conducting their regular inspections, reviews, or other regulatory activities, it could significantly impact the ability of the FDA or other regulatory authorities to timely review and process our regulatory submissions, which could have a material adverse effect on our business.

Even if we obtain FDA approval of any of our product candidates, we may never obtain approval or commercialize such products outside of the United States, which would limit our ability to realize their full market potential.

In order to market any products outside of the United States, we must establish and comply with numerous and varying regulatory requirements of other countries regarding safety and efficacy. Clinical trials conducted in one country may not be accepted by regulatory authorities in other countries, and regulatory approval in one country does not mean that regulatory approval will be obtained in any other countries. Approval procedures vary among countries and can involve additional product testing

and validation and additional administrative review periods. Seeking regulatory approvals in various jurisdictions could result in significant delays, difficulties and costs for us and may require additional preclinical studies or clinical trials which would be costly and time consuming. Regulatory requirements can vary widely from country to country and could delay or prevent the introduction of our products in those countries. Satisfying these and other regulatory requirements is costly, time consuming, uncertain and subject to unanticipated delays. In addition, our failure to obtain regulatory approval in any country may delay or have negative effects on the process for regulatory approval in other countries. We do not have any product candidates approved for sale in any jurisdiction, including international markets, and we do not have experience in obtaining regulatory approval in international markets. If we fail to comply with regulatory requirements in international markets or to obtain and maintain required approvals, our ability to realize the full market potential of our products will be harmed.

We are conducting clinical trials and may in the future conduct additional clinical trials for our product candidates outside the United States and/or China, and FDA, NMPA and similar foreign regulatory authorities may not accept data from such trials.

We are conducting clinical trials and may in the future conduct additional clinical trials for our product candidates outside the United States, including in Australia, Europe or other foreign jurisdictions. The acceptance of trial data from clinical trials conducted outside the United States by the FDA may be subject to certain conditions. In cases where data from clinical trials conducted outside the United States are intended to serve as the sole basis for marketing approval in the United States, the FDA will generally not approve the application on the basis of foreign data alone unless (i) the data are applicable to the United States population and United States medical practice; (ii) the trials were performed by clinical investigators of recognized competence and (iii) the data may be considered valid without the need for an on-site inspection by the FDA or, if the FDA considers such an inspection to be necessary, the FDA is able to validate the data through an on-site inspection or other appropriate means. Otherwise, for studies that are conducted at sites outside of the United States and not subject to an IND and which are intended to support a marketing application (but which are not intended to serve as the sole basis for marketing approval), the FDA requires the clinical trial to have been conducted in accordance with good clinical practice, or GCP, requirements and the FDA must be able to validate the data from the clinical trial through an on site inspection if it deems such inspection necessary. Additionally, the FDA's clinical trial requirements, including sufficient size of patient populations and statistical powering, must be met. Many foreign regulatory bodies, such as NMPA, have similar approval requirements. In addition, such foreign trials would be subject to the applicable local laws of the foreign jurisdictions where the trials are conducted. There can be no assurance that the FDA, NMPA or any similar foreign regulatory authority will accept data from trials conducted outside of the United States or the applicable jurisdiction. If the FDA, NMPA or any similar foreign regulatory authority does not accept such data, it would result in the need for additional trials, which would be costly and time-consuming and delay aspects of our business plan, and which may result in our product candidates not receiving approval or clearance for commercialization in the applicable jurisdiction.

Our product candidates may cause undesirable adverse events or have other properties that could delay or prevent their regulatory approval, limit the commercial profile of an approved label, or result in significant negative consequences following any regulatory approval.

Undesirable adverse events caused by our product candidates could cause us or regulatory authorities to interrupt, delay or halt clinical trials and could result in a more restrictive label or the delay or denial of regulatory approval by the FDA, NMPA or other comparable regulatory authority. Results of our trials could reveal a high and unacceptable severity or prevalence of adverse events. In such an event, our trials could be suspended or terminated and the FDA, NMPA or other comparable regulatory authorities could order us to cease further development of, or deny approval of, our product

candidates for any or all targeted indications. Drug-related adverse events could affect patient recruitment or the ability of enrolled subjects to complete the trial, and could result in potential product liability claims. Moreover, such events may require us to amend our trials, including reducing the dosage in our clinical trials.

For instance, in September 2019, in our Phase I trial of ADG116 in the United States, we reported to the FDA the death of the only patient dosed in the trial. The FDA subsequently placed the trial on clinical hold on September 30, 2019. The FDA noted that there was insufficient information to assess the risk of drug-induced liver toxicity in patients receiving ADG116 but requested us to revise the study protocol to mitigate the risk of drug-induced liver toxicity. We submitted to the FDA an amendment to the study protocol, which significantly lowered the starting dose of ADG116, tightened the inclusion and exclusion criteria to exclude, among others, patients with liver dysfunction, history of alcohol abuse, or poorly controlled diabetes, and stipulated additional post-dosing monitoring to closely follow treated patients for safety or toxicity signals. The FDA removed the clinical hold on December 5, 2019. We have obtained authorization from the Australian Therapeutic Goods Administration under a CTN to start a Phase I clinical trial of ADG116. We have decided to focus on conducting the Australian clinical trial of ADG116 at a higher starting dose than currently permitted in the United States, and consequently, we have not recommenced patient enrollment in the United States. If the data from the Australian clinical trial are favorable, we may consider using such data for re-submission of an IND application to the NMPA. The NMPA in August 2020 did not approve our application for a clinical trial in China at a higher starting dose than currently permitted in Australia and requested additional patient safety data prior to considering approving the IND at the proposed starting dose.

We cannot provide any assurance that there will not be treatment-related severe adverse events with our product candidates, that the trials for our product candidates will not be suspended in the future, or that patient recruitment for trials with our product candidates will not be adversely impacted by the ADG116 related adverse events, any of which could materially and adversely affect our business and prospects. Further, clinical trials by their nature utilize a sample of the potential patient population. With a limited number of patients and limited duration of exposure, rare and severe side effects of our product candidates may only be uncovered with a significantly larger number of patients exposed to the product candidate. In the event that any of our product candidates receives marketing approval and we or others later identify undesirable or unacceptable side effects caused by such products, a number of potentially significant negative consequences could result, including:

- we may suspend marketing of such products;
- regulatory authorities may withdraw or limit approvals of such products or require us to take an approved product off the market;
- regulatory authorities may require the addition of labeling statements, specific warnings, a contraindication or field alerts to physicians and pharmacies, or issue other communications containing warnings or other safety information about the product;
- regulatory authorities may require a medication guide outlining the risks of such side effects for distribution to patients, or that we implement a REMS plan to ensure that the benefits of the product outweigh its risks or to develop a similar strategy as required by a comparable regulatory authority, that may, for instance, restrict distribution of our drugs and impose burdensome implementation requirements on us;
- we may be required to conduct post-market studies;
- we may be subject to limitations on how we may promote or manufacture the product;
- sales of the product may decrease significantly;
- we could be sued and held liable for harm caused to subjects or patients; and

- our reputation may suffer.

Any of these events could prevent us from achieving or maintaining market acceptance of the particular product candidate, if approved, and could significantly harm our business, results of operations and prospects.

Further, combination therapy involves unique adverse events that could be exacerbated compared to adverse events from monotherapies. These types of adverse events could be caused by our product candidates and could also cause us or regulatory authorities to interrupt, delay or halt clinical trials and could result in a more restrictive label or the delay or denial of regulatory approval by the FDA, NMPA or other comparable regulatory authority. Results of our trials could reveal a high and unacceptable severity or prevalence of adverse events.

We may seek Orphan Drug Designation for some of our product candidates, and we may be unsuccessful.

Regulatory authorities in some jurisdictions, including the United States, may designate drugs for relatively small patient populations as orphan drugs. Under the Orphan Drug Act of 1983, the FDA may designate a drug as an orphan drug if it is a drug intended to treat a rare disease or condition, which is generally defined as a disease with a patient population of fewer than 200,000 individuals in the United States, or a patient population of greater than 200,000 individuals in the United States, but for which there is no reasonable expectation that the cost of developing the drug will be recovered from sales in the United States.

Generally, if a drug with an Orphan Drug Designation subsequently receives the first FDA approval for the indication for which it has such designation, the drug is entitled to a period of marketing exclusivity, which precludes the FDA from approving another marketing application for the same drug for the same indication during the period of exclusivity. The applicable period is seven years in the United States. Orphan Drug exclusivity may be lost if the FDA determines that the request for designation was materially defective or if the manufacturer is unable to assure sufficient quantity of the drug to meet the needs of patients with the rare disease or condition.

Even if we obtain Orphan Drug exclusivity for a product candidate, that exclusivity may not effectively protect the product candidate from competition because different drugs can be approved for the same condition and the same drugs can be approved for a different condition but used off-label for any orphan indication we may obtain. Even after an orphan drug is approved, the FDA can subsequently approve a different drug for the same condition if the FDA concludes that the later drug is clinically superior in that it is shown to be safer, more effective or makes a major contribution to patient care. Orphan drug designation neither shortens the development time or regulatory review time of a drug nor gives the drug any advantage in the regulatory review or approval process.

We may attempt to secure approval from the FDA or comparable foreign regulatory authorities through the use of accelerated approval pathways. If we are unable to obtain such approval, we may be required to conduct additional preclinical studies or clinical trials beyond those that we contemplate, which could increase the expense of obtaining, and delay the receipt of, necessary marketing approvals. Even if we receive accelerated approval from the FDA, if our confirmatory trials do not verify clinical benefit, or if we do not comply with rigorous post-marketing requirements, the FDA may seek to withdraw accelerated approval.

We may in the future seek an accelerated approval for our one or more of our product candidates. Under the accelerated approval program, the FDA may grant accelerated approval to a product candidate designed to treat a serious or life-threatening condition that provides meaningful therapeutic benefit over available therapies upon a determination that the product candidate has an effect on a surrogate endpoint or intermediate clinical endpoint that is reasonably likely to predict clinical benefit. The FDA considers a clinical benefit to be a positive therapeutic effect that is clinically meaningful in the context of a given disease, such as irreversible morbidity or mortality. For the purposes of

accelerated approval, a surrogate endpoint is a marker, such as a laboratory measurement, radiographic image, physical sign, or other measure that is thought to predict clinical benefit, but is not itself a measure of clinical benefit. An intermediate clinical endpoint is a clinical endpoint that can be measured earlier than an effect on irreversible morbidity or mortality that is reasonably likely to predict an effect on irreversible morbidity or mortality or other clinical benefit. The accelerated approval pathway may be used in cases in which the advantage of a new drug over available therapy may not be a direct therapeutic advantage, but is a clinically important improvement from a patient and public health perspective. If granted, accelerated approval is usually contingent on the sponsor's agreement to conduct, in a diligent manner, additional post-approval confirmatory studies to verify and describe the drug's clinical benefit. If such post-approval studies fail to confirm the drug's clinical benefit, the FDA may withdraw its approval of the drug.

Prior to seeking accelerated approval for any of our product candidates, we intend to seek feedback from the FDA and will otherwise evaluate our ability to seek and receive accelerated approval. There can be no assurance that after our evaluation of the feedback and other factors we will decide to pursue or submit a BLA for accelerated approval or any other form of expedited development, review or approval. Similarly, there can be no assurance that after subsequent FDA feedback we will continue to pursue or apply for accelerated approval or any other form of expedited development, review or approval, even if we initially decide to do so. Furthermore, if we decide to submit an application for accelerated approval or receive an expedited regulatory designation (e.g., breakthrough therapy designation) for our product candidates, there can be no assurance that such submission or application will be accepted or that any expedited development, review or approval will be granted on a timely basis, or at all. The FDA or other comparable foreign regulatory authorities could also require us to conduct further studies prior to considering our application or granting approval of any type. A failure to obtain accelerated approval or any other form of expedited development, review or approval for our product candidate would result in a longer time period to commercialization of such product candidate, could increase the cost of development of such product candidate and could harm our competitive position in the marketplace.

Even if we receive regulatory approvals for our product candidates, we will be subject to ongoing regulatory obligations and continued regulatory reviews, which may result in significant additional expense and we may be subject to penalties if we fail to comply with regulatory requirements or experience unanticipated problems with our product candidates.

If our product candidates are approved, they will be subject to ongoing regulatory requirements for manufacturing, labeling, packaging, storage, import, export, adverse event reporting, advertising, promotion, sampling, record-keeping, conduct of post-marketing studies, and submission of safety, efficacy, and other post-market information, including both federal and state requirements in the United States and requirements of comparable regulatory authorities.

Manufacturers and manufacturers' facilities are required to comply with extensive FDA, NMPA and comparable regulatory authority requirements, including, in the United States, ensuring that quality control and manufacturing procedures conform to current cGMP regulations. As such, we and our contract manufacturers will be subject to continual review and inspections to assess compliance with cGMP and adherence to commitments made in any BLA, other marketing application, and previous responses to inspection observations. Accordingly, we and others with whom we work must continue to expend time, money and effort in all areas of regulatory compliance, including manufacturing, production and quality control.

Any regulatory approvals that we receive for our product candidates may be subject to limitations on the approved indicated uses for which the drug may be marketed or to the conditions of approval, or contain requirements for potentially costly post-marketing testing, including Phase IV clinical trials and surveillance to monitor the safety and efficacy of the product candidate. The FDA may also

require a REMS program as a condition of approval of our product candidates, which could entail requirements for long-term patient follow-up, a medication guide, physician communication plans or additional elements to ensure safe use, such as restricted distribution methods, patient registries and other risk minimization tools. In addition, if the FDA, NMPA, or a comparable regulatory authority approves our product candidates, we will have to comply with requirements including, for example, submissions of safety and other post-marketing information and reports, registration, as well as continued compliance with cGMPs and Good Clinical Practices, or cGCPs, for any clinical trials that we conduct post approval.

The FDA and NMPA may impose consent decrees or withdraw approval if compliance with regulatory requirements and standards is not maintained or if problems occur after the drug reaches the market. Later discovery of previously unknown problems with our product candidates, including adverse events of unanticipated severity or frequency, or with our third-party manufacturers or manufacturing processes, or failure to comply with regulatory requirements, may result in revisions to the approved labeling to add new safety information, imposition of post-market studies or clinical trials to assess new safety risks, or imposition of distribution restrictions or other restrictions under a REMS program. Other potential consequences include, among other things:

- restrictions on the marketing or manufacturing of our drugs, withdrawal of the product from the market, or voluntary or mandatory product recalls;
- fines, untitled or warning letters, or holds on clinical trials;
- refusal by the FDA, NMPA or comparable regulatory authorities to approve pending applications or supplements to approved applications filed by us or suspension or revocation of license approvals;
- product seizure or detention, or refusal to permit the import or export of our product candidates; and
- injunctions or the imposition of civil or criminal penalties.

The FDA strictly regulates marketing, labeling, advertising and promotion of products that are placed on the market. Drugs may be promoted only for the approved indications and in accordance with the provisions of the approved label. The FDA, NMPA and other regulatory authorities actively enforce the laws and regulations prohibiting the promotion of off-label uses, and a company that is found to have improperly promoted off-label uses may be subject to significant liability. The policies of the FDA, NMPA and of other regulatory authorities may change and additional government regulations may be enacted that could prevent, limit or delay regulatory approval of our product candidates. We cannot predict the likelihood, nature or extent of government regulation that may arise from future legislation or administrative action, either in the United States or abroad. If we are slow or unable to adapt to changes in existing requirements or the adoption of new requirements or policies, or if we are not able to maintain regulatory compliance, we may lose any regulatory approval that we may have obtained and we may not achieve or sustain profitability.

All material aspects of the research, development and commercialization of pharmaceutical products are heavily regulated.

All jurisdictions in which we intend to conduct our pharmaceutical-industry activities regulate these activities in great depth and detail. We intend to focus our activities in the major markets such as the United States and China. These jurisdictions strictly regulate the pharmaceutical industry, and in doing so they employ broadly similar regulatory strategies, including regulation of product development and approval, manufacturing, and marketing, sales and distribution of products. However, there are differences in the regulatory regimes that make for a more complex and costly regulatory compliance burden for a company like ours that plans to operate in these regions.

The process of obtaining regulatory approvals and compliance with appropriate laws and regulations require substantial time and financial resources. Failure to comply with the applicable requirements at any time during the product development process and approval process, or after approval, may subject an applicant to administrative or judicial sanctions. These sanctions could include: refusal to approve pending applications; withdrawal of an approval; license revocation; clinical hold; voluntary or mandatory product recalls; product seizures; total or partial suspension of production or distribution; injunctions; fines; refusals of government contracts; providing restitution; undergoing disgorgement; or other civil or criminal penalties. Failure to comply with these regulations could have a material adverse effect on our business.

Future strategic partnerships may be important to us. We will face significant competition in seeking new strategic partners.

We do not yet have any capability for manufacturing, sales, marketing or distribution. For some of our product candidates, we may in the future determine to collaborate with pharmaceutical and biotechnology companies for development and potential commercialization of therapeutic products. The competition for strategic partners is intense. Our ability to reach a definitive agreement for collaboration will depend, among other things, upon our assessment of the strategic partner's resources and expertise, the terms and conditions of the proposed collaboration and the proposed strategic partner's evaluation of a number of factors. These factors may include the design or results of clinical trials, the likelihood of approval by the FDA or comparable foreign regulatory authorities, the potential market for the subject product candidate, the costs and complexities of manufacturing and delivering such product candidate to patients, the potential of competing products, the existence of uncertainty with respect to our ownership of technology, which can exist if there is a challenge to such ownership without regard to the merits of the challenge, and industry and market conditions generally. The strategic partner may also consider alternative product candidates or technologies for similar indications that may be available for collaboration and whether such collaboration could be more attractive than the one with us for our product candidate.

Strategic partnerships are complex and time-consuming to negotiate and document. In addition, there have been a significant number of recent business combinations among large pharmaceutical companies that have resulted in a reduced number of potential future strategic partners. Even if we are successful in entering into collaboration, the terms and conditions of that collaboration may restrict us from entering into future agreements with other potential collaborators.

If we are unable to reach agreements with suitable strategic partners on a timely basis, on acceptable terms, or at all, we may have to curtail the development of a product candidate, reduce or delay one or more of our other development programs, delay its potential commercialization or reduce the scope of any sales or marketing activities, or increase our expenditures and undertake development or commercialization activities at our own expense. If we elect to fund and undertake development or commercialization activities on our own, we may need to obtain additional expertise and additional capital, which may not be available to us on acceptable terms or at all. If we fail to enter into strategic partnerships and do not have sufficient funds or expertise to undertake the necessary development and commercialization activities, we may not be able to further develop our product candidates or bring them to market or continue to develop our technology platform and our business may be materially and adversely affected. Any collaboration may be on terms that are not optimal for us, and we may not be able to maintain any new collaboration if, for example, development or approval of a product candidate is delayed, sales of an approved product candidate do not meet expectations or the partner terminates the collaboration. Any such collaboration, or other strategic transaction, may require us to incur non-recurring or other charges, and increase our near- and long-term expenditures and pose significant integration or implementation challenges or disrupt our management or business. Accordingly, although there can be no assurance that we will undertake or successfully complete any

transactions of the nature described above, any transactions that we do complete may be subject to the foregoing or other risks and have a material and adverse effect on our business, financial condition, results of operations, and prospects. Conversely, any failure to enter any collaboration or other strategic transaction that would be beneficial to us could delay the development and potential commercialization of our product candidates and have a negative impact on the competitiveness of any product candidate that reaches the market.

We may not be able to enter into additional collaboration agreements beyond our existing strategic partnerships or collaborations with NIH, Exelixis, ADC Therapeutics and Guilin Sanjin. If we are unable to maintain existing and future strategic partnerships or collaborations, or if these strategic partnerships or collaborations are not successful, our business could be adversely affected.

Our existing strategic partnerships, collaborations and any future strategic partnerships we enter into may pose a number of risks, including the following:

- we may not be able to enter into critical strategic partnerships or enter into them on favorable terms;
- strategic partners or collaborators have significant discretion in determining the effort and resources that they will apply to such a partnership, and they may not perform their obligations as agreed, expected, or in compliance with applicable legal requirements;
- strategic partners or collaborators may not pursue development and commercialization of any product candidates that achieve regulatory approval or may elect not to continue or renew development or commercialization programs based on clinical trial results, changes in the partners' strategic focus or available funding, or external factors, such as an acquisition that diverts resources or creates competing priorities;
- strategic partners or collaborators may delay clinical trials, provide insufficient funding for a clinical trial program, stop a clinical trial or abandon a product candidate, repeat or conduct new clinical trials or require a new formulation of a product candidate for clinical testing;
- strategic partners or collaborators could independently develop, or develop with third parties, products that compete directly or indirectly with our product candidates if the strategic partners or collaborators believe that competitive products are more likely to be successfully developed or can be commercialized under terms that are more economically attractive than our product candidates;
- product candidates discovered in collaboration with us may be viewed by our strategic partners or collaborators as competitive with their own product candidates or products, which may cause strategic partners or collaborators to cease to devote resources to the commercialization of our product candidates;
- a strategic partner or collaborator with marketing and distribution rights to one or more of our product candidates that achieve regulatory approval may not commit sufficient resources to the marketing and distribution of such product candidates;
- disagreements with strategic partners or collaborators, including disagreements over proprietary rights, contract interpretation or the preferred course of development, might cause delays or termination of the research, development or commercialization of product candidates, might lead to additional responsibilities for us with respect to product candidates, or might result in litigation or arbitration, any of which would be time-consuming and expensive;
- strategic partners or collaborators may not properly maintain or defend our intellectual property rights or may use our proprietary information in such a way as to invite litigation that could

jeopardize or invalidate our intellectual property or proprietary information or expose us to potential litigation;

- strategic partners or collaborators may infringe the intellectual property rights of third parties, which may expose us to litigation and potential liability;
- strategic partners or collaborators may claim for a substantial compensation for our failure of development of the product candidates specified under the relevant out-licensing agreements that solely arose out of problems of our previous R&D basis; and
- strategic partnerships or collaborations may be terminated for the convenience of the partner or the collaborator and, if terminated, we could be required to raise additional capital to pursue further development or commercialization of the applicable product candidates.

If any partnerships or collaboration we enter into do not result in the successful development of our product candidates or if one of our partners or collaborator terminates the agreement with us, our continued development of our product candidates could be delayed and our business may be materially and adversely affected.

We will need to grow our organization, and we may experience difficulties in managing this growth, which could disrupt our operations.

As of December 31, 2020, we had 198 full-time employees. As our development and commercialization plans and strategies develop, and as we transition into operating as a public company, we expect to expand our employee base for managerial, operational, financial and other resources. As our product candidates enter and advance through preclinical studies and clinical trials, we will need to expand our development, regulatory and manufacturing capabilities or contract with other organizations to provide these capabilities for us. In the future, we expect to enter into additional relationships with collaborators or partners, suppliers and other organizations and establish a sales and marketing team in preparation for commercialization activities. Our ability to manage our operations and future growth will require us to continue to improve our operational, financial and management controls, reporting systems and procedures. We may not be able to implement improvements to our management information and control systems in an efficient or timely manner and may discover deficiencies in existing systems and controls. Our inability to successfully manage our growth and expand our operations could have a material and adverse effect on our business, financial condition, results of operations and prospects.

Disruptions in the financial markets and economic conditions could affect our ability to raise capital.

Global economies could suffer dramatic downturns as the result of a deterioration in the credit markets and related financial crisis as well as a variety of other factors including, extreme volatility in security prices, severely diminished liquidity and credit availability, ratings downgrades of certain investments and declining valuations of others. In the past, governments have taken unprecedented actions in an attempt to address and rectify these extreme market and economic conditions by providing liquidity and stability to the financial markets. If these actions are not successful, the return of adverse economic conditions may cause a significant impact on our ability to raise capital, if needed, on a timely basis and on acceptable terms or at all.

In addition, there is considerable uncertainty over the long-term effects of the expansionary monetary and fiscal policies adopted by the central banks and financial authorities of some of the world's leading economies, including the United States and China. There have been concerns over unrest and terrorist threats in the Middle East, Europe and Africa and over the conflicts involving Ukraine, Syria and North Korea. There have also been concerns on the relationship among China and other Asian countries, which may result in or intensify potential conflicts in relation to territorial

disputes or the trade related disputes between the United States and China. In addition, the U.K. held a referendum on June 23, 2016 on its membership in the EU, in which voters approved an exit from the EU, commonly referred to as "Brexit"; the U.K. formally left the EU on January 31, 2020. The U.K. is currently in a transition period which is expected to continue through December 31, 2020, when agreements surrounding trade and other aspects of the U.K.'s future relationship with the EU will need to be finalized. Brexit could adversely affect European and worldwide economic and market conditions and could contribute to instability in global financial and foreign exchange markets. It is unclear whether these challenges and uncertainties will be contained or resolved, and what effects they may have on the global political and economic conditions in the long term.

If we face allegations of non-compliance with laws and encounter sanctions, our reputation, revenues and liquidity may suffer, and our product candidates and approved products, if any, could be subject to restrictions or withdrawal from the market.

Any government investigation of alleged violations of laws or regulations could require us to expend significant time and resources in response, and could generate negative publicity. Any failure to comply with ongoing regulatory requirements may significantly and adversely affect our ability to obtain approvals commercialize and generate revenues from our drugs. If regulatory sanctions are applied or if regulatory approval is withdrawn, the value of our company and our operating results will be adversely affected. Additionally, if we are unable to generate revenues from our product sales, our potential for achieving profitability will be diminished and the capital necessary to fund our operations will be increased.

Our reputation is important to our success. Negative publicity may adversely affect our reputation and business prospects.

Any negative publicity concerning us, our affiliates or any entity that shares the "Adagene" name, even if untrue, could adversely affect our reputation and business prospects. There can be no assurance that negative publicity about us or any of our affiliates or any entity that shares the "Adagene" name would not damage our brand image or have a material adverse effect on our business, results of operations and financial condition.

Potential future acquisitions or strategic collaborations may increase our capital requirements, dilute our shareholders, cause us to incur debt or assume contingent liabilities and subject us to other risks.

We may evaluate various acquisitions and strategic collaborations, including licensing or acquiring complementary products, intellectual property rights, technologies or businesses. Any potential acquisition or strategic collaboration may entail numerous risks, including:

- increased operating expenses and cash requirements;
- the assumption of additional indebtedness or contingent liabilities;
- assimilation of operations, intellectual property and products of an acquired company, including difficulties associated with integrating new personnel;
- the diversion of our management's attention from our existing product programs and initiatives in pursuing such a strategic merger or acquisition;
- retention of key employees, the loss of key personnel, and uncertainties in our ability to maintain key business relationships;
- risks and uncertainties associated with the other party to such a transaction, including the prospects of that party and their existing products or product candidates and regulatory approvals; and/or

- our inability to generate revenue from acquired technology and/or products sufficient to meet our objectives in undertaking the acquisition or even to offset the associated acquisition and maintenance costs.

In addition, if we undertake acquisitions, we may issue dilutive securities, assume or incur debt obligations, incur large one-time expenses and acquire intangible assets that could result in significant future amortization expense. Moreover, we may not be able to locate suitable acquisition opportunities and this inability could impair our ability to grow or obtain access to technology or products that may be important to the development of our business.

Our business operations and current or future relationships with healthcare professionals, principal investigators, consultants, customers and third-party payors in the United States and elsewhere may be subject, directly or indirectly, to applicable anti-kickback, fraud and abuse, false claims, physician payment transparency and other healthcare laws and regulations. If we or our employees, independent contractors, consultants, commercial partners, or vendors violate these laws, we could face substantial penalties.

Our business operations and current or future arrangements with investigators, healthcare professionals, consultants, third-party payors, patient organizations and customers may expose us to broadly applicable fraud and abuse and other healthcare laws and regulations. These laws may constrain the business or financial arrangements and relationships through which we conduct our operations, including how we research, market, sell and distribute our product candidates, if approved. Such laws include, without limitation:

- the U.S. federal civil and criminal Anti-Kickback Statute, which prohibits, among other things, persons or entities from knowingly and willfully soliciting, offering, receiving or providing any remuneration (including any kickback, bribe, or certain rebate), directly or indirectly, overtly or covertly, in cash or in kind, to induce or reward, or in return for, either the referral of an individual for, or the purchase, lease, order or recommendation of, any good, facility, item or service, for which payment may be made, in whole or in part, under U.S. federal and state healthcare programs such as Medicare and Medicaid. A person or entity does not need to have actual knowledge of the statute or specific intent to violate it in order to have committed a violation;
- the U.S. federal false claims laws, including the False Claims Act, which can be enforced through whistleblower actions, and civil monetary penalties laws, which, among other things, impose criminal and civil penalties against individuals or entities for knowingly presenting, or causing to be presented, to the U.S. federal government, claims for payment or approval that are false or fraudulent, knowingly making, using or causing to be made or used, a false record or statement material to a false or fraudulent claim, or from knowingly making a false statement to avoid, decrease or conceal an obligation to pay money to the U.S. federal government. In addition, the government may assert that a claim including items and services resulting from a violation of the U.S. federal Anti-Kickback Statute constitutes a false or fraudulent claim for purposes of the False Claims Act;
- the federal Health Insurance Portability and Accountability Act of 1996, or HIPAA, which imposes criminal and civil liability for, among other things, knowingly and willfully executing, or attempting to execute, a scheme to defraud any healthcare benefit program, or knowingly and willfully falsifying, concealing or covering up a material fact or making any materially false statement, in connection with the delivery of, or payment for, healthcare benefits, items or services; similar to the U.S. federal Anti-Kickback Statute, a person or entity does not need to have actual knowledge of the statute or specific intent to violate it in order to have committed a violation;

- federal consumer protection and unfair competition laws, which broadly regulate marketplace activities and activities that potentially harm consumers;
- the United States Physician Payments Sunshine Act and its implementing regulations, which require certain manufacturers of drugs, devices, biologics and medical supplies that are reimbursable under Medicare, Medicaid or the Children's Health Insurance Program to report annually to the government information related to certain payments and other transfers of value to physicians, as defined by such law, and teaching hospitals, as well as ownership and investment interests held by the physicians described above and their immediate family members. Effective January 1, 2022, the U.S. federal physician transparency reporting requirements will extend to include transfers of value made during the previous year to certain non-physician providers such as physician assistants and nurse practitioners;
- analogous U.S. state laws and regulations, including: state anti-kickback and false claims laws, which may apply to our business practices, including but not limited to, research, distribution, sales;
- the U.S. Foreign Corrupt Practices Act of 1977, as amended, which prohibits, among other things, U.S. companies and their employees and agents from authorizing, promising, offering or providing, directly or indirectly, corrupt or improper payments or anything else of value to foreign government officials, employees of public international organizations and foreign government owned or affiliated entities, candidates for foreign political office and foreign political parties or officials thereof; and
- marketing arrangements and claims involving healthcare items or services reimbursed by any third-party payor, including private insurers; state laws that require pharmaceutical companies to comply with the pharmaceutical industry's voluntary compliance guidelines and the relevant compliance guidance promulgated by the U.S. federal government, or otherwise restrict payments that may be made to healthcare providers and other potential referral sources; state and local laws that require the registration of pharmaceutical sales representatives; and state laws and regulations that require drug manufacturers to file reports relating to pricing and marketing information, which requires tracking gifts and other remuneration and items of value provided to healthcare professionals and entities. For detailed discussion on material applicable PRC regulation, see "Regulation—PRC Regulation"

Ensuring that our internal operations and future business arrangements with third parties comply with applicable healthcare laws and regulations will involve substantial costs. It is possible that governmental authorities will conclude that our business practices do not comply with current or future statutes, regulations, agency guidance or case law involving applicable fraud and abuse or other healthcare laws and regulations. If our operations are found to be in violation of any of the laws described above or any other governmental laws and regulations that may apply to us, we may be subject to significant penalties, including civil, criminal and administrative penalties, damages, fines, exclusion from government-funded healthcare programs, such as Medicare and Medicaid or similar programs in other countries or jurisdictions, disgorgement, imprisonment, contractual damages, reputational harm, diminished profits and the curtailment or restructuring of our operations. Further, defending against any such actions can be costly and time-consuming and may require significant personnel resources. Even if we are successful in defending against any such actions that may be brought against us, our business may be impaired.

Recently enacted and future legislation in the United States and other countries may affect the prices we may obtain for our product candidates and increase the difficulty and cost for us to obtain marketing approval of and commercialize our product candidates.

In the United States and many other countries, rising healthcare costs have been a concern for governments, patients and the health insurance sector, which resulted a number of changes to laws and regulations, and may result in further legislative and regulatory action regarding the healthcare and health insurance systems that could affect our ability to profitably sell any product candidates for which we obtain marketing approval.

For example, the Patient Protection and Affordable Care Act, as amended by the Health Care and Education Reconciliation Act, or collectively, the ACA, was enacted in the United States in March 2010 with the stated goals of containing healthcare costs, improving quality and expanding access to healthcare, and includes measures to change health care delivery, increase the number of individuals with insurance, ensure access to certain basic health care services, and contain the rising cost of care. Since its enactment, there have been numerous judicial, administrative, executive, and legislative challenges to certain aspects of the ACA, and we expect there will be additional challenges and amendments to the ACA in the future. For example, various portions of the ACA are currently facing legal and constitutional challenges in the Fifth Circuit Court of Appeals and the United States Supreme Court. Additionally, the current administration has issued various Executive Orders which eliminated cost sharing subsidies and various provisions that would impose a fiscal burden on states or a cost, fee, tax, penalty or regulatory burden on individuals, healthcare providers, health insurers, or manufacturers of pharmaceuticals or medical devices, and Congress has introduced several pieces of legislation aimed at significantly revising or repealing the ACA. It is unclear whether the ACA will be overturned, repealed, replaced, or further amended.

In addition, other federal health reform measures have been proposed and adopted in the United States. For example, as a result of the Budget Control Act of 2011, providers are subject to Medicare payment reductions of 2% per fiscal year through 2030, with the exception of a temporary suspension from May 1, 2020 through December 31, 2020, unless additional Congressional action is taken. Further, the American Taxpayer Relief Act of 2012 reduced Medicare payments to several providers and increased the statute of limitations period for the government to recover overpayments from providers from three to five years. The Medicare Access and CHIP Reauthorization Act of 2015 also introduced a quality payment program under which certain individual Medicare providers will be subject to certain incentives or penalties based on new program quality standards. Payment adjustments for the Medicare quality payment program began in 2019. At this time, it is unclear how the introduction of the quality payment program will impact overall physician reimbursement under the Medicare program. Any reduction in reimbursement from Medicare or other government programs may result in a similar reduction in payments from private payors.

Further, there has been heightened governmental scrutiny in the United States of pharmaceutical pricing practices in light of the rising cost of prescription drugs and biologics.

Since January 2017, there has been legislation considered in Congress to restrict the pricing of drug products to a governmental negotiated rate or to other similar rates that would reduce the costs to government, commercial payers and individuals. In July 2020, President Trump signed four Executive Orders directing the Department of Health and Human Services and other agencies to take specific actions to reduce prescription drug prices. The first order directs federally qualified health centers to pass along significant discounts on insulin and epinephrine from drug companies to low-income individuals. The second order would allow the importation of prescription drugs from Canada into the United States where the prices are deemed to be lower. The third order would eliminate safe harbor protections under the federal Anti-Kickback Statute that currently covers rebates paid by manufacturers to Medicare Part D plans and Medicaid managed care organizations, either directly or through

pharmacy benefit managers under contract with such plans or organizations, so long as such actions are not projected to increase federal spending, Medicare beneficiary premiums or patients' total out-of-pocket cases. The fourth order will reduce the payment for Medicare part B drugs to be paid at the same rate as other developed nations, thereby reducing the reimbursement. All of these Executive Orders require rulemaking prior to implementation and could be stalled by Congress or the next election.

The combination of healthcare cost containment measures, increased health insurance costs, reduction of the number of people with health insurance coverage, as well as future legislation and regulations focused on reducing healthcare costs by reducing the cost of or reimbursement and access to pharmaceutical products, may limit or delay our ability to generate revenue, attain profitability, or commercialize our pipeline products, if approved.

We face intense competitions and rapid technological changes, as well as the possibility that our competitors may develop therapies that are similar, more advanced, or more effective than ours, which may adversely affect our ability to successfully commercialize our product candidates and our financial condition.

The biotechnology and pharmaceutical industries are intensely competitive and subject to rapid and significant technological change. We are currently aware of various existing therapies and development candidates that may compete with ADG106, ADG126, ADG116 and ADG104 for the potential treatment of similar targets, such as urelumab and utomilumab targeting CD137 and ipilimumab targeting CTLA-4. While we are developing ADG126 and ADG116, both targeting CTLA-4, we intend to focus on and prioritize ADG126 as it is generated by our SAFEbody technology designed to enhance its safety profile. We have competitors in the United States, China and internationally, including major multinational pharmaceutical companies, specialty pharmaceutical companies, and biotechnology companies. Many of our competitors have substantially greater financial, technical, and other resources, such as larger research and development, marketing and manufacturing organizations. Additional mergers and acquisitions in the biotechnology and pharmaceutical industries may result in even more resources being concentrated in our competitors. As a result, these companies may obtain regulatory approval more rapidly than we are able to and may be more effective in selling and marketing their products as well. Smaller or early-stage companies may also prove to be significant competitors, particularly through collaborative arrangements with large, established companies. Competition may increase further as a result of advances in the commercial applicability of technologies and greater availability of capital for investment in these industries. Our competitors may succeed in developing, acquiring or licensing on an exclusive basis, products that are more effective or less costly than any product candidate that we may develop, or achieve earlier patent protection, regulatory approval, product commercialization and market penetration than we do. Additionally, technologies developed by our competitors may render our potential product candidates uneconomical or obsolete, and we may not be successful in marketing our product candidates against competitors.

We have no experience in launching and marketing product candidates. We may not be able to effectively build and manage our sales network, or benefit from third-party collaborators' sales network.

We currently have no manufacturing, sales, marketing or commercial product distribution capabilities and have no experience in marketing drugs. We intend to develop an in-house marketing organization and sales force, which will require significant capital expenditures, management resources and time. We will have to compete with other biopharmaceutical companies to recruit, hire, train and retain marketing and sales personnel.

If we are unable or decide not to establish internal sales, marketing and commercial distribution capabilities for any or all of the drugs we develop, we will likely pursue collaborative arrangements regarding the sales and marketing of our product candidates, if approved. However, there can be no assurance that we will be able to establish or maintain such collaborative arrangements, or, if we are

able to do so, that they will have effective sales forces. Any revenue we receive will depend on the efforts of such third parties, which may not be successful. We may have little or no control over the marketing and sales efforts of such third parties, and our revenue from product sales may be lower than if we had commercialized our product candidates ourselves. We will also face competition in our search for third parties to assist us with the sales and marketing efforts of our product candidates, if approved.

There can be no assurance that we will be able to develop in-house sales and commercial distribution capabilities or establish or maintain relationships with third-party collaborators to successfully commercialize any product candidate, if approved, and as a result, we may not be able to generate product sales revenue.

Even if we are able to commercialize any approved product candidates, coverage and reimbursement may be limited or unavailable in certain market segments for our product candidates, and we may be subject to unfavorable pricing regulations, which could harm our business.

The regulations that govern regulatory approvals, pricing and reimbursement for new therapeutic products vary widely from country to country. Some countries require approval of the sale price of a drug before it can be marketed. In many countries, the pricing review period begins after marketing or licensing approval is granted. In some non-U.S. markets, prescription pharmaceutical pricing remains subject to continuing governmental control even after initial approval is granted. As a result, we might obtain regulatory approval for a drug in a particular country, but then be subject to price regulations that delay our commercial launch of the drug and negatively impact our revenues.

A primary trend in the global healthcare industry is cost containment. Government authorities and third-party payors have attempted to control costs by limiting coverage and the amount of reimbursement for particular medications. Our ability to commercialize any drugs successfully also will depend in part on the extent to which coverage and reimbursement for these drugs and related treatments will be available on adequate terms, or at all, from government health administration authorities, private health insurers and other organizations.

In the United States, no uniform policy of coverage and reimbursement for biopharmaceutical products exists among third-party payors. As a result, obtaining coverage and reimbursement approval of a drug from a government or other third-party payor is a time-consuming and costly process that could require us to provide to each payor supporting scientific, clinical and cost-effectiveness data for the use of our drugs on a payor-by-payor basis, with no assurance that coverage and adequate reimbursement will be obtained. Even if we obtain coverage for a given drug, the resulting reimbursement rates might not be adequate for us to achieve or sustain profitability or may require co-payments that patients find unacceptably high. Increasingly, third-party payors are requiring that companies provide them with predetermined discounts from list prices and are challenging the prices charged for medical products. Additionally, third-party payors may not cover, or provide adequate reimbursement for, long-term follow-up evaluations required following the use of our genetically modified drugs. Patients are unlikely to use our drugs and any approved product candidates unless coverage is provided and reimbursement is adequate to cover a significant portion of the cost of the drug. Because some of our product candidates have a higher cost of goods than conventional therapies, may require long-term follow up evaluations, and will likely be administered under the supervision of a physician, the risk that coverage and reimbursement rates may be inadequate for us to achieve profitability may be greater.

If reimbursement is not available or is available only to limited levels, we may not be able to successfully commercialize any product candidate that we successfully develop.

There may be significant delays in obtaining reimbursement for approved drugs, and coverage may be more limited than the purposes for which the drug is approved by the FDA or other comparable

regulatory authorities outside the United States. Moreover, eligibility for reimbursement does not imply that any drug will be paid for in all cases or at a rate that covers our costs, including research, development, manufacture, sale and distribution. Interim payments for new drugs, if applicable, may also not be sufficient to cover our costs and may not be made permanent. Payment rates may vary according to the use of the drug and the clinical setting in which it is used, may be based on payments allowed for lower cost drugs that are already reimbursed, and may be incorporated into existing payments for other services. Net prices for drugs may be reduced by mandatory discounts or rebates required by government healthcare programs or private payors and by any future weakening of laws that presently restrict imports of drugs from countries where they may be sold at lower prices than in the United States. Our inability to promptly obtain coverage and profitable payment rates from both government-funded and private payors for our drugs and any new product candidates that we develop could have a material adverse effect on our business, our operating results, and our overall financial condition.

In China, the Ministry of Human Resources and Social Security of China or provincial or local human resources and social security authorities, together with other government authorities, review the inclusion or removal of drugs from China's National Drug Catalog for Basic Medical Insurance, Work-related Injury Insurance and Maternity Insurance, or the National Reimbursement Drug List, or the NRDL, or provincial or local medical insurance catalogues for the National Medical Insurance Program regularly, and the tier under which a drug will be classified, both of which affect the amounts reimbursable to program participants for their purchases of those drugs. There can be no assurance that our drugs and any approved product candidates will be included in the NRDL or provincial reimbursements lists. Products included in the NRDL have been typically generic and essential drugs. Innovative drugs similar to our product candidates have historically been more limited on their inclusion in the NRDL due to the affordability of the government's Basic Medical Insurance, although this has been changing in recent years. According to currently effective PRC laws and regulations, the prices of approved drugs are determined by market competition. The government regulate prices mainly by establishing a consolidated procurement mechanism, revising the NRDL and strengthening regulation of medical and pricing practices. We cannot predict the extent to which our business may be affected by potential future legislative or regulatory developments. Changes in pricing regulation could restrict the amount that we are able to charge for our future approved drugs, which would adversely affect our revenue, profitability and results of operations.

We intend to seek approval to market our product candidates in the United States, China and in other jurisdictions. In some non-U.S. countries, the pricing of drugs and biologics is subject to governmental control, which can take considerable time even after obtaining regulatory approval. Market acceptance and sales of our drugs will depend significantly on the availability of adequate coverage and reimbursement from third-party payors for drugs and may be affected by existing and future health care reform measures.

As we engage in collaboration worldwide, we may be exposed to specific risks of conducting our business and operations in international markets.

We are a biopharmaceutical company with global footprints. We are currently building our clinical and technology infrastructures to support our future global operations and prepare to serve global markets. If we fail to obtain applicable licenses or fail to enter into strategic collaboration arrangements with third parties in these markets, or if these collaboration arrangements turn out unsuccessful, our revenue-generating growth potential will be adversely affected.

Moreover, international business relationships subject us to additional risks that may materially adversely affect our ability to attain or sustain profitable operations, including:

- efforts to enter into collaboration or licensing arrangements with third parties in connection with our international sales, marketing and distribution efforts may increase our expenses or divert our management's attention from the acquisition or development of product candidates;
- changes in a specific country's or region's political and cultural climate or economic condition;
- differing regulatory requirements for drug approvals and marketing internationally;
- difficulty of effective enforcement of contractual provisions in local jurisdictions;
- potentially reduced protection for intellectual property rights;
- potential third-party patent rights;
- unexpected changes in tariffs, trade barriers and regulatory requirements;
- economic weakness, including inflation or political instability;
- compliance with tax, employment, immigration and labor laws for employees traveling abroad;
- the effects of applicable tax structures and potentially adverse tax consequences;
- currency fluctuations, which could result in increased operating expenses and reduced revenue, and other obligations incidental to doing business in another country;
- workforce uncertainty and labor unrest and failure to comply with the applicable laws and regulations in relation to management of the employment of foreigners within the PRC;
- the potential for so-called parallel importing, which is what happens when a local seller, faced with high or higher local prices, opts to import goods from an international market with low or lower prices rather than buying them locally;
- failure of our employees and contracted third parties to comply with Office of Foreign Assets Control rules and regulations and the Foreign Corrupt Practices Act of the United States, and other applicable rules and regulations;
- production shortages resulting from any events affecting raw material supply or manufacturing capabilities abroad; and
- business interruptions resulting from geo-political actions, including war and terrorism, or natural disasters, including earthquakes, volcanoes, typhoons, floods, hurricanes and fires.

These and other risks may materially adversely affect our ability to attain or sustain revenue from international markets.

Building our commercialization capabilities will require significant investment of time and money. There would be no assurance that we will successfully set up our commercialization capabilities in any of the proposed jurisdictions or at all, or that we will successfully commercialize any of our product candidates in the future.

We are currently in the early stages of building and expanding our commercial capabilities to allow us to market our own products, if approved, in the future. We will need to set up full commercialization capabilities in the jurisdictions, including China and the United States, which would require substantial investment of time and money and will divert significant management focus and resources. We also face competition with multinational and local pharmaceutical and biotechnology companies with established commercialization capabilities in terms of marketing and attracting talents.

Therefore, there can be no assurance that our efforts to set up commercialization capabilities will be successful in any of the proposed jurisdictions or at all.

Even if ADG106 or one of our other proprietary product candidates obtains regulatory approval, we may determine that commercializing such product candidate ourselves would not be the most effective way to create value for our shareholders or holders of ADSs. In addition, if we choose to commercialize any of our product candidates, our marketing efforts may be unsuccessful as a result of unfavorable pricing or reimbursement limitations, delays, competition or other factors. Failure to successfully market one or more of our approved products, or delays in our commercialization efforts, may diminish the commercial prospects for such products and may result in financial losses or damage to our reputation, each of which may have a negative impact on the market price of our ADSs and our financial condition, results of operations and future growth prospects.

We may continue to pursue collaborations or licensing arrangements, joint ventures, strategic alliances, partnerships or other strategic investment or arrangements, which may fail to produce anticipated benefits and adversely affect our operations.

We may continue to pursue opportunities for collaboration, out-license, joint ventures, acquisitions of products, assets or technology, strategic alliances, or partnerships that we believe would advance our development. We may consider pursuing growth through the acquisition of technology, assets or other businesses that may enable us to enhance our technologies and capabilities. Proposing, negotiating and implementing these opportunities may be a lengthy and complex process. Other companies, including those with substantially greater financial, marketing, technology, or other business resources, may compete with us for these opportunities or arrangements. We may not be able to identify, secure, or complete any such transactions or arrangements in a timely manner, on a cost-effective basis, on acceptable terms, or at all.

We have limited experience with respect to these business development activities. Management and integration of a licensing arrangement, collaboration, joint venture or other strategic arrangement may disrupt our current operations, decrease our profitability, result in significant expenses, or divert management resources that otherwise would be available for our existing business. We may not realize the anticipated benefits of any such transaction or arrangement.

Furthermore, partners, collaborators, or other parties to such transactions or arrangements may fail to fully perform their obligations or meet our expectations or cooperate with us satisfactorily for various reasons and subject us to potential risks, including the followings:

- partners, collaborators, or other parties have significant discretion in determining the efforts and resources that they will apply to a transaction or arrangement;
- partners, collaborators, or other parties could independently develop, or develop with third parties, services and products that compete directly or indirectly with our product candidates;
- partners, collaborators, or other parties may stop, delay or discontinue clinical trials as well as repeat clinical trials or conduct new clinical trials by using our intellectual property or proprietary information;
- partners, collaborators, or other parties may not properly maintain or defend our intellectual property rights or may use our intellectual property or proprietary information in a way that gives rise to actual or threatened litigation that could jeopardize or invalidate our intellectual property or proprietary information or expose us to potential liabilities;
- disputes may arise between us and partners, collaborators, or other parties that cause the delay or termination of the research, development or commercialization of our product candidates, or that result in costly litigation or arbitration that diverts management's attention and resources;

- partners, collaborators, or other parties may be terminated and, if terminated, may result in a need for additional capital to pursue further development or commercialization of the applicable services and products; and
- partners, collaborators, or other parties may own or co-own intellectual properties covering our product candidates that results from our collaborating with them, and in such cases, we would not have the exclusive right to commercialize such intellectual properties.

Any such transactions or arrangements may also require actions, consents, approval, waiver, participation or involvement of various degrees from third parties, such as regulators, government authorities, creditors, licensors or licensees, related individuals, suppliers, distributors, shareholders or other stakeholders or interested parties. There is no assurance that such third parties will be cooperative as we desire, or at all, in which case we may be unable to carry out the relevant transactions or arrangements.

We rely on third parties to support, conduct and monitor our preclinical studies and clinical trials. Therefore, we may not be able to directly control the timing, process, expense and quality of our clinical trials and we cannot assure these third parties can duly perform their obligations as agreed and expected.

We have relied upon and plan to continue to rely upon third parties, including independent clinical investigators and third-party CROs, to conduct our preclinical studies and clinical trials and to monitor and manage data for our ongoing preclinical and clinical programs. We rely on these parties for execution of our preclinical studies and clinical trials, and control only certain aspects of their activities. Nevertheless, we are responsible for ensuring that each of our studies and trials is conducted in accordance with the applicable protocol, legal, regulatory and scientific standards, and our reliance on these third parties does not relieve us of our regulatory responsibilities. We and our third-party contractors and CROs are required to comply with GCP requirements, which are regulations and guidelines enforced by the FDA and comparable foreign regulatory authorities for all of our products candidates in clinical development. Regulatory authorities enforce these GCPs through periodic inspections of trial sponsors, principal investigators and trial sites. If we or any of our CROs fail to comply with applicable GCPs, the clinical data generated in our clinical trials may be deemed unreliable and the FDA or comparable foreign regulatory authorities may require us to perform additional clinical trials before approving our marketing applications. We cannot assure you that upon inspection by a given regulatory authority, such regulatory authority will determine that any of our clinical trials comply with GCP regulations. In addition, our clinical trials must be conducted with product produced under cGMP regulations. Our failure to comply with these regulations may require us to repeat or suspend clinical trials, which would delay the regulatory approval process.

Further, these investigators and CROs are not our employees and we will not be able to control, other than by contract, the amount of resources, including time, which they devote to our product candidates and clinical trials. If independent investigators or CROs fail to devote sufficient resources to the development of our product candidates, or if their performance is substandard, it may delay or compromise the prospects for approval and commercialization of any product candidates that we develop. In addition, the use of third-party service providers may require us to disclose our proprietary information to these parties, which could increase the risk that this information will be misappropriated.

Our CROs have the right to terminate their agreements with us in the event of an uncured material breach. In addition, some of our CROs have an ability to terminate their respective agreements with us if it can be reasonably demonstrated that the safety of the subjects participating in our clinical trials warrants such termination, if we make a general assignment for the benefit of our creditors or if we are liquidated.

If any of our relationships with these third-party CROs terminate, we may not be able to enter into arrangements with alternative CROs or to do so on commercially reasonable terms. If CROs do not successfully carry out their contractual duties or obligations or meet expected deadlines, if they need to be replaced or if the quality or accuracy of the clinical data they obtain is compromised due to the failure to adhere to our clinical protocols, regulatory requirements or for other reasons, our clinical trials may be extended, delayed or terminated and we may not be able to obtain regulatory approval for or successfully commercialize our product candidates. As a result, our results of operations and the commercial prospects for our product candidates would be harmed, our costs could increase and our ability to generate revenues could be delayed.

Switching or adding additional CROs involves additional cost and requires management time and focus. In addition, there is a natural transition period when a new CRO commences work. As a result, delays occur, which can materially impact our ability to meet our desired clinical development timelines. Additionally, CROs may lack the capacity to absorb higher workloads or take on additional capacity to support our needs. Though we carefully manage our relationships with our CROs, there can be no assurance that we will not encounter similar challenges or delays in the future or that these delays or challenges will not have a material adverse impact on our business, financial condition and prospects.

We currently rely on a third-party manufacturer to produce our product candidates. Any failure by the third-party manufacturer to produce acceptable product candidates for us pursuant to our specifications and regulatory standards may delay or impair our ability to initiate or complete our clinical trials, obtain and maintain regulatory approvals or commercialize approved products, if any.

We currently rely on a third-party manufacturer and expect to continue to rely for some time on third parties to manufacture our product candidates for preclinical testing and clinical trials, in compliance with applicable regulatory and quality standards, and may do so for the commercial manufacture of some of our product candidates, if approved. To date, we have obtained bulk drug substance for ADG106, ADG126 and ADG116 from a single-source third-party contract manufacturer. Any reduction or halt in supply of the drug substance from such contract manufacturer could severely constrain our ability to develop our product candidates until a replacement contract manufacturer is found and qualified. If we are unable to arrange for and maintain such third-party manufacturing sources that are capable of meeting regulatory standards, or fail to do so on commercially reasonable terms, we may not be able to successfully produce sufficient supply of product candidate or we may be delayed in doing so. If we were to experience an unexpected loss of supply of our product candidates, for any reason, whether as a result of manufacturing, supply or storage issues or otherwise, we could experience delays, disruptions, suspensions or terminations of, or be required to restart or repeat, any pending or ongoing clinical trials. Such failure or substantial delay or loss of supply could materially harm our business. We are continuously evaluating multiple vendors both in China and abroad to ensure that we have a continuous supply of products for global trials.

We may have little to no control regarding the occurrence of third-party manufacturer incidents. Any failure by our third-party manufacturer to comply with good manufacturing practices, or GMPs, or failure to scale up manufacturing processes, including any failure to deliver sufficient quantities of product candidates in a timely manner, would lead to a delay in, or failure to seek or obtain, regulatory approval of any of our product candidates. Furthermore, any change in manufacturer of our product candidates or approved products, if any, would require new regulatory approvals, which could delay completion of clinical trials or disrupt commercial supply of approved products.

Our current and anticipated future dependence upon others for the manufacture of our product candidates may adversely affect our future profit margins and our ability to commercialize any product candidates that receive marketing approval on a timely and competitive basis.

In some cases, the technical skills or technology required to manufacture our product candidates may be unique or proprietary to the original manufacturer, we may have difficulty transferring such skills or technology to another third party and a feasible alternative may not exist. These factors would increase our reliance on such manufacturer or require us to obtain a license from such manufacturer in order to have another third party manufacture our product candidates. If we are required to change manufacturers for any reason, we will be required to verify that the new manufacturer maintains facilities and procedures that comply with quality standards and with all applicable regulations and guidelines. The delays associated with the verification of a new manufacturer could negatively affect our ability to develop product candidates in a timely manner or within budget.

If our current research collaborators or scientific advisors and employees terminate their relationships with us or develop relationships with our competitors, our ability to discover antibodies and to conduct research and development could be adversely affected.

The responsibility of overseeing research and development of our product candidates is concentrated among a number of key research collaborators and/or scientific advisors. There can be no assurance that there will not be a detrimental impact on us if one or more of these key research collaborators and/or scientific advisors were to cease relationship or employment with us, potentially as a result of lateral recruitment by existing or new competitors. As a result, this may adversely affect our ability to conduct research and development on antibody product candidates.

Furthermore, our ability to continue to conduct and expand operations depends on our ability to attract and retain a large and growing number of personnel. The ability to meet our expertise needs, including the ability to find qualified personnel to fill positions that become vacant at our research and development department or to collaborate with us in research and development efforts, while controlling our costs, is generally subject to numerous external factors, including the availability of a sufficient number of qualified persons in the biopharmaceutical industry, the unemployment levels within those markets, prevailing wage rates, changing demographics, health and other insurance costs and adoption of new or revised employment and labor laws and regulations. If we are unable to locate, to attract or to retain qualified personnel, the quality of services and products provided to customers may decrease and our financial performance may be adversely affected. In addition, if costs of labor or related costs to maintain relationships with research collaborators increase for other reasons or if new or revised labor laws, rules or regulations or healthcare laws are adopted or implemented that further increase labor costs, our business, financial condition and results of operations could be materially adversely affected.

We may not be able to attract and retain key senior management members or research and development personnel.

Our future success depends upon the continuing services of members of our senior management team and key research and development personnel and consultants. In particular, Peter Luo, our Chief Executive Officer, Fangyong (Felix) Du, our Chief Technology Officer, Hua Gong, our Chief Operating Officer and Head of Clinical Development and Precision Medicine, JC Xu, our Chief Scientific Officer, Qinghai Zhao, our Chief Manufacturing Officer and Yu (Albert) Ren, our Chief Strategy and Corporate Development Officer are crucial to our research and development and operations. Although we typically require our key personnel to enter into non-compete and confidentiality agreements with us, we cannot prevent them from joining our competitors after the non-compete period. The loss of their services could adversely impact our ability to achieve our business objectives. If one or more of our senior management or key clinical and scientific personnel are unable or unwilling to continue in their present positions or joins a competitor or forms a competing company, we may not be able to replace them in a timely manner or at all, which will have a material and adverse effect on our business, financial condition and results of operations. We do not maintain "key person" insurance for any of our executives or other employees.

In addition, the continued growth of our business depends on our ability to hire additional qualified personnel with expertise in molecular biology, chemistry, biological information processing, computational biology, software, engineering, sales, marketing, and technical support. We compete for qualified management and scientific personnel with other life science and technology companies, universities, and research institutions in China and overseas. Competition for these individuals is intense, and the turnover rate can be high. Failure to attract and retain management and scientific and engineering personnel could prevent us from pursuing collaborations or developing our product candidates or technologies.

We face risks related to health epidemics, severe weather conditions and other outbreaks.

China has in the past experienced significant natural disasters, including earthquakes, extreme weather conditions, as well as health scares related to epidemic diseases, and any similar event could materially impact our business in the future. If a disaster or other disruption were to occur in the future that affects the regions where we operate our business, our operations could be materially and adversely affected due to loss of personnel and damage to property. Even if we are not directly affected, such a disaster or disruption could affect the operations or financial conditions of our customers, which could harm our results of operations.

In addition, our business could be affected by public health epidemics and pandemics, such as the outbreak of avian influenza, severe acute respiratory syndrome, or SARS, Zika virus, Ebola virus or other diseases. In late December 2019, a strain of SARS-CoV-2, which causes the COVID-19 disease, was reported to have surfaced in Wuhan, China. On January 30, 2020, the World Health Organization reportedly declared this COVID-19 outbreak a health emergency of international concern. On February 28, 2020, the World Health Organization reportedly increased the assessment of the risk of spread and the risk of impact of COVID-19 to very high at global level. In March 2020, the World Health Organization declared the COVID-19 a pandemic. As COVID-19 has evolved into a worldwide health crisis, it has resulted in adverse effects in the global economy and financial markets, such as significant declines in the global stock markets. If the COVID-19 outbreak is not effectively controlled globally, our business and results of operations could be adversely affected to the extent the COVID-19 outbreak harms the Chinese or world economy generally. The extent to which the COVID-19 outbreak impacts our financial condition and results of operations for the full year of 2020 cannot be reasonably estimated at this time and will depend on future developments that currently cannot be predicted, including the development of a COVID-19 vaccine and the actions taken to contain the COVID-19 outbreak, among others. Any future outbreak of public health epidemics may restrict economic activities in affected regions, disrupt our business operations and adversely affect our results of operations.

The COVID-19 pandemic could adversely impact our business, including our clinical trials.

The spread of the COVID-19 coronavirus in many countries continues to adversely impact global economic activity and has contributed to significant volatility and negative pressure in financial markets and supply chains. The pandemic has had, and could have a significantly greater, material adverse effect on the global economy. The pandemic has resulted, and may continue to result for an extended period, in significant disruption of global financial markets, which may reduce our ability to access capital in the future, which could negatively affect our liquidity.

The COVID-19 pandemic has adversely affected the clinical development of our product candidates. Our clinical development program timelines could continue to be negatively affected by COVID-19, which could materially and adversely affect our business, financial condition, and results of operations. Further, due to "shelter in place" orders and other public health guidance measures, we may be required to implement a work-from-home policy for all staff members excluding those necessary to maintain minimum basic operations. In such an instance, our increased reliance on personnel

working from home may negatively impact productivity, or disrupt, delay or otherwise adversely impact our business. For example, with our personnel working from home, some of our research activities that require our personnel to be in our laboratories may be delayed.

The COVID-19 pandemic may impact our workforce, supply chains or distribution networks or otherwise impact our ability to restock our medical device and supply inventories and depending upon the severity of the COVID-19 pandemic's continued spread in the United States and other countries, we may experience disruptions that could severely impact our business and clinical trials, including:

- limitation of company operations, including work from home policies and office closures;
- one or more key officers and/or employees could contract COVID-19 or otherwise be adversely affected by the virus;
- delays or difficulties in receiving deliveries of critical experimental materials;
- delays or difficulties in enrolling patients in our clinical trials;
- delays or difficulties in clinical site initiation or expansion, including difficulties in recruiting clinical site investigators and clinical site staff;
- increased rates of patients withdrawing from our clinical trials following enrollment as a result of contracting COVID-19 or other health conditions or being forced to quarantine;
- delays or disruptions in preclinical experiments and IND-enabling studies due to restrictions of on-site staff and unforeseen circumstances at contract research organizations, or CROs, and vendors;
- interruption or delays in the operations of the FDA or other regulatory authorities, which may impact review and approval timelines;
- diversion of healthcare resources away from the conduct of clinical trials, including the diversion of hospitals serving as our clinical trial sites and hospital staff supporting the conduct of our clinical trials;
- interruption of key clinical trial activities, such as clinical trial site monitoring, due to limitations on travel imposed or recommended by federal or state governments, employers and others;
- changes in regulations as part of a response to the COVID-19 pandemic, which may require us to change the ways in which our clinical trials are conducted, which may result in unexpected costs, or to discontinue the clinical trials altogether;
- delays in necessary interactions with regulators, ethics committees and other important agencies and contractors due to limitations in employee resources or forced furlough of government or contractor personnel;
- refusal of the FDA or other regulatory authorities to accept data from clinical trials in affected geographies; and
- limitations in employee resources that would otherwise be focused on our business, including the conduct of our clinical trials, such as because of sickness of employees or their families or the desire of employees to avoid contact with large groups of people.

The global outbreak of the COVID-19 coronavirus continues to rapidly evolve. The extent to which the COVID-19 coronavirus may impact our business and clinical trials will depend on future developments, which are highly uncertain and cannot be predicted with confidence, such as the ultimate geographic spread of the disease, the duration of the outbreak, travel restrictions and social distancing in China, the United States and other countries, business closures or business disruptions and the effectiveness of actions taken in the United States and other countries to contain and treat the disease.

We have limited insurance coverage, and any claims beyond our insurance coverage may result in us incurring substantial costs and a diversion of resources.

We maintain insurance policies that are required under PRC laws and regulations as well as insurance based on our assessment of our operational needs and industry practice. We also maintain liability insurance covering our clinical trials. In line with industry practice in the PRC, we have elected not to maintain certain types of insurances, such as business interruption insurance or key-man insurance. Our insurance coverage may be insufficient to cover any claim for product liability, damage to our fixed assets or employee injuries. Any uninsured risks may result in substantial costs and the diversion of resources, which could adversely affect our results of operations and financial condition.

We have adopted a share incentive plan and will continue to grant share-based awards in the future, which may increase expenses associated with share-based compensation. Exercise of the awards granted will increase the number of our outstanding ordinary shares, which may adversely affect the market price of our ADSs.

We adopted the Second Amended and Restated Share Incentive Plan in December 2019 and the 2021 Performance Incentive Plan in January 2021, which we refer to as the 2019 Plan and the 2021 Plan, respectively, in this prospectus, to enhance our ability to attract and retain exceptionally qualified individuals and to encourage them to acquire a proprietary interest in the growth and performance of us. Upon the effectiveness of our registration statement on Form F-1, we will terminate the authority to grant additional awards under the 2019 Plan and all future awards will be granted under the 2021 Plan. Therefore, the effective maximum number of shares issuable under the 2019 Plan will be 10,728,501. A total of 2,994,000 of our ordinary shares is authorized for issuance with respect to awards granted under the 2021 Plan. As of the date of this prospectus, the aggregate number of our ordinary shares underlying our outstanding awards under the 2019 Plan is 3,601,522. We have not granted any awards under the 2021 Plan. See "Management—Share Incentive Plan."

We believe the granting of share-based awards is of significant importance to our ability to attract and retain key personnel and employees, and we will continue to grant share-based compensation to employees in the future. As a result, our expenses associated with share-based compensation may increase, which may have an adverse effect on our results of operations.

Our employees, third-party suppliers, consultants and commercial partners may engage in misconduct or other improper activities, including noncompliance with regulatory standards and requirements, and insider trading.

We are exposed to the risk of fraud or other misconduct by our employees, third-party suppliers, consultants and commercial partners. Misconduct by these parties could include intentional failures to comply with the regulations of the NMPA and overseas regulators that have jurisdictions over us, comply with healthcare fraud and abuse laws and regulations in China and abroad, report financial information or data accurately or disclose unauthorized activities to us. In particular, sales, marketing, and business arrangements in the healthcare industry are subject to extensive laws and regulations intended to prevent fraud, misconduct, kickbacks, self-dealing and other abusive practices. These laws and regulations may restrict or prohibit a wide range of pricing, discounting, marketing and promotion, sales commission, customer incentive programs and other business arrangements. Such misconduct could also involve the improper use of information obtained in the course of clinical trials, which could result in regulatory sanctions and cause serious harm to our reputation. We currently have a code of conduct applicable to all of our employees, but it is not always possible to identify and deter employee misconduct, and our code of conduct and the other precautions we take to detect and prevent this activity may not be effective in controlling unknown or unmanaged risks or losses, or in protecting us from governmental investigations or other actions or lawsuits stemming from a failure to comply with these laws or regulations. If any such actions are instituted against us, and we are not successful in defending ourselves or asserting our rights, those actions could result in the imposition of significant civil, criminal and administrative penalties, including, without limitation, damages, monetary fines,

individual imprisonment, disgorgement of profits, contractual damages, reputational harm, diminished profits and future earnings, additional reporting or oversight obligations if we become subject to a corporate integrity agreement or other agreement to resolve allegations of noncompliance with the law and curtailment or restructuring of our operations, which could have a significant impact on our business. Whether or not we are successful in defending against such actions or investigations, we could incur substantial costs, including legal fees and divert the attention of management in defending ourselves against any of these claims or investigations.

We may be subject to liability lawsuits arising from our clinical trials.

We currently carry liability insurance covering our clinical trials. Although we maintain such insurance, any claim that may be brought against us could result in a court judgment or settlement in an amount that is not covered, in whole or in part, by our insurance or which is in excess of the limits of our insurance coverage. Our insurance policies also contain various exclusions, and we may be subject to particular liability claims for which we have no coverage. We will have to pay any amount awarded by a court or negotiated in a settlement that exceed our coverage limitations or that are not covered by our insurance, and we may not have, or be able to obtain, sufficient capital to pay such amounts. In addition, if we cannot successfully defend ourselves against such claims, we may incur substantial liabilities and be required to suspend or delay our ongoing clinical trials. Even a successful defense would require significant financial and management resources.

Regardless of the merits or eventual outcome, liability claims may result in significant negative consequences to our business and prospects, including, but not limited to:

- decreased demand for our product candidates or any resulting products;
- damage to our reputation;
- withdrawal of other clinical trial participants;
- costs to defend the related litigation;
- a diversion of our management's time and resources;
- substantial monetary awards to trial participants or patients;
- inability to commercialize our product candidates; and
- a decline in the market price of our ADSs.

We are subject to changing law and regulations regarding regulatory matters, corporate governance and public disclosure that have increased both our costs and the risk of noncompliance.

We are subject to rules and regulations by various governing bodies, including, for example, the FDA, the NMPA, the SEC, which is charged with the protection of investors and the oversight of companies whose securities are publicly traded, and the various regulatory authorities in China, the United States, the EU, the Cayman Islands, and to new and evolving regulatory measures under applicable law. Our efforts to comply with new and changing laws and regulations have resulted in and are likely to continue to result in, increased general and administrative expenses and a diversion of management time and attention from revenue-generating activities to compliance activities.

Moreover, because these laws, regulations and standards are subject to varying interpretations, their application in practice may evolve over time as new guidance becomes available. This evolution may result in continuing uncertainty regarding compliance matters and additional costs necessitated by ongoing revisions to our disclosure and governance practices. If we fail to address and comply with these regulations and any subsequent changes, we may be subject to penalties and our business may be harmed.

We may be exposed to liabilities under the U.S. Foreign Corrupt Practices Act and Chinese anti-corruption laws, and any determination that we have violated these laws could have a material adverse effect on our business or our reputation.

We are subject to anti-bribery laws in China that generally prohibit companies and their intermediaries from making payments to government officials for the purpose of obtaining or retaining business or securing any other improper advantage. In addition, although currently our primary operating business is in China, we are subject to the Foreign Corrupt Practices Act, or the FCPA. The FCPA generally prohibits us from making improper payments to non-U.S. officials for the purpose of obtaining or retaining business. We are also subject to the anti-bribery laws of other jurisdictions, particularly China. As our business expands, the applicability of the FCPA and other anti-bribery laws to our operations will increase. Our procedures and controls to monitor anti-bribery compliance may fail to protect us from reckless or criminal acts committed by our employees or agents. If we, due to either our own deliberate or inadvertent acts or those of others, fail to comply with applicable anti-bribery laws, our reputation could be harmed and we could incur criminal or civil penalties, other sanctions and/or significant expenses, which could have a material adverse effect on our business, including our financial condition, results of operations, cash flows and prospects.

Any failure to comply with applicable regulations and industry standards or obtain various licenses and permits could harm our reputation, business, results of operations and prospects.

A number of governmental agencies or industry regulatory bodies in China, the United States and other applicable jurisdictions impose strict rules, regulations and industry standards governing biopharmaceutical research and development activities, which apply to us. Our or our CROs' failure to comply with such regulations could result in the termination of ongoing research, administrative penalties imposed by regulatory bodies or the disqualification of data for submission to regulatory authorities. This could harm our business, reputation, prospects for future work and results of operations. For example, if we or our CROs were to treat research animals inhumanely or in violation of international standards set out by the Association for Assessment and Accreditation of Laboratory Animal Care, it could revoke any such accreditation and the accuracy of our animal research data could be questioned.

If we or our third-party research collaborators or other contractors or consultants fail to comply with environmental, fire protection, drainage or health and safety laws and regulations, we could become subject to fines or penalties or incur costs that could have a material adverse effect on the success of our business.

We and third parties, such as our CROs, are subject to numerous environmental, fire protection, drainage or health and safety laws and regulations, including but not limited to those governing laboratory procedures and the handling, use, storage, treatment and disposal of hazardous materials and wastes discharge of stationary pollution sources. The cost of compliance with health and safety regulations is substantial. Our business activities involve the controlled use of hazardous materials. Our research and development activities involve the controlled storage, use and disposal of hazardous materials, including the components of our product candidates and other hazardous compounds. We generally contract with third parties for the disposal of these materials and wastes. We cannot eliminate the risk of contamination or injury from these materials, which could cause an interruption of our commercialization efforts, research and development efforts and business operations. We cannot guarantee that the safety procedures utilized by our partners and by third-party manufacturers and suppliers with whom we may contract will comply with the standards prescribed by laws and regulations or will eliminate the risk of accidental contamination or injury from these materials. In such an event, we could be held liable for any resulting damages, and such liability could exceed our resources. In addition, we may be required to incur substantial costs to comply with current or future environmental, health and safety laws and regulations which are complex, change frequently and have tended to

become more stringent. We do not currently carry biological or hazardous waste insurance coverage. In the event of an accident or environmental discharge, we may be held liable for any consequential damage and any resulting claims for damages, which may exceed our financial resources and may materially adversely affect our business, financial condition, results of operations and future growth prospects, and the value of our ADSs.

Security breaches, loss of data, and other disruptions could compromise sensitive information related to our business or prevent us from accessing critical information and expose us to liability, which could adversely affect our business and our reputation.

In the ordinary course of our business, we collect and store preclinical trial data and clinical trial data which could be sensitive, including research and development information, health-related information, personally identifiable information, intellectual property, and proprietary business information owned or controlled by ourselves or other parties. We manage and maintain our applications and data utilizing a combination of on-site systems and cloud-based application systems. We utilize external security and infrastructure vendors to manage parts of our data centers. We also communicate sensitive data with third parties. We face a number of risks relative to protecting this critical information, including material system failure or security breach, loss of access risk, inappropriate use or disclosure, inappropriate modification, and the risk of our being unable to adequately monitor, audit, and modify our controls over our critical information. This risk extends to the third-party vendors and subcontractors we use to manage this sensitive data and third-party collaborators who share with us sensitive data.

Despite the implementation of security measures to protect sensitive data from unauthorized access, use or disclosure, our information technology and infrastructure may be vulnerable to attacks by hackers or viruses or breached due to employee error, malfeasance, or other malicious or inadvertent disruptions that could result in unauthorized access to, use or disclosure of, corruption of, or loss of sensitive, and/ or proprietary data, including health-related and other personal information, and could subject us to significant liabilities and regulatory and enforcement actions, and reputational damage. The COVID-19 pandemic is generally increasing the attack surface available to criminals, as more companies and individuals work online and work remotely, and as such, the risk of a cybersecurity incident potentially occurring, and our investment in risk mitigations against such an incident, is increasing. For example, there has been an increase in phishing and spam emails as well as social engineering attempts from "hackers" hoping to use the COVID-19 pandemic to their advantage. In addition, while we have implemented security measures to prevent unauthorized access to patient data, such data is currently accessible through multiple channels, and there is no guarantee we can protect our data from breach. Unauthorized access, loss, or dissemination could also result in delays of our product development and regulatory approval efforts as well as damage our reputation.

For example, the loss of clinical trial data from completed or ongoing clinical trials could result in delays in any regulatory approval or clearance efforts and significantly increase our costs to recover or reproduce the data, and subsequently commercialize the product. If we or our third-party collaborators, consultants, contractors, suppliers, or service providers were to suffer an attack or breach, for example, that resulted in the unauthorized access to or use or disclosure of health-related or other personal information, we may have to notify consumers, partners, collaborators, government authorities, and the media, and may be subject to investigations, civil penalties, administrative and enforcement actions, and litigation, any of which could harm our business and reputation. Likewise, we rely on our third-party research institution collaborators and other third parties to conduct clinical trials, and similar events relating to their computer systems could also have a material adverse effect on our business.

Our insurance policies may not be adequate to compensate us for the potential losses arising from such disruptions, failure, or security breach. In addition, such insurance may not be available to us in the future on economically reasonable terms, or at all. Further, our insurance may not cover all claims

made against us and defending a suit, regardless of its merit, could be costly, divert management attention, and harm our reputation.

Failure to comply with existing or future laws and regulations related to privacy or data security could lead to government enforcement actions, which could include civil or criminal fines or penalties, private litigation, other liabilities, and/or adverse publicity. Compliance or the failure to comply with such laws could increase the costs of our products and services, limit their use or adoption, and otherwise negatively affect our operating results and business.

The regulatory framework for the collection, use, safeguarding, sharing, transfer and other processing of personal information worldwide is rapidly evolving and is likely to remain uncertain for the foreseeable future. Regulatory authorities in virtually every jurisdiction in which we operate have implemented and are considering a number of legislative and regulatory proposals concerning personal data protection.

Regulatory authorities in China have implemented and are considering a number of legislative and regulatory proposals concerning data protection. For example, China's Cyber Security Law, which became effective in June 2017, created China's first national-level data protection for "network operators," which may include all organizations in China that provide services over the internet or another information network. Numerous regulations, guidelines and other measures are expected to be adopted under the umbrella of the Cyber Security Law. Drafts of some of these measures have now been published, including (i) the draft rules on cross-border data transfers of personal information and important data published by the China Cyberspace Administration in 2017, and draft rules on measures for security assessment for cross-border transfer of personal information published by China Cyberspace Administration in 2019, which may, upon enactment, require security review before transferring human health-related data out of China, and (ii) the Draft Data Security Law promulgated by the Standing Committee of PRC National People's Congress in 2020, which outlines the main system framework of data security protection. In addition, certain industry-specific laws and regulations affect the collection and transfer of personal data in China. For example, the PRC State Council promulgated Regulations on the Administration of Human Genetic Resources (effective in July 2019), which require approval from or filings with the Science and Technology Administration Department of the State Council where human genetic resources, or HGR, are involved in any international collaborative project and additional approval for any export or cross-border transfer of the HGR samples or human genetic resource information. It is possible that these laws may be interpreted and applied in a manner that is inconsistent with our practices, potentially resulting in confiscation of HGR samples and human genetic resource information and administrative fines or in worst cases, criminal penalties. In addition, the interpretation and application of data and personal information protection laws in China and elsewhere are often uncertain and in flux.

In the United States, we and our partners may be subject to state and federal laws and regulations that govern data privacy, protection and security. Numerous laws and regulations, including security breach notification laws, health information privacy laws, and consumer protection laws, govern the collection, use, disclosure and protection of health-related and other personal information. In addition, we may obtain health information from third parties (including research institutions from which we obtain clinical trial data) that are subject to privacy and security requirements under the Health Insurance Portability and Accountability Act of 1996, as amended by the Health Information Technology for Economic and Clinical Health Act, and regulations implemented thereunder (collectively, "HIPAA"). Depending on the facts and circumstances, we could be subject to criminal penalties if we knowingly obtain, use, or disclose individually identifiable health information maintained by a HIPAA-covered entity in a manner that is not authorized or permitted by HIPAA.

Even when HIPAA does not apply, according to the Federal Trade Commission, or the FTC, violating consumers' privacy rights or failing to take appropriate steps to keep consumers' personal

information secure may constitute unfair acts or practices in or affecting commerce in violation of Section 5(a) of the Federal Trade Commission Act. The FTC expects a company's data security measures to be reasonable and appropriate in light of the sensitivity and volume of consumer information it holds, the size and complexity of its business, and the cost of available tools to improve security and reduce vulnerabilities. Individually identifiable health information is considered sensitive data that merits stronger safeguards.

In addition, certain state and non-U.S. laws, such as the European Union General Data Protection Regulation, or the GDPR, govern the privacy and security of health information and other personal information in certain circumstances, many of which differ from each other in significant ways and may not have the same effect, thus complicating compliance efforts. Failure to comply with these laws, where applicable, can result in the imposition of significant civil and/or criminal penalties and private litigation. For example, California recently enacted legislation, the California Consumer Privacy Act, or CCPA, which went into effect January 1, 2020. The CCPA, among other things, creates new data privacy obligations for covered companies and provides new privacy rights to California residents, including the right to opt out of certain disclosures of their information. The CCPA also creates a private right of action with statutory damages for certain data breaches, thereby potentially increasing risks associated with a data breach. In Europe, the GDPR went into effect in May 2018 and introduces strict requirements for processing the personal data of individuals within the EU, and the European Economic Area, or the EEA. In addition, the GDPR increases the scrutiny of transfers of personal data from clinical trial sites located in the EEA to the United States and other jurisdictions that the European Commission does not recognize as having "adequate" data protection laws, and recent legal developments in Europe have created complexity and compliance uncertainty regarding certain transfers of personal data from the EEA. For example, on July 16, 2020, the Court of Justice of the European Union, or the CJEU, invalidated the EU-US Privacy Shield Framework, or Privacy Shield, under which personal data could be transferred from the EU and the EEA to United States entities who had self-certified under the Privacy Shield scheme. Moreover, it is uncertain whether the standard contractual clauses will also be invalidated by the European courts or legislature. Companies that must comply with the GDPR face increased compliance obligations and risk, including more robust regulatory enforcement of data protection requirements and potential fines for noncompliance of up to €20 million or 4% of the annual global revenues of the noncompliant company, whichever is greater. Additionally, following Brexit, companies will have to comply with the GDPR and the GDPR as incorporated into United Kingdom national law, the latter regime having the ability to separately fine up to the greater of £17.5 million or 4% of global turnover. The relationship between the U.K. and the EU in relation to certain aspects of data protection law remains unclear, for example around how data can lawfully be transferred between each jurisdiction, which exposes us to further compliance risk.

Given the variability and evolving state of these laws, we face uncertainty as to the exact interpretation of the new requirements, and we may be unsuccessful in implementing all measures required by regulators or courts in their interpretation.

We expect that we will continue to face uncertainty as to whether our efforts to comply with evolving obligations under global data protection, privacy and security laws will be sufficient. Any failure or perceived failure by us to comply with applicable laws and regulations could result in reputational damage or proceedings or actions against us by governmental entities, individuals or others. These proceedings or actions could subject us to significant civil or criminal penalties and negative publicity, result in the delayed or halted transfer or confiscation of certain personal information, require us to change our business practices, increase our costs and materially harm our business, prospects, financial condition and results of operations. In addition, our current and future relationships with customers, vendors, pharmaceutical partners and other third parties could be negatively affected by any proceedings or actions against us or current or future data protection obligations imposed on them under applicable law, including the GDPR. In addition, a data breach

affecting personal information, including health information, could result in significant legal and financial exposure and reputational damage that could potentially have an adverse effect on our business.

Business disruptions could seriously harm our future revenue, increase our costs and expenses, and have adverse effect on our financial condition.

Our operations and third parties with which we have collaborations could be subjected to earthquakes, power shortages, telecommunications failures, water shortages, floods, hurricanes, typhoons, fires, extreme weather conditions, medical epidemics and other natural or man-made disasters or business interruptions, for which we are predominantly self-insured. In addition, we partially rely on our CROs for conducting research and development, and they may be affected by government shutdowns or withdrawn funding. The occurrence of any of these business disruptions could seriously harm our operations and financial condition and increase our costs and expenses.

Although we maintain incident management and disaster response plans, in the event of a major disruption caused by a natural disaster or man-made problem, such as power disruptions, computer viruses, data security breaches or terrorism, we may be unable to continue our operations and may endure system interruptions, reputational harm, delays in our development activities, lengthy interruptions in service, breaches of data security and loss of critical data, any of which could adversely affect our business, results of operations and financial condition.

If we fail to implement and maintain an effective system of internal controls, we may be unable to accurately or timely report our results of operations or prevent fraud, and investors' confidence and the market price of our ADSs may be materially and adversely affected.

Prior to this offering, we were a private company with limited accounting personnel and other resources with which to address our internal controls and procedures. Our management has not completed an assessment of the effectiveness of our internal control over financial reporting, and our independent registered public accounting firm has not conducted an audit of our internal control over financial reporting. In the course of auditing our consolidated financial statements as of December 31, 2019 and for the year ended December 31, 2019, we and our independent registered public accounting firm identified two material weaknesses in our internal control over financial reporting and other control deficiencies as of December 31, 2019. A "material weakness" is a deficiency, or a combination of deficiencies, in internal control over financial reporting, such that there is a reasonable possibility that a material misstatement of the company's annual or interim financial statements will not be prevented or detected on a timely basis. The material weaknesses identified relate to:

- Our lack of sufficient and competent financial reporting and accounting personnel with appropriate knowledge of U.S. GAAP and SEC reporting and compliance requirements; and
- Our lack of sufficient documented financial closing policies and procedures, specifically those related to period end expenses cut-off and accruals.

We have taken measures and plan to continue to take measures to remedy the material weaknesses. For details, please refer to "Management's Discussion and Analysis of Financial Condition and Results of Operations—Internal Control over Financial Reporting." The implementation of these measures may not fully address the material weaknesses in our internal control over financial reporting, and we cannot conclude that they have been fully remedied. Our failure to correct these material weaknesses or our failure to discover and address any other material weaknesses could result in inaccuracies in our financial statements and could also impair our ability to comply with applicable financial reporting requirements and related regulatory filings on a timely basis.

Upon the completion of this offering, we will become a public company in the United States subject to the Sarbanes-Oxley Act of 2002. Section 404 of the Sarbanes-Oxley Act of 2002, or Section 404, will require that we include a report from management on our internal control over financial reporting in our annual report on Form 20-F beginning with our annual report for the fiscal year ending December 31, 2021. In addition, once we cease to be an "emerging growth company" as such term is defined in the JOBS Act, our independent registered public accounting firm must attest to and report on the effectiveness of our internal control over financial reporting. Our management may conclude that our internal control over financial reporting is not effective. Moreover, even if our management concludes that our internal control over financial reporting is effective, our independent registered public accounting firm, after conducting its own independent testing, may issue a report that is qualified if it is not satisfied with our internal controls or the level at which our controls are documented, designed, operated or reviewed, or if it interprets the relevant requirements differently from us. In addition, after we become a public company, our reporting obligations may place a significant strain on our management, operational and financial resources and systems for the foreseeable future. We may be unable to timely complete our evaluation testing and any required remediation.

During the course of documenting and testing our internal control procedures, in order to satisfy the requirements of Section 404, we may identify weaknesses and deficiencies in our internal control over financial reporting. In addition, if we fail to maintain the adequacy of our internal control over financial reporting, as these standards are modified, supplemented or amended from time to time, we may not be able to conclude on an ongoing basis that we have effective internal control over financial reporting in accordance with Section 404. Generally speaking, if we fail to achieve and maintain an effective internal control environment, we could suffer material misstatements in our financial statements and fail to meet our reporting obligations, which would likely cause investors to lose confidence in our reported financial information. This could, in turn, limit our access to capital markets, harm our results of operations and lead to a decline in the trading price of our ADSs. Additionally, ineffective internal control over financial reporting could expose us to increased risk of fraud or misuse of corporate assets and subject us to potential delisting from the stock exchange on which we list, regulatory investigations and civil or criminal sanctions.

Risks Related to Our Intellectual Property

It is difficult and costly to protect our proprietary rights and technology, and we may not be able to protect our intellectual property rights throughout the world.

Our commercial success will depend, in part, on our ability to obtain, maintain and defend patent and other intellectual property protection (including trademarks and trade secrets) with respect to our product candidates. We cannot be certain that patents will be issued or granted with respect to our patent applications that are currently pending, or that issued or granted patents will not later be found to be invalid and/or unenforceable, be interpreted in a manner that does not adequately protect our product candidates, or otherwise provide us with any competitive advantage. Additionally, the patent applications in respect of patents licensed under our in-license arrangements may not be issued or granted, and as a result, we may not be able to have adequate protection with respect to such patents.

The patent position of biotechnology and pharmaceutical companies is generally uncertain because it involves complex legal and factual considerations. Patent applications we have filed may not be granted or issued as valid enforceable patents. Moreover, some of our patents and patent applications may in the future be, co-owned with third parties. If we are unable to obtain an exclusive license to any such third-party co-owned interest in such patents or patent applications, such co-owners may be able to license or transfer their rights to other third parties, including our competitors, and our competitors could market competing products and technology. In addition, we may need the cooperation of any such co-owners of our patents in order to enforce such patents against third parties, and such

cooperation may not be provided to us. Any of the foregoing could have a material adverse effect on our competitive position, business, financial conditions, results of operations and prospects. As such, we do not know the degree of future protection that we will have on our product candidates and technology, if any, and a failure to obtain adequate intellectual property protection with respect to our product candidates could have a material adverse impact on our business.

Composition-of-matter patents on the active pharmaceutical ingredient are generally considered to be the strongest form of intellectual property protection for pharmaceutical products, as such patents provide protection without regard to any method of use. Method-of-use patents protect the use of a product for the specified method. This type of patent does not prevent a competitor from making and marketing a product that is identical to our products for an indication that is outside the scope of the patented method. Moreover, even if competitors do not actively promote their product for our targeted indications, physicians may prescribe these products "off-label." Although off-label prescriptions may infringe or contribute to the infringement of method-of use patents, the practice is common and such infringement is difficult to prevent or prosecute. We endeavor to seek composition-of-matter patent protection for all of our product candidates. Where appropriate, we also seek method-of-use patents and patents protecting other aspects of our product candidates, including processes for discovery and manufacturing.

Even if our patent applications issue as patents, they may not issue in a form that will provide us with any meaningful protection, prevent competitors from competing with us or otherwise provide us with any competitive advantage. Our competitors may be able to circumvent our owned or licensed patents by developing similar or alternative technologies or products in a non-infringing manner. The issuance of a patent is not conclusive as to its scope, validity or enforceability, and our owned and in-licensed patents may be challenged in the courts or patent offices in the PRC and abroad. For example, we may become involved in opposition, interference, derivation, inter partes review or other similar proceedings challenging our patent rights, and the outcome of any proceedings are highly uncertain. Such challenges may result in the patent claims of our owned or in-licensed patents being narrowed, invalidated or held unenforceable, which could limit our ability to stop or prevent us from stopping others from using or commercializing similar or identical technology and products, or limit the duration of the patent protection of our technology and products. Given the amount of time required for the development, testing and regulatory review of new product candidates, patents protecting such candidates might expire before or shortly after such candidates are commercialized. As a result, our patent portfolio may not provide us with sufficient rights to exclude others from commercializing products similar or identical to ours or otherwise provide us with a competitive advantage.

Despite the measures we can take to increase our likelihood of obtaining patent and other intellectual property protections with respect to our product candidates, there can be no assurance that the existence, validity, enforceability, or scope of our intellectual property rights will not be challenged by a third party, or that we can obtain sufficient scope of claim in those patents to prevent a third party from competing against our product candidates. For example, in an infringement proceeding, a court may hold that patent rights or other intellectual property rights owned by us are invalid or unenforceable, or may refuse to stop the other party from using the technology at issue on the ground that our patent rights or other intellectual property rights do not cover the technology in question. An adverse result in any litigation proceedings could put our patent, as well as any patents that may issue in the future from our pending patent applications, at risk of being invalidated, held unenforceable or interpreted narrowly. Furthermore, because of the substantial amount of discovery required in connection with intellectual property litigation, there is a risk that some of our confidential information could be compromised by disclosure during this type of litigation.

In addition, if we were to initiate legal proceedings against a third party to enforce a patent covering one of our product candidates, the defendant could counterclaim that our patent is invalid and/or unenforceable. Grounds for a validity challenge could be an alleged failure to meet any of

several statutory requirements, for example, lack of novelty, obviousness or non-enablement. Grounds for an unenforceability assertion could be an allegation that someone connected with prosecution of the patent withheld relevant information from the National Intellectual Property Administration of China, or NIPA, or the applicable foreign counterpart, or made a misleading statement, during prosecution. Although we believe that we have conducted our patent prosecution in accordance with all applicable duty of candor and in good faith, the outcome following legal assertions of invalidity and unenforceability during patent litigation is unpredictable. With respect to the validity question, for example, we cannot be certain that there is no invalidating prior art, of which we and the patent examiner were unaware during prosecution. If a defendant were to prevail on a legal assertion of invalidity and/or unenforceability, we would lose at least part, and perhaps all, of the patent protection on our product candidates. Even if a defendant does not prevail on a legal assertion of invalidity and/or unenforceability, our patent claims may be construed in a manner that would limit our ability to enforce such claims against the defendant and others.

Third parties may also raise similar claims before administrative bodies in the PRC or abroad, even outside the context of litigation. Such mechanisms include ex parte re-examination, inter partes review, post-grant review, derivation and equivalent proceedings, such as opposition proceedings. Such legal proceedings could result in revocation or amendment to our patents in such a way that they no longer cover and protect our product candidates. The outcome following legal assertions of invalidity and unenforceability can be unpredictable. With respect to the validity of our patents, for example, we cannot be certain that there is no invalidating prior art of which we and the patent examiner were unaware during prosecution. If a defendant were to prevail on a legal assertion of invalidity and/or unenforceability, we may lose part or all of the patent protection on our product candidates. Any loss of patent protection could have a material adverse impact on one or more of our product candidates and our business.

In addition to the protection afforded by patents, we seek to rely on trade secret protection and confidentiality agreements to protect proprietary know-how that is not patentable, processes for which patents are difficult to enforce and any other elements of our drug discovery and development processes that involve proprietary know-how, information or technology that is not covered by patents. We may also rely on trade secret protection as temporary protection for concepts that may be included in a future patent filing. However, trade secret protection will not protect us from innovations that a competitor develops independently of our proprietary know-how. If a competitor independently develops a technology that we protect as a trade secret and files a patent application on that technology, then we may not be able to patent that technology in the future, may require a license from the competitor to use our own know-how, and if the license is not available on commercially viable terms, then we may not be able to launch our product. Although we require our employees to assign their inventions to us, and require our employees, consultants, advisors and any third parties who have access to our proprietary know-how, information or technology to enter into confidentiality agreements, we cannot be certain that our trade secrets and other confidential proprietary information will not be disclosed or that competitors will not otherwise gain access to our trade secrets or independently develop substantially equivalent information and techniques. Furthermore, the laws of different countries do not protect proprietary rights to the same extent or in the same manner as the laws of the PRC. We may encounter significant problems in protecting and defending our intellectual property both in the PRC and abroad. If we are unable to prevent unauthorized material disclosure of our intellectual property to third parties, we will not be able to establish or maintain a competitive advantage in our market, and this scenario could materially adversely affect our business, financial condition and results of operations.

Moreover, trade secrets are difficult to protect, and we have limited control over the protection of trade secrets used by our collaborators and suppliers. Although we use reasonable efforts to protect our trade secrets, our employees, consultants, contractors, outside scientific collaborators and other advisors

may unintentionally or willfully disclose our information to competitors or use such information to compete with us. Moreover, our competitors may independently develop equivalent knowledge, methods and know-how. If our confidential or proprietary information is divulged to or acquired by third parties, including our competitors, our competitive position in the marketplace will be adversely affected and this would have a material adverse effect on our business.

Many companies have encountered significant problems in protecting and defending intellectual property rights in certain countries. The legal systems of certain countries do not favor the enforcement or protection of patents, trade secrets and other intellectual property, particularly those relating to pharmaceutical and biotechnology products, which could make it difficult for us to stop the infringement of our patents or marketing of competing products in violation of our proprietary rights generally. Proceedings to enforce our patent rights in certain countries could result in substantial costs and divert our efforts and attention from other aspects of our business, could put our patents at risk of being invalidated or interpreted narrowly and our patent applications at risk of not issuing and could provoke third parties to assert claims against us. We may not prevail in any lawsuits that we initiate and the damages or other remedies awarded, if any, may not be commercially meaningful. We and our contractors and partners operate in certain countries that are at heightened risk of theft of technology, data and intellectual property through direct or indirect intrusion by private parties or international actors, including those affiliated with or controlled by state actors. Accordingly, our efforts to protect or enforce our intellectual property rights around the world may be inadequate to obtain a significant commercial advantage from the intellectual property that we develop or license.

Changes in patent law could diminish the value of patents in general, thereby impairing our ability to protect our product candidates.

As is the case with other biopharmaceutical companies, our success is heavily dependent on intellectual property, particularly patents. Obtaining and enforcing patents in the pharmaceutical industry involves both technological and legal complexity and is therefore costly, time consuming and inherently uncertain. Changes in either the patent laws or interpretation of the patent laws in various countries could increase the uncertainties and costs surrounding the prosecution of patent applications and the enforcement or defense of issued patents. For example, patent reform legislation in the United States includes provisions that affect the way patent applications are prosecuted, redefine prior art and provide more efficient and cost-effective avenues for competitors to challenge the validity of patents via post-grant proceedings. The Leahy-Smith Act and any continuing changes in patent laws and regulations in various patent jurisdictions could make it more difficult to obtain patent protection for our inventions and increase the uncertainties and costs surrounding the prosecution of our or our collaboration partners' patent applications and the enforcement or defense of our or our collaboration partners' issued patents, all of which could harm our business, results of operations, financial condition and prospects.

In addition, the patent positions of companies in the development and commercialization of biologics and pharmaceuticals are particularly uncertain due to changes in law and courts' interpretation of the law. For example, the U.S. Supreme Court has ruled on several patent cases in recent years, either narrowing the scope of patent protection available in certain circumstances or weakening the rights of patent owners in certain situations. Other courts in the United States, for example, have heightened the bar for broadly claiming antibodies. This combination of events has created uncertainty with respect to the validity and enforceability of patents, once obtained. Additionally, there are periodic proposals for changes to the patent laws of China, United States and other countries that, if adopted, could impact our ability to enforce our proprietary technology. Depending on future actions by the relevant law-making bodies in other countries, the laws and regulations governing patents could change in unpredictable ways that could have a material adverse effect on our existing patent portfolio and weaken our ability to obtain new patents or to enforce our existing patents and patents that we might obtain in the future.

Similarly in China, intellectual property laws are constantly evolving, with efforts being made to improve intellectual property protection in China. For example, a Draft Amendment to the PRC Patent Law was released in January 2019 and updated in July 2020, which proposes introduction of patent term extensions to eligible innovative drug patents. If adopted, the terms of our Chinese patents may be eligible for extension and allow us to extend patent protection of our products, and the terms of the patents owned by third parties may also be extended, which may in turn affect our ability to commercialize our products candidates, if and when approved, without facing infringement risks. The length of any such patent term extension is uncertain. If we are required to delay commercialization for an extended period of time, technological advances may develop and new competitor products may be launched, which may render our product non-competitive. We also cannot guarantee that other changes to Chinese intellectual property laws would not have a negative impact on our intellectual property protection.

We may not be successful in obtaining or maintaining necessary rights for our development pipeline through acquisitions and licensing deals.

Because our programs may involve additional product candidates that may require the use of proprietary rights held by third parties, the growth of our business may depend in part on our ability to acquire and maintain licenses or other rights to use these proprietary rights. We may be unable to acquire or in-license any compositions, methods of use, or other intellectual property rights from third parties that we identify. The licensing and acquisition of third-party intellectual property rights is a competitive area, and a number of more established companies are also pursuing strategies to license or acquire third-party intellectual property rights that we may consider attractive or necessary. These established companies may have a competitive advantage over us due to their size, cash resources and greater clinical development and commercialization capabilities. In addition, companies that perceive us to be a competitor may be unwilling to assign or license rights to us. We also may be unable to license or acquire third-party intellectual property rights on terms that would allow us to make an appropriate return on our investment or at all. If we are unable to successfully obtain rights to required third-party intellectual property rights or maintain the existing intellectual property rights we have, we may have to abandon development of the relevant program or product candidate, which could have a material adverse effect on our business, financial condition, results of operations and prospects for growth.

Additionally, we may sometimes collaborate with academic institutions to accelerate our preclinical research or development under written agreements with these institutions. These institutions may provide us with an option to negotiate a license to any of the institution's rights in technology resulting from the collaboration. Regardless of such option, we may be unable to negotiate a license within the specified timeframe or under terms that are acceptable to us. If we are unable to do so, the institution may offer the intellectual property rights to others, potentially blocking our ability to pursue our program. If we are unable to successfully obtain rights to required third-party intellectual property or to maintain the existing intellectual property rights we have, we may have to abandon development of such program and our business and financial condition could suffer.

Licensing of intellectual property involves complex legal, business and scientific issues. Disputes may arise between us and our licensors regarding intellectual property subject to a license agreement, including:

- the scope of rights granted under the license agreement and other interpretation-related issues;
- whether and the extent to which our technology and processes infringe on intellectual property of the licensor that is not subject to the licensing agreement;
- our right to sublicense patents and other rights to third parties;
- our diligence obligations with respect to the use of the licensed technology in relation to our development and commercialization of our product candidates, and what activities satisfy those diligence obligations;

- our right to transfer or assign the license; and
- the ownership of inventions and know-how resulting from the joint creation or use of intellectual property by our licensors and us and our partners.

If disputes over intellectual property that we license in the future prevent or impair our ability to maintain our licensing arrangements on acceptable terms, we may not be able to successfully develop and commercialize the affected product candidates, which would have a material adverse effect on our business.

In addition, certain of our future agreements with third parties may limit or delay our ability to consummate certain transactions, may impact the value of those transactions, or may limit our ability to pursue certain activities. For example, we may in the future enter into license agreements that are not assignable or transferable, or that require the licensor's express consent in order for an assignment or transfer to take place.

We may become involved in lawsuits to protect or enforce our patents or other intellectual property, which could be expensive, time consuming and unsuccessful, and any unfavorable outcome from such litigation could limit our research and development activities and/or our ability to commercialize our product candidates.

Competitors may infringe our patent rights or misappropriate or otherwise violate our intellectual property rights. To counter infringement or unauthorized use, we may be required to file infringement claims, which can be expensive and time consuming and divert the time and attention of our management and scientific personnel. Any claims we assert against perceived infringers could provoke these parties to assert counterclaims against us, alleging that we infringed their patents. In addition, in a patent infringement proceeding, there is a risk that a court will decide that our asserted patents are invalid or unenforceable, in whole or in part, and that we do not have the right to stop the others from using the invention at issue. There is also a risk that, even if the validity of such patents is upheld, the court will construe the patent's claims narrowly or decide that we do not have the right to stop the others from using the invention at issue on the grounds that our patents do not cover the alleged infringing activity or product. An adverse outcome in a litigation or proceeding involving our patents could limit our ability to assert our patents against those parties and other competitors, and may curtail or preclude our ability to exclude third parties from making and selling similar or competitive products. Any of these occurrences could adversely affect our competitive business position, business prospects, and financial condition. Similarly, if we assert trademark infringement claims, a court may determine that our asserted marks are invalid or unenforceable, or that the party against whom we have asserted trademark infringement has superior rights to the marks in question. In this case, we could ultimately be forced to cease use of our trademarks.

In any litigation involving our intellectual property, the award of monetary damages we receive may not be commercially valuable or even sufficient to cover our cost of bringing such action. Furthermore, because of the substantial amount of discovery required in connection with intellectual property litigation, there is a risk that some of our confidential information could be compromised by disclosure during litigation. There could also be public announcements of the results of hearings, motions or other interim proceedings or developments, which could have a material adverse effect on the price of our ADSs. Moreover, there can be no assurance that we will have sufficient financial or other resources to file and pursue such infringement claims, which typically last for years before they are concluded. Even if we ultimately prevail in such claims, the monetary cost of such litigation and the diversion of the attention of our management and scientific personnel could outweigh any benefit we receive as a result of the proceedings.

Our commercial success depends significantly on our ability to operate without infringing upon, misappropriating or otherwise violating the intellectual property rights of third parties.

The life sciences industry is subject to rapid technological change and substantial litigation regarding patent and other intellectual property rights. Our potential competitors in both the PRC and abroad, may have substantially greater resources than us and are likely to make substantial investments in patent portfolios and competing technologies, and may apply for or obtain patents that could prevent, limit or otherwise interfere with our ability to make, use and sell our products. Numerous third-party patents exist in fields relating to our products and technologies, and it is difficult for industry participants, including us, to identify all third-party patent rights relevant to our products and technologies. Moreover, because some patent applications are maintained as confidential for a certain period of time, we cannot be certain that third parties have not filed patent applications that will cover our products and technologies if they issue as patents.

Patents could be issued to third parties that we may ultimately be found to infringe. Third parties may have or obtain valid and enforceable patents or proprietary rights that could block us from using our technology. Our failure to obtain or maintain a license from third parties to any technology that we require may materially harm our business, financial condition and results of operations. Furthermore, we would be exposed to a threat of litigation.

Third-party intellectual property right holders may also actively bring infringement or other intellectual property-related claims against us, even if we have received patent protection for our technologies, products, and services. Regardless of the merit of third parties claims against us for infringement, misappropriation or violations of their intellectual property rights, such third parties may seek and obtain injunctive or other equitable relief, which could effectively block our ability to perform clinical trials or develop, manufacture or sell our products. Further, if a patent infringement suit were brought against us, we could be forced to temporarily or permanently stop or delay our development or regulatory approval process or other activities that are the subject of such suit. Defense of these claims, even if such claims are resolved in our favor, could cause us to incur substantial expenses and be a substantial diversion of our employee resources even if we are ultimately successful. Any adverse ruling or perception of an adverse ruling in defending ourselves could have a material adverse impact on our cash position and stock price. Such litigation or proceedings could substantially increase our operating losses and reduce the resources available for development activities or any future sales, marketing, or distribution activities. We may not have sufficient financial or other resources to conduct such litigation or proceedings adequately. Some of our competitors may be able to sustain the costs of such litigation or proceedings more effectively than we can because of their greater financial resources and more mature and developed intellectual property portfolios.

Furthermore, because of the substantial amount of discovery required in connection with intellectual property litigation, there is a risk that some of our confidential information could be compromised by disclosure during this type of litigation. There could also be public announcements of the results of hearings, motions or other interim proceedings or developments, which could have a material adverse effect on the price of our ADSs. If securities analysts or investors perceive these results to be negative, it could have a material adverse effect on the price of our common stock. The occurrence of any of these events may have a material adverse effect on our business, results of operation, financial condition or cash flows.

Obtaining and maintaining patent protection depend on compliance with various procedural, document submission, fee payment and other requirements imposed by governmental patent authorities, and our patent protection could be reduced or eliminated for non-compliance with these requirements.

The NIPA, and various foreign governmental patent agencies including the USPTO, JPO, and EPO require compliance with a number of procedural, documentary, fee payment and other provisions

during the patent application and prosecution process. Periodic maintenance fees, renewal fees, annuity fees, and various other governmental fees on patents and/or applications will be due to be paid to the NIPA and various other governmental patent agencies outside of China in several stages over the lifetime of the patents and/or applications. We employ reputable professionals and rely on such third parties to help us comply with these requirements and effect payment of these fees with respect to the patents and patent applications that we own. Noncompliance events that could result in abandonment or lapse of a patent or patent application include failure to respond to official communications within prescribed time limits, non-payment of fees and failure to properly legalize and submit formal documents. In many cases, an inadvertent lapse can be cured by payment of a late fee or by other means in accordance with the applicable rules. However, there are situations in which noncompliance can result in abandonment or lapse of a patent or patent application, resulting in loss of patent rights in the relevant jurisdiction. In such an event, competitors might be able to enter the market earlier than would otherwise have been the case, which could have a material adverse effect on our competitive position, business, financial condition, results of operations, and prospects.

We may not enter into invention assignment and confidentiality agreements with all of our employees and third parties and such agreements may not prevent ownership disputes or unauthorized disclosure of trade secrets and other proprietary information.

We rely in part upon unpatented or unpatentable trade secrets, know-how and continuing technological innovation to develop and maintain our competitive position, which we seek to protect, in part, by entering into agreements, including confidentiality agreements and non-disclosure agreements, with parties that have a need for access to them, such as certain of our employees, consultants, academic institutions, corporate partners and, other third-party service providers. Nevertheless, there can be no guarantee that an employee or a third party will not make an unauthorized use or disclosure of our proprietary confidential information. This might happen intentionally or inadvertently. It is possible that a competitor will gain access to such information and make use of such information, and that our competitive position will be compromised, in spite of any legal action we might take against persons making such unauthorized disclosures. In addition, to the extent that our employees, consultants or contractors use intellectual property owned by others in their work for us, disputes may arise as to the rights in related or resulting know-how and inventions.

Trade secrets are difficult to protect. Although we use reasonable efforts to protect our trade secrets, our employees, consultants, contractors or business partners might intentionally or inadvertently disclose our trade secret information to competitors or our trade secrets may otherwise be misappropriated. Enforcing a claim that a third party illegally obtained and is using any of our trade secrets is expensive and time consuming, and the outcome is unpredictable.

We sometimes engage individuals or research institutions to conduct research relevant to our business. The ability of these individuals or research institutions to publish or otherwise publicly disclose data and other information generated during the course of their research is subject to certain contractual limitations. These contractual provisions may be insufficient or inadequate to protect our confidential information. If we do not apply for patent protection prior to such publication, or if we cannot otherwise maintain the confidentiality of our proprietary technology and other confidential information, then our ability to obtain patent protection or to protect our trade secret information may be jeopardized, which could adversely affect our business, financial condition and results of operations.

We also seek to enter agreements with our employees and consultants that obligate them to assign any inventions created during their work for us to us. However, we may not obtain these agreements in all circumstances and the assignment of intellectual property under such agreements may not be self-executing. And it is possible that technology relevant to our business will be independently developed by a person that is not a party to such an agreement. Furthermore, if the employees and consultants who are parties to these agreements breach or violate the terms of these agreements, we

may not have adequate remedies for any such breach or violation, and we could lose our trade secrets and inventions through such breaches or violations. Any of the foregoing could have a material and adverse effect on our business, financial condition and results of operations.

We may be subject to claims that our employees have wrongfully used or disclosed alleged trade secrets of their former employers.

Some of our employees and consultants were previously employed at universities or other biotechnology or pharmaceutical companies, including our competitors or potential competitors. Although we try to ensure that our employees do not use the proprietary information or know-how of others in their work for us, we may be subject to claims that we or our employees have used or disclosed intellectual property, including trade secrets or other proprietary information, of any such employee's former employer. Litigation may be necessary to defend against these claims. If we fail in defending any such claims, in addition to paying monetary damages, we may lose valuable intellectual property rights or personnel. Even if we are successful in defending against such claims, litigation could result in substantial costs and be a distraction to management and our specific personnel.

We may be subject to claims challenging the inventorship of our patents and other intellectual property.

We may be subject to claims that former employees, collaborators or other third parties have an interest in our patents, trade secrets, or other intellectual property as an inventor or co-inventor. For example, we may have inventorship disputes arise from conflicting obligations of employees, consultants or others who are involved in developing our product candidates. Also, former employees may become employed by competitors who develop similar technology, and could assist the competitor in designing around our patents. While it is our policy to require our employees and contractors who may be involved in the development of intellectual property to execute agreements assigning such intellectual property to us, we may be unsuccessful in executing such an agreement with each party who in fact develops intellectual property that we regard as our own. Our and their assignment agreements may not be self-executing or may be breached, and litigation may be necessary to defend against these and other claims challenging inventorship or our ownership of our patents, trade secrets or other intellectual property. If we fail in defending any such claims, in addition to paying monetary damages, we may lose valuable intellectual property rights, such as exclusive ownership of, or right to use, intellectual property that is important to our product candidates. Even if we are successful in defending against such claims, litigation could result in substantial costs and be a distraction to management and other employees. Any of the foregoing could have a material adverse effect on our business, financial condition, results of operations and prospects.

Intellectual property rights do not necessarily protect us from all potential threats to our competitive advantages.

The degree of future protection afforded by our intellectual property rights is uncertain because intellectual property rights have limitations, and may not adequately protect our business, or permit us to maintain our competitive advantage. The following examples are illustrative:

- others may be able to independently develop similar or alternative technologies or designs that are similar to our product candidates but that are not covered by the claims of the patents that we own or have exclusively licensed;
- we might not have been the first to make the inventions covered by the issued patents or pending patent applications that we own or may in the future exclusively license, which could result in the patent applications not issuing or being invalidated after issuing;
- we might not have been the first to file patent applications covering certain of our inventions, which could result in the patent applications not issuing or being invalidated after issuing;

- others may independently develop similar or alternative technologies or duplicate any of our technologies without infringing our intellectual property rights;
- it is possible that our pending patent applications will not lead to issued patents;
- issued patents that we own or have exclusively licensed may not provide us with any competitive advantages, or may be held invalid or unenforceable, as a result of legal challenges by our competitors;
- our competitors might conduct research and development activities in countries where we do not have patent rights and then use the information learned from such activities to develop competitive services and products for commercialization in our major markets;
- we may fail to develop additional proprietary technologies that are patentable;
- we may fail to apply for or obtain adequate intellectual property protection in all the jurisdictions in which we operate; and
- the patents of others may have an adverse effect on our business, for example by preventing us from commercializing one or more of our product candidates for one or more indications.

Any of the aforementioned threats to our competitive advantage could have a material adverse effect on our business.

Patent terms may not be sufficient to effectively protect our product candidates.

In most countries in which we plan to file applications for patents, the term of an issued patent is generally 20 years from the earliest claimed filing date of the priority application to which a non-provisional patent application in the applicable country claims priority. Although various extensions may be available in various countries, the life of a patent and the protection it affords are limited. Even if patents covering our product candidates are obtained, we may be open to competition from other companies once our patent rights expire. Accordingly, given the amount of time required for the development, testing and regulatory review of new product candidates, patents protecting such candidates might expire before or shortly after such candidates are commercialized. Excluding any patent term adjustment and patent term extension, our currently issued patents are expected to expire from 2033 to 2034. As a result, our patent portfolio may not provide us with sufficient rights over a sufficient length of time to exclude others from commercializing products similar or identical to ours.

Uncertainty of the length of patent term extensions and data and market exclusivities for our pharmaceutical products could increase the risk of generic competition.

In the United States, the Federal Food, Drug, and Cosmetic Act, as amended by the law generally referred to as the "Hatch-Waxman Amendments," provides the opportunity for patent-term restoration of up to five years to reflect patent term lost during certain portions of product development and the FDA regulatory review process. The Hatch-Waxman Amendments also have a process for patent linkage, pursuant to which FDA will stay approval of certain follow-on applications during the pendency of litigation between the follow-on applicant and the patent holder or licensee, generally for a period of 30 months. Finally, the Hatch-Waxman Amendments provide for statutory exclusivities that can prevent submission or approval of certain follow-on marketing applications. For example, federal law provides a five-year period of exclusivity within the United States to the first applicant to obtain approval of a new chemical entity (as defined) and three years of exclusivity protecting certain innovations to previously approved active ingredients where the applicant was required to conduct new clinical investigations to obtain approval for the modification. Similarly, the Orphan Drug Act provides seven years of market exclusivity for certain drugs to treat rare diseases, where FDA designates the product candidate as an orphan drug and the drug is approved for the designated orphan indication.

These provisions, designed to promote innovation, can prevent competing products from entering the market for a certain period of time after FDA grants marketing approval for the innovative product.

In China, however, there is no currently effective law or regulation providing patent term extension, patent linkage, or data exclusivity (referred to as regulatory data protection). Therefore, a lower-cost generic drug can emerge onto the market much more quickly. Chinese regulators have set forth a framework for integrating patent linkage and data exclusivity into the Chinese regulatory regime, as well as for establishing a pilot program for patent term extension. To be implemented, this framework will require adoption of regulations. To date, the NMPA has issued several draft implementing regulations in this regard for public comment but no regulations have been formally issued. These factors result in weaker protection for us against generic competition in China than could be available to us in the United States until the relevant implementing regulations for extension, patent linkage, or data exclusivity are put into effect officially in China.

We may not be able to protect our intellectual property rights throughout the world.

Filing, prosecuting and defending patents on our products in all countries throughout the world would be prohibitively expensive. We may also encounter difficulties in protecting and defending such rights in foreign jurisdictions. Consequently, we may not be able to prevent third parties from practicing our inventions in all countries outside the jurisdictions of the registration of our intellectual properties. Competitors may use our technologies in jurisdictions where we have not obtained patent protection to develop their own products. Our patents or other intellectual property rights may not be effective or sufficient to prevent them from competing.

Many companies have encountered significant problems in protecting and defending intellectual property rights in certain jurisdictions. The legal systems of many other countries do not favor the enforcement of patents and other intellectual property protection, which could make it difficult for us to stop the infringement of our patents in such countries.

Proceedings to enforce our patent rights in foreign jurisdictions could result in substantial cost and divert our efforts and attention from other aspects of our business, could put our patents at risk of being invalidated or interpreted narrowly and our patent applications at risk of not issuing, and could provoke third parties to assert claims against us. We may not prevail in any lawsuits that we initiate and the damages or other remedies awarded, if any, may not be commercially meaningful. Accordingly, our efforts to enforce our intellectual property rights around the world may be inadequate to obtain a significant commercial advantage from the intellectual property that we develop or license.

We may not be able to protect and enforce our trademarks.

We currently hold issued trademark registrations and have trademark applications pending, any of which may be the subject of a governmental or third-party objection, which could prevent the registration or maintenance of the same. If we are unsuccessful in obtaining trademark protection for our primary brands, we may be required to change our brand names, which could materially adversely affect our business. Moreover, as our products mature, our reliance on our trademarks to differentiate us from our competitors will increase, and as a result, if we are unable to prevent third parties from adopting, registering or using trademarks and trade dress that infringe, dilute or otherwise violate our trademark rights, or engaging in conduct that constitutes unfair competition, defamation or other violation of our rights, our business could be materially adversely affected. Over the long term, if we are unable to establish name recognition based on our trademarks and trade names, then we may not be able to compete effectively and our business may be adversely affected. We may license our trademarks and trade names to third parties, such as distributors. Though these license agreements may provide guidelines for how our trademarks and trade names may be used, a breach of these agreements or misuse of our trademarks and tradenames by our licensees may jeopardize our rights in or diminish

the goodwill associated with our trademarks and trade names. Our efforts to enforce or protect our proprietary rights related to trademarks, trade names, trade secrets, domain names, copyrights or other intellectual property may be ineffective and could result in substantial costs and diversion of resources and could adversely affect our financial condition or results of operations.

Risks Related to Doing Business in the PRC

Uncertainties with respect to the PRC legal system and changes in laws and regulations in China could adversely affect us.

Our operations in China are governed by the PRC laws and regulations. The PRC legal system is a civil law system based on written statutes. Unlike the common law system, prior court decisions under the civil law system may be cited for reference but have limited precedential value. In addition, any new PRC laws or changes in PRC laws and regulations related to, among other things, foreign investment and manufacturing in China could have a material adverse effect on our business and our ability to operate our business in China.

From time to time, we may have to resort to administrative and court proceedings to enforce our legal rights. Any administrative and court proceedings in China may be protracted, resulting in substantial costs and diversion of resources and management attention. Since PRC administrative and court authorities have significant discretion in interpreting and implementing statutory provisions and contractual terms, it may be more difficult to evaluate the outcome of administrative and court proceedings and the level of legal protection we enjoy, than in more developed legal systems. These uncertainties may impede our ability to enforce contracts in China and could materially and adversely affect our business and results of operations.

Furthermore, the PRC legal system is based in part on government policies and internal rules, some of which are not published on a timely basis, or at all, and may have retroactive effect. As a result, we may not be aware of our violation of any of these policies and rules until sometime after the violation. Such unpredictability towards our contractual, property and procedural rights could adversely affect our business, and impede our ability to continue our operations and proceed with our future business plans.

We may be restricted from transferring our scientific data abroad.

On March 17, 2018, the General Office of the PRC State Council promulgated the Measures for the Management of Scientific Data, or the Scientific Data Measures, which provide a broad definition of scientific data and relevant rules for the management of scientific data. According to the Scientific Data Measures, enterprises in China must seek governmental approval before any scientific data involving a state secret may be transferred abroad or to foreign parties. Further, any researcher conducting research funded, at least in part, by the PRC government is required to submit relevant scientific data for management by the entity to which such researcher is affiliated before such data may be published in any foreign academic journal. Currently, as the term "state secret" is not clearly defined, there is no assurance that we can always obtain relevant approvals for sending scientific data (such as the results of our preclinical studies or clinical trials conducted within China) abroad, or to our foreign partners in China.

If we are unable to obtain the necessary approvals in a timely manner, or at all, our research and development of product candidates may be hindered, which may materially and adversely affect our business, results of operations, financial conditions and prospects. If relevant government authorities consider the transmission of our scientific data to be in violation of the requirements under the Scientific Data Measures, we may be subject to specific administrative penalties imposed by those government authorities

Recent litigation and negative publicity surrounding China-based companies listed in the United States may result in increased regulatory scrutiny of us and negatively impact the trading price of the ADSs and could have a material adverse effect upon our business, including our results of operations, financial condition, cash flows and prospects.

We believe that litigation and negative publicity surrounding companies with operations in China that are listed in the United States have negatively impacted stock prices for these companies. Various equity-based research organizations have published reports on China-based companies after examining their corporate governance practices, related party transactions, sales practices and financial statements, and these reports have led to special investigations and listing suspensions on U.S. national exchanges. Any similar scrutiny of us, regardless of its lack of merit, could result in a diversion of management resources and energy, potential costs to defend ourselves against rumors, decreases and volatility in the ADS trading price, and increased directors and officers insurance premiums and could have an adverse effect upon our business, including our results of operations, financial condition, cash flows and prospects.

Changes in U.S. and international trade policies, particularly with regard to China, may adversely impact our business and operating results.

The U.S. government has recently made statements and taken certain actions that may lead to potential changes to U.S. and international trade policies, including imposing several rounds of tariffs affecting certain products manufactured in China. In March 2018, U.S. President Donald J. Trump announced the imposition of tariffs on steel and aluminum entering the United States and in June 2018 announced further tariffs targeting goods imported from China. Recently both China and the United States have each imposed tariffs indicating the potential for further trade barriers. It is unknown whether and to what extent new tariffs (or other new laws or regulations) will be adopted, or the effect that any such actions would have on us or our industry. While we have not started commercialization of product candidates, any unfavorable government policies on international trade, such as capital controls or tariffs, may affect the demand for our drug products, the competitive position of our drug products, the hiring of scientists and other research and development personnel, and import or export of raw materials in relation to drug development, or prevent us from selling our drug products in certain countries. If any new tariffs, legislation and/or regulations are implemented, or if existing trade agreements are renegotiated or, in particular, if the U.S. government takes retaliatory trade actions due to the recent U.S.-China trade tension, such changes could have an adverse effect on our business, financial condition and results of operations.

You may be subject to PRC income tax on dividends from us or on any gain realized on the transfer of our ADSs.

Under the Enterprise Income Tax Law of the PRC, or the EIT Law, and its implementation rules, PRC withholding tax at the rate of 10% is generally applicable to dividends from PRC sources paid to investors that are resident enterprises outside of China and that do not have an establishment or place of business in China, or that have an establishment or place of business in China but the relevant income is not effectively connected with the establishment or place of business. Any gain realized on the transfer of shares by such investors is subject to 10% PRC income tax if this gain is regarded as income derived from sources within China. Under the PRC Individual Income Tax Law and its implementation rules, dividends from sources within China paid to foreign individual investors who are not PRC residents are generally subject to a PRC withholding tax at a rate of 20% and gains from PRC sources realized by these investors on the transfer of shares are generally subject to 20% PRC income tax. Any such PRC tax liability may be reduced by the provisions of an applicable tax treaty.

Although substantially all of our business operations are in China, it is unclear whether the dividends we pay with respect to our shares or ADSs, or the gains realized from the transfer of our

shares or ADSs, would be treated as income derived from sources within China and as a result be subject to PRC income tax if we are considered a PRC resident enterprise. If PRC income tax is imposed on gains realized through the transfer of our ADSs or on dividends paid to our non-resident investors, the value of your investment in our ADSs may be adversely affected. Furthermore, our shareholders whose jurisdictions of residence have tax treaties or arrangements with China may not qualify for benefits under these tax treaties or arrangements.

In addition, pursuant to the Double Tax Avoidance Arrangement between Hong Kong and China, or the Double Tax Avoidance Treaty, and the Notice on Certain Issues with Respect to the Enforcement of Dividend Provisions in Tax Treaties, or the Notice on Tax Treaties, issued on February 20, 2009 by the State Administration of Taxation, or the SAT, if a Hong Kong resident enterprise owns more than 25% of the equity interest of a PRC company at all times during the twelve-month period immediately prior to obtaining a dividend from such company, the 10% withholding tax on such dividend is reduced to 5%, provided that certain other conditions and requirements under the Double Tax Avoidance Treaty and other applicable PRC laws are satisfied at the discretion of the relevant PRC tax authority. However, based on the Notice on Tax Treaties, if the relevant PRC tax authorities determine, in their discretion, that a company benefits from such reduced income tax rate due to a structure or arrangement that is primarily tax-driven, the PRC tax authorities may adjust the preferential tax treatment. Based on the Notice on Issues concerning Beneficial Owner in Tax Treaties, or Circular 9, issued on February 3, 2018 by the SAT and effective on April 1, 2018, when determining the applicant's status as a "beneficial owner" for purpose of tax treatments in connection with dividends, interests or royalties in the tax treaties, several factors will be taken into account, and it will be analyzed according to the actual circumstances of the specific cases. If our Hong Kong subsidiary is determined by PRC government authorities as receiving benefits from reduced income tax rates due to a structure or arrangement that is primarily tax-driven, the dividends paid by our PRC subsidiary to our Hong Kong subsidiary will be taxed at a higher rate, which will have an adverse effect on our financial and operational conditions.

The biopharmaceutical industry in China is highly regulated and such regulations are subject to changes which may affect approval and commercialization of our product candidates.

Part of our research and development operations are in China, which we believe confers clinical, commercial and regulatory advantages. The biopharmaceutical industry in China is subject to comprehensive government regulation and supervision, encompassing the approval, registration, manufacturing, packaging, licensing and marketing of new product candidates. See "Regulation" for a discussion of the regulatory requirements that are applicable to our current and planned business activities in China. In recent years, the regulatory framework in China regarding the biopharmaceutical industry has undergone significant changes, and we expect that it will continue to undergo significant changes. Any such changes or amendments may result in increased compliance costs on our business or cause delays in or prevent the successful development or commercialization of our product candidates in China and reduce the current benefits we believe are available to us from developing and manufacturing drugs in China. PRC authorities have become increasingly vigilant in enforcing laws in the biopharmaceutical industry and any failure by us or our partners to maintain compliance with applicable laws and regulations or obtain and maintain required licenses and permits may result in the suspension or termination of our business activities fines, warnings, administrative or criminal penalties in China. We believe our strategy and approach are aligned with the PRC government's regulatory policies, but we cannot ensure that our strategy and approach will continue to be aligned.

Substantial uncertainties exist with respect to the interpretation and implementation of the newly enacted Foreign Investment Law and how it may impact the viability of our current corporate structure, corporate governance and business operations.

On March 15, 2019, the PRC National People's Congress approved the Foreign Investment Law, which came into effect on January 1, 2020 and replaces the trio of existing laws regulating foreign investment in the PRC, namely, the Sino-Foreign Equity Joint Venture Enterprise Law, the Sino-Foreign Cooperative Joint Venture Enterprise Law and the Wholly Foreign-Invested Enterprise Law, together with their implementation rules and ancillary regulations and become the legal foundation for foreign investment in the PRC. Meanwhile, the *Implementation Regulation of the Foreign Investment Law and the Measures for Reporting of Information on Foreign Investment* came into effect as of January 1, 2020, which clarified and elaborated the relevant provisions of the *Foreign Investment Law*.

The Foreign Investment Law sets out the basic regulatory framework for foreign investments and proposes to implement a system of pre-entry national treatment with a negative list for foreign investments, pursuant to which (i) foreign entities and individuals are prohibited from investing in the areas that are not open to foreign investments, (ii) foreign investments in the restricted industries must satisfy certain requirements under the law, and (iii) foreign investments in business sectors outside of the negative list will be treated equally with domestic investments. The Foreign Investment Law also sets forth necessary mechanisms to facilitate, protect and manage foreign investments and proposes to establish a foreign investment information reporting system, through which foreign investors or foreign-invested enterprises are required to submit initial report, report of changes, report of deregistration and annual report relating to their investments to the Ministry of Commerce, or MOFCOM, or its local branches.

You may experience difficulties in effecting service of legal process, enforcing foreign judgments or bringing actions in China against us or our management named in the prospectus based on foreign laws.

We are a company incorporated under the laws of the Cayman Islands, we conduct most of our operations in China, and substantially all of our assets are located in China. As a result, it may be difficult for our shareholders to effect service of process upon us or those persons inside China. In addition, China does not have treaties providing for the reciprocal recognition and enforcement of judgments of courts with the Cayman Islands and many other countries and regions. Therefore, recognition and enforcement in China of judgments of a court in any of these non-PRC jurisdictions in relation to any matter not subject to a binding arbitration provision may be difficult or impossible.

Shareholder claims that are common in the United States, including securities law class actions and fraud claims, generally are difficult to pursue as a matter of law or practicality in China. For example, in China, there are significant legal and other obstacles to obtaining information needed for shareholder investigations or litigation outside China or otherwise with respect to foreign entities. Although the local authorities in China may establish a regulatory cooperation mechanism with the securities regulatory authorities of another country or region to implement cross-border supervision and administration, such regulatory cooperation with the securities regulatory authorities in the United States have not been efficient in the absence of mutual and practical cooperation mechanism. According to Article 177 of the PRC Securities Law which became effective in March 2020, no overseas securities regulator is allowed to directly conduct investigation or evidence collection activities within the territory of the PRC. Accordingly, without the consent of the competent PRC securities regulators and relevant authorities, no organization or individual may provide the documents and materials relating to securities business activities to overseas parties. See also "—Risks Related to the ADSs and this Offering—You may face difficulties in protecting your interests, and your ability to protect your rights through U.S. courts may be limited, because we are incorporated under Cayman Islands law" for risks associated with investing in us as a Cayman Islands company.

Our business may be negatively affected by the potential obligations to make additional social insurance and housing fund contributions.

We are required by PRC labor laws and regulations, such as the Social Insurance Law, Administrative Regulations on the Housing Provident Fund and other related rules, to pay various statutory employee benefits, including pensions insurance, medical insurance, work-related injury insurance, unemployment insurance, maternity insurance and housing fund, to designated government agencies for the benefit of our employees. The relevant government agencies may examine whether an employer has made adequate and timely payments of the requisite statutory employee benefits, and employers who fail to make adequate and timely payments may be subject to supplemental contributions, late payment fees, fines compulsory enforcement and/or other penalties. If the relevant PRC authorities determine that we shall make supplemental social insurance and housing fund contributions or that we are subject to fines and legal sanctions in relation to our failure to make social insurance and housing fund contributions in full for our employees, our business, financial condition and results of operations may be adversely affected.

The lease agreements of our leased properties have not been registered with the relevant PRC government authorities as required by PRC law, which may expose us to potential fines.

Under PRC law, lease agreements of commodity housing tenancy are required to be registered with the local construction (real estate) departments. Although failure to do so does not in itself invalidate the leases, the parties of the lease agreements may be exposed to potential fines if they fail to rectify such non-compliance within the prescribed time frame after receiving notice from the relevant PRC government authorities. The penalty ranges from RMB1,000 to RMB10,000 for each unregistered lease, at the discretion of the relevant authority. As of the date of this Prospectus, the lease agreements for our leased properties in China have not been registered with the relevant PRC government authorities. As of the date of this prospectus, we are not aware of any regulatory or governmental actions, claims or investigations being contemplated or any challenges by third parties to our use of our leased properties that the lease agreements of which have not been registered with the government authorities. However, we cannot assure you that the government authorities will not impose fines on us due to our failure to register any of our lease agreements, which may negatively impact our financial condition.

Any failure to comply with PRC regulations regarding the registration requirements for employee stock incentive plans may subject the PRC plan participants or us to fines and other legal or administrative sanctions.

In February 2012, SAFE promulgated the Notices on Issues Concerning the Foreign Exchange Administration for Domestic Individuals Participating in Stock Incentive Plan of Overseas Publicly Listed Company, replacing earlier rules promulgated in 2007. Pursuant to these rules, PRC citizens and non-PRC citizens who reside in China for a continuous period of not less than one year who participate in any stock incentive plan of an overseas publicly listed company, subject to a few exceptions, are required to register with SAFE through a domestic qualified agent, which could be the PRC subsidiary of such overseas-listed company, and complete certain other procedures. In addition, an overseas-entrusted institution must be retained to handle matters in connection with the exercise or sale of stock options and the purchase or sale of shares and interests. We and our executive officers and other employees who are PRC citizens or who reside in the PRC for a continuous period of not less than one year and who have been granted options will be subject to these regulations when our company becomes an overseas-listed company upon completion of this offering. Failure to complete the SAFE registrations may subject them to fines and legal sanctions, there may be additional restrictions on the ability of them to exercise their stock options or remit proceeds gained from the sale of their stock into

the PRC. We also face regulatory uncertainties that could restrict our ability to adopt incentive plans for our directors, executive officers and employees under PRC law.

If we are classified as a PRC resident enterprise for PRC income tax purposes, such classification could result in unfavorable tax consequences to us and our non-PRC shareholders or ADS holders.

Under the PRC Enterprise Income Tax Law and its implementation rules, an enterprise established outside of the PRC with a "de facto management body" within the PRC is considered a "resident enterprise" and will be subject to the enterprise income tax on its global income at the rate of 25%. The implementation rules define the term "de facto management body" as the body that exercises full and substantial control over and overall management of the business, productions, personnel, accounts and properties of an enterprise. In 2009, SAT issued a circular, known as SAT Circular 82, which provides certain specific criteria for determining whether the "de facto management body" of a PRC-controlled enterprise that is incorporated offshore is located in China. Although this circular only applies to offshore enterprises controlled by PRC enterprises or PRC enterprise groups, not those controlled by PRC individuals or foreigners, the criteria set forth in the circular may reflect the SAT's general position on how the "de facto management body" test should be applied in determining the tax resident status of all offshore enterprises. According to SAT Circular 82, an offshore incorporated enterprise controlled by a PRC enterprise or a PRC enterprise group will be regarded as a PRC tax resident by virtue of having its "de facto management body" in China and will be subject to PRC enterprise income tax on its global income only if all of the following conditions are met: (i) the primary location of the day-to-day operational management and the places where they perform their duties are in the PRC; (ii) decisions relating to the enterprise's financial and human resource matters are made or are subject to approval by organizations or personnel in the PRC; (iii) the enterprise's primary assets, accounting books and records, company seals, and board and shareholder resolutions, are located or maintained in the PRC; and (iv) at least 50% of voting board members or senior executives habitually reside in the PRC.

We believe that we are not a PRC resident enterprise for PRC tax purposes. See "Taxation—People's Republic of China Taxation." However, the tax resident status of an enterprise is subject to determination by the PRC tax authorities and uncertainties remain with respect to the interpretation of the term "de facto management body." If the PRC tax authorities determine that we or any of our non-PRC subsidiaries are a PRC resident enterprise for enterprise income tax purposes, we may be required to withhold a 10% withholding tax from dividends we pay to our shareholders that are non-resident enterprises, including the holders of the ADSs. In addition, non-resident enterprise shareholders (including ADS holders) may be subject to PRC tax on gains realized on the sale or other disposition of ADSs or ordinary shares, if such income is treated as sourced from within the PRC. Furthermore, if we are deemed a PRC resident enterprise, dividends payable to our non-PRC individual shareholders (including ADS holders) and any gain realized on the transfer of ADSs or ordinary shares by such shareholders may be subject to PRC tax at a rate of 20% (which, in the case of dividends, may be withheld at source by us). Any PRC tax liability may be reduced under applicable tax treaties. However, it is unclear whether in practice our non-PRC shareholders would be able to obtain the benefits of any tax treaties between their countries of tax residence and the PRC in the event that we are treated as a PRC resident enterprise. Any such tax may reduce the returns on your investment in the ADSs or our ordinary shares.

We face uncertainty with respect to indirect transfers of equity interests in PRC resident enterprises by their non-PRC holding companies.

Pursuant to the Notice on Strengthening Administration of Enterprise Income Tax for Share Transfers by Non-PRC Resident Enterprises, or SAT Circular 698, issued by the SAT in 2009 with retroactive effect from January 1, 2008, where a non-resident enterprise transfers the equity interests of

a PRC resident enterprise indirectly by disposition of the equity interests of an overseas holding company, or an Indirect Transfer, and such overseas holding company is located in a tax jurisdiction that: (i) has an effective tax rate less than 12.5% or (ii) does not tax foreign income of its residents, the non-resident enterprise, being the transferor, shall report to the competent tax authority of the PRC resident enterprise this Indirect Transfer.

On February 3, 2015, the SAT issued the Announcement of the State Administration of Taxation on Several Issues Concerning the Enterprise Income Tax on Indirect Property Transfer by Non-Resident Enterprises, or SAT Bulletin 7. SAT Bulletin 7 supersedes the rules with respect to the Indirect Transfer under SAT Circular 698. SAT Bulletin 7 has introduced a new tax regime that is significantly different from the previous one under SAT Circular 698. SAT Bulletin 7 extends the PRC's tax jurisdiction to not only Indirect Transfers set forth under SAT Circular 698 but also transactions involving a transfer of other taxable assets through an offshore transfer of a foreign intermediate holding company. In addition, SAT Bulletin 7 provides clearer criteria than SAT Circular 698 for assessment of reasonable commercial purposes and has introduced safe harbors for internal group restructurings and the purchase and sale of equity through a public securities market. SAT Bulletin 7 also brings challenges to both foreign transferor and transferee (or another person who is obligated to pay for the transfer) of taxable assets. Where a non-resident enterprise transfers taxable assets indirectly by disposing of the equity interests of an overseas holding company, which is an Indirect Transfer, the non-resident enterprise, being the transferor, or the transferee, or the PRC entity that directly owns the taxable assets, may report such Indirect Transfer to the relevant tax authority. Using a "substance over form" principle, the PRC tax authority may disregard the existence of the overseas holding company if it lacks a reasonable commercial purpose and was established for the purpose of reducing, avoiding or deferring PRC tax. As a result, gains derived from such Indirect Transfer may be subject to PRC enterprise income tax, and the transferee or another person who is obligated to pay for the transfer is obligated to withhold the applicable taxes, currently at a rate of 10% for the transfer of equity interests in a PRC resident enterprise. Both the transferor and the transferee may be subject to penalties under PRC tax laws if the transferee fails to withhold the taxes and the transferor fails to pay the taxes.

On October 17, 2017, the SAT issued the Announcement of the State Administration of Taxation on Matters Concerning Withholding of Income Tax of Non-resident Enterprises at Source, or SAT Bulletin 37, which, among others, repealed the SAT Circular 698 on December 1, 2017. SAT Bulletin 37 further details and clarifies the tax withholding methods in respect of income of non-resident enterprises under SAT Circular 698. And certain rules stipulated in SAT Bulletin 7 are replaced by SAT Bulletin 37. Where the non-resident enterprise fails to declare the tax payable pursuant to Article 39 of the PRC Enterprise Income Tax Law, the tax authority may order it to pay the tax due within required time limits, and the non-resident enterprise shall declare and pay the tax payable within such time limits specified by the tax authority; however, if the non-resident enterprise voluntarily declares and pays the tax payable before the tax authority orders it to do so within required time limits, it shall be deemed that such enterprise has paid the tax in time.

We face uncertainties as to the reporting and other implications of certain past and future transactions where PRC taxable assets are involved, such as offshore restructuring, sale of the shares in our offshore subsidiaries and investments. Our company may be subject to filing obligations or taxed if our company is a transferor in such transactions, and may be subject to withholding obligations if our company is a transferee in such transactions, under SAT Bulletin 7 and SAT Bulletin 37. For transfer of shares in our company by investors who are non-PRC resident enterprises, our PRC subsidiary may be requested to assist in the filing under SAT Bulletin 7 and SAT Bulletin 37. As a result, we may be required to expend valuable resources to comply with SAT Bulletin 7 and SAT Bulletin 37 or to request the relevant transferors from whom we purchase taxable assets to comply with these circulars, or to

establish that our company should not be taxed under these circulars, which may have a material adverse effect on our financial condition and results of operations.

If our preferential tax treatments are revoked, become unavailable or if the calculation of our tax liability is challenged by the PRC tax authorities, we may be required to pay tax, interest and penalties in excess of our tax provisions, and our results of operations could be materially and adversely affected.

The Chinese government has provided various tax incentives to our subsidiaries in China. These incentives include reduced enterprise income tax rates. For example, under the Enterprise Income Tax Law and its implementation rules, the statutory enterprise income tax rate is 25%. However, the income tax of an enterprise that has been determined to be a technologically advanced service enterprise can be reduced to a preferential rate of 15%. Any increase in the enterprise income tax rate applicable to our PRC subsidiary, or any discontinuation or retroactive or future reduction of any of the preferential tax treatments currently enjoyed by our PRC subsidiary, could adversely affect our business, financial condition and results of operations. In addition, in the ordinary course of our business, we are subject to complex income tax and other tax regulations and significant judgment is required in the determination of a provision for income taxes. Although we believe our tax provisions are reasonable, if the PRC tax authorities successfully challenge our position and we are required to pay tax, interest and penalties in excess of our tax provisions, our financial condition and results of operations would be materially and adversely affected.

Certain PRC regulations may make it more difficult for us to pursue growth through acquisitions.

Among other things, the Regulations on Mergers and Acquisitions of Domestic Enterprises by Foreign Investors, or the M&A Rules, adopted by six PRC regulatory agencies in 2006 and amended in 2009, established additional procedures and requirements that could make merger and acquisition activities by foreign investors more time-consuming and complex. Such regulation requires, among other things, the MOFCOM be notified in advance or its approval be obtained in certain situations, such as any change-of-control transaction in which a foreign investor acquires control of a PRC domestic enterprise of Undertakings, issued by the State Council in 2008 and amended in 2018, were triggered. Moreover, the Anti-Monopoly Law promulgated by the Standing Committee of the PRC National People's Congress, or NPC, which became effective in 2008 requires that transactions which are deemed concentrations and involve parties with specified turnover thresholds must be cleared by the MOFCOM before they can be completed. In addition, PRC national security review rules which became effective in September 2011 require acquisitions by foreign investors of PRC companies engaged in military-related or certain other industries that are crucial to national security be subject to security review before consummation of any such acquisition. We may pursue potential strategic acquisitions that are complementary to our business and operations. Complying with the requirements of these regulations to complete such transactions could be time-consuming, and any required approval processes, including obtaining approval or clearance from the MOFCOM, may delay or inhibit our ability to complete such transactions, which could affect our ability to expand our business or maintain our market share.

The approval of the China Securities Regulatory Commission may be required in connection with this offering, and, if required, we cannot predict whether we will be able to obtain such approval.

The M&A Rules requires an overseas special purpose vehicle formed for listing purposes through acquisitions of PRC domestic companies and controlled by PRC companies or individuals to obtain the approval of the CSRC prior to the listing and trading of such special purpose vehicle's securities on an overseas stock exchange. However, the application of the M&A Rules remains unclear. If CSRC approval is required, it is uncertain whether it would be possible for us to obtain the approval, and any

failure to obtain or delay in obtaining CSRC approval for this offering would subject us to sanctions imposed by the CSRC and other PRC regulatory agencies.

Our PRC legal counsel has advised us based on their understanding of the current PRC laws, rules and regulations that the CSRC's approval may not be required for the listing and trading of the ADSs on the Nasdaq Global Market in the context of this offering, given that: (i) the CSRC currently has not issued any definitive rule or interpretation concerning whether offering such as this offering contemplated by our Company are subject to the M&A Rules; and (ii) our PRC subsidiary was incorporated as wholly foreign-owned enterprises by means of direct investment rather than by merger or acquisition of equity interest or assets of a PRC domestic company owned by PRC companies or individuals as defined under the M&A Rules that are our beneficial owners. However, there is uncertainty as to how the M&A Rules will be interpreted or implemented and we cannot assure you that relevant PRC governmental authorities, including CSRC, would reach the same conclusion as our PRC Legal Counsel.

PRC regulations relating to offshore investment activities by PRC residents may limit our PRC subsidiaries' ability to change their registered capital or distribute profits to us or otherwise expose us or our PRC resident beneficial owners to liability and penalties under PRC laws.

In July 2014, SAFE promulgated the Circular on Relevant Issues Concerning Foreign Exchange Control on Domestic Residents' Offshore Investment and Financing and Roundtrip Investment Through Special Purpose Vehicles, or SAFE Circular 37. SAFE Circular 37 requires PRC residents (including PRC individuals and PRC corporate entities as well as foreign individuals that are deemed as PRC residents for foreign exchange administration purpose) to register with SAFE or its local branches in connection with their direct or indirect offshore investment activities. SAFE Circular 37 is applicable to our shareholders who are PRC residents and may be applicable to any offshore acquisitions that we make in the future.

Under SAFE Circular 37, PRC residents who make, or have prior to the implementation of SAFE Circular 37 made, direct or indirect investments in offshore special purpose vehicles, or SPVs, will be required to register such investments with SAFE or its local branches. In addition, any PRC resident who is a direct or indirect shareholder of an SPV, is required to update its filed registration with the local branch of SAFE with respect to that SPV, to reflect any material change, including, among other things, any major change of a PRC resident shareholder, name or term of operation of the SPVs, or any increase or reduction of the SPVs' registered capital, share transfer or swap, merger or division. Moreover, any subsidiary of such SPV in China is required to urge the PRC resident shareholders to update their registration with the local branch of SAFE. If any PRC shareholder of such SPV fails to make the required registration or to update the previously filed registration, the subsidiary of such SPV in China may be prohibited from distributing its profits or the proceeds from any capital reduction, share transfer or liquidation to the SPV, and the SPV may also be prohibited from making additional capital contributions into its subsidiary in China. On February 13, 2015, SAFE promulgated a Notice on Further Simplifying and Improving Foreign Exchange Administration Policy on Direct Investment, or SAFE Notice 13, which became effective on June 1, 2015. Under SAFE Notice 13, applications for foreign exchange registration of inbound foreign direct investments and outbound overseas direct investments, including those required under SAFE Circular 37, will be filed with qualified banks instead of SAFE or its branches. The qualified banks will directly examine the applications and accept registrations under the supervision of SAFE.

Some of our existing shareholders, each of whom owns our ordinary shares, including but not limited to as a result of exercising share options, are PRC residents under SAFE Circular 37. However, we cannot provide any assurance that these PRC residents comply with our request to make or obtain any applicable registrations or change registration or comply with all of the requirements under SAFE Circular 37 or other related rules. Furthermore, we may not be informed of the identities of all the

PRC residents holding direct or indirect interest in our company. The failure or inability of our PRC resident shareholders to comply with the registration procedures set forth in these regulations may subject us to fines and legal sanctions, restrict our cross-border investment activities, limit the ability of our wholly foreign-owned subsidiary in China to distribute dividends and the proceeds from any reduction in capital, share transfer or liquidation to us, and we may also be prohibited from injecting additional capital into the subsidiary. Moreover, failure to comply with the various foreign exchange registration requirements described above could result in liability under PRC law for circumventing applicable foreign exchange restrictions. As a result, our business operations and our ability to distribute profits to you could be materially and adversely affected.

Furthermore, as these foreign exchange regulations are still relatively new and their interpretation and implementation has been constantly evolving, it is unclear how these regulations, and any future regulation concerning offshore or cross-border transactions, will be interpreted, amended and implemented by the relevant government authorities. For example, we may be subject to a more stringent review and approval process with respect to our foreign exchange activities, such as remittance of dividends and foreign-currency-denominated borrowings, which may adversely affect our financial condition and results of operations. In addition, if we decide to acquire a PRC domestic company, we cannot assure you that we or the owners of such company, as the case may be, will be able to obtain the necessary approvals or complete the necessary filings and registrations required by the foreign exchange regulations. This may restrict our ability to implement our acquisition strategy and could adversely affect our business and prospects.

We may be materially adversely affected if our shareholders and beneficial owners who are PRC entities fail to comply with the relevant PRC overseas investment regulations.

On December 26, 2017, the NDRC promulgated the Administrative Measures on Overseas Investments, or NDRC Order No. 11, which took effect as of March 1, 2018. According to NDRC Order No. 11, non-sensitive overseas investment projects are subject to record-filing requirements with the local branch of the NDRC. On September 6, 2014, MOFCOM promulgated the *Administrative Measures on Overseas Investments*, which took effect as of October 6, 2014. According to this regulation, overseas investments of PRC enterprises that involve non-sensitive countries and regions and non-sensitive industries are subject to record-filing requirements with a local MOFCOM branch. According to the *Circular of the State Administration of Foreign Exchange on Issuing the Regulations on Foreign Exchange Administration of the Overseas Direct Investment of Domestic Institutions*, which was promulgated by SAFE on July 13, 2009 and took effect on August 1, 2009, PRC enterprises must register for overseas direct investment with a local SAFE branch.

We may not be fully informed of the identities of all our shareholders or beneficial owners who are PRC entities, and we cannot provide any assurance that all of our shareholders and beneficial owners who are PRC entities has or will comply with our request to complete the overseas direct investment procedures under the aforementioned regulations or other related rules in a timely manner, or at all. If they fail to complete the filings or registrations required by the overseas direct investment regulations, the relevant authorities may order them to suspend or cease the implementation of such investment impose warnings and sanctions and make corrections within a specified time, or limit our ability to distribute dividends and proceeds to our PRC subsidiary, which may adversely affect our business, financial condition and results of operations.

PRC regulation of loans to and direct investment in PRC entities by offshore holding companies and governmental control of currency conversion may delay or prevent us from using the proceeds of this offering to make loans or additional capital contributions to our PRC subsidiary, which could materially and adversely affect our liquidity and our ability to fund and expand our business.

We are an offshore holding company conducting our operations in China through our PRC subsidiary. We also hold certain intellectual properties and outsource certain research and development activities related to these intellectual properties to our subsidiaries. We may in the future make loans or provide guarantee to our PRC subsidiary subject to the approval or registration from governmental authorities and limitation of amount, or we may make additional capital contributions to our wholly foreign-owned subsidiary in China. Any loans to our wholly foreign-owned subsidiary in China, which are treated as foreign-invested enterprises under PRC law, are subject to foreign exchange loan registrations. In addition, a foreign-invested enterprise, or FIE, shall use its capital pursuant to the principle of authenticity and self-use within its business scope. The capital of an FIE shall not be used for the following purposes: (i) directly or indirectly used for payment beyond the business scope of the enterprises or the payment prohibited by relevant laws and regulations; (ii) directly or indirectly used for investment in securities or investments other than banks' principal-secured products unless otherwise provided by relevant laws and regulations; (iii) the granting of loans to non-affiliated enterprises, except where it is expressly permitted in the business license; and (iv) paying the expenses related to the purchase of real estate that is not for self-use (except for the foreign-invested real estate enterprises).

In light of the various requirements imposed by PRC regulations on loans to and direct investment in PRC entities by offshore holding companies, we cannot assure you that we will be able to complete the necessary government registrations or obtain the necessary government approvals on a timely basis, if at all, with respect to future loans by us to our PRC subsidiary or with respect to future capital contributions by us to our PRC subsidiary. If we fail to complete such registrations or obtain such approvals, our ability to use the proceeds from this offering and to capitalize or otherwise fund our PRC operations may be negatively affected, which could materially and adversely affect our liquidity and our ability to fund and expand our business.

We may rely on dividends and other distributions on equity paid by our PRC subsidiary to fund any cash and financing requirements we may have, and any limitation on the ability of our PRC subsidiary to make payments to us could have a material and adverse effect on our ability to conduct our business.

We are a Cayman Islands holding company and we rely principally on dividends and other distributions on equity from our PRC subsidiary for our cash requirements, including the funds necessary to pay dividends and other cash distributions to our shareholders for services of any debt we may incur. If our PRC subsidiary incurs debt on its own behalf in the future, the instruments governing the debt may restrict its ability to pay dividends or make other distributions to us. Under PRC laws and regulations, our PRC subsidiary, which is a wholly foreign-owned enterprise, may pay dividends only out of its respective accumulated profits as determined in accordance with PRC accounting standards and regulations. In addition, a wholly foreign-owned enterprise is required to set aside at least 10% of its accumulated after-tax profits each year, if any, to fund a certain statutory reserve fund, until the aggregate amount of such fund reaches 50% of its registered capital. Such reserve funds cannot be distributed to us as dividends. At its discretion, a wholly foreign-owned enterprise may allocate a portion of its after-tax profits based on PRC accounting standards to an enterprise expansion fund, or a staff welfare and bonus fund.

A portion of our revenue was generated by our PRC subsidiary in Renminbi, which is not freely convertible into other currencies. As a result, any restriction on currency exchange may limit the ability of our PRC subsidiary to use its Renminbi revenues to pay dividends to us.

The PRC government may continue to strengthen its capital controls, and more restrictions and substantial vetting process may be put forward by SAFE for cross-border transactions falling under both the current account and the capital account. Any limitation on the ability of our PRC subsidiary to pay dividends or make other kinds of payments to us could materially and adversely limit our ability to grow, make investments or acquisitions that could be beneficial to our business, pay dividends, or otherwise fund and conduct our business.

In addition, the Enterprise Income Tax Law and its implementation rules provide that a withholding tax rate of up to 10% will be applicable to dividends payable by Chinese companies to non-PRC-resident enterprises unless otherwise exempted or reduced according to treaties or arrangements between the PRC central government and governments of other countries or regions where the non-PRC-resident enterprises are incorporated.

Fluctuations in exchange rates could have a material adverse effect on our results of operations and the value of your investment.

The value of the Renminbi against the U.S. dollar and other currencies may fluctuate and is affected by, among other things, changes in political and economic conditions in China and by China's foreign exchange policies. On July 21, 2005, the PRC government changed its policy of pegging the value of the Renminbi to the U.S. dollar, and the Renminbi appreciated more than 20% against the U.S. dollar over the following three years. Between July 2008 and June 2010, this appreciation halted and the exchange rate between the Renminbi and the U.S. dollar remained within a narrow band. Since June 2010, the Renminbi has fluctuated against the U.S. dollar, at times significantly and unpredictably. Since October 1, 2016, Renminbi has joined the International Monetary Fund's basket of currencies that make up the Special Drawing Right, or SDR, along with the U.S. dollar, the Euro, the Japanese yen and the British pound. In the fourth quarter of 2016, the Renminbi has depreciated significantly in the backdrop of a surging U.S. dollar and persistent capital outflows of China. With the development of the foreign exchange market and progress towards interest rate liberalization and Renminbi internationalization, the PRC government may in the future announce further changes to the exchange rate system, and we cannot assure you that the Renminbi will not appreciate or depreciate significantly in value against the U.S. dollar in the future. It is difficult to predict how market forces or PRC or U.S. government policy may impact the exchange rate between the Renminbi and the U.S. dollar in the future.

Significant revaluation of the Renminbi may have a material and adverse effect on your investment. For example, to the extent that we need to convert U.S. dollars we receive from this offering into Renminbi for our operations, appreciation of the Renminbi against the U.S. dollar would have an adverse effect on the Renminbi amount we would receive from the conversion. Conversely, if we decide to convert our Renminbi into U.S. dollars for the purpose of making payments for dividends on our ordinary shares or the ADSs or for other business purposes, appreciation of the U.S. dollar against the Renminbi would have a negative effect on the U.S. dollar amount available to us.

Very limited hedging options are available in China to reduce our exposure to exchange rate fluctuations. To date, we have not entered into any hedging transactions in an effort to reduce our exposure to foreign currency exchange risk. While we may decide to enter into hedging transactions in the future, the availability and effectiveness of these hedges may be limited and we may not be able to adequately hedge our exposure or at all. In addition, our currency exchange losses may be magnified by PRC exchange control regulations that restrict our ability to convert Renminbi into foreign currency.

Governmental control of currency conversion may limit our ability to utilize our cash balance effectively and affect the value of your investment.

The PRC government imposes controls on the convertibility of the Renminbi into foreign currencies and, in certain cases, the remittance of currency out of China. In the years ended December 31, 2018 and 2019 and nine months ended September 30, 2020, we received a portion of our revenues in Renminbi. Under our current corporate structure, our Cayman Islands holding company primarily relies on previous rounds of private financing to fund any cash and financing requirements we may have. Under existing PRC foreign exchange regulations, payments of current account items, including profit distributions, interest payments and trade and service-related foreign exchange transactions, can be made in foreign currencies without prior approval of SAFE by complying with applicable laws and regulations, as well as certain procedural requirements. Specifically, under the existing exchange restrictions, without prior approval of SAFE, cash generated from the operations of our PRC subsidiary may be used to pay dividends to our company. However, approval from or registration with appropriate government authorities is required where Renminbi is to be converted into foreign currency and remitted out of China to pay capital expenses such as the repayment of loans denominated in foreign currencies. As a result, we need to obtain SAFE approval to use the cash generated from the operations of our PRC subsidiary to pay off their respective debt in a currency other than Renminbi owed to entities outside China, or to make other capital expenditure payments outside China in a currency other than Renminbi. The PRC government may at its discretion restrict access to foreign currencies for current account transactions in the future. If the foreign exchange control system prevents us from obtaining sufficient foreign currencies to satisfy our foreign currency demands, we may not be able to pay dividends in foreign currencies to our shareholders, including holders of the ADSs.

The audit report included in this prospectus is prepared by an auditor that is not inspected by the Public Company Accounting Oversight Board and, as such, our investors are deprived of the benefits of such inspection. In addition, various legislative and regulatory developments related to U.S.-listed China based companies due to lack of PCAOB inspection and other developments due to political tensions between the United States and China may have a material adverse impact on our listing and trading in the U.S. and the trading prices of our ADSs, and we could be delisted if we are unable to meet the PCAOB inspection requirements in time.

Our auditor, the independent registered public accounting firm that issued the audit reports included elsewhere in this prospectus filed with the U.S. Securities and Exchange Commission, or SEC, as an auditor of companies that are traded publicly in the United States and a firm registered with the Public Company Accounting Oversight Board (United States), or PCAOB, is subject to laws in the United States pursuant to which the PCAOB conducts regular inspections to assess its compliance with applicable professional standards. Our auditor is located in, and organized under the laws of, the PRC, which is a jurisdiction where the PCAOB has been unable to conduct inspections without the approval of the Chinese authorities.

On May 24, 2013, the PCAOB announced that it had entered into a Memorandum of Understanding on Enforcement Cooperation with the China Securities Regulatory Commission, or CSRC and the PRC Ministry of Finance, which establishes a cooperative framework between the parties for the production and exchange of audit documents relevant to investigations undertaken by the PCAOB, the CSRC or the PRC Ministry of Finance in the United States and the PRC, respectively. The PCAOB continues to be in discussions with the CSRC and the PRC Ministry of Finance to permit joint inspections in the PRC of audit firms that are registered with the PCAOB and audit Chinese companies that trade on U.S. exchanges. On December 7, 2018, the SEC and the PCAOB issued a joint statement highlighting continued challenges faced by the U.S. regulators in their oversight of financial statement audits of U.S.-listed companies with significant operations in China. On April 21,

2020, the Chairman of the SEC, the Chairman of the PCAOB and certain other SEC divisional heads jointly issued a public statement highlighting the significant disclosure, financial reporting and other risks associated with emerging market investments, including the PCAOB's continued inability to inspect audit work papers in China. The 2018 joint statement and the 2020 public statement reflect a heightened regulatory interest in this issue. In response to the U.S. President Trump's Memorandum on Protecting United States Investors from Significant Risks from Chinese Companies, on August 6, 2020, the U.S. President's Working Group on Financial Markets, or the PWG, released a report where it recommends that the SEC take steps to enhanced listing requirements on companies from certain jurisdictions, such as China, that do not provide the PCAOB with sufficient access to audit working papers. The proposed enhanced listing standards require, as a condition to initial and continued exchange listing, unrestricted PCAOB access to work papers of the principal audit firm for the audit of the listed company. Companies that are unable to satisfy this standard as a result of governmental restrictions may satisfy this standard by providing a co-audit from an audit firm with comparable resources and experience where the PCAOB determines it has sufficient access to audit work papers and practices to conduct an appropriate inspection of the co-audit firm. The proposed new listing standards provide for a transition period until January 1, 2022 for currently listed companies. After this transition period, if currently listed companies were unable to meet the enhanced listing standards, then they would become subject to securities exchange rules and processes that could lead to possible de-listing if not cured. The measures in the PWG report are presumably subject to the standard SEC rulemaking process before becoming effective. On August 10, 2020, the SEC announced that SEC Chairman Jay Clayton had directed the SEC staff to prepare proposals in response to the PWG report, and that the SEC was soliciting public comments and information with respect to these proposals. The PCAOB's inspections of other firms outside China have identified deficiencies in those firms' audit procedures and quality control procedures, which may be addressed as part of the inspection process to improve future audit quality. However, it remains unclear what additional actions the SEC and the stock exchanges will take in response to the PWG report.

This lack of PCAOB inspections in China prevents the PCAOB from fully evaluating audits and quality control procedures of our independent registered public accounting firm. As a result, we and investors in our ADSs are deprived of the benefits of such PCAOB inspections. The inability of the PCAOB to conduct inspections of auditors in China makes it more difficult to evaluate the effectiveness of our independent registered public accounting firm's audit procedures or quality control procedures as compared to auditors outside of China that are subject to PCAOB inspections, which could cause investors and potential investors in our ADSs to lose confidence in our audit procedures and reported financial information and the quality of our financial statements.

In addition, as part of a continued regulatory focus in the United States on access to audit and other information currently protected by national law, in particular China's, in June 2019, a bipartisan group of lawmakers introduced bills in both houses of the U.S. Congress, which if passed, would require the SEC to maintain a list of issuers for which PCAOB is not able to inspect or investigate an auditor report issued by a foreign public accounting firm. The proposed Ensuring Quality Information and Transparency for Abroad-Based Listings on our Exchanges (EQUITABLE) Act prescribes increased disclosure requirements for these issuers and, beginning in 2025, the delisting from U.S. national securities exchanges such as the NYSE of issuers included on the SEC's list for three consecutive years. On May 20, 2020, the U.S. Senate passed S. 945, the Holding Foreign Companies Accountable Act, or the Kennedy Bill. On July 21, 2020, the U.S. House of Representatives approved its version of the National Defense Authorization Act for Fiscal Year 2021, which contains provisions comparable to the Kennedy Bill. As part of a continued regulatory focus in the United States on access to audit and other information currently protected by national law, in particular China's, U.S. President Donald J. Trump signed into law on December 18, 2020 the Holding Foreign Companies Accountable Act, which will require the SEC to propose rules within 90 days after its enactment to prohibit securities of any registrant from being listed on any of the U.S. securities exchanges or traded "over the

counter" if the auditor of the registrant's financial statements is not subject to PCAOB inspection for three consecutive years after the law becomes effective. Enactment of this legislation or other efforts to increase U.S. regulatory access to audit information could cause investor uncertainty for affected issuers, including us, and the market price of the ADSs could be adversely affected, and we could be delisted if we are unable to cure the situation to meet the PCAOB inspection requirement in time. It is unclear if and when any of such proposed legislations will be enacted. See "—We could be delisted if we are unable to meet the PCAOB inspection requirements in time."

We could be delisted if we are unable to meet the PCAOB inspection requirements in time.

On December 18, 2020, the Holding Foreign Companies Accountable Act was enacted. In essence, the act requires the SEC to prohibit securities of any foreign companies from being listed on U.S. securities exchanges or traded "over-the-counter" if a company retains a foreign accounting firm that cannot be inspected by the PCAOB for three consecutive years, beginning in 2021. Our independent registered public accounting firm is located in and organized under the laws of the PRC, a jurisdiction where the PCAOB is currently unable to conduct inspections without the approval of the Chinese authorities, and therefore our auditors are not currently inspected by the PCAOB.

The enactment of the Holding Foreign Companies Accountable Act and any additional rulemaking efforts to increase U.S. regulatory access to audit information in China could cause investor uncertainty for affected SEC registrants, including us, and the market price of our ADSs could be materially adversely affected. Additionally, whether the PCAOB will be able to conduct inspections of our auditors in the next three years, or at all, is subject to substantial uncertainty and depends on a number of factors out of our control. If we are unable to meet the PCAOB inspection requirement in time, we could be delisted from the Nasdaq Global Market and our ADSs will not be permitted for trading "over-the-counter" either. Such a delisting would substantially impair your ability to sell or purchase our ADSs when you wish to do so, and the risk and uncertainty associated with delisting would have a negative impact on the price of our ADSs. Also, such a delisting would significantly affect our ability to raise capital on terms acceptable to us, or at all, which would have a material adverse impact on our business, financial condition and prospects.

Proceedings instituted by the SEC against certain PRC-based accounting firms, including the affiliate of our independent registered public accounting firm, or any related adverse regulatory development in the PRC, could result in our financial statements being determined to not be in compliance with the requirements of the Exchange Act.

In December 2012, the SEC instituted administrative proceedings against the Big Four PRC-based accounting firms, including our independent registered public accounting firm, alleging that these firms had violated U.S. securities laws and the SEC's rules and regulations thereunder by failing to provide to the SEC the firms' audit work papers with respect to certain PRC-based companies that are publicly traded in the United States.

On January 22, 2014, the administrative law judge presiding over the matter rendered an initial decision that each of the firms had violated the SEC's rules of practice by failing to produce audit papers and other documents to the SEC. The initial decision censured each of the firms and barred them from practicing before the SEC for a period of six months.

On February 6, 2015, the four China-based accounting firms each agreed to a censure and to pay a fine to the SEC to settle the dispute and avoid suspension of their ability to practice before the SEC and audit U.S.-listed companies. The settlement required the firms to follow detailed procedures and to seek to provide the SEC with access to Chinese firms' audit documents via the CSRC. Under the terms of the settlement, the underlying proceeding against the four China-based accounting firms was deemed

dismissed with prejudice four years after entry of the settlement. The four-year mark occurred on February 6, 2019.

While we cannot predict if the SEC will further challenge the four China-based accounting firms' compliance with U.S. law in connection with U.S. regulatory requests for audit work papers or if the results of such a challenge would result in the SEC imposing penalties such as suspensions, if the accounting firms are subject to additional remedial measures, our ability to file our financial statements in compliance with SEC requirements could be impacted. A determination that we have not timely filed financial statements in compliance with SEC requirements could ultimately lead to the delisting of our ordinary shares or ADSs or the termination of the registration of our ordinary shares or ADSs under the Exchange Act, or both, which would substantially reduce or effectively terminate the trading of our ordinary shares or ADSs in the United States.

Risks Related to the ADSs and This Offering

You may be subject to limitations on transfer of your ADSs.

Your ADSs are transferable on the books of the depository. However, the depository may close its transfer books at any time or from time to time when it deems expedient in connection with the performance of its duties. In addition, the depository may refuse to deliver, transfer or register transfers of ADSs generally when our books or the books of the depository are closed, or at any time if we or the depository deems it advisable to do so because of any requirement of law or of any government or governmental body, or under any provision of the deposit agreement, or for any other reason in accordance with the terms of the deposit agreement.

An active trading market for our ordinary shares or the ADSs may not develop and the trading price for the ADSs may fluctuate significantly.

We have applied to list the ADSs on the Nasdaq Global Market. We have no current intention to seek a listing for our ordinary shares on any stock exchange. Prior to the completion of this offering, there has been no public market for the ADSs or our ordinary shares, and we cannot assure you that a liquid public market for the ADSs will develop. If an active public market for the ADSs does not develop following the completion of this offering, the market price and liquidity of the ADSs may be materially and adversely affected. The initial public offering price for the ADSs will be determined by negotiation between us and the underwriters based upon several factors, and we can provide no assurance that the trading price of the ADSs after this offering will not decline below the initial public offering price. As a result, investors in our securities may experience a significant decrease in the value of their ADSs.

The trading price of the ADSs is likely to be volatile, which could result in substantial losses to investors.

The trading price of the ADSs is likely to be volatile and could fluctuate widely due to factors beyond our control. This may happen because of broad market and industry factors, including the performance and fluctuation of the market prices of other companies with business operations located mainly in China that have listed their securities in the United States. In addition to market and industry factors, the price and trading volume for the ADSs may be highly volatile for factors specific to our own operations, including the following:

- variations in our net revenues, earnings and cash flow;
- announcements of new investments, acquisitions, strategic partnerships, or joint ventures by us or our competitors;
- announcements of new products and services and expansions by us or our competitors;

- changes in financial estimates by securities analysts;
- fluctuations in operating metrics;
- failure on our part to realize monetization opportunities as expected;
- changes in revenues generated from our significant business partners;
- additions or departures of key personnel;
- release of lock-up or other transfer restrictions on our outstanding equity securities or sales of additional equity securities;
- detrimental negative publicity about us, our management, our competitors or our industry;
- regulatory developments affecting us or our industry; and
- potential litigation or regulatory investigations.

Any of these factors may result in large and sudden changes in the trading volume and price of the ADSs.

In the past, shareholders of public companies have often brought securities class action suits against those companies following periods of instability in the market price of their securities. If we were involved in a class action suit, it could divert a significant amount of our management's attention and other resources from our business and operations and require us to incur significant expenses to defend the suit, which could harm our results of operations. Any such class action suit, whether or not successful, could harm our reputation and restrict our ability to raise capital in the future. In addition, if a claim is successfully made against us, we may be required to pay significant damages, which could have a material adverse effect on our financial condition and results of operations.

We are an emerging growth company within the meaning of the Securities Act and may take advantage of certain reduced reporting requirements.

We are an "emerging growth company," as defined in the JOBS Act, and we may take advantage of certain exemptions from requirements applicable to other public companies that are not emerging growth companies including, most significantly, not being required to comply with the auditor attestation requirements of Section 404 of the Sarbanes-Oxley Act of 2002 for so long as we remain an emerging growth company. As a result, if we elect not to comply with such auditor attestation requirements, our investors may not have access to certain information they may deem important.

If securities or industry analysts cease to publish research or reports about our business, or if they adversely change their recommendations regarding the ADSs, the market price for the ADSs and trading volume could decline.

The trading market for the ADSs will be influenced by research or reports that industry or securities analysts publish about our business. If one or more analysts who cover us downgrade the ADSs, the market price for the ADSs would likely decline. If one or more of these analysts cease to cover us or fail to regularly publish reports on us, we could lose visibility in the financial markets, which, in turn, could cause the market price or trading volume for the ADSs to decline.

The sale or availability for sale, or perceived sale or availability for sale, of substantial amounts of the ADSs could adversely affect their market price.

Sales of substantial amounts of the ADSs in the public market after the completion of this offering, or the perception that these sales could occur, could adversely affect the market price of the ADSs and could materially impair our ability to raise capital through equity offerings in the future. The

ADSs sold in this offering will be freely tradable without restriction or further registration under the Securities Act, and shares held by our existing shareholders may also be sold in the public market in the future subject to the restrictions in Rule 144 and Rule 701 under the Securities Act and the applicable lock-up agreements. There will be ADSs (equivalent to ordinary shares) outstanding immediately after this offering, or ADSs (equivalent to ordinary shares) if the underwriters exercise their over-allotment option in full. [In connection with this offering, we, our directors and executive officers and our existing shareholders have agreed not to sell any ordinary shares or ADSs for 180 days after the date of this prospectus without the prior written consent of the underwriters, subject to certain exceptions. However, the underwriters may release these securities from these restrictions at any time, subject to applicable regulations of the Financial Industry Regulatory Authority, Inc.] We cannot predict what effect, if any, market sales of securities held by our significant shareholders or any other shareholder or the availability of these securities for future sale will have on the market price of the ADSs. See "Underwriting" and "Shares Eligible for Future Sale" for a more detailed description of the restrictions on selling these securities after this offering.

Our memorandum and articles of association contain anti-takeover provisions that could have a material adverse effect on the rights of holders of our ordinary shares and the ADSs.

We will adopt amended and restated memorandum and articles of association that will become effective immediately prior to the completion of this offering. Our new memorandum and articles of association contain provisions to limit the ability of others to acquire control of our company or cause us to engage in change-of-control transactions. These provisions could have the effect of depriving our shareholders of an opportunity to sell their shares at a premium over prevailing market prices by discouraging third parties from seeking to obtain control of our company in a tender offer or similar transaction. Our board of directors has the authority, without further action by our shareholders, to issue preferred shares in one or more series and to fix their designations, powers, preferences, privileges, and relative participating, optional or special rights and the qualifications, limitations or restrictions, including dividend rights, conversion rights, voting rights, terms of redemption and liquidation preferences, any or all of which may be greater than the rights associated with our ordinary shares, including ordinary shares represented by ADSs. Preferred shares could be issued quickly with terms calculated to delay or prevent a change in control of our company or make removal of management more difficult. If our board of directors decides to issue preferred shares, the price of the ADSs may fall and the voting and other rights of the holders of our ordinary shares and the ADSs may be materially and adversely affected.

We have not determined a specific use for a portion of the net proceeds from this offering, and we may use these proceeds in ways with which you may not agree, and such use may not produce income or increase the ADS price.

We have not determined a specific use for a portion of the net proceeds of this offering, and our management will have considerable discretion in deciding how to apply these proceeds. You will not have the opportunity to assess whether the proceeds are being used appropriately before you make your investment decision. You must rely on the judgment of our management regarding the application of the net proceeds of this offering. We cannot assure you that the net proceeds will be used in a manner that would improve our results of operations or increase the ADS price, nor that these net proceeds will be placed only in investments that generate income or appreciate in value. Currently, we do not have any plans, commitments or understandings to acquire complementary business, assets and technologies.

We are entitled to amend the deposit agreement and to change the rights of ADS holders under the terms of such agreement, or to terminate the deposit agreement, without the prior consent of the ADS holders.

We are entitled to amend the deposit agreement and to change the rights of the ADS holders under the terms of such agreement, without the prior consent of the ADS holders. We and the depositary may agree to amend the deposit agreement in any way we decide is necessary or advantageous to us. Amendments may reflect, among other things, operational changes in the ADS program, legal developments affecting ADSs or changes in the terms of our business relationship with the depositary. In the event that the terms of an amendment are disadvantageous to ADS holders, ADS holders will only receive 30 days' advance notice of the amendment, and no prior consent of the ADS holders is required under the deposit agreement. Furthermore, we may decide to terminate the ADS facility at any time for any reason. For example, terminations may occur when we decide to list our shares on a non-U.S. securities exchange and determine not to continue to sponsor an ADS facility or when we become the subject of a takeover or a going-private transaction. If the ADS facility will terminate, ADS holders will receive at least 90 days' prior notice, but no prior consent is required from them. Under the circumstances that we decide to make an amendment to the deposit agreement that is disadvantageous to ADS holders or terminate the deposit agreement, the ADS holders may choose to sell their ADSs or surrender their ADSs and become direct holders of the underlying ordinary shares, but will have no right to any compensation whatsoever.

ADSs holders may not be entitled to a jury trial with respect to claims arising under the deposit agreement, which could result in less favorable outcomes to the plaintiff(s) in any such action.

The deposit agreement governing the ADSs representing our ordinary shares provides that, to the fullest extent permitted by law, ADS holders waive the right to a jury trial of any claim they may have against us or the depositary arising out of or relating to our shares, the ADSs or the deposit agreement, including any claim under the U.S. federal securities laws.

If we or the depositary opposed a jury trial demand based on the waiver, the court would determine whether the waiver was enforceable based on the facts and circumstances of that case in accordance with the applicable state and federal law. To our knowledge, the enforceability of a contractual pre-dispute jury trial waiver in connection with claims arising under the federal securities laws has not been finally adjudicated by the United States Supreme Court. However, we believe that a contractual pre-dispute jury trial waiver provision is generally enforceable, including under the laws of the State of New York, which govern the deposit agreement, by a federal or state court in the City of New York, which has non-exclusive jurisdiction over matters arising under the deposit agreement. In determining whether to enforce a contractual pre-dispute jury trial waiver provision, courts will generally consider whether a party knowingly, intelligently and voluntarily waived the right to a jury trial. We believe that this is the case with respect to the deposit agreement and the ADSs. It is advisable that you consult legal counsel regarding the jury waiver provision before entering into the deposit agreement.

If you or any other holders or beneficial owners of ADSs bring a claim against us or the depositary in connection with matters arising under the deposit agreement or the ADSs, including claims under federal securities laws, you or such other holder or beneficial owner may not be entitled to a jury trial with respect to such claims, which may have the effect of limiting and discouraging lawsuits against us or the depositary. If a lawsuit is brought against us or the depositary under the deposit agreement, it may be heard only by a judge or justice of the applicable trial court, which would be conducted according to different civil procedures and may result in different outcomes than a trial by jury would have had, including results that could be less favorable to the plaintiff(s) in any such action.

Nevertheless, if this jury trial waiver provision is not permitted by applicable law, an action could proceed under the terms of the deposit agreement with a jury trial.

No condition, stipulation or provision of the deposit agreement or ADSs serves as a waiver by any holder or beneficial owner of ADSs or by us or the depositary of compliance with any substantive provision of the U.S. federal securities laws and the rules and regulations promulgated thereunder.

The voting rights of holders of ADSs are limited by the terms of the deposit agreement, and you may not be able to exercise your right to direct the voting of the underlying ordinary shares represented by your ADSs.

Holders of ADSs do not have the same rights as our registered shareholders. As a holder of ADSs, you will not have any direct right to attend general meetings of our shareholders or to cast any votes at such meetings. You will only be able to exercise the voting rights attached to the ordinary shares underlying your ADSs indirectly by giving voting instructions to the depositary in accordance with the provisions of the deposit agreement. Where any matter is to be put to a vote at a general meeting, then upon receipt of your voting instructions, the depositary will try, as far as is practicable, to vote the underlying ordinary shares represented by your ADSs in accordance with your instructions. You will not be able to directly exercise your right to vote with respect to the underlying ordinary shares unless you cancel and withdraw the shares and become the registered holder of such shares prior to the record date for the general meeting.

When a general meeting is convened, you may not receive sufficient advance notice of the meeting to withdraw the ordinary shares represented by your ADSs and become the registered holder of such shares to allow you to attend the general meeting and to vote directly with respect to any specific matter or resolution to be considered and voted upon at the general meeting. In addition, under our post-offering memorandum and articles of association that will become effective immediately prior to completion of this offering, for the purposes of determining those shareholders who are entitled to attend and vote at any general meeting, our directors may close our register of members and/or fix in advance a record date for such meeting, and such closure of our register of members or the setting of such a record date may prevent you from withdrawing the underlying ordinary shares represented by your ADSs and from becoming the registered holder of such shares prior to the record date, so that you would not be able to attend the general meeting or to vote directly. Where any matter is to be put to a vote at a general meeting, upon our instruction the depositary will notify you of the upcoming vote and will arrange to deliver our voting materials to you. We cannot assure you that you will receive the voting materials in time to ensure that you can instruct the depositary to vote the underlying ordinary shares represented by your ADSs.

In addition, the depositary and its agents are not responsible for failing to carry out voting instructions or for their manner of carrying out your voting instructions. This means that you may not be able to exercise your right to direct how the underlying ordinary shares represented by your ADSs are voted and you may have no legal remedy if the underlying ordinary shares represented by your ADSs are not voted as you requested. In addition, in your capacity as an ADS holder, you will not be able to call a shareholders' meeting.

Under the deposit agreement, if you do not vote, the depositary may give us a discretionary proxy to vote the ordinary shares underlying the ADSs at shareholders' meetings if we have timely provided the depositary with notice of meeting and related voting materials and (i) we have instructed the depositary that we wish a discretionary proxy to be given, (ii) we have informed the depositary that there is no substantial opposition as to a matter to be voted on at the meeting, and (iii) a matter to be voted on at the meeting would not have a material adverse impact on shareholders.

The effect of this discretionary proxy is that you cannot prevent the underlying ordinary shares represented by the ADSs from being voted, except under the circumstances described above. This may make it more difficult for ADS holders to influence the management of the company. Holders of ordinary shares are not subject to this discretionary proxy.

Because the initial public offering price is substantially higher than the pro forma net tangible book value per share, you will experience immediate and substantial dilution.

If you purchase the ADSs in this offering, you will pay more for each ADS than the corresponding amount paid by existing shareholders for their ordinary shares. As a result, you will experience immediate and substantial dilution of approximately US\$ per ADS, assuming that no outstanding options to acquire ordinary shares are exercised. This number represents the difference between the assumed initial public offering price of US\$ per ADS, being the mid-point of the estimated range of the initial offering price shown on the front cover of this prospectus, and our pro forma net tangible book value per ADS as of , 2019, after giving effect to this offering. You may experience further dilution to the extent that our ordinary shares are issued upon exercise of any share options. See "Dilution" for a more complete description of how the value of your investment in ADSs will be diluted upon completion of this offering.

Because we do not expect to pay dividends in the foreseeable future after this offering, you must rely on price appreciation of the ADSs for return on your investment.

We currently intend to retain most, if not all, of our available funds and any future earnings after this offering to fund the development and growth of our business. As a result, we do not expect to pay any cash dividends in the foreseeable future. Therefore, you should not rely on an investment in the ADSs as a source for any future dividend income.

Our board of directors has complete discretion as to whether to distribute dividends, subject to certain requirements of Cayman Islands law. In addition, our shareholders may, subject to the provisions of our articles of association, by ordinary resolution declare a dividend, but no dividend may exceed the amount recommended by our directors. Under Cayman Islands law, a Cayman Islands company may pay a dividend out of either profit or share premium account, provided that in no circumstances may a dividend be paid if this would result in the company being unable to pay its debts as they fall due in the ordinary course of business. Even if our board of directors decides to declare and pay dividends, the timing, amount and form of future dividends, if any, will depend on our future results of operations and cash flow, our capital requirements and surplus, the amount of distributions, if any, received by us from our subsidiary, our financial condition, contractual restrictions and other factors deemed relevant by our board of directors. Accordingly, the return on your investment in the ADSs will likely depend entirely upon any future price appreciation of the ADSs. There is no guarantee that the ADSs will appreciate in value after this offering or even maintain the price at which you purchased the ADSs. You may not realize a return on your investment in the ADSs and you may even lose your entire investment in the ADSs.

You may not receive dividends or other distributions on our ordinary shares and you may not receive any value for them, if it is illegal or impractical to make them available to you.

The depositary has agreed to pay to you the cash dividends or other distributions it or the custodian receives on ordinary shares or other deposited securities underlying the ADSs, after deducting its fees and expenses. You will receive these distributions in proportion to the number of ordinary shares your ADSs represent. However, the depositary is not responsible if it decides that it is unlawful or impractical to make a distribution available to any holders of ADSs. For example, it would be unlawful to make a distribution to a holder of ADSs if it consists of securities that require registration under the Securities Act of 1933 but that are not properly registered or distributed under an applicable exemption from registration. The depositary may also determine that it is not feasible to distribute certain property through the mail. Additionally, the value of certain distributions may be less than the cost of mailing them. In these cases, the depositary may determine not to distribute such property. We have no obligation to register under U.S. securities laws any ADSs, ordinary shares, rights or other securities received through such distributions. We also have no obligation to take any other

action to permit the distribution of ADSs, ordinary shares, rights or anything else to holders of ADSs. This means that you may not receive distributions we make on our ordinary shares or any value for them if it is illegal or impractical for us to make them available to you. These restrictions may cause a material decline in the value of the ADSs.

You may experience dilution of your holdings due to the inability to participate in rights offerings.

We may, from time to time, distribute rights to our shareholders, including rights to acquire securities. Under the deposit agreement, the depositary will not distribute rights to holders of ADSs unless the distribution and sale of rights and the securities to which these rights relate are either exempt from registration under the Securities Act with respect to all holders of ADSs, or are registered under the provisions of the Securities Act. The depositary may, but is not required to, attempt to sell these undistributed rights to third parties, and may allow the rights to lapse. We may be unable to establish an exemption from registration under the Securities Act, and we are under no obligation to file a registration statement with respect to these rights or underlying securities or to endeavor to have a registration statement declared effective. Accordingly, holders of ADSs may be unable to participate in our rights offerings and may experience dilution of their holdings as a result.

We will incur increased costs as a result of being a public company, particularly after we cease to qualify as an "emerging growth company."

Upon completion of this offering, we will become a public company and expect to incur significant legal, accounting and other expenses that we did not incur as a private company. The Sarbanes-Oxley Act of 2002, as well as rules subsequently implemented by the Securities and Exchange Commission, or the SEC, and the Nasdaq Global Market, impose various requirements on the corporate governance practices of public companies. As a company with less than US\$1.07 billion in revenues for our last fiscal year, we qualify as an "emerging growth company" pursuant to the JOBS Act. An emerging growth company may take advantage of specified reduced reporting and other requirements that are otherwise applicable generally to public companies. These provisions include exemption from the auditor attestation requirement under Section 404 of the Sarbanes-Oxley Act of 2002, or Section 404, in the assessment of the emerging growth company's internal control over financial reporting and permission to delay adopting new or revised accounting standards until such time as those standards apply to private companies.

We expect these rules and regulations to increase our legal and financial compliance costs and to make some corporate activities more time-consuming and costly. After we are no longer an "emerging growth company," we expect to incur significant expenses and devote substantial management effort toward ensuring compliance with the requirements of Section 404 of the Sarbanes-Oxley Act of 2002 and the other rules and regulations of the SEC. For example, as a result of becoming a public company, we will need to increase the number of independent directors and adopt policies regarding internal controls and disclosure controls and procedures. We also expect that operating as a public company will make it more difficult and more expensive for us to obtain director and officer liability insurance, and we may be required to accept reduced policy limits and coverage or incur substantially higher costs to obtain the same or similar coverage. In addition, we will incur additional costs associated with our public company reporting requirements. It may also be more difficult for us to find qualified persons to serve on our board of directors or as executive officers. We are currently evaluating and monitoring developments with respect to these rules and regulations, and we cannot predict or estimate with any degree of certainty the number of additional costs we may incur or the timing of such costs.

You may face difficulties in protecting your interests, and your ability to protect your rights through U.S. courts may be limited, because we are incorporated under Cayman Islands law.

We are an exempted company incorporated under the laws of the Cayman Islands with limited liability. Our corporate affairs are governed by our memorandum and articles of association, the Companies Act (as amended) of the Cayman Islands and the common law of the Cayman Islands. The rights of shareholders to take action against our directors, actions by our minority shareholders and the fiduciary duties of our directors to us under Cayman Islands law are to a large extent governed by the common law of the Cayman Islands. The common law of the Cayman Islands is derived in part from comparatively limited judicial precedent in the Cayman Islands as well as from the common law of England, the decisions of whose courts are of persuasive authority, but are not binding, on a court in the Cayman Islands. The rights of our shareholders and the fiduciary duties of our directors under Cayman Islands law are not as clearly established as they would be under statutes or judicial precedent in some jurisdictions in the United States. In particular, the Cayman Islands has a less developed body of securities laws than the United States. Some U.S. states, such as Delaware, have more fully developed and judicially interpreted bodies of corporate law than the Cayman Islands. In addition, Cayman Islands companies may not have the standing to initiate a shareholder derivative action in a federal court of the United States.

Shareholders of Cayman Islands exempted companies like us have no general rights under Cayman Islands law to inspect corporate records or to obtain copies of lists of shareholders of these companies. Our directors have discretion under our articles of association that will become effective immediately prior to completion of this offering to determine whether or not, and under what conditions, our corporate records may be inspected by our shareholders, but are not obliged to make them available to our shareholders. This may make it more difficult for you to obtain the information needed to establish any facts necessary for a shareholder motion or to solicit proxies from other shareholders in connection with a proxy contest.

As a result of all of the above, our public shareholders may have more difficulties in protecting their interests in the face of actions taken by management, members of our board of directors or controlling shareholders than they would as public shareholders of a company incorporated in the United States. For a discussion of significant differences between the provisions of the Companies Act of the Cayman Islands and the laws applicable to companies incorporated in the United States and their shareholders, see "Description of Share Capital—Differences in Corporate Law."

Certain judgments obtained against us by our shareholders may not be enforceable.

We are a Cayman Islands company and substantially all of our assets are located outside of the United States. Our current operations are primarily conducted in China. In addition, some of our current directors and officers are nationals and residents of countries other than the United States. Substantially all of the assets of these persons are located outside the United States. As a result, it may be difficult or impossible for you to bring an action against us or against these individuals in the United States in the event that you believe that your rights have been infringed under the U.S. federal securities laws or otherwise. Even if you are successful in bringing an action of this kind, the laws of the Cayman Islands and of China may render you unable to enforce a judgment against our assets or the assets of our directors and officers. For more information regarding the relevant laws of the Cayman Islands and China, see "Enforceability of Civil Liabilities."

As a company incorporated in the Cayman Islands, we are permitted to adopt certain home country practices in relation to corporate governance matters that differ significantly from the Nasdaq listing standards; these practices may afford less protection to shareholders than they would enjoy if we complied fully with the Nasdaq listing standards.

As a Cayman Islands company listed on the Nasdaq Global Market, we are subject to the Nasdaq listing standards. However, Nasdaq rules permit a foreign private issuer like us to follow the corporate governance practices of its home country. Certain corporate governance practices in the Cayman Islands, which is our home country, may differ significantly from the Nasdaq listing standards. We may elect to rely on home country practice to be exempted from the corporate governance requirements. As a result, our shareholders may be afforded less protection than they would otherwise enjoy under the Nasdaq listing standards applicable to U.S. domestic issuers.

We are a foreign private issuer within the meaning of the rules under the Exchange Act, and as such we are exempt from certain provisions applicable to U.S. domestic public companies.

Because we qualify as a foreign private issuer under the Exchange Act, we are exempt from certain provisions of the securities rules and regulations in the United States that are applicable to U.S. domestic issuers, including:

- the rules under the Exchange Act requiring the filing with the SEC of quarterly reports on Form 10-Q or current reports on Form 8-K;
- the sections of the Exchange Act regulating the solicitation of proxies, consents or authorizations in respect of a security registered under the Exchange Act;
- the sections of the Exchange Act requiring insiders to file public reports of their stock ownership and trading activities and liability for insiders who profit from trades made in a short period of time; and
- the selective disclosure rules by issuers of material nonpublic information under Regulation FD.

We will be required to file an annual report on Form 20-F within four months of the end of each fiscal year. In addition, we intend to publish our results on a quarterly basis as press releases, distributed pursuant to the rules and regulations of the Nasdaq Global Market. Press releases relating to financial results and material events will also be furnished to the SEC on Form 6-K. However, the information we are required to file with or furnish to the SEC will be less extensive and less timely compared to that required to be filed with the SEC by U.S. domestic issuers. As a result, you may not be afforded the same protections or information that would be made available to you were you investing in a U.S. domestic issuer.

There can be no assurance that we will not be a passive foreign investment company, or PFIC, for U.S. federal income tax purposes for any taxable year, which could subject U.S. investors in our ADSs or ordinary shares to significant adverse U.S. federal income tax consequences.

In general, a non-U.S. corporation will be a PFIC for any taxable year in which (i) 75% or more of its gross income consists of passive income, or the income test, or (ii) 50% or more of the average value of its assets (generally determined on a quarterly basis) consists of assets that produce, or are held for the production of, passive income, or the asset test. For purposes of the above calculations, a non-U.S. corporation that directly or indirectly owns at least 25% by value of the ordinary shares of another corporation is treated as if it held its proportionate share of the assets of the other corporation and received directly its proportionate share of the income of the other corporation. Passive income generally includes interest, dividends, gains from certain property transactions, rents and royalties (other than certain rents or royalties derived in the active conduct of a trade or business). Cash is a

passive asset for PFIC purposes. Goodwill is an active asset under the PFIC rules to the extent attributable to activities that produce active income.

The assets shown on our balance sheet are expected to consist primarily of cash and cash equivalents for the foreseeable future. Therefore, whether we will satisfy the asset test for the current or any future taxable year will depend largely on the value of our goodwill and on how quickly we utilize the cash in our business. We cannot give any assurance as to whether we will be a PFIC for the current or any future taxable year because (i) the value of our goodwill may be determined by reference to the market price of our ADSs, which may be volatile given the nature and early stage of our business, (ii) we expect to hold a significant amount of cash, and (iii) a company's PFIC status is an annual determination that can be made only after the end of each taxable year. In addition, prior to commercialization of our product candidates, we could be a PFIC for any taxable year in which our interest and other investment income constitutes 75% or more of our total gross income.

If we were a PFIC for any taxable year during which a U.S. investor owns our ADSs or ordinary shares, certain adverse U.S. federal income tax consequences could apply to such U.S. investor. See "Taxation—Material U.S. Federal Income Tax Consequences—Passive Foreign Investment Company Rules."

SPECIAL NOTE REGARDING FORWARD-LOOKING STATEMENTS

This prospectus contains forward-looking statements that involve risks and uncertainties. All statements other than statements of historical facts are forward-looking statements. These statements involve known and unknown risks, uncertainties and other factors that may cause our actual results, performance or achievements to be materially different from those expressed or implied by the forward-looking statements.

You can identify these forward-looking statements by words or phrases such as "may," "will," "expect," "anticipate," "aim," "estimate," "intend," "plan," "believe," "likely to" or other similar expressions. We have based these forward-looking statements largely on our current expectations and projections about future events and financial trends that we believe may affect our financial condition, results of operations, business strategy and financial needs. These forward-looking statements include, but are not limited to, statements about:

- our goals and growth strategies;
- our future business development, results of operations and financial condition;
- the timing of the initiation, progress and potential results of our preclinical studies, clinical trials and our discovery programs;
- our ability to utilize our proprietary Dynamic Precision Library platform, or DPL, to design, construct and develop next-generation antibodies;
- our ability to advance product candidates into, and successfully complete, clinical trials;
- the timing or likelihood of regulatory filings and approvals;
- our estimates of the number of patients who suffer from the diseases we are targeting and the number of patients that may enroll in our clinical trials;
- the commercializing of our product candidates, if approved;
- our ability and the potential to successfully manufacture and supply our product candidates for clinical trials and for commercial use, if approved;
- future strategic arrangements and/or collaborations and the potential benefits of such arrangements;
- our anticipated use of our existing resources and the proceeds from this offering;
- our estimates regarding expenses, future revenue, capital requirements and needs for additional financing and our ability to obtain additional capital;
- the sufficiency of our existing cash and cash equivalents to fund our future operating expenses and capital expenditure requirements;
- our ability to retain the continued service of our key personnel and to identify, hire and retain additional qualified professionals;
- the implementation of our business model, strategic plans for our business and product candidates;
- the scope of protection we are able to establish and maintain for intellectual property rights, such as our proprietary DPL, which includes NEObody platform, SAFEbody platform and POWERbody platform, product candidates and discovery programs;
- our potential to enter into new collaborations;

- our ability to contract with third-party suppliers and manufacturers and their ability to perform adequately;
- the pricing, coverage and reimbursement of our product candidates, if approved;
- developments relating to our competitors and our industry, including competing product candidates and therapies;
- relevant government policies and regulations relating to our business and industry;
- general economic and business condition in the markets we have businesses; and
- assumptions underlying or related to any of the foregoing.

You should read thoroughly this prospectus and the documents that we refer to in this prospectus with the understanding that our actual future results may be materially different from and worse than what we expect. Other sections of this prospectus include additional factors which could adversely impact our business and financial performance. Moreover, we operate in an evolving environment. New risk factors and uncertainties emerge from time to time and it is not possible for our management to predict all risk factors and uncertainties, nor can we assess the impact of all factors on our business or the extent to which any factor, or combination of factors, may cause actual results to differ materially from those contained in any forward-looking statements. We qualify all of our forward-looking statements by these cautionary statements.

You should not rely upon forward-looking statements as predictions of future events. We undertake no obligation to update or revise any forward-looking statements, whether as a result of new information, future events or otherwise.

USE OF PROCEEDS

We estimate that we will receive net proceeds from this offering of approximately US\$ million, or approximately US\$ million if the underwriters exercise their option to purchase additional ADSs in full, after deducting underwriting discounts and commissions and the estimated offering expenses payable by us.

We intend to use the net proceeds from this offering for the following purposes:

- approximately 95% for research and development;

- approximately 26% to fund ADG106, covering both Phase I trial and advancement to Phase II trial;
- approximately 26% to fund both ADG126 and ADG116, covering Phase I trial and advancement to Phase II trial;
- approximately 43% for our technical and platform development and development of our preclinical candidates; and

- approximately 5% for working capital and other general corporate purposes.

The net proceeds from this offering, together with our cash and cash equivalents, will not be sufficient for us to fund any of our product candidates through regulatory approval, our preclinical development pipeline through Phase I clinical trials, or any product candidates resulting from our discovery pipeline into clinical trials. As a result, we will need to raise additional capital to complete the development and commercialization of our products candidates.

If an unforeseen event occurs or business conditions change, we may use the proceeds of this offering differently than as described in this prospectus. In utilizing the proceeds from this offering, we are permitted under PRC laws and regulations to provide funding to our PRC subsidiaries only through loans or capital contributions, and only if we satisfy the applicable government registration and approval requirements. We cannot assure you that we will be able to meet these requirements on a timely basis, if at all. See "Risk Factors—Risks Related to Doing Business in China—PRC regulation of loans to and direct investment in PRC entities by offshore holding companies and governmental control of currency conversion may delay or prevent us from using the proceeds of this offering to make loans or additional capital contributions to our PRC subsidiary, which could materially and adversely affect our liquidity and our ability to fund and expand our business."

DIVIDEND POLICY

We have not previously declared or paid cash dividends and we have no plan to declare or pay any dividends in the near future on our shares or the ADSs representing our ordinary shares. We currently intend to retain most, if not all, of our available funds and any future earnings to operate and expand our business.

We are a holding company incorporated in the Cayman Islands. In the future, we may rely on dividends from our subsidiaries, including our PRC subsidiary, for our cash requirements, including any payment of dividends to our shareholders. PRC regulations may restrict the ability of our PRC subsidiary to pay dividends to us. See "PRC Regulation—Regulations on Foreign Exchange and Dividend Distribution."

Our board of directors has discretion as to whether to distribute dividends, subject to certain requirements of Cayman Islands law. In addition, our shareholders may, subject to the provisions of our post-offering articles of association, by ordinary resolution declare a dividend, but no dividend may exceed the amount recommended by our board of directors. Under Cayman Islands law, a Cayman Islands company may pay a dividend out of either profit or share premium account, provided that in no circumstances may a dividend be paid if this would result in the company being unable to pay its debts as they fall due in the ordinary course of business. Even if our board of directors decides to pay dividends, the form, frequency and amount will depend upon our future operations and earnings, capital requirements and surplus, general financial condition, contractual restrictions and other factors that the board of directors may deem relevant. If we pay any dividends on our ordinary shares, we will pay those dividends which are payable in respect of the ordinary shares underlying the ADSs to the depository, as the registered holder of such ordinary shares, and the depository then will pay such amounts to the ADS holders in proportion to the ordinary shares underlying the ADSs held by such ADS holders, subject to the terms of the deposit agreement, including the fees and expenses payable thereunder. See "Description of American Depositary Shares."

CAPITALIZATION

The following table sets forth our capitalization as of September 30, 2020:

- on an actual basis;
- on a pro forma basis to reflect the automatic conversion of all of our outstanding preferred shares into 27,249,824 ordinary shares immediately prior to the completion of this offering; and
- on a pro forma as adjusted basis to reflect (i) the conversion of all of our issued and outstanding preferred shares into 27,249,824 ordinary shares upon completion of this offering, (ii) the recognition of share-based compensation expense of US\$0.2 million associated with share options for which the service condition was satisfied as of September 30, 2020 and the performance condition will be satisfied upon the completion of this offering, which has been reflected as an increase to additional paid-in capital and accumulated deficit and (iii) the issuance and sale of ordinary shares in the form of ADSs by us in this offering at an assumed initial public offering price of US\$ per ADS being the mid-point of the estimated range of the initial offering price shown on the front cover of this prospectus, after deducting the underwriting discounts and commissions and estimated offering expenses payable by us (assuming the underwriters do not exercise their option to purchase additional ADSs).

You should read this table together with our consolidated financial statements and the related notes included elsewhere in this prospectus and the information under "Management's Discussion and Analysis of Financial Condition and Results of Operations."

	As of September 30, 2020	
	Actual US\$	Pro forma US\$ as adjusted ⁽¹⁾
		(in thousands)
Current portion of long-term borrowings	943	943
Long-term borrowings, less current portion	3,135	3,135
Mezzanine equity		
Series A-1 convertible redeemable preferred shares (par value of US\$0.0001 per share; 5,473,957 shares authorized, issued and outstanding on an actual basis; and nil outstanding on a pro-forma and pro forma as adjusted basis as of June 30, 2020)	5,474	—
Series A-2 convertible redeemable preferred shares (par value of US\$0.0001 per share; 2,370,414 shares authorized, issued and outstanding on an actual basis; and nil outstanding on a pro-forma and pro forma as adjusted basis as of June 30, 2020)	3,000	—
Series B convertible redeemable preferred shares (par value of US\$0.0001 per share; 7,494,537 shares authorized, issued and outstanding on an actual basis; and nil outstanding on a pro-forma and pro forma as adjusted basis as of June 30, 2020)	28,000	—
Series C-1 convertible redeemable preferred shares (par value of US\$0.0001 per share; 5,597,354 shares authorized, issued and outstanding on an actual basis; and nil outstanding on a pro-forma and pro forma as adjusted basis as of June 30, 2020)	48,913	—
Series C-2 convertible redeemable preferred shares (par value of US\$0.0001 per share; 1,861,121 shares authorized, issued and outstanding on an actual basis; and nil outstanding on a pro-forma and pro forma as adjusted basis as of June 30, 2020)	19,000	—
Series C-3 convertible redeemable preferred shares (par value of US\$0.0001 per share; 4,452,441 shares authorized, issued and outstanding on an actual basis; and nil outstanding on a pro-forma and pro forma as adjusted basis as of June 30, 2020)	50,000	—
Total mezzanine equity	<u>154,387</u>	<u>—</u>
Shareholders' equity (deficit)		
Ordinary shares (par value of US\$0.0001 per share; 500,000,000 shares authorized; 16,466,897 shares issued and outstanding as of and September 30, 2020;	2	4
Subscriptions receivable from shareholders	(1,975)	(1,975)
Additional paid-in capital	16,500	170,884
Accumulated other comprehensive loss	(560)	(560)
Accumulated deficit	(92,887)	(92,887)
Total shareholders' equity (deficit)	<u>(78,921)</u>	<u>75,466</u>
Total mezzanine equity and shareholders' equity (deficit)	<u>75,466</u>	<u>75,466</u>
Total capitalization	<u>79,545</u>	<u>79,545</u>

Notes:

- (1) The pro forma as adjusted information does not include the impact of: (i) the modification and exercise of 2,375,000 share options on November 9, 2020 and (ii) the surrender of 491,119 ordinary shares on January 16, 2021.

- (2) The pro forma as adjusted information discussed above is illustrative only. Our additional paid-in capital and total shareholders' equity (deficit) following the completion of this offering are subject to adjustment based on the actual initial public offering price and other terms of this offering determined at pricing. Assuming the number of ADSs offered by us as set forth on the cover page of this prospectus remains the same, and after deduction of underwriting discounts and commissions and the estimated offering expenses payable by us, a US\$1.00 change in the assumed initial public offering price of US\$ per ADS would, in the case of an increase, increase and, in the case of a decrease, decrease each of additional paid-in capital and total shareholders' equity (deficit) by US\$ million.

DILUTION

If you invest in our ADSs, your interest will be diluted to the extent of the difference between the initial public offering price per ADS and our net tangible book value per ADS after this offering. Dilution results from the fact that the initial public offering price per ordinary share is substantially in excess of the book value per ordinary share attributable to the existing shareholders for our presently outstanding ordinary shares.

Our net tangible book value as of September 30, 2020 was approximately US\$ million. Net tangible book value represents the amount of our total consolidated tangible assets, less the amount of our total consolidated liabilities and mezzanine equity. Dilution is determined by subtracting pro forma as adjusted net tangible book value per ordinary share from the public offering price per ordinary share.

Without taking into account any other changes in such net tangible book value after September 30, 2020, other than to give effect to (i) the conversion of all of our preferred shares into ordinary shares on a one-to-one basis, which will occur automatically immediately prior to the completion of this offering, (ii) the recognition of share-based compensation expense of US\$0.2 million associated with share options for which the service condition was satisfied as of September 30, 2020 and the performance condition will be satisfied upon the completion of this offering, which has been reflected as an increase to additional paid-in capital and accumulated deficit and (iii) our issuance and sale of ADSs offered in this offering at an initial public offering price of US\$ per ADS, after deduction of the underwriting discounts and commissions and estimated offering expenses payable by us, our pro forma as adjusted net tangible book value as of September 30, 2020 would have been approximately US\$ million, or US\$ per ordinary share and US\$ per ADS, to existing shareholders and an immediate dilution in net tangible book value of US\$ per ordinary share, or US\$ per ADS, to purchasers of ADSs in this offering.

The following table illustrates the dilution on a per ordinary share basis at the initial public offering price per ordinary share is US\$ and all ADSs are exchanged for ordinary shares:

	<u>Per Share</u>	<u>Per ADS</u>
Assumed initial public offering price per ordinary share	US\$	US\$
Pro forma net tangible book value per ordinary share after giving effect to the automatic conversion of all of our outstanding preferred shares	US\$	US\$
Pro forma as adjusted net tangible book value per ordinary share to give effect to the automatic conversion of all of our outstanding preferred shares and this offering	US\$	US\$
Amount of dilution in pro forma net tangible book value per share to new investors in the offering	US\$	US\$

The pro forma information discussed above is illustrative only.

The following table summarizes, on a pro forma as adjusted basis as of September 30, 2020, the differences between the existing shareholders and the new investors with respect to the number of ordinary shares (in the form of ADSs or ordinary shares) purchased from us in this offering, the total consideration paid and the average price per ordinary share paid and per ADS at the initial public offering price of US\$ per ADS before deducting estimated underwriting discounts and commissions and estimated offering expenses. The total number of ordinary shares does not include

ordinary shares underlying the ADSs issuable upon the exercise of the over-allotment option granted to the underwriters.

	Ordinary Shares Purchased		Total Consideration		Average Price Per Ordinary Share	Average Price Per ADS
			Amount (in thousands of US\$)			
	Number	Percent	US\$	Percent	US\$	US\$
Existing shareholders		%		%		
New investors		%		%		
Total		100.0%		100.0%		

A US\$1.00 increase (decrease) in the assumed public offering price of US\$ per ADS (the mid-point of the estimated initial public offering price range shown on the front cover page of this prospectus) would increase (decrease) our pro forma as adjusted net tangible book value after giving effect to the offering by US\$ million, the pro forma as adjusted net tangible book value per ordinary share and per ADS after giving effect to this offering by US\$ per ordinary share and US\$ per ADS and the dilution in pro forma as adjusted net tangible book value per ordinary share and per ADS to new investors in this offering by US\$ per ordinary share and US\$ per ADS, assuming no change to the number of ADS offered by us as set forth on the front cover page of this prospectus, and after deducting underwriting discounts and commissions and estimated offering expenses payable by us.

The discussion and tables above also assume no exercise of any awards outstanding as of the date of this prospectus. As of the date of this prospectus, the aggregate number of our ordinary shares underlying our outstanding awards under the 2019 Plan is 3,601,522, excluding awards that were forfeited, cancelled or exercised after the relevant grant dates. We have not granted any awards under the 2021 Plan. To the extent that any outstanding share options are exercised, or new awards are issued under our equity incentive plans, or we issue additional equity or convertible debt securities in the future, there will be further dilution to investors participating in this offering. In addition, we may choose to raise additional capital because of market conditions or strategic considerations, even if we believe that we have sufficient funds for our current or future operating plans. If we raise additional capital through the sale of equity or convertible debt securities, the issuance of these securities could result in further dilution to our shareholders and/or holders of ADSs.

ENFORCEABILITY OF CIVIL LIABILITIES

Cayman Islands

We are incorporated under the laws of the Cayman Islands as an exempted company with limited liability. We enjoy the following benefits:

- political and economic stability;
- an effective judicial system;
- a favorable tax system;
- the absence of exchange control or currency restrictions; and
- the availability of professional and support services.

However, certain disadvantages accompany incorporation in the Cayman Islands. These disadvantages include, but are not limited to, the following:

- the Cayman Islands has a less developed body of securities laws as compared to the United States and these securities laws provide significantly less protection to investors; and
- Cayman Islands companies may not have standing to sue before the federal courts of the United States.

Our constitutional documents do not contain provisions requiring that disputes, including those arising under the securities laws of the United States, between us, our officers, directors and shareholders, be arbitrated.

Substantially all of our operations are conducted in China, and a significant portion of our assets are located in China. A majority of our directors and executive officers are nationals or residents of jurisdictions other than the United States and a substantial portion of their assets are located outside the United States. As a result, it may be difficult for a shareholder to effect service of process within the United States upon these persons, or to enforce against us or them judgments obtained in United States courts, including judgments predicated upon the civil liability provisions of the securities laws of the United States or any state in the United States.

We have appointed Cogency Global Inc., located at 10E 40th Street, 10th Floor, New York, New York 10016, as our agent upon whom process may be served in any action brought against us under the securities laws of the United States.

Walkers (Hong Kong), our counsel as to Cayman Islands law, and Tian Yuan Law Firm, our counsel as to PRC law, have advised us, respectively, that there is uncertainty as to whether the courts of the Cayman Islands and China, respectively, would:

- recognize or enforce judgments of United States courts obtained against us or our directors or officers predicated upon the civil liability provisions of the securities laws of the United States or any state in the United States; or
- entertain original actions brought in each respective jurisdiction against us or our directors or officers predicated upon the securities laws of the United States or any state in the United States.

We have been advised by our Cayman Islands legal counsel, Walkers (Hong Kong), that the courts of the Cayman Islands are unlikely (i) to recognize or enforce against us judgments of courts of the United States predicated upon the civil liability provisions of the securities laws of the United States or any State; and (ii) in original actions brought in the Cayman Islands, to impose liabilities against us predicated upon the civil liability provisions of the securities laws of the United States or any State, so

far as the liabilities imposed by those provisions are penal in nature. In those circumstances, although there is no statutory enforcement in the Cayman Islands of judgments obtained in the United States, the courts of the Cayman Islands, will, at common law, recognize and enforce a foreign money judgment of a foreign court of competent jurisdiction without retrial on the merits of the underlying dispute, based on the principle that a judgment of a competent foreign court imposes upon the judgment debtor an obligation to pay the sum for which judgment has been given provided certain conditions are met. For such a foreign judgment to be enforced in the Cayman Islands, such judgment must be final and conclusive and for a liquidated sum, and must not be in respect of taxes or a fine or penalty nor inconsistent with a Cayman Islands judgment in respect of the same matter, impeachable on the grounds of fraud or obtained in a manner, and be of a kind the enforcement of which is, contrary to natural justice or the public policy of the Cayman Islands (awards of punitive or multiple damages may well be held to be contrary to public policy). A Cayman Islands court may stay enforcement proceedings if concurrent proceedings are being brought elsewhere.

PRC

We have been advised by Tian Yuan Law Firm, our PRC legal counsel, that there is uncertainty as to whether the courts of the PRC would enforce judgments of United States courts or Cayman courts obtained against us or these persons predicated upon the civil liability provisions of the U.S. federal and state securities laws. Tian Yuan Law Firm has further advised us that the recognition and enforcement of foreign judgments are provided for under the PRC Civil Procedures Law. PRC courts may recognize and enforce foreign judgments in accordance with the requirements of the PRC Civil Procedures Law and other applicable laws and regulations based either on treaties between China and the country where the judgment is made or on principles of reciprocity between jurisdictions. China does not have any treaties or other form of reciprocity with the United States or the Cayman Islands that provide for the reciprocal recognition and enforcement of foreign judgments. In addition, according to the PRC Civil Procedures Law, courts in the PRC will not enforce a foreign judgment against us or our directors and officers if they decide that the judgment violates the basic principles of PRC law or national sovereignty, security or public interest. As a result, it is uncertain whether and on what basis a PRC court would enforce a judgment rendered by a court in the United States or in the Cayman Islands. Under the PRC Civil Procedures Law, foreign shareholders may originate actions based on PRC law against a company in the PRC, if they can establish sufficient nexus to the PRC for a PRC court to have jurisdiction, and meet other procedural requirements. However, it would be difficult for foreign shareholders to establish sufficient nexus to the PRC by virtue only of holding the ADSs or ordinary shares.

CORPORATE HISTORY AND STRUCTURE

Corporate History

In February 2011, Adagene Inc. was incorporated under the laws of the Cayman Islands as our offshore holding company.

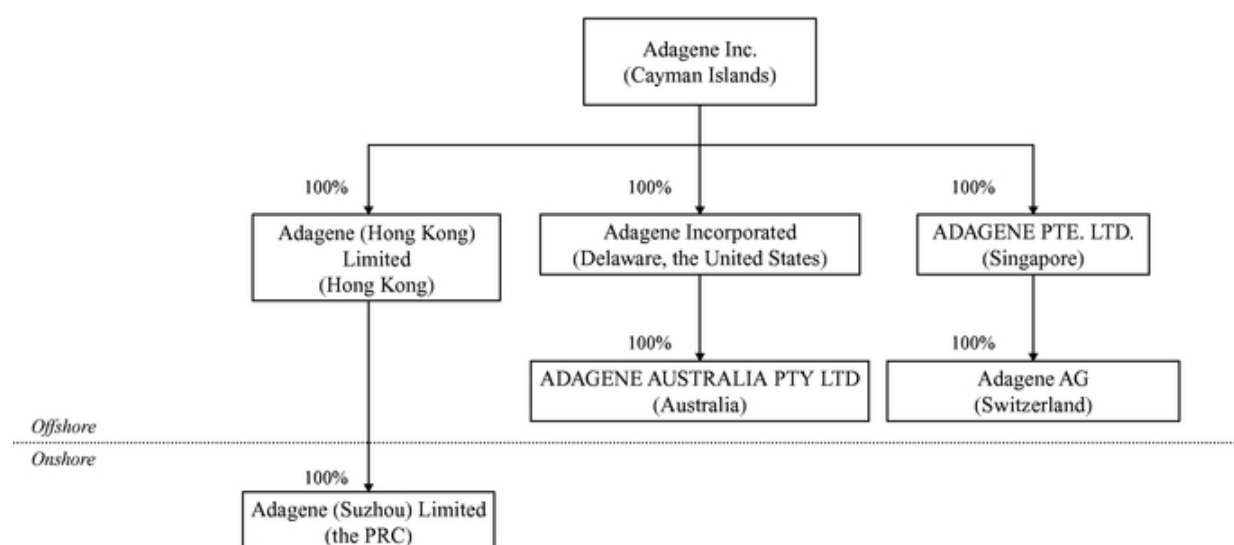
In December 2011, we established Adagene (Hong Kong) Limited, or Adagene Hong Kong, a wholly-owned subsidiary incorporated under the laws of Hong Kong, as our intermediary holding company. In February 2012, Adagene Hong Kong incorporated Adagene (Suzhou) Limited, or Adagene Suzhou, in China, through which we commenced our research and development activities in China.

In September 2017, we established a wholly-owned subsidiary in the state of Delaware, the United States, Adagene Incorporated, to conduct our research and development activities in the United States to facilitate the discovery and development of product candidates and expand our global presence, we have further incorporated several subsidiaries overseas, such as Australia, Singapore and Switzerland.

We are a holding company and do not directly own any substantive business operations in the PRC. We currently focus our business operations within the PRC through Adagene Suzhou. We (Adagene Inc.), however, hold certain intellectual properties and outsource certain research and development activities related to these intellectual properties to our subsidiaries. See "Risk Factors—Risks Related to Doing Business in the PRC."

Corporate Structure

The following diagram illustrates our corporate structure as of the date of this prospectus, including our material subsidiaries:



SELECTED CONSOLIDATED FINANCIAL DATA

The following selected consolidated statements of operations data for the years ended December 31, 2018 and 2019, summary consolidated balance sheet data as of December 31, 2018 and 2019 and summary consolidated cash flow data for the years ended December 31, 2018 and 2019 have been derived from our audited consolidated financial statements included elsewhere in this prospectus. Our consolidated financial statements are prepared in accordance with U.S. GAAP. The following selected consolidated statements of comprehensive loss for the nine months ended September 30, 2019 and 2020, selected consolidated balance sheet data as of September 30, 2020 and selected consolidated cash flows data for the nine months ended September 30, 2019 and 2020 have been derived from our unaudited interim condensed consolidated financial statements included elsewhere in this prospectus and have been prepared on the same basis as our audited consolidated financial statements and include all adjustments, consisting only of normal and recurring adjustments, that we consider necessary for a fair statement of our financial position and operating results for the periods presented. Our historical results are not necessarily indicative of results expected for future periods. You should read this Selected Consolidated Financial Data section together with our consolidated financial statements and the related notes and "Management's Discussion and Analysis of Financial Condition and Results of Operations" included elsewhere in this prospectus.

Selected Consolidated Statements of Comprehensive Loss Data

The following table presents our selected consolidated statements of comprehensive loss data for the years ended December 31, 2018 and 2019 and our selected unaudited interim condensed

consolidated statements of comprehensive loss data for the nine months ended September 30, 2019 and 2020.

	For the Year Ended		For the	
	December 31,		Nine Months	
	2018	2019	2019	2020
	US\$	US\$	US\$	US\$
	(in thousands)			
Revenue:				
Licensing revenue	1,511	480	—	310
Expenses:				
Research and development expenses	(16,081)	(16,212)	(11,518)	(23,568)
Administrative expenses	(2,765)	(3,438)	(2,268)	(7,448)
Total operating expenses	(18,846)	(19,650)	(13,786)	(31,016)
Loss from operations	(17,335)	(19,170)	(13,786)	(30,706)
Interest income	711	923	712	613
Interest expense	(91)	(138)	(102)	(114)
Other income	902	723	217	655
Foreign exchange gain (loss), net	13	22	(68)	(77)
Change in fair value of warrant liabilities	534	1,207	1,207	—
Loss before income tax	(15,266)	(16,432)	(11,818)	(29,629)
Income tax expense	—	—	—	—
Net loss attributable to Adagene Inc.'s shareholders	(15,266)	(16,432)	(11,818)	(29,629)
Other comprehensive income (loss):				
Foreign currency translation adjustments, net of nil tax	(11)	66	281	(215)
Total comprehensive loss attributable to Adagene Inc.'s shareholders	(15,277)	(16,367)	(11,537)	(29,844)
Net loss attributable to Adagene Inc.'s shareholders	(15,266)	(16,432)	(11,818)	(29,629)
Deemed contribution from convertible redeemable preferred shareholders	1,186	—	—	—
Accretion of convertible redeemable preferred shares to redemption value	(223)	(246)	(184)	(186)
Net loss attributable to ordinary shareholders	(14,303)	(16,678)	(12,002)	(29,815)
Weighted average number of ordinary shares used in per share calculation:				
—Basic	15,159	15,178	15,173	15,757
—Diluted	15,159	15,178	15,173	15,757
Net loss per ordinary share				
—Basic	(0.94)	(1.10)	(0.79)	(1.89)
—Diluted	(0.94)	(1.10)	(0.79)	(1.89)

Selected Consolidated Balance Sheet Data

The following table presents our selected consolidated balance sheet data as of December 31, 2018 and 2019 and our selected unaudited interim condensed consolidated balance sheet data as of September 30, 2020:

	As of December 31,		As of
	2018	2019	September 30,
(in USD thousands)			
Current assets:			
Cash and cash equivalents	16,058	92,533	82,895
Short-term investments	33,000	8,000	—
Total current assets	51,817	103,923	86,606
Total assets	54,417	105,889	89,077
Current liabilities:			
Amounts due to related parties	3,674	1,896	2,387
Accruals and other current liabilities	2,574	2,540	2,789
Short-term borrowings	2,331	717	2,203
Total current liabilities	10,346	7,181	10,417
Long-term borrowings	—	1,516	3,135
Total liabilities	10,488	8,697	13,610
Total mezzanine equity	84,955	154,201	154,387
Shareholders' deficit:			
Ordinary shares (par value of US\$0.0001 per share; 500,000,000 and 500,000,000 shares authorized; 15,159,136, 15,193,136 and 16,466,897 shares issued and outstanding as of December 31, 2018 and 2019 and September 30, 2020, respectively)	2	2	2
Subscriptions receivable from shareholders	(197)	(197)	(1,975)
Additional paid-in capital	6,405	6,790	16,500
Accumulated other comprehensive loss	(411)	(345)	(560)
Accumulated deficit	(46,826)	(63,258)	(92,887)
Total shareholders' deficit	(41,027)	(57,009)	(78,921)

Selected Consolidated Cash Flow Data

The following table presents our selected consolidated cash flow data for the years ended December 31, 2018 and 2019 and our selected unaudited interim condensed consolidated cash flow data the nine months ended September 30, 2019 and 2020.

	Year Ended		Nine Months	
	December 31,	December 31,	Ended	Ended
	2018	2019	September 30,	September 30,
(in USD thousands)				
Net cash used in operating activities	(14,265)	(18,154)	(11,791)	(19,988)
Net cash generated (used in) from investing activities	(29,510)	24,856	18,942	7,434
Net cash generated from financing activities	51,058	69,694	16,310	3,097
Effect of exchange rate on cash and cash equivalents	39	78	153	(180)
Net increase in cash and cash equivalents	7,322	76,474	23,614	(9,638)
Cash and cash equivalents at the beginning of year/period	8,736	16,058	16,058	92,533
Cash and cash equivalents at the end of year/period	16,058	92,533	39,673	82,895

MANAGEMENT'S DISCUSSION AND ANALYSIS OF FINANCIAL CONDITION AND RESULTS OF OPERATIONS

You should read the following discussion and analysis of our financial condition and results of operations in conjunction with the section entitled "Selected Consolidated Financial Data" and our consolidated financial statements and the related notes included elsewhere in this prospectus. This discussion contains forward-looking statements that involve risks and uncertainties. Our actual results and the timing of events could differ materially from those anticipated in these forward-looking statements as a result of various factors, including those set forth under "Risk Factors" and elsewhere in this prospectus.

Overview

We are a platform-driven, clinical-stage biopharmaceutical company committed to transforming the discovery and development of novel antibody-based cancer immunotherapies. Our platform is designed to generate therapeutic antibody candidates with unique functional epitopes and species cross-reactivity. These features enable our novel drug discovery strategy to advance from lead identification through vigorous preclinical modeling to biomarker-guided mono- and combination immunotherapy development in clinical settings. We have pioneered a dynamic interface design to harness the conformational diversity of antibodies, which enlarges epitope sampling of a given drug target for differentiated therapeutic antibody development. Our platform is designed to enable the rapid development of precision immunotherapy candidates, through the identification of predictive biomarkers for patient stratification and preselection. We aim to push the boundaries of antibody discovery and engineering through the precise design, construction, and selection of antibody product candidates intractable to traditional antibody technology.

We have developed our proprietary Dynamic Precision Library, or DPL, platform to explore the dynamic conformational diversity of protein sequences, and the flexible binding sites of antibody sequences in particular, as a new paradigm for antibody drug discovery. Our DPL platform samples a potentially infinite number of dynamic binding interface structures arising from the conformational diversity of a finite number of antibody amino acid sequences, allowing us to exponentially expand the universe of candidate antibody binding sites far beyond conventional natural or synthetic antibody repertoires. By exploiting conformational diversity through the combination of our proprietary computational algorithms and artificial intelligence, we have designed and precisely constructed approximately one trillion (10^{12}) antibody sequences in our DPL. These antibodies feature broad epitope (the portion of an antigen that are recognized by an antibody) coverage and robust chemistry, manufacturing, and control, or CMC, attributes. Our DPL platform is designed to enable high fidelity translation from preclinical to clinical studies by identifying antibodies well suited for broad species cross-reactivity against the transiently accessible epitopes of challenging targets.

By leveraging our proprietary DPL platform, we have developed a robust pipeline of innovative product candidates in various stages of development, ranging from research and discovery to preclinical and clinical development. Our highly differentiated clinical-stage pipeline consists of ADG106, ADG126 and ADG116. We also have a robust preclinical pipeline in various stages of development.

Since our inception in 2011, our operations have focused on organizing and staffing our company, conducting preclinical studies and clinical trials, business planning, establishing our intellectual property portfolio and raising capital. We do not have any product candidates approved for sale and have not generated any revenue from product sales. We have financed operations mainly through the private placements of our preferred shares. From November 2014 to December 2019, we raised an aggregate of US\$155.5 million of gross proceeds from issuance of our preferred shares.

Since inception, we have incurred significant operating losses. Our net losses were US\$15.3 million and US\$16.4 million for the years ended December 31, 2018 and 2019, respectively. For the nine months ended September 30, 2020, our net loss was US\$29.6 million. As of September 30, 2020, we

had accumulated deficit of US\$92.9 million. We expect to continue to incur significant expenses and operating losses for the foreseeable future. We anticipate that our expenses will increase significantly in connection with our ongoing activities, as we:

- continue advancement of and investment in our proprietary DPL platform;
- advance the development of ADG106, ADG126, ADG116 and other preclinical drug candidates;
- continue our ongoing and planned research and development of other lead product candidates;
- discover and develop additional antibody product candidates and further expand our preclinical and clinical product pipeline;
- maintain, expand and protect our intellectual property portfolio;
- expand our collaborations with contract manufacturing organizations and contract research organizations;
- seek regulatory approvals for any product candidates that successfully complete clinical trials;
- establish sales and marketing teams and distribution network to commercialize any product candidate for which we may obtain regulatory approval;
- attract, hire and retain additional clinical, scientific, management and administrative personnel;
- expand our operations globally; and
- incur additional costs associated with operating as a public company upon the completion of this offering.

Beginning in January 2020, the emergence and wide spread of COVID-19 has resulted in quarantines, travel restrictions, and the temporary closure of stores and facilities in the United States and China and elsewhere. Substantially all of our operating and workforce are based in the United States and China. Consequently, the COVID-19 outbreak could potentially delay patient's access to hospital and the progress of our clinical trials, including patient enrollment, which may adversely affect our business operations, financial condition and operating results for 2020. The extent of the impact of the COVID-19 pandemic on our business, operations and regulatory and commercialization timelines will depend on certain developments, including the duration and spread of the outbreak and its impact on clinical trials, regulatory authorities and our key scientific and management personnel as well as its impact on our partners, laboratory sites, and other third parties with whom we collaborate. See "Risk Factors—Risks relating to obtaining regulatory approval of our drug candidates—The COVID-19 pandemic could adversely impact our business, including our clinical trials." We will continue to actively monitor the rapidly evolving situation related to COVID-19 and may take further actions that alter our business operations, including those that may be required by government authorities, or that we determine are in the best interests of our employees, partners and shareholders. At this point, the extent to which the COVID-19 pandemic may impact our business, operations and regulatory and commercialization timelines remains uncertain.

Key Components of Results of Operations

Revenue

Licensing revenue. Our licensing revenue is currently comprised of royalties, milestone payments, license fees and reimbursement income. Our licensing and collaboration revenue for the years ended December 31, 2018 and 2019 and the nine months ended September 30, 2020 was primarily derived from granting licenses to use and otherwise exploit certain of our intellectual properties. To date, we have not generated any revenue from the sale of products and do not expect to generate any revenue from product sales in the near future.

Expenses

Research and Development Expenses. Our research and development expenses consist principally of (1) payroll and other related costs of personnel engaged in research and development activities, (2) costs related to pre-clinical testing of our technologies under development and clinical trials such as payments to contract research organizations, or CRO, and contract manufacturing organization, or CMO, investigators and clinical trial sites that conduct the clinical studies; (3) costs to develop the product candidates, including raw materials and supplies, product testing, depreciation and amortization, and facility related expenses, (4) other research and development expenses.

For the years ended December 31, 2018 and 2019 and the nine months ended September 30, 2019 and 2020, our research and development expenses were US\$16.1 million, US\$16.2 million, US\$11.5 million and US\$23.6 million, respectively. The increase is primarily due to increased share-based compensation and expansion of staff and an increase in contract manufacturing costs in light of the progression of the programs.

Our research and development expenses may vary substantially from period to period according to the status of our research and development activities. The timing of expenses is impacted by the commencement of clinical trials and enrollment of patients in clinical trials. Research and development expenses are expected to increase as we advance the clinical development of ADG106, ADG126 and ADG116, and further advance the research and development of our other product candidates. The successful development of our product candidates is uncertain.

The following table summarizes our research and development expenses for our clinical-stage product candidates, preclinical product candidates and research pipeline for the years ended December 31, 2018 and 2019 and the nine months ended September 30, 2019 and 2020, respectively.

	For the Year Ended December 31,		For the Nine Months Ended September 30,	
	2018	2019	2019	2020
	US\$	US\$	US\$	US\$
	(in thousands)			
ADG106	5,145	5,742	3,609	6,528
ADG126	1,210	2,326	1,169	11,315
ADG116	6,812	5,329	4,630	2,537
Preclinical product candidates, research pipeline and others	2,914	2,814	2,110	3,188
Total	16,081	16,212	11,518	23,568

At this time, we cannot reasonably estimate the nature, timing and estimated costs of the efforts that will be necessary to complete the development of, or the period, if any, in which material net cash inflows may commence from, any of our product candidates. This is due to numerous risks and uncertainties associated with developing drugs, including the uncertainty of:

- the scope, rate of progress, results and cost of our clinical trials, preclinical studies and other related activities;
- the cost of manufacturing clinical supplies, and establishing commercial supplies, of any product candidates;
- the number and characteristics of product candidates that we pursue;
- the cost, timing and outcomes of regulatory approvals;
- the cost and timing of establishing sales, marketing and distribution capabilities; and

- the terms and timing of any collaboration, licensing or other arrangements that we may establish, including any required milestone and royalty payments thereunder.

A change in the outcome of any of these variables with respect to the development of our product candidates or any other current or future product candidates could mean a significant change in the costs and timing associated with the development of such product candidate. For example, if we are required to conduct additional clinical trials or other testing of any of our product candidates beyond those that are contemplated or if we experience significant delays in enrollment in any clinical trials, we could incur significant additional costs and the clinical development timeline for our product candidates may be delayed.

Administrative expenses. Our administrative expenses consist primarily of wages, salaries and benefits for personnel other than research and development staff. We expect our administrative expenses to increase in absolute amount in the foreseeable future as we incur additional costs as a result of operating as a public company and as we advance our product candidates through clinical development, which will also increase our general and administrative expenses.

Interest income

Interest income consists primarily of interest income derived from our term deposit.

Other income

Other income primarily includes government subsidies that Adagene Suzhou received from local government in the PRC. The receipt of such government subsidies is not dependent on our performance of any obligations.

Taxation

Cayman Islands

We were incorporated in the Cayman Islands. Under the current laws of the Cayman Islands, we are not subject to income, corporation or capital gains tax in the Cayman Islands. In addition, our payment of dividends, if any, is not subject to withholding tax in the Cayman Islands.

Hong Kong

Our subsidiaries in Hong Kong, including Adagene (Hong Kong) Limited, our wholly-owned subsidiary, are subject to Hong Kong profits tax on their activities conducted in Hong Kong at a uniform tax rate of 16.5%. Under Hong Kong tax law, our subsidiary in Hong Kong is exempted from income tax on their foreign-derived income and there is no withholding tax in Hong Kong on remittance of dividends. No provision for Hong Kong profits tax was made as we had no estimated assessable profit that was subject to Hong Kong profits tax during fiscal years 2019 or 2020.

PRC

Our subsidiaries in China are companies incorporated under PRC law and, as such, are subject to PRC enterprise income tax on their taxable income in accordance with the relevant PRC income tax laws. Pursuant to the PRC Enterprise Income Tax Law, or EIT Law, which became effective on January 1, 2008, a uniform 25% enterprise income tax rate is generally applicable to both foreign-invested enterprises and domestic enterprises, except where a special preferential rate applies. In accordance with the implementation rules of EIT Law, a qualified Technology Advanced Service Enterprises, or TASE, is eligible for a preferential tax rate of 15%. The TASE certificate is effective for three years. An entity must file required supporting documents with the tax authority and ensure fulfillment of the relevant TASE criteria before using the preferential rate. An entity could apply for

the TASE certificate every year. Adagene Suzhou was first recognized as a qualified TASE in March 2015 and renewed in December 2018. Adagene Suzhou can enjoy the preferential tax rate of 15% from 2015 to 2021. In addition, the research and development expenses of Adagene Suzhou are subject to a 75% super-deduction for the income tax. The enterprise income tax is calculated based on the entity's global income as determined under PRC tax laws and accounting standards.

We are subject to VAT at a rate of 3%, 6%, or 13% on the services we provide and related surcharges. We are also subject to surcharges on VAT payments in accordance with PRC law.

As a Cayman Islands holding company, we may receive dividends from Adagene Suzhou. The PRC EIT Law and its implementing rules provide that dividends paid by a PRC entity to a nonresident enterprise for income tax purposes is subject to PRC withholding tax at a rate of 10%, subject to reduction by an applicable tax treaty with China. Pursuant to the Arrangement between Mainland China and the Hong Kong Special Administrative Region for the Avoidance of Double Taxation and Tax Evasion on Income, the withholding tax rate in respect to the payment of dividends by a PRC enterprise to a Hong Kong enterprise may be reduced to 5% from a standard rate of 10% if the Hong Kong enterprise directly holds at least 25% of the PRC enterprise. Pursuant to the Notice of the State Administration of Taxation on the Issues concerning the Application of the Dividend Clauses of Tax Agreements, or SAT Circular 81, a Hong Kong resident enterprise must meet the following conditions, among others, in order to apply the reduced withholding tax rate: (i) it must be a company; (ii) it must directly own the required percentage of equity interests and voting rights in the PRC resident enterprise; and (iii) it must have directly owned such required percentage in the PRC resident enterprise throughout the 12 months prior to receiving the dividends. In August 2015, the State Administration of Taxation promulgated the Administrative Measures for Nonresident Taxpayers to Enjoy Treatment under Tax Treaties, or SAT Circular 35, which became effective on January 1, 2020, 2015. SAT Circular 35 provides that nonresident enterprises are not required to obtain pre-approval from the relevant tax authority in order to enjoy the reduced withholding tax. Instead, nonresident enterprises and their withholding agents may, by self-assessment and on confirmation that the prescribed criteria to enjoy the tax treaty benefits are met, directly apply the reduced withholding tax rate, and file necessary forms and supporting documents when performing tax filings, which will be subject to post-tax filing examinations by the relevant tax authorities. Accordingly, Adagene (Hong Kong) Limited may be able to benefit from the 5% withholding tax rate for the dividends it receives from its PRC subsidiaries, if it satisfies the conditions prescribed under SAT Circular 81 and other relevant tax rules and regulations. However, according to SAT Circular 81 and SAT Circular 35, if the relevant tax authorities consider the transactions or arrangements we have are for the primary purpose of enjoying a favorable tax treatment, the relevant tax authorities may adjust the favorable withholding tax in the future.

If our holding company in the Cayman Islands or any of our subsidiaries outside of China were deemed to be a "resident enterprise" under the PRC EIT Law, it would be subject to enterprise income tax on its worldwide income at a rate of 25%. See "Risk Factors—Risks Related to Doing Business in China—If we are classified as a PRC resident enterprise for PRC income tax purposes, such classification could result in unfavorable tax consequences to us and our non-PRC shareholders or ADS holders."

Results of Operations

The following table summarizes our consolidated results of operations for the periods presented. This information should be read together with our consolidated financial statements and related notes

included elsewhere in this prospectus. The operating results in any period are not necessarily indicative of the results that may be expected for any future period.

	For the Year Ended		For the Nine Months	
	December 31,		Ended	
	2018	2019	2019	2020
	US\$	US\$	US\$	US\$
	(in thousands)			
Revenue:				
Licensing revenue	1,511	480	—	310
Expenses:				
Research and development expenses	(16,081)	(16,212)	(11,518)	(23,568)
Administrative expenses	(2,765)	(3,438)	(2,268)	(7,448)
Total operating expenses	(18,846)	(19,650)	(13,786)	(31,016)
Loss from operations	(17,335)	(19,170)	(13,786)	(30,706)
Interest income	711	923	712	613
Interest expense	(91)	(138)	(102)	(114)
Other income	902	723	217	655
Foreign exchange gain (loss), net	13	22	(68)	(77)
Change in fair value of warrant liabilities	534	1,207	1,207	—
Loss before income tax	(15,266)	(16,432)	(11,818)	(29,629)
Income tax expense	—	—	—	—
Net loss attributable to Adagene Inc.'s shareholders	(15,266)	(16,432)	(11,818)	(29,629)
Other comprehensive income (loss):				
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Total comprehensive loss attributable to Adagene Inc.'s shareholders	(15,277)	(16,367)	(11,537)	(29,844)
Net loss attributable to Adagene Inc.'s shareholders	(15,266)	(16,432)	(11,818)	(29,629)
Deemed contribution from convertible redeemable preferred shareholders	1,186	—	—	—
Accretion of convertible redeemable preferred shares to redemption value	(223)	(246)	(184)	(186)
Net loss attributable to ordinary shareholders	(14,303)	(16,678)	(12,002)	(29,815)
Weighted average number of ordinary shares used in per share calculation:				
—Basic	15,159	15,178	15,173	15,757
—Diluted	15,159	15,178	15,173	15,757
Net loss per ordinary share				
—Basic	(0.94)	(1.10)	(0.79)	(1.89)
—Diluted	(0.94)	(1.10)	(0.79)	(1.89)

Nine Months Ended September 30, 2020 Compared to Nine Months Ended September 30, 2019

Licensing revenue

Our licensing revenue increased from nil for the nine months ended September 30, 2019 to US\$0.3 million for the nine months ended September 30, 2020. Our licensing revenue generated in the nine months ended September 30, 2020 are a payment from Celgene (now Bristol-Myers Squibb), in connection with providing certain sequences related to antibody discovery services to Celgene.

Research and development expenses

The following table sets forth a breakdown of the major components of our research and development expenses in absolute amounts and as a percentage of our total research and development expenses for the periods indicated:

	For the Nine Months Ended September 30			
	2019		2020	
	US\$	%	US\$	%
Research and development expenses	(in thousands, except percentages)			
Payroll and other related costs of personnel	3,349	29.1%	9,847	41.8%
Costs related to clinical trials	1,869	16.2%	3,110	13.2%
Costs related to preclinical testing	3,780	32.8%	7,143	30.3%
Costs to develop the product candidates	1,173	10.2%	1,414	6.0%
Other research and development expenses	1,347	11.7%	2,054	8.7%
Total	11,518	100.0%	23,568	100.0%

Our research and development expenses increased by 104.6% from US\$11.5 million for the nine months ended September 30, 2019 to US\$23.6 million for the nine months ended September 30, 2020, primarily attributable to (i) an increase of US\$6.5 million in payroll and other related costs of personnel primarily due to increased share-based compensation and expansion of staff and (ii) an increase of US\$4.6 million in costs related to preclinical testing and clinical trials primarily due to increased contract manufacturing costs in light of the progression of the programs.

Administrative expenses

Our administrative expenses increased by 228.4% from US\$2.3 million for the nine months ended September 30, 2019 to US\$7.4 million for the nine months ended September 30, 2020, primarily attributable to (i) an increase of US\$1.5 million in employee compensations due to an increase in average payroll and an increase in the number of employees and (ii) an increase of US\$2.9 million in the share-based compensation expenses.

Loss from operations

As a result of the foregoing, our loss from operations increased by 122.7% from approximately US\$13.8 million in the nine months ended September 30, 2019 to approximately US\$30.7 million in the nine months ended September 30, 2020.

Interest income

Our interest income was US\$0.6 million for the nine months ended September 30, 2020, as compared to US\$0.7 million for the nine months ended September 30, 2019, representing a decrease of US\$0.1 million. This decrease was primarily attributable to a decrease of cash balance and amount of short-term investments.

Other income

Our other income increased from US\$0.2 million for the nine months ended September 30, 2019 to US\$0.7 million for the nine months ended September 30, 2020, primarily attributable to (i) an increase in government subsidies received by Adagene Suzhou to support our ongoing operations in Jiangsu Province during the nine months ended September 30, 2020 and (ii) an exclusivity payment from ADC Therapeutics pursuant to our collaboration arrangement with ADC Therapeutics, which is not related to our major operation activities.

Foreign exchange loss, net

Our net foreign exchange loss decreased by 13.8% from US\$67.7 thousand for the nine months ended September 30, 2019 to US\$77.0 thousand for the nine months ended September 30, 2020. This loss of foreign exchange was primarily attributable to the weakening of Renminbi against U.S. dollars which negatively impacted our Renminbi denominated portfolio held by Adagene Suzhou. This improvement was primarily due to the positive effect of the fluctuation of Reminbi against U.S. dollars.

Change in fair value of warrant liabilities

We recorded a gain from change in the fair value of warrant liabilities of US\$1.2 million for the nine months ended September 30, 2019 and of nil for the nine months ended September 30, 2020. The change was primarily because the warrants to subscribe Series C-2 Preferred Shares expired on April 1, 2019.

Net loss attributable to Adagene Inc.'s shareholders

As a result of the foregoing, our net loss for the period increased by 150.7% from US\$11.8 million for the nine months ended September 30, 2019 to US\$29.6 million for the nine months ended September 30, 2020.

Year Ended December 31, 2019 Compared to Year Ended December 31, 2018**Licensing revenue**

Our licensing revenue decreased by 68.2% from US\$1.5 million in 2018 to US\$0.5 million in 2019. For the year ended December 31, 2018, we received a payment of US\$1.5 million from Guilin Sanjin Group Co., Ltd., or Sanjin, pursuant to our out-licensing arrangement. The relevant rights under the license were granted to Sanjin in 2018. For the year ended December 31, 2019, we recognized a US\$0.5 million service fee received from Celgene in connection with providing certain sequences related to antibody discovery services.

Research and development expenses

The following table sets forth a breakdown of the major components of our research and development expenses in absolute amounts and as a percentage of our total research and development expenses for the periods indicated:

	For the year ended December 31,			
	2018		2019	
	US\$	%	US\$	%
	(in thousands, except percentages)			
Research and development expenses				
Payroll and other related costs of personnel	4,039	25.1%	4,880	30.1%
Costs related to clinical trials	997	6.1%	3,361	20.7%
Costs related to preclinical testing	8,020	49.9%	4,726	29.2%
Costs to develop the product candidates	1,790	11.1%	1,599	9.9%
Other research and development expenses	1,255	7.8%	1,645	10.1%
Total	16,081	100%	16,212	100%

Our research and development expenses remained relatively stable and increased by 0.8% from US\$16.1 million in 2018 to US\$16.2 million in 2019, which was mainly attributable to an increase of clinical costs associated with the Phase I clinical trial of ADG106 and ADG116, partially offset by a decrease of contract manufacturing costs, due to ADG106's advance from preclinical development stage into clinical trial stage.

Administrative expenses

Our administrative expenses increased by 24.3% from US\$2.8 million in 2018 to US\$3.4 million in 2019. The increase was primarily due to the increase in the employee compensation due to a rise in average payroll and an increase in the number of employees.

Loss from operations

As a result of the foregoing, our loss from operations increased by 10.6% from approximately US\$17.3 million in 2018 to approximately US\$19.2 million in 2019.

Interest income

Our interest income was US\$0.9 million in 2019, as compared to US\$0.7 million in 2018, representing an increase of US\$0.2 million. This increase was primarily attributable to the increase of the balance of fixed deposits.

Other income

Our other income was US\$0.7 million in 2019, as compared to US\$0.9 million in 2018, representing a decrease of US\$0.2 million. This decrease was primarily due to the decrease in government subsidies during the year of 2019.

Foreign exchange gain, net

Our net foreign exchange gain increased by 72.2% from US\$12.7 thousand to US\$21.9 thousand, primarily attributable to further appreciation of Renminbi against U.S. dollars in 2019 following 2018.

Change in fair value of warrant liabilities

We recorded a gain from change in the fair value of warrant liabilities of US\$0.5 million and US\$1.2 million in 2018 and 2019, respectively. The change in the fair value of warrant liabilities was primarily attributable to the change in fair value of warrants due to the passage of time as approaching to the expiration date of the warrants.

Net Loss attributable to Adagene Inc.'s shareholders

As a result of the foregoing, our net loss increased by 7.6% from US\$15.3 million in 2018 to US\$16.4 million in 2019.

Liquidity and Capital Resources

Since the inception, we have incurred net losses and negative cash flow from our operations. We do not expect to generate any revenue from product sales unless and until we obtain regulatory approval and subsequently commercialize one of our current or future drug candidates. We expect to

incur additional operating losses in the near future and our operating expenses will increase as we continue to expand our research and development capabilities, invest in preclinical tests and clinical trials and increase our efforts in obtaining regulatory approvals. In addition, subject to obtaining regulatory approval of any of our product candidates, we expect to incur significant commercialization expenses for product sales, marketing and manufacturing. Moreover, following the completion of this offering, we expect to incur additional costs associated with operating as a public company, including expenses related to legal, accounting, regulatory, maintaining compliance with exchange listing and SEC requirements, director and officer insurance premiums, and investor relations.

Historically, we have financed our operations principally through proceeds from the issuance and sale of preferred shares in private placement transactions. For more information of our equity financing, see "Description of Share Capital—History of Securities Issuances." As of September 30, 2020, we had US\$82.9 million in cash and cash equivalents. Our cash and cash equivalents consist primarily of bank deposits and highly liquid investments, which have original maturities of three months or less when purchased. In February 2019, we entered into a three-year bank facility agreement with Shanghai Pudong Development Bank, pursuant to which we are entitled to borrow up to RMB7.5 million at a fixed annual interest rate of 5.46%, and as of September 30, 2020, the outstanding balance due under such facility was RMB5.9 million. In June 2019, we entered into another three-year bank facility agreement with the same bank, pursuant to which we are entitled to borrow up to RMB6.0 million at a fixed annual interest rate of 5.23%, and as of September 30, 2020, the outstanding balance due under such facility was RMB5.4 million. In addition, in June 2020, we borrowed a loan with amount of RMB10.0 million from Agricultural Bank of China Limited for a term of one year and at the interest rate of 4.20% per annum. In September 2020, we borrowed a one-year term loan with amount of RMB5.0 million from Bank of Ningbo Co., Ltd. at the interest rate of 4.20% per annum. In September 2020, we also entered into a three-year bank facility agreement with Shanghai Pudong Development Bank, pursuant to which we are entitled to borrow up to RMB16.5 million at a fixed annual interest rate of 4.27%, as of September 30, 2020, the outstanding balance due under such facility was RMB16.5 million. In November 2020, we entered into two one-year term loans with amount of RMB5.0 million each from China Merchant Bank Co., Ltd. at the interest rate of 4.10% per annum, as of the date of this prospectus, the outstanding balance due under such facilities was RMB10.0 million.

We intend to finance our future working capital requirements and capital expenditures primarily from funds raised from financing activities, including the net proceeds we will receive from this offering.

Based on our current operating plan, we believe that our current cash and cash equivalents and proceeds from this offering will be sufficient to meet our current and anticipated working capital requirements and capital expenditures for at least the next 12 months. In that time, we expect that our expenses will increase substantially as we fund new and ongoing research and development activities and working capital needs. The assumptions on which our estimates are based may prove to be wrong, and we may use our available capital resources sooner than we currently expect. Because of the numerous risks and uncertainties associated with the development and commercialization of our product candidates, we are unable to estimate the amounts of increased capital outlays and operating expenditures necessary to complete the development and commercialization of our product candidates.

In utilizing the proceeds that we expect to receive from this offering, we may make capital contributions to our subsidiaries in PRC, the United States, Australia, Hong Kong, Singapore and Switzerland, acquire or establish new subsidiaries, or give loans to our subsidiaries. However, uses of the proceeds by our PRC subsidiary are subject to PRC regulations. See "Risk Factors—Risks Relating to Doing Business in China—PRC regulation of loans to and direct investment in PRC entities by offshore holding companies and governmental control of currency conversion may delay or prevent us

from using the proceeds of this offering to make loans or additional capital contributions to our PRC subsidiary, which could materially and adversely affect our liquidity and our ability to fund and expand our business." and "Use of Proceeds."

Our operations are primarily based in China. A significant portion of our transactions are settled in Renminbi and our financial statements are presented in U.S. dollars. Under existing PRC foreign exchange regulations, Renminbi may be converted into foreign currencies for current account items, including profit distributions, interest payments and trade- and service-related foreign exchange transactions, without prior SAFE approval as long as certain routine procedural requirements are fulfilled. Our PRC subsidiary is allowed to pay dividends in foreign currencies to us without prior SAFE approval by following certain routine procedural requirements. However, approval from or registration with competent government authorities is required where the Renminbi is to be converted into foreign currency and remitted out of China to pay capital expenses such as the repayment of loans denominated in foreign currencies. The PRC government may at its discretion restrict access to foreign currencies for current account transactions in the future.

The following table presents our selected consolidated cash flow data for the years ended December 31, 2018 and 2019 and the nine months ended September 30, 2019 and 2020.

	For the Year Ended December 31,		For the Nine Months Ended September 30,	
	2018	2019	2019	2020
	US\$	US\$	US\$	US\$
	(in thousands)			
Net cash used in operating activities	(14,265)	(18,154)	(11,791)	(19,988)
Net cash (used in) generated from investing activities	(29,510)	24,856	18,942	7,434
Net cash generated from financing activities	51,058	69,694	16,310	3,097
Effect of exchange rate on cash and cash equivalents	39	78	153	(180)
Net increase in cash and cash equivalents	7,322	76,474	23,614	(9,638)
Cash and cash equivalents at the beginning of year/period	8,736	16,058	16,058	92,533
Cash and cash equivalents at the end of year/period	16,058	92,533	39,673	82,895

Operating activities

Net cash used in operating activities was US\$20.0 million in the nine months ended September 30, 2020. The difference between our net loss of US\$29.6 million and the net cash used in operating activities was mainly due to (i) an increase in prepayments and other current assets of US\$1.8 million due to our prepaid service fees to our vendor, and (ii) a decrease in contract liabilities of US\$0.1 million, partially offset by (i) an increase in share-based compensation of US\$8.0 million, (ii) an increase in accounts payable of US\$0.5 million (iii) a decrease in accounts receivable of US\$0.5 million and (iv) an increase in accruals and other current liabilities of US\$0.2 million.

Net cash used in operating activities was US\$18.2 million in 2019. The difference between our net loss of US\$16.4 million and the net cash used in operating activities was mainly due to (i) a decrease in amount due to related parties of US\$1.8 million due to our payment of services fees to Wuxi Biologics, (ii) an increase in amount due from related parties of US\$0.7 million, and (iii) change in fair value of warrant liabilities of US\$1.2 million, partially offset by depreciation and amortization of US\$0.8 million.

Net cash used in operating activities was US\$14.3 million in 2018. The difference between our net loss of US\$15.3 million and the net cash used in operating activities was mainly due to (i) an increase in prepayments and other current assets of US\$1.3 million, (ii) a decrease in accruals and other current liabilities of US\$0.9 million, and (iii) a decrease in fair value of warrant liabilities of US\$0.5 million,

partially offset by (i) an increase in amount due to related parties of US\$2.1 million, (ii) an increase in accounts payable of US\$0.4 million, and (iii) depreciation and amortization of US\$0.9 million.

Investing activities

Net cash generated from investing activities was US\$7.4 million in the nine months ended September 30, 2020, which was primarily attributable to (i) withdrawal of short-term investments of US\$8.0 million and (ii) proceeds from disposal of property, equipment and software of US\$0.01 million, partially offset by purchase of property, equipment and software of US\$0.6 million.

Net cash generated from investing activities was US\$24.9 million in 2019, which was primarily attributable to withdrawal of short-term investments of US\$44.0 million, partially offset by placement of short-term investments of US\$19.0 million.

Net cash used in investing activities was US\$29.5 million in 2018, which was primarily attributable to placement of short-term investments of US\$58.0 million, partially offset by withdrawal of short-term investments of US\$29.0 million.

Financing activities

Net cash generated from financing activities was US\$3.1 million in the nine months ended September 30, 2020, which was mainly attributable to proceeds from borrowings of US\$4.5 million, partially offset by repayment of borrowings of US\$0.9 million.

Net cash generated from financing activities was US\$69.7 million in 2019, which was mainly attributable to proceeds from issuance of convertible redeemable preferred shares and warrants of US\$69.0 million, partially offset by repayment of borrowings of US\$2.4 million.

Net cash generated from financing activities was US\$51.1 million in 2018, which was mainly attributable to proceeds from issuance of convertible redeemable preferred shares and warrants of US\$50.0 million, partially offset by repayment of borrowings of US\$1.4 million.

Capital Expenditures

Our capital expenditures are incurred primarily in connection with research and development equipment. Our capital expenditures were US\$0.5 million, US\$0.2 million and US\$0.6 million, respectively, in 2018, 2019 and the nine months ended September 30, 2020. We intend to fund our future capital expenditures with our existing cash balance and proceeds from this offering. We will continue to make capital expenditures to meet the expected growth of our business.

Commitments

The following table sets forth our commitments as of December 31, 2019.

	Payments Due by Years Ending				
	Total	Less than 1 year	1 - 3 years (in thousands)	3 - 5 years	More than 5 years
Short-term and long-term borrowings	2,555	1,039	1,516	—	—
Operating Leases ⁽¹⁾	257	174	83	—	—
Total commitments	2,812	1,213	1,599	—	—

(1) Operating leases relate to certain office buildings under non-cancellable operating lease agreements.

Holding Company Structure

Adagene Inc. is a holding company with no material operations of its own. Adagene Inc. holds certain intellectual properties and outsources certain research and development activities related to these intellectual properties to its subsidiaries. We conduct our operations primarily through our subsidiaries. As a result, our ability to pay dividends depends upon dividends paid by our subsidiaries. If our subsidiaries or any newly formed subsidiaries incur debt on their own behalf in the future, the instruments governing their debt may restrict their ability to pay dividends to us.

In addition, our subsidiary in China is permitted to pay dividends to us only out of their retained earnings, if any, as determined in accordance with the Accounting Standards for Business Enterprise as promulgated by the Ministry of Finance of the PRC, or PRC GAAP. Pursuant to the law applicable to China's foreign investment enterprise, our subsidiaries that are foreign investment enterprises in the PRC have to make appropriation from their after-tax profit, as determined under PRC GAAP, to reserve funds including (i) general reserve fund, (ii) enterprise expansion fund and (iii) staff bonus and welfare fund. The appropriation to the general reserve fund must be at least 10% of the after-tax profits calculated in accordance with PRC GAAP. Appropriation is not required if the reserve fund has reached 50% of the registered capital of our subsidiary. Appropriation to the other two reserve funds are at our subsidiary's discretion.

As an offshore holding company, we are permitted under PRC laws and regulations to provide funding from the proceeds of our offshore fund raising activities to our PRC subsidiaries only through loans or capital contributions, and to our consolidated affiliated entity only through loans, in each case subject to the satisfaction of the applicable government registration and approval requirements. See "Risk Factors—Risks Related to Doing Business in China—PRC regulation of loans to and direct investment in PRC entities by offshore holding companies and governmental control of currency conversion may delay or prevent us from using the proceeds of this offering to make loans or additional capital contributions to our PRC subsidiary, which could materially and adversely affect our liquidity and our ability to fund and expand our business." As a result, there is uncertainty with respect to our ability to provide prompt financial support to our PRC subsidiaries when needed.

Off-Balance Sheet Commitments and Arrangements

We have not entered into any financial guarantees or other commitments to guarantee the payment obligations of any third parties. We have not entered into any derivative contracts that are indexed to our shares and classified as shareholder's equity or that are not reflected in our consolidated financial statements. Furthermore, we do not have any retained or contingent interest in assets transferred to an unconsolidated entity that serves as credit, liquidity or market risk support to such entity. We do not have any variable interest in any unconsolidated entity that provides financing, liquidity, market risk or credit support to us or engages in leasing, hedging or product development services with us.

Critical Accounting Policies, Judgments and Estimates

Basis of presentation

Our consolidated financial statements have been prepared in accordance with U.S. generally accepted accounting principles, or U.S. GAAP.

Principles of Consolidation

Our consolidated financial statements include the financial statements of Adagene Inc. and its subsidiaries. All significant intercompany balances and transactions have been eliminated upon consolidation.

Use of estimates

The preparation of consolidated financial statements in conformity with U.S. GAAP requires management to make estimates and assumptions that affect the reported amounts of assets and liabilities, disclosures of contingent assets and liabilities at the balance sheet dates and the reported amounts of expenses during the reporting periods. Significant estimates and assumptions reflected in our consolidated financial statements include, but are not limited to, the useful lives and impairment of long-lived assets, tax valuation allowance, share-based compensation expenses and the fair value of warrant liabilities. Management bases the estimates on historical experience and various other assumptions that are believed to be reasonable, the results of which form the basis for making judgments about the carrying values of assets and liabilities. Actual results could materially differ from those estimates.

Foreign currency translation

The functional currency of Adagene Inc., Adagene (Hong Kong) Limited and Adagene Incorporated is the U.S. dollar, or US\$. The functional currency of our PRC subsidiary is Renminbi, or RMB. The functional currency of our Australia subsidiary is Australian dollar, or AU\$. The determination of the respective functional currency is based on the criteria stated in Accounting Standard Codification, or ASC, 830, Foreign Currency Matters. We use US\$ as our reporting currency. The financial statements of our PRC subsidiary and Australia subsidiaries are translated from the functional currency to the reporting currency.

Transactions denominated in foreign currencies are re-measured into the functional currency at the exchange rates quoted by the People's Bank of China, or the PBOC, prevailing on the transaction dates. Monetary assets and liabilities denominated in foreign currencies are re-measured at the exchange rates prevailing at the balance sheet date. Non-monetary items that are measured in terms of historical costs in foreign currency are re-measured using the exchange rates at the dates of the initial transactions. Exchange gains and losses are included in the consolidated statements of comprehensive loss.

Assets and liabilities are translated at the exchange rates at the balance sheet date, equity accounts are translated at historical exchange rates and revenues, expenses, gains and losses are translated using the average rate for the year. Translation adjustments are reported as accumulated comprehensive loss and are shown as a separate component of other comprehensive loss in the consolidated statements of comprehensive loss.

Revenue recognition

At contract inception of collaboration and out-licensing arrangement, we analyze the arrangements to assess whether they are within the scope of ASC 808, *Collaborative Arrangements* or ASC 808, to determine whether such arrangements involve joint operating activities performed by parties that are both active participants in the activities and exposed to significant risks and rewards dependent on the commercial success of such activities. For collaboration arrangements within the scope of ASC 808 that contain multiple elements, we first determine which elements of the collaboration are deemed to be within the scope of ASC 808 and those that are reflective of a vendor-customer relationship and therefore within the scope of ASC 606. For elements of collaboration arrangements that are accounted for pursuant to ASC 808, an appropriate recognition method is determined and applied consistently.

Under the criteria of *Accounting Standard Codification, or ASC, 606, Revenue from Contracts with Customers* (Topic 606), or ASC 606, we recognize revenue to depict the transfer of control of promised goods or services to customers in an amount that reflects the consideration to which the entity expects to receive in exchange for those goods or services. We adopted ASC 606 for all periods presented. To determine revenue recognition for arrangements that an entity determines are within the scope of

ASC 606, the entity performs the following five steps: (i) identify the contract(s) with a customer; (ii) identify the performance obligations in the contract; (iii) determine the transaction price, including variable consideration, if any; (iv) allocate the transaction price to the performance obligations in the contract; and (v) recognize revenue when (or as) the entity satisfies a performance obligation. We only apply the five-step model to contracts when it is probable that the entity will collect substantially all the consideration to which it is entitled in exchange for the goods or services it transfers to the customer. We review the contract to determine which performance are distinct and represent a promise to provide distinct goods or services or a series of distinct goods or services as defined by the standard. We recognize as revenue the amount of the transaction price that is allocated to each performance obligation as and when that performance obligation is satisfied.

Licenses of Intellectual Property: Upfront non-refundable payments for licensing our intellectual property are evaluated to determine if the license is distinct from the other performance obligations identified in the arrangement. For licenses determined to be distinct, we recognize revenues from non-refundable, up-front fees allocated to the license at a point in time, when the transfer of control of the license to the licensee occurs and the licensee is able to use and benefit from the license.

Milestone Payments: At the inception of each arrangement that includes development, commercialization, and regulatory milestone payments, we evaluate whether the milestones are considered probable of being reached and to the extent that a significant reversal of cumulative revenue would not occur in future periods, estimates the amount to be included in the transaction price using the most likely amount method. The transaction price is then allocated to each performance obligation on a relative stand-alone selling price basis, for which we recognize revenue as or when the performance obligations under the contract are satisfied. At the end of each subsequent reporting period, we re-evaluate the probability of achieving such development milestones and any related constraint, and if necessary, adjust the estimate of the overall transaction price. Any such adjustments are recorded on a cumulative catch-up basis, which would affect revenues and earnings in the period of adjustment.

Royalties: For arrangements that include sales-based royalties, including milestone payments based on the level of sales, and the license is deemed to be the predominant item to which the royalties relate, we recognize revenue at the later of (i) when the related sales occur, or (ii) when the performance obligation to which some or all of the royalty has been allocated has been satisfied (or partially satisfied).

To date, no milestone payments or royalty payments were received. Substantially all of our revenue has been derived from its out-licensing agreements with respect to licensed products such as DNA sequences, cell lines, etc., and such revenues are recognized when the customer obtains control of the licensed product, which occurs at a point in time, upon delivery to the customer.

Contract assets and contract liabilities: When a customer pays consideration before we transfer products or services, we record our obligation as a contract liability; when we satisfy our performance obligations by providing products or services to a customer before the customer pays consideration and before payment is due, we recognize our rights to consideration as a contract asset.

Fair value measurements

We apply ASC 820, *Fair Value Measurements and Disclosures*. ASC 820 defines fair value, establishes a framework for measuring fair value and expands disclosures about fair value measurements. ASC 820 requires disclosures to be provided for fair value measurements. ASC 820

establishes a three-tier fair value hierarchy, which prioritizes the inputs used in measuring fair value as follows:

Level 1—Observable inputs that reflect quoted prices (unadjusted) for identical assets or liabilities in active markets.

Level 2—Includes other inputs that are directly or indirectly observable in the marketplace.

Level 3—Unobservable inputs which are supported by little or no market activity.

ASC 820 describes three main approaches to measuring the fair value of assets and liabilities: (1) market approach; (2) income approach; and (3) cost approach. The market approach uses prices and other relevant information generated from market transactions involving identical or comparable assets or liabilities. The income approach uses valuation techniques to convert future amounts to a single present value amount. The measurement is based on the value indicated by current market expectations about those future amounts. The cost approach is based on the amount that would currently be required to replace an asset.

The carrying amounts of cash and cash equivalent, short-term investments, accounts receivable, amounts due to related parties and other current assets, accounts payable, amounts due to related parties, accrued liabilities and other current liabilities and short-term borrowings approximate their fair values because of their generally short maturities. The carrying amount of long-term borrowings approximate their fair values since they bear interest rates which approximate market interest rates.

We measured our warrant liabilities at fair value on a recurring basis. As our warrants are not traded in an active market with readily observable prices, we use significant unobservable inputs to measure the fair value of warrant liabilities. These instruments are categorized in the Level 3 valuation hierarchy based on the significance of unobservable factors in the overall fair value measurement.

The following table presents a reconciliation of all financial instruments measured at fair value on a recurring basis using Level 3 unobservable inputs:

	<u>Warrant liabilities</u> US\$
Initial recognition during the year ended December 31, 2018	1,741,720
Fair value change	(534,305)
Balance as of December 31, 2018	1,207,415
Fair value change	(1,207,415)
Balance as of December 31, 2019	—

We did not transfer any assets or liabilities in or out of Level 3 during the year ended December 31, 2018 and 2019.

We had no financial assets and liabilities measured and recorded at fair value on a nonrecurring basis as of December 31, 2018 and 2019.

Research and development expenses

Elements of research and development expenses primarily include (1) payroll and other related costs of personnel engaged in research and development activities, (2) costs related to pre-clinical testing of our technologies under development and clinical trials such as payments to contract research organizations, or CRO, and contract manufacturing organizations, or CMO, investigators and clinical trial sites that conduct the clinical studies; (3) costs to develop the product candidates, including raw materials and supplies, product testing, depreciation and amortization, and facility related expenses,

(4) other research and development expenses. Research and development expenses are charged to expense as incurred when these expenditures relate to our research and development services and have no alternative future uses. As of December 31, 2019, we have several ongoing clinical studies in various clinical trial stages. The contracts with CRO and CMO are generally cancellable, with notice, at our option. We did not record any accrued expenses related to cancellation of CRO or CMO contracts as of December 31, 2019 as we did not have any plan to cancel the existing CRO or CMO contracts.

Income taxes

We follow the liability method of accounting for income taxes in accordance with ASC 740, *Income Taxes*, or ASC 740. Under this method, deferred tax assets and liabilities are determined based on the difference between the financial reporting and tax bases of assets and liabilities using enacted tax rates that will be in effect in the period in which the differences are expected to reverse. We record a valuation allowance to offset deferred tax assets if based on the weight of available evidence, it is more likely than not that some portion, or all, of the deferred tax assets will not be realized. The effect on deferred taxes of a change in tax rate is recognized in tax expense in the period that includes the enactment date of the change in tax rate.

We evaluate our uncertain tax positions using the provisions of ASC 740, which prescribes a recognition threshold that a tax position is required to meet before being recognized in the consolidated financial statements.

We recognize in the consolidated financial statements the benefit of a tax position which is "more likely than not" to be sustained under examination based solely on the technical merits of the position assuming a review by tax authorities having all relevant information. Tax positions that meet the recognition threshold are measured using a cumulative probability approach, at the largest amount of tax benefit that has a greater than fifty percent likelihood of being realized upon settlement. It is our policy to recognize interest and penalties related to unrecognized tax benefits, if any, as a component of income tax expense.

We have incurred net accumulated operating losses for income tax purposes since our inception. We believe that it is more likely than not that these net accumulated operating losses will not be utilized in the future. Therefore, we have provided full valuation allowances for the deferred tax assets as of December 31, 2018 and 2019. There was no income tax expense for the years ended December 31, 2018 and 2019, as our subsidiaries did not have any taxable profits.

Share-based compensation

We grant restricted shares and stock options to eligible employees and nonemployees and accounts for share-based compensation in accordance with ASC 718, *Compensation—Stock Compensation*.

Share-based compensation awards are measured at the grant date fair value of the awards and recognized as expenses a) immediately at the grant date if no vesting conditions are required; or b) for share-based awards granted with only service conditions, using the straight-line method over the vesting period; or c) for share-based awards granted with service conditions and performance conditions, using the graded vesting method over the vesting period if and when the company concludes that it is probable that the performance conditions will be achieved.

A change in any of the terms or conditions of share-based awards is accounted for as a modification of the awards. We calculate incremental compensation expense of a modification as the excess of the fair value of the modified awards over the fair value of the original awards immediately before its terms are modified at the modification date. For vested awards, we recognize incremental compensation cost in the period when the modification occurs. For awards not being fully vested, we recognize the sum of the incremental compensation expense and the remaining unrecognized

compensation expense for the original awards over the remaining requisite service period after modification.

The fair value of share options was determined using the binomial option valuation model, with the assistance from an independent third-party appraiser. The binomial model requires the input of highly subjective assumptions, including the expected volatility, the exercise multiple, the risk-free rate and the dividend yield. For expected volatility, we have made reference to historical volatility of several comparable companies in the same industry. The exercise multiple was estimated as the average ratio of the stock price to the exercise price of when employees would decide to voluntarily exercise their vested share options. The risk-free rate for periods within the contractual life of the share options is based on the market yield of U.S. Treasury Bonds in effect at the time of grant. The dividend yield is based on the expected dividend policy over the contractual life of the share options. The estimated fair value of the ordinary shares, at the share option grant dates, was determined with the assistance from an independent third-party appraiser. Our management is ultimately responsible for the determination of the estimated fair value of its ordinary shares.

The assumptions used to estimate the fair value of the share options granted are as follows:

	For the year ended December 31, 2019	For the nine months ended September 30	
		2019	2020
Risk-free interest rate	1.78%-2.73%	2.73%	0.71%-0.83%
Dividend yield	0%	—	—
Expected volatility range	67.5%-71.0%	71.0%	72.3%-73.1%
Exercise multiple	2.2-2.8	2.2	2.2-2.8
Contractual life	10 years	10 years	10 years

Borrowings

Borrowings are recognized initially at fair value, net of transaction costs incurred. Borrowings are subsequently stated at amortized cost; any difference between the proceeds (net of transaction costs) and the redemption value is recognized in the consolidated statements of comprehensive loss over the period of the borrowings using the effective interest method.

Internal Control Over Financial Reporting

Prior to this offering, we have been a private company with limited accounting personnel and other resources with which to address our internal control over financial reporting. In connection with the audit of our consolidated financial statements as of and for the three fiscal years ended December 31, 2018 and 2019, our independent registered public accounting firm identified two material weaknesses in our internal control over financial reporting as at December 31, 2019. As defined in standards established by the PCAOB, a "material weakness" is a deficiency, or combination of deficiencies, in internal control over financial reporting, such that there is a reasonable possibility that a material misstatement of the annual or interim financial statements will not be prevented or detected on a timely basis.

The two material weaknesses that have been identified related to:

- Our lack of sufficient and competent financial reporting and accounting personnel with appropriate knowledge of U.S. GAAP and SEC reporting and compliance requirements; and
- Our lack of sufficient documented financial closing policies and procedures, specifically those related to period end expenses cut-off and accruals.

Such material weaknesses, if not timely remedied, may lead to significant misstatements in our consolidated financial statements in the future.

To remediate our identified material weakness, we are in the process of adopting several measures to improve our internal control over financial reporting, including (i) hiring more qualified accounting personnel, with relevant U.S. GAAP and SEC reporting experience and qualifications to strengthen the financial reporting function and setting up a financial and system control framework; (ii) implementing regular and continuous U.S. GAAP accounting and financial reporting training programs for our accounting and financial reporting personnel; and (iii) preparing comprehensive accounting policies, manuals and closing procedures to improve the quality and accuracy of our period-end financial closing process.

We expect that we will incur significant costs in the implementation of such measures. However, we cannot assure you that all these measures will be sufficient to remediate our material weakness in time, or at all. See "Risk factors—Risks Relating to Our Operations—If we fail to implement and maintain an effective system of internal controls, we may be unable to accurately or timely report our results of operations or prevent fraud, and investor confidence and the market price of our ADSs may be materially and adversely affected."

As a company with less than US\$1.07 billion in revenue for our last fiscal year, we qualify as an "emerging growth company" pursuant to the JOBS Act. An emerging growth company may take advantage of specified reduced reporting and other requirements that are otherwise applicable generally to public companies. These provisions include exemption from the auditor attestation requirement under Section 404 of the Sarbanes-Oxley Act of 2002, in the assessment of the emerging growth company's internal control over financial reporting.

Quantitative and Qualitative Disclosure about Market Risk

Interest rate risk

Our exposure to interest rate risk primarily relates to the interest income generated by excess cash, which is mostly held in interest-bearing bank deposits. We have not used any derivative financial instruments to manage our interest risk exposure. Interest-earning instruments carry a degree of interest rate risk. We have not been exposed, nor do we anticipate being exposed, to material risks due to changes in interest rates. However, our future interest income may be lower than expected due to changes in market interest rates.

Foreign exchange risk

We are a global business enterprise with part of our operations based in the PRC. A part of our transactions were settled in Renminbi, and our financial statements are presented in U.S. dollars. We do not believe that we currently have any significant direct foreign exchange risk and have not used any derivative financial instruments to hedge our exposure to such risk. Although, in general, our exposure to foreign exchange risks should be limited, the value of your investment in the ADSs will be affected by the exchange rate between U.S. dollar and Renminbi because a portion of value of our business is effectively denominated in Renminbi, while the ADSs representing our ordinary shares will be traded in U.S. dollars.

The value of the Renminbi against the U.S. dollar and other currencies is affected by changes in China's political and economic conditions and by China's foreign exchange policies, among other things. In July 2005, the PRC government changed its decades-old policy of pegging the value of the Renminbi to the U.S. dollar, and the Renminbi appreciated more than 20% against the U.S. dollar over the following three years. Between July 2008 and June 2010, this appreciation subsided and the exchange rate between the Renminbi and the U.S. dollar remained within a narrow band. Since June 2010, the

Renminbi has fluctuated against the U.S. dollar, at times significantly and unpredictably. While appreciating approximately by 7% against the U.S. dollar in 2017, the Renminbi in 2018 depreciated approximately by 5% against the U.S. dollar. Since October 1, 2016, the US\$ has joined the International Monetary Fund (IMF)'s basket of currencies that make up the Special Drawing Right (SDR), along with the U.S. dollar, the Euro, the Japanese yen and the British pound. With the development of the foreign exchange market and progress towards interest rate liberalization and Renminbi internationalization, the PRC government may in the future announce further changes to the exchange rate system and there is no guarantee that the RMB will not appreciate or depreciate significantly in value against the U.S. dollar in the future. It is difficult to predict how market forces or PRC or U.S. government policy may impact the exchange rate between the Renminbi and the U.S. dollar in the future.

To the extent that we need to convert U.S. dollars into Renminbi for our operations, appreciation of Renminbi against the U.S. dollar would reduce the Renminbi amount we receive from the conversion. Conversely, if we decide to convert Renminbi into U.S. dollars for the purpose of making payments for dividends on our ordinary shares or ADSs, servicing our outstanding debt, or for other business purposes, appreciation of the U.S. dollar against the Renminbi would reduce the U.S. dollar amounts available to us.

We estimate that we will receive net proceeds of approximately US\$ million from this offering, after deducting underwriting discounts and commissions and the estimated offering expenses payable by us, based on the assumed initial offering price of US\$ per ADS, the midpoint of the estimated initial public offering price range set forth on the front cover of this prospectus.

Inflation risk

Since our inception, inflation in China has not materially impacted our results of operations. According to the National Bureau of Statistics of China, the year-over-year percent changes in the consumer price index for December 2018 and 2019 were increases of 1.9% and 4.5%, respectively. Although we have not in the past been materially affected by inflation since our inception, we can provide no assurance that we will not be affected in the future by higher rates of inflation in China.

Recently Issued Accounting Pronouncements

For detailed discussion on recent accounting pronouncements, see Note 2 to our Consolidated Financial Statements.

BUSINESS

OVERVIEW

We are a platform-driven, clinical-stage biopharmaceutical company committed to transforming the discovery and development of novel antibody-based cancer immunotherapies. Our platform is designed to generate therapeutic antibody candidates with unique functional epitopes and species cross-reactivity as highlighted by our immunotherapy pipeline. These features enable our novel drug discovery strategy to advance from lead identification through vigorous preclinical modeling to biomarker-guided mono- and combination immunotherapy development in clinical settings. We have pioneered a dynamic interface design to harness the conformational diversity of antibodies, which enlarges epitope sampling of a given drug target for differentiated therapeutic antibody development. We aim to push the boundaries of antibody discovery and engineering through the precise design, construction, and selection of antibody product candidates intractable to traditional antibody technology.

Life is motion. The motion of proteins and their dynamic interactions trigger a cascade of complex biological and pharmacological effects. Our core technology is built upon our fundamental understanding of the role that protein folding and the motion of molecules play in giving rise to dynamic conformational diversity, where an amino acid sequence can adopt multiple structures and functions. Our approach recognizes that a protein's native state is not accurately represented by a single static structure but rather by a variety of structures in dynamic equilibrium, resulting in a high level of functional diversity, in contrast to the conventional static antibody drug discovery paradigm of "one sequence, one structure and one function."

We have developed our proprietary AI-Powered DPL platform to explore the dynamic conformational diversity of protein sequences, and the flexible binding sites of antibody sequences in particular, as a new paradigm for antibody drug discovery. Our DPL platform combines artificial intelligence, or AI, algorithms and ever-increasing big data in antibody sequence, structure, binding epitope and affinity from public and our own proprietary databases to design, construct and screen high-quality proprietary antibody libraries with well-defined sequence, scaffold and biophysical attributes for antibody drug discovery. Powered by computational physics in combination with AI and big data, our DPL platform samples a potentially infinite number of dynamic binding interface structures arising from the conformational diversity of a finite number of antibody amino acid sequences, allowing us to exponentially expand the universe of candidate antibody binding sites far beyond conventional natural or synthetic antibody repertoires. By exploiting conformational diversity, we have designed and precisely constructed approximately one trillion (10^{12}) antibody sequences in our DPL. These antibodies feature broad epitope (the portion of an antigen that are recognized by an antibody) coverage and robust chemistry, manufacturing, and control, or CMC, attributes.

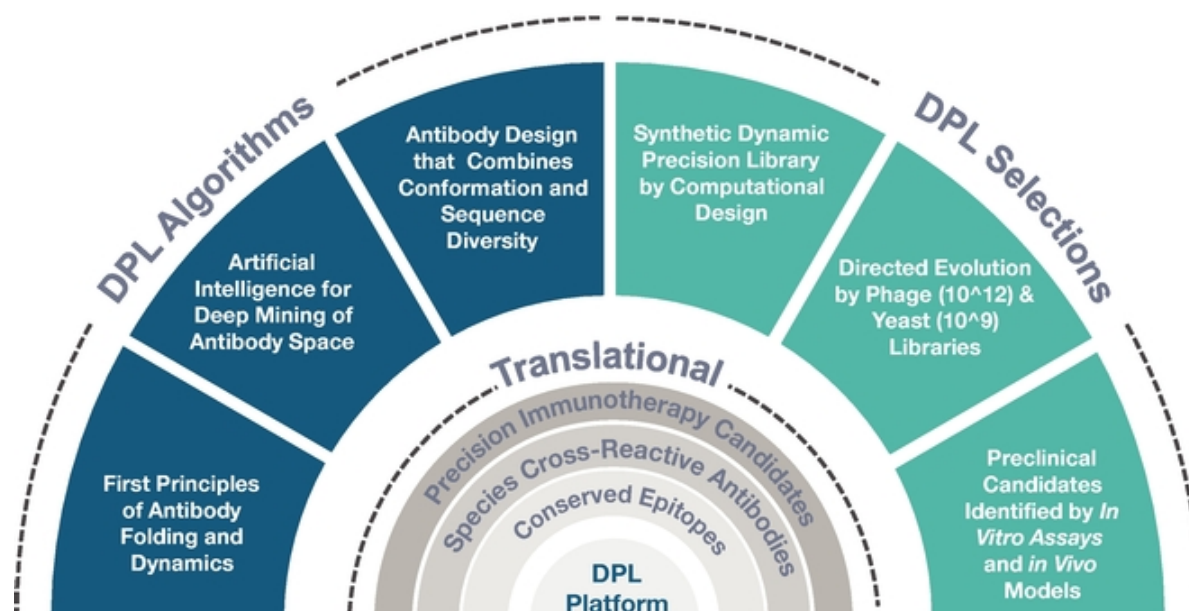
We believe our AI-powered antibody discovery and engineering DPL platform significantly increase R&D productivity for antibody drug discovery, as illustrated by our clinical anti-CD137 and anti-CTLA-4 programs. DPL library screening against CD137 or CTLA-4 antigens has yielded a large number of high affinity primary hits. The abundant discovery hits with diversified binding epitopes show the power of our AI-powered DPL platform not only in creating novel antibodies in targeting different epitopes of a given antigen, but also in targeting the conserved epitope across different species of a given antigen with broad species cross-reactivity from human, monkey to mouse, which enables us to study their efficacy and safety in extensive immuno-competent or syngeneic animal models, to explore their pharmacodynamics and predictive biomarkers in responding vs nonresponding tumor models in vivo, and to understand their deep target biology and novel MOA before testing them in human clinical trials to look for their clinical signals consistent with their MOA. Our DPL platform empowers us to engage the dynamic epitope of the conformationally dynamic target which might be challenging using conventional antibody discovery approaches. We believe that the high-affinity and cross-reactive primary hits from our AI-Powered DPL library screening save time and cost from discovery to early clinical

proof of concept. The broad species cross-reactivity of the primary hits also streamlines lead identification and potentially enables high fidelity translation from preclinical to clinical studies. Furthermore, we believe our AI-powered dynamic precision library expands the diversity at the start of discovery to maximize the chance that suitable leads are found on the first pass.

Translational fidelity from preclinical modeling to informed clinical development is one of the top challenges to developing cancer immunotherapies. Most traditionally developed antibodies do not cross react between their human and murine targets due to their limited species cross-reactivity, making it very difficult to reliably evaluate the same antibody in both the preclinical and clinical settings. Some of the most contentious issues related to preclinical and clinical studies of CD137 and CTLA-4, the targets of our lead product candidates, immunotherapies are traceable to the differences between the antibodies used for preclinical and clinical studies. For example, according to Frost & Sullivan, two of the leading clinical anti-CD137 agonist antibodies bind to different epitopes and exhibit dramatic differences in their respective clinical safety and efficacy results, underscoring the importance of finding suitable species cross-reactive antibodies like those we have utilized for comprehensive preclinical evaluation before entering clinical trials.

We believe that it is essential to model the interactions between tumors and an intact host immune system *in vivo* to evaluate the therapeutic potential of antibodies in preclinical studies. The flexibility of antibody binding interface is fundamental to the NEObody technology of our DPL Platform and allows us to generate species cross-reactive antibodies to assess the safety and efficacy potential of mono- and combination therapy candidates in syngeneic animal models before launching clinical trials. We use syngeneic animal models which are known for their intact *in vivo* immune systems to provide the original proof of concept for cancer immunotherapies by blocking immune check points with monoclonal antibodies, or mAbs. We believe that the use of species cross-reactive antibodies, rather than surrogate antibodies used in traditional syngeneic mouse models, should facilitate the translational relevance and clinical utility of these well-established preclinical models for determining optimal dose, schedule, sequencing, combination synergy, risk and benefit features. The results from the assessment of new species cross-reactive antibodies in rigorous preclinical models may allow us to control the scope and cost of clinical trials, enable the identification of potential clinical biomarkers useful to monitor clinical pharmacological and safety signals, and help preselect patients for precision mono- and combination therapies.

The figure below illustrates how our DPL platform integrates our computational algorithm-enabled high-throughput screening and functional antibody evaluation for preclinical candidates suitable for clinical development as explained above.

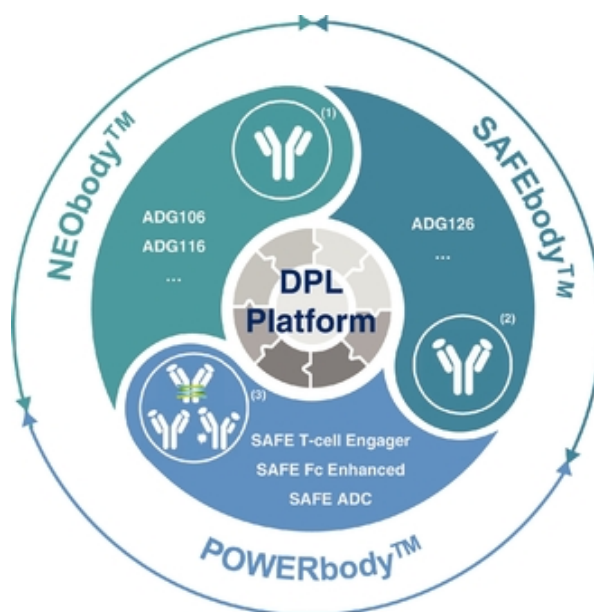


Our DPL platform is further composed of three proprietary enabling technologies tailored to three key attributes of antibody-based therapeutic modalities:

- NEObody technology is a fully synthetic phage display and yeast display-based antibody discovery technology, which we believe is differentiated from other synthetic antibody technologies through its innovative designs and precise constructions. NEObody technology enables the generation of antibodies designed with dynamic binding sites that adapt kinetically to unique epitopes, triggering a novel MOA. The species cross-reactive antibodies generated by NEObody technology not only have the potential to reveal new biological functions of the targets, but also facilitate preclinical studies using various immune system intact animal models, resulting in high fidelity translation from preclinical to clinical studies. We refer to antibodies generated by our NEObody technology as NEObodies.
- SAFEbody technology is designed to mask an antibody binding interface with a masking motif, which then prevents an antibody from binding to its target in healthy tissues. The masking motif is designed to activate, or unmask, the antibody to allow binding in the tumor microenvironment, or TME, where certain activation conditions such as a protease is upregulated as compared to healthy tissues, allowing the antibody to bind to its target for tumor killing. Our SAFEbody enabled therapeutic candidates are therefore designed to be activated predominantly in the TME while remaining largely in an inactive state in healthy tissues. Our SAFEbody technology can be applied to mask the binding sites of any antibodies including but not limited to NEObodies. We refer to such masked antibodies including NEObodies as SAFEbodies.
- POWERbody technology, which enables the creation of new versions of bispecific T-cell engagers, or TCEs, with enhanced safety profiles, antibody-drug conjugates, or ADCs, or antibodies that are designed to reach beyond the therapeutic potency of traditional monospecific antibodies.
- SAFEbody technology can be applied to our NEObodies, such as what we did with ADG116 to convert into ADG126. POWERbody is created by applying the SAFEbody technology to a bispecific TCE in either CD3 or both in CD3 and antigen binding arms to enhance the safety profile. Our POWERbody technology is designed to improve antitumor activity while maintaining the enhanced safety profile.

We believe that comprehensive *in vivo* preclinical evaluations are the key to assess the efficacy and safety potential of tailor-made antibody candidates before progressing them into lengthy and costly clinical trials. NEObody, SAFEbody and POWERbody technologies are all designed to facilitate favorable druggability, manageable CMC attributes, and reduced immunogenicity. As highlighted by our lead product candidates, such as ADG106, ADG126 and ADG116, our NEObody technology allows us to engineer and select species cross-reactive NEObodies designed to dynamically adapt to unique and evolutionally conserved epitopes. ADG126 has been further engineered using our SAFEbody technology to address the safety concerns associated with existing CTLA-4 therapeutics.

The figure below shows how our NEObody, SAFEbody, and POWERbody technologies on the one hand and DPL platform on the other hand are interconnected and utilized for the building of our product pipeline of mono- and combination immunotherapies.



Notes:

- (1) Antibodies generated by NEObody technology, which are designed with dynamic binding sites that adapt kinetically to unique epitopes, triggering a novel MOA.
- (2) NEObodies or traditional antibodies are masked in their binding sites by SAFEbody technology as shown by the blocking bar in the antibody binding sites in the figure, which are designed to be selectively activated in the TME, potentially limiting on-target off-tumor toxicity in normal tissues.
- (3) Multiple potent modalities, including bispecific T-cell engagers, ADC, and Fc engineering for enhanced ADCC, etc., are masked in their binding sites by SAFEbody to create a POWERbody.

Our most advanced NEObody product candidate, ADG106, is a fully human ligand-blocking agonistic anti-CD137 mAb currently being evaluated in Phase Ib clinical trials in the United States and China. ADG106 is designed to target a unique epitope of CD137 that is different from other anti-CD137 antibodies currently under clinical development. Epitope mapping and X-ray structural analysis of ADG106 with CD137 have shown in preclinical studies that ADG106 is capable of binding to CD137 in a fashion similar to its natural ligand, CD137L. Our first SAFEbody product candidate, ADG126, is a fully human anti-CTLA-4 masked antibody designed to address the safety concerns associated with existing CTLA-4 therapeutics. It is designed to enhance the safety features by masking the antibody binding site of ADG126, which would be unmasked in the TME, where the activated ADG126 would block CTLA-4 and deplete regulatory T-cells by means of enhanced antibody-

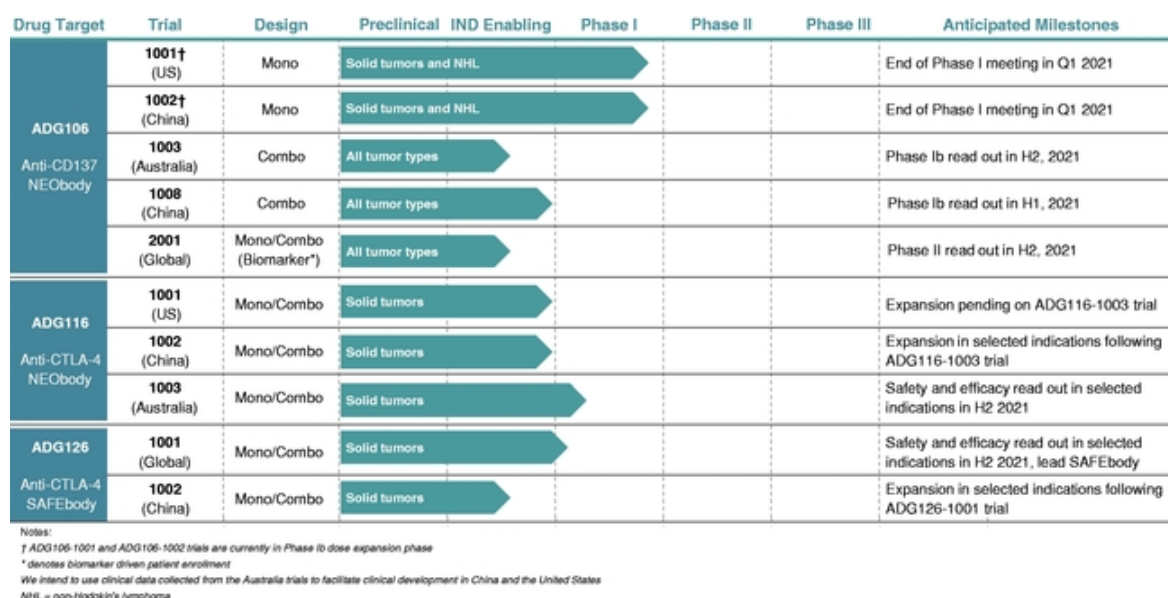
dependent cellular cytotoxicity, or ADCC. In preclinical studies, ADG126 was tolerated at doses of up to 200 mg/kg in nonhuman primate models. As ADG126 is also species cross-reactive in humans, cynomolgus monkeys and mice, we believe that preclinical studies of ADG126 will support the rational design of clinical trials to expedite its development. Our third product candidate, ADG116, is a fully human anti-CTLA-4 NEObody. Epitope mapping and X-ray structural analysis have shown that in preclinical studies, ADG116 is capable of binding to a novel epitope of CTLA-4 different from ipilimumab, the only CTLA-4 mAb approved globally. The dynamic interface of ADG116 enabled not only its species cross-reactivity with human, cynomolgus monkey, and mouse CTLA-4 for preclinical studies, but also its dynamic engagement on a unique epitope of CTLA-4 to trigger a novel MOA distinct from ipilimumab by softer ligand blocking and stronger regulatory T-cell depletion via strong ADCC.

Our species cross-reactive ADG106, ADG126 and ADG116 have enabled a deep understanding of the interaction between tumor and host immune system *in vivo* in syngeneic animal models. This understanding has been utilized to design and guide the clinical development of rational, mechanism-based mono- and combination therapies using ADG106, ADG126 and ADG116. Because there is limited clinical safety and efficacy data available for anti-CD137 agonists, we have followed our preclinical and mechanistic study for the clinical development of ADG106. We observed three patients with Grade 3 liver enzyme increase; among them, one patient with abnormal liver enzyme baseline showed a Grade 3 AST increase. ADG106 showed preliminary clinical antitumor activity in patients who have progressed after several lines of treatment in our completed Phase Ia dose escalation and ongoing Phase Ib dose expansion trials in the United States and China. It is very encouraging to observe the clinical response in connection with the changes in PD biomarkers due to target engagement in a dose dependent manner. We have also observed a correlation with a potential predictive biomarker for patient selection in three patients with more than 30% tumor shrinkage across different indications in retrospective analysis in our ongoing Phase I trials. This predictive biomarker is related to the CD137 pathway. We intend to further explore this predictive biomarker in order to guide our development of precision mono- and combination immunotherapies based on our preclinical and preliminary clinical data.

Our Pipeline

By leveraging our proprietary DPL platform, we have developed a robust pipeline of innovative product candidates in various stages of development, ranging from research and discovery to preclinical and clinical development. Our highly differentiated clinical-stage pipeline consists of ADG106, ADG126 and ADG116. We also have a robust preclinical pipeline in various stages of development. In addition, we have out-licensed the Greater China rights of ADG104, a PD-L1 mAb under clinical development, to our partner, Sanjin. We have the right to apply for the patents derived from our core and key technologies pertaining to ADG104 in the rest of the world and we retain a majority of the economic benefits derived from ADG104 in the rest of the world.

The following chart provides an overview of the status of each of our clinical-stage and IND-enabling study stage programs at clinical or IND-enabling stages, for which we have global rights.



ADG106: Novel agonistic anti-CD137 NEObody candidate

ADG106, is a fully human ligand-blocking, agonistic anti-CD137 immunoglobulin G4, or IgG4, mAb generated using our NEObody technology. ADG106 is being developed for the treatment of advanced solid tumors and non-Hodgkin's lymphoma, or NHL. CD137 stimulates the immune system to attack cancer cells and is a key driver for T-cell and natural killer cell, or NK cell, proliferation. ADG106 is designed to target a unique conserved epitope of CD137 with a novel MOA for CD137 agonism by its natural ligand-like binding and cross-linking by Fcγ receptors. The broad species cross-reactivity of ADG106 observed in preclinical studies has enabled us to explore robust translational studies concerning the biology of CD137 in mouse, rat, nonhuman primate, and human, especially for anti-CD137 as a single agent and in combination therapies in immuno-competent syngeneic tumor models. In preclinical studies to date, we observed that ADG106 had encouraging antitumor activity and was well tolerated as a monotherapy and in combination with the existing standard-of-care, or SOC, and other immuno-oncology therapies, which was confirmed in the Phase I clinical trials in cancer patients as a monotherapy. These data indicate that ADG106 has the potential to address the limitations of other existing anti-CD137 therapies.

As of the November 30, 2020, the Data Cut-off Date, we have completed the Phase Ia dose escalation in each of our Phase I studies of ADG106 in both the United States and China as a monotherapy in patients with advanced or metastatic solid tumors and/or NHL. We are currently in the Phase Ib dose expansion phase for both trials in the United States and China. ADG106 was dose escalated up to 10 mg/kg and is well tolerated at dose expansion at 3 mg/kg, 5 mg/kg and at 300mg and 400 mg flat doses. A total number of 92 patients have been dosed. Both clinical trials, namely ADG106-1001 and ADG106-1002, have limited Treatment Emergent Adverse Effects, or TEAEs, liver toxicity or hematologic abnormalities. We observed three patients with Grade 3 liver enzyme increase; among them, one patient with abnormal liver enzyme baseline showed a Grade 3 AST increase. Two patients developed hemolytic anemia in ADG106-1002 which were controlled; one of the two patients received prior therapies which were known to cause autoimmune hemolytic anemia. One patient with a solid tumor who previously failed chemotherapies, radiotherapy, and an anti-PD-L1 related antibody treatment showed a partial response to ADG106 treatment with a 40% tumor size reduction after two

treatment cycles. In addition, two NHL patients showed more than a 30% tumor size reduction after one and two ADG106 treatment cycles, respectively. Furthermore, biomarker studies showed target engagement with specific PD biomarkers indicative of immune system activation, and clinical response correlated with changes in CD137 target engagement. These data are encouraging given that the enrolled population was not preselected and was heavily pretreated. We have identified a potential predictive biomarker which correlates with patient response to ADG106 treatment from a retrospective analysis of the ongoing Phase I clinical trial. Based on this finding, we are in the process of preparing an additional Phase II trial which we expect to initiate in 2021 and for which we intend to stratify and preselect patients using this predictive biomarker to potentially enhance clinical response of patients to ADG106 treatment. We also plan to pursue potential registrational trials evaluating ADG106 in biomarker enriched patient populations.

We have also evaluated ADG106 in combination with other therapies including chemotherapies, immune modulators, and immuno-oncology therapies in preclinical studies. Data from combination studies in tumor bearing mice showed that the combination of ADG106 with immune checkpoint inhibitors, including an anti-PD-1/L1 mAb or anti-CTLA-4 mAb, enhanced *in vivo* antitumor activity. We plan to explore the combination of ADG106 with other targeted antibody therapies for the treatment of hematologic malignancies and solid tumors. We also have identified tumor-specific biomarkers that we believe may correlate with ADG106 antitumor activity in multiple mouse tumor models. These preclinical study findings are consistent with the interim results from our ongoing Phase Ib clinical trials.

Going forward, we are focusing on evaluating combinations of ADG106 with various therapies in clinical trials. We have received NMPA approval for the ADG106-1008 study in China, which is a Phase Ib/II combination trial of ADG106 with anti-PD-1 in advanced solid tumors and hematological malignancies. We are also in the preparation for a submission of a Clinical Trial Notification, or CTN, for our ADG106-1003 clinical trial in Australia, which will combine ADG106 with anti-PD-1, and other therapies in advanced solid tumors and hematological malignancies.

We have observed responses to ADG106 in all three T cell lymphoma patients enrolled in the ADG106-1002 Phase I trial. We are enrolling additional patients in our ADG106-1002 trial and will conduct an investigator initiated trial in the United States to further explore ADG106 as a monotherapy and in combination with other therapies in T cell lymphoma. We expect to submit an Orphan Drug Designation application to the FDA for ADG106 requesting for an expedited regulatory pathway for this indication.

ADG126: Novel anti-CTLA-4 SAFEbody candidate

Our most advanced SAFEbody program, ADG126, is a masked fully-human anti-CTLA-4 mAb engineered to address the safety concerns associated with existing CTLA-4 therapeutics, while maintaining potency when locally activated in the TME. The FDA approval of ipilimumab validated anti-CTLA-4 for cancer treatment. However, due to its on-target off-tumor toxicity, the approved indications for ipilimumab are limited, which we believe has caused sales of ipilimumab to trail other immuno-oncology therapies such as anti-PD-1/L1 antibodies.

ADG126 is designed to address the toxicity and efficacy issues related to the MOA of existing approved CTLA-4 immuno-oncology therapies and to expand the potential of CTLA-4 as a target for the treatment of cancer. In preclinical studies, ADG126 was tolerated at doses of up to 200 mg/kg in nonhuman primate models. We believe the encouraging preclinical tolerability of ADG126 suggests its potential in combination with other immunotherapies such as an anti-PD-1/PD-L1 antibody or an anti-CD137 antibody, including ADG106.

To better address the unmet clinical need for a safe and potent anti-CTLA-4 antibody for chemotherapy-free mono- and combination immunotherapy, we have submitted a CTN and received

HREC approval for an ADG126 Phase I dose escalation trial in Australia and are expecting to commence patient enrollment by February 2021. We also have received approval from the FDA to initiate the Phase I clinical trial of ADG126 in the United States in 2021. These two Phase I dose escalation trials will test five doses at 0.1, £0.3, £1, £3, and £10 mg/kg, based on our preclinical primate toxicity study in comparison with ipilimumab. We anticipate that these clinical trials will allow us to determine the ADG126 effective dose in late 2021. Meanwhile, we are preparing IND submissions to initiate ADG126 clinical trials in China.

ADG116: Novel anti-CTLA-4 NEObody candidate

ADG116 is a fully-human ligand-blocking anti-CTLA-4 mAb generated using our NEObody technology. ADG116 is designed to target a unique conserved epitope of CTLA-4. In preclinical studies, ADG116 was observed to have softer CTLA-4 ligand blocking and stronger ADCC for depleting regulatory T-cells than ipilimumab. In a head-to-head *in vivo* efficacy study, ADG116 was observed to have a five-fold greater anti-tumor activities in comparison with ipilimumab. In addition, ADG116 was observed to induce antitumor responses concomitant with reduced immunosuppressive regulatory T-cell and enhanced cytotoxic T lymphocyte (CD8⁺ T-cells) activities in the TME. We believe that these preclinical results support the further clinical evaluation of ADG116 both as monotherapy and combination therapy for a wide range of tumor types.

As of January 15, 2021, we have successfully completed dose escalation at the first three doses in ADG116-1003 clinical trial in Australia, including dose level at 0.03mg/kg, the dose level used for the first patient who suffered a fatal incident in the ADG116 Phase I trial in the United States. As of January 15, 2021, no dose limiting toxicity or treatment related serious adverse events have been observed in any of the patients treated. There was no treatment induced liver toxicity and all three patients treated at 0.03mg/kg showed normal liver enzyme level post ADG116 treatment. The Safety Review Committee approved dose escalation for ADG116-1003 clinical trial to 0.1mg/kg. The ADG116-1001 Phase I trial in the United States was placed on clinical hold on September 30, 2019 by the FDA, after we reported to the FDA the death of the only patient dosed in the trial. The FDA removed the clinical hold on December 5, 2019 after we submitted an amendment to the study protocol.

Our Global Partnerships and Collaborations

We have a successful track record of collaborations and partnerships with global biopharmaceutical companies and academic institutions. So far, we have established multiple collaboration programs and will continue to seek partnership opportunities where we can leverage our proprietary technology platform to develop novel antibodies to address unmet medical needs. Over the past two years, we have established partnerships and collaborations with multiple biopharmaceutical companies.

- We entered into a material transfer and collaboration and license agreement with ADC Therapeutics SA, or ADC Therapeutics, under which ADC Therapeutics intends to use our SAFEbody technology to generate a masked antibody that could be combined with the pyrrolbenzodiazepine cytotoxic payload technology used in ADC Therapeutics' ADCs for the development of a novel ADC against a solid tumor target. Under the agreement, we could be eligible to receive royalty payments and could have an exclusive option to negotiate a license to develop and commercialize co-developed assets in certain territories; and
- We are also collaborating with Guilin Sanjin Pharmaceutical Co., Ltd., or Sanjin, and its affiliates to develop two different monoclonal antibodies with the first being ADG104, a monospecific antibody that targets PD-L1 and is in Phase Ib and Phase II clinical trials concurrently in China, and the second being an undisclosed monoclonal antibody, which Sanjin and its affiliates have recently filed an IND application that has been accepted by the NMPA;

- We have recently extended the term of our research collaboration agreement, or the NIH RCA, with the National Heart Lung and Blood Institute, part of the National Institutes of Health, or NIH, to 2023. We first entered into the NIH RCA in March 2018 to collaborate with Dr. Richard Childs, Chief of the Laboratory of Transplantation Immunotherapy at NHLBI, who has used antibodies discovered by us to develop a CAR-T cell therapy candidate targeting a human endogenous retrovirus expressed in the majority of clear cell kidney tumors. The extension of the NIH RCA will allow us to use our proprietary antibody technology to further collaborate with Dr. Richard Childs and his team to develop other antibodies; and
- We entered into a collaboration and license agreement with Exelixis, Inc., or Exelixis, under which Exelixis will utilize our SAFEbody technology platform to generate masked versions of mAbs from Exelixis' growing preclinical pipeline for the development of ADCs or other innovative biologics against Exelixis-nominated targets. Under the agreement, we could be eligible to receive upfront payment as well as milestone and royalty payments on net sales of any potential products developed around each of these targets.

We are also working with global biopharmaceutical companies to potentially develop additional strategic partnerships.

- We had recently worked with Celgene (now Bristol-Myers Squibb) to discover antibodies targeting novel antigens using our proprietary DPL platform;
- Under a material transfer agreement, we are developing SAFEbody drug conjugates against a tumor target selected by Tanabe Research Laboratories, Inc., or TRL, with potential for negotiating a future license agreement with TRL if our pilot work proves successful;
- We recently entered into an evaluation agreement with a global biotechnology company, or the Evaluation Agreement, under which we are developing bispecific antibodies leveraging our SAFEbody and bispecific technologies against tumor targets provided by the counterparty. The Evaluation Agreement could facilitate the parties' interest in a further collaboration concerning the development of a new antibody therapeutic; and
- We had previously collaborated with GlaxoSmithKline (China), or GSK China, where we were engaged to generate high affinity antibodies against multiple epitopes of multi-transmembrane targets; and we previously worked with Jiangsu Hengrui Medicine Company Limited, or Jiangsu Hengrui, where we were able to discover cross-reactive agonistic antibody for immuno-oncology.

Our Team and Investors

We were founded in 2011 by Dr. Peter Luo and is led by an experienced management team. Dr. Luo, who previously founded the biopharmaceutical company Abmaxis which was subsequently acquired by Merck, has a proven track record of more than two decades in antibody discovery and engineering using a multidisciplinary approach that combines computational and experimental technology based on physical, chemical, and biological sciences. Our management team is composed of industry veterans with extensive experience in therapeutic antibody research and development and collectively has decades of experience in molecular biology, immunotherapy, immunology, antibody discovery, protein engineering, and clinical development. Our management team brings a strong history of leadership, innovation, and research and development experience at leading companies, including Merck/Abmaxis, Affomix/Illumina, Amgen, AnaptysBio, Bristol-Myers Squibb, Celgene, Corixa, Forty Seven Inc., Xencor, Novartis, Pfizer, Prometheus, Qantical, and Teva Biopharmaceuticals. Our company is further supported by a strong group of investors that share our commitment to developing next-generation immuno-oncology therapies for the treatment of cancers. Our investors include leading institutional investors such as F-Prime, Eight Roads, General Atlantic, GP Healthcare Capital, Sequoia China and Wuxi AppTec.

OUR STRATEGIES

We are utilizing our proprietary DPL platform to design, construct and develop novel immunotherapies and precision antibodies to address unmet patient needs globally. Our strategy encompasses the following key elements:

- **Advance clinical development of our lead product candidates, ADG106, ADG126 and ADG116, as monotherapies and in combination with other therapies.** We have completed the Phase Ia dose escalation of our ADG106 Phase Ia clinical trials in both the United States and China and are expanding our clinical trials in Phase Ib to patients in sensitive tumor types based on clinical and preclinical observations. We have also secured regulatory approval for combination studies in China and are in process of preparing for biomarker-driven Phase II global trials. We have obtained authorization from the Australian Therapeutic Goods Administration under a CTN for ADG126, our second-generation CTLA-4 antibody that is designed to activate in the TME, for a Phase I trial in Australia and expect to commence patient enrollment by February 2021. We also have received approval from the FDA to initiate the Phase I clinical trial of ADG126 in the United States in 2021. We expect both trials to support our global development for our SAFEbody ADG126. In addition, we obtained authorization from the Australian Therapeutic Goods Administration under a CTN to start a Phase I trial of ADG116 in July 2020. Five patients at three different dose levels were subsequently dosed. We intend to use clinical data collected from the Australia trial to facilitate effective clinical development in the United States. We intend to continue to explore the use of our product candidates as monotherapies by focusing on different tumor types and as combination therapies by conducting combination studies with various immuno-oncology agents, such as an anti-PD-1/L1 mAb. We are planning to expand the cohorts in our clinical trials with patients with responsive cancer types or biomarker enriched patients to seek expedited development in the United States and China. We plan to further assess the combination of ADG106 and an anti-PD-1/L1 mAb, or anti-CTLA-4 mAb for the treatment of solid tumors and hematologic malignancies in a clinical setting. We intend to build a CTLA-4 franchise by exploring the use of our product candidates as tolerable and potent monotherapies and in combination therapies with other cancer therapies, such as chemotherapy, targeted small molecule drugs, multiple-tyrosine kinase inhibitor drugs and other immune checkpoint inhibitors.
- **Develop a deep and broad pipeline of promising antibodies using NEObodyTM, SAFEbodyTM and POWERbodyTM technologies and advance into preclinical and clinical development.** We plan to continue to leverage our proprietary DPL platform to generate and select a broad clinical pipeline of novel and potentially differentiated product candidates with the goal of submitting more than ten INDs or equivalent applications in the next three to five years. We are also planning to conduct a basket trial to explore the synergistic effects of our product candidates in combination with other therapeutic agents, such as PD-1 or PD-L1 antibodies, to expand the market opportunity of our pipeline. We intend to prioritize product candidates based on a range of factors, including strength of preclinical data, potential for development as both a monotherapy and in combination with other therapies, clinical benefit, efficiency of clinical development paths and commercial market opportunities. In particular, we plan to leverage our differentiated NEObody, SAFEbody and POWERbody technologies to build a rich pipeline of product candidates with improved product profiles (e.g., enhanced safety and/or efficacy and/or dosing convenience such as less frequent subcutaneous injections) over existing therapies or targeting novel MOA.
- **Leverage our technology to develop our pipeline and strengthen our DPL platform.** We intend to exploit the conformational diversity of antibody discovery through the combination of our proprietary computational biology algorithms and artificial intelligence to design and discover leading oncology immunotherapies with differentiated product profiles. Leveraging our

technology together with our expertise in immuno-oncology and targeted therapy, we plan to build a rich pipeline of immuno-oncology antibody candidates directed against both clinically validated as well as potentially novel targets designed to overcome safety or efficacy deficiencies in the current standard-of-care. For example, we are advancing multiple NEObody and SAFEbody programs with potential unique MOA, which we intend to develop as monotherapies and/or as combination therapies. We intend to also devote resources to further strengthen our DPL platform, including enrichment of our DPL and masking peptide libraries and optimization of our computational biology algorithms. Through leveraging our enabling DPL technologies for high fidelity translational studies, we aim to accelerate the discovery and development of a deep, broad and differentiated pipeline of antibody-based therapeutic candidates in precision immuno-oncology.

- ***Continue to collaborate with leading biopharmaceutical companies and academic institutions to discover and develop novel candidates based upon our DPL platform.*** Collaboration is a key part of our growth strategy and we have established multiple collaboration programs in validating our platforms. We intend to continue to seek collaboration and partnership opportunities where we can leverage our proprietary technology platform to develop novel antibodies to address unmet medical needs. We also plan to explore our antibody programs in combination trials, and will opportunistically evaluate strategic collaborations or other partnerships to further these potential future clinical trials. Given the species cross-reactive design of our antibodies, we are able to take advantage of highly predictive animal models where we believe there is a potential for strong additive or synergistic behavior into the clinical setting.
- ***Maximize value creation by advancing our product candidates to potential commercialization in key markets alone or with strategic partners.*** We have retained exclusive worldwide rights to all of our product candidates other than ADG104, for which we out-licensed the Greater China rights to our partner Sanjin, and intend to pursue clinical development programs with the goal of obtaining regulatory approvals in the United States, China and major international markets. We intend to directly commercialize our product candidates, if approved, but may opportunistically enter into strategic collaborations or other partnerships with leading biopharmaceutical companies to accelerate our development timelines and maximize the global commercial potential of our product candidates.
- ***Build global operations for global markets, while leveraging a global supply chain and China cost effectiveness.*** We are a clinical stage biopharmaceutical company with global footprints. We have established subsidiaries in the United States, China, Hong Kong, Australia, Singapore and Switzerland to conduct and/or support preclinical studies and clinical trials and are seeking patent protection in a number of jurisdictions worldwide. In particular, our presence in mainland China enables us to access a large pool of target patients and creates the potential to progress our product candidates through clinical development in an efficient and cost-effective manner. We employ a global clinical development strategy and leverage our global supply chain to comply with the requirements applicable to clinical trials globally in accordance with the requirements of the FDA, the European Medicines Agency, or EMA, the Japanese Pharmaceuticals and Medical Devices Agency, or PMDA, and other comparable regulatory authorities. We are currently building our clinical and technology infrastructures to support our future global operations and prepare to serve global markets.

ANTIBODIES FOR THE TREATMENT OF CANCER

Cancer treatment has traditionally included chemotherapy, radiation, surgery or a combination of these approaches. Small molecule target therapy can be effective in certain types of cancer, but they can also cause toxicities that may have life-threatening consequences, lower quality of life or untimely termination of treatment. Furthermore, these small molecule agents offer limited efficacy in many types

of cancer. Over the last 20 years, a new paradigm of cancer research and treatment has emerged that involves targeted therapies, including mAbs. Monoclonal antibodies are proteins derived from living organisms that bind to targets, called antigens, on tumor cells to inhibit tumor growth. As a product class, immunotherapeutic mAbs have transformed oncology treatment and represent some of the most effective and top selling immunotherapies currently approved by the FDA and available on the market. The success of conventional immunotherapeutic mAbs has been hindered by limited efficacy and by safety and tolerability concerns. Administration of these mAbs may cause systemic side effects, as well as localized, organ-specific damage. Much of this toxicity is a direct consequence of the fact that healthy tissues express some of the same antigens that conventional immunotherapeutic mAbs target on cancerous cells.

More recently, immuno-oncology has emerged as a promising new field of cancer therapy that aims to enhance antitumor immune responses by, for example, overcoming mechanisms that cancer cells have developed to evade the immune system. Some cancer cells overly express proteins, called immune checkpoints, which apply brakes to the immune system, and enable the tumor cells to evade destruction. Immune checkpoint inhibitors nivolumab, pembrolizumab and ipilimumab—antibodies targeting these immune inhibitory proteins—release these brakes to allow the immune system to destroy tumor cells. These drugs have shown promising efficacy and are currently being explored for multiple solid and hematological cancer indications. Although these drugs have demonstrated promising results, only a minority of patients receive durable benefit from treatment with these mAbs alone. Most recently, combination regimens of immunotherapy agents have demonstrated signs of improved efficacy in larger numbers of patients. However, many of these combinations have significant toxicity and tolerability issues, due in part to the activation of the immune system in both healthy and cancerous environments.

In the past decade, new modalities of highly potent mAb-based therapies have emerged. ADCs represent one such modality. These agents are comprised of two functional units chemically fused or conjugated to each other: a cytotoxic drug payload and a mAb. ADCs combine the targeting abilities of the antibody with the cancer-killing ability of cytotoxic drugs, leading to better specificity in targeting tumor cells compared to traditional chemotherapy. Bispecific antibodies, another class of second-generation biologics, have the ability to simultaneously bind a cancer cell and a T-cell, leading to the destruction of the cancerous cell by the T-cell. This ability improves the potency of bispecific antibodies compared to first-generation immunotherapeutic mAbs.

While all of these potent new therapies have shown promise, none addresses a key limitation of antibody-based therapeutics—expression of targets in healthy tissue, which leads to off-tumor toxicity and limits clinical use.

OUR PIPELINE

By leveraging our proprietary DPL platform, we have developed a robust pipeline of innovative product candidates in various stages of development, ranging from research and discovery to preclinical and clinical development. Our highly differentiated pipeline consists of our lead clinical-stage candidates, ADG106, ADG126 and ADG116. We also have a robust preclinical pipeline in various stages of development. In addition, we have out-licensed the Greater China rights of ADG104, a PD-L1 mAb in Phase Ib and Phase II trials concurrently in China to our partner Sanjin. We have the right to apply for the patents derived from our core and key technologies pertaining to ADG104 in the rest of the world and we retain a majority of the economic benefits derived from ADG104 in the rest of the world.

ADG106: Novel Agonistic Anti-CD137 NEObody Candidate

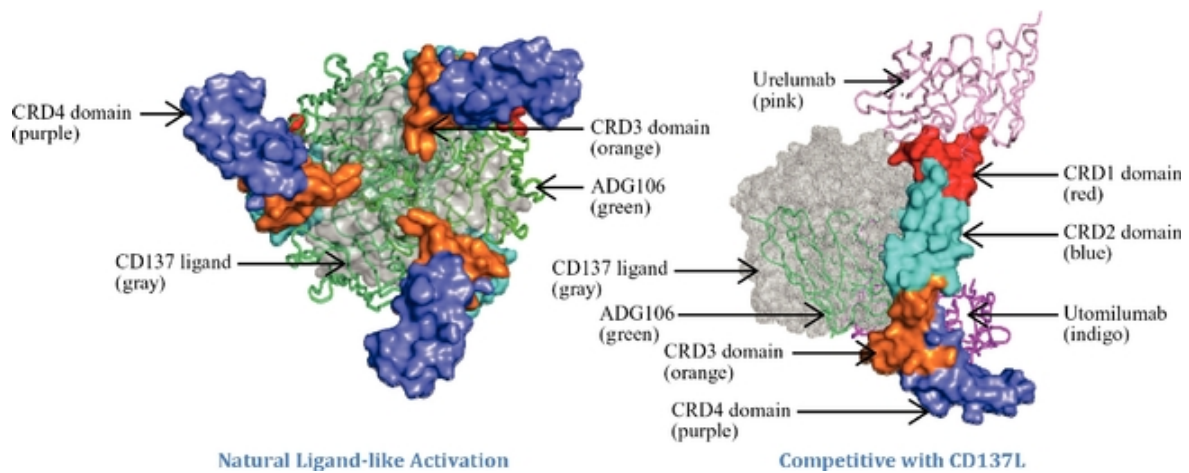
Summary

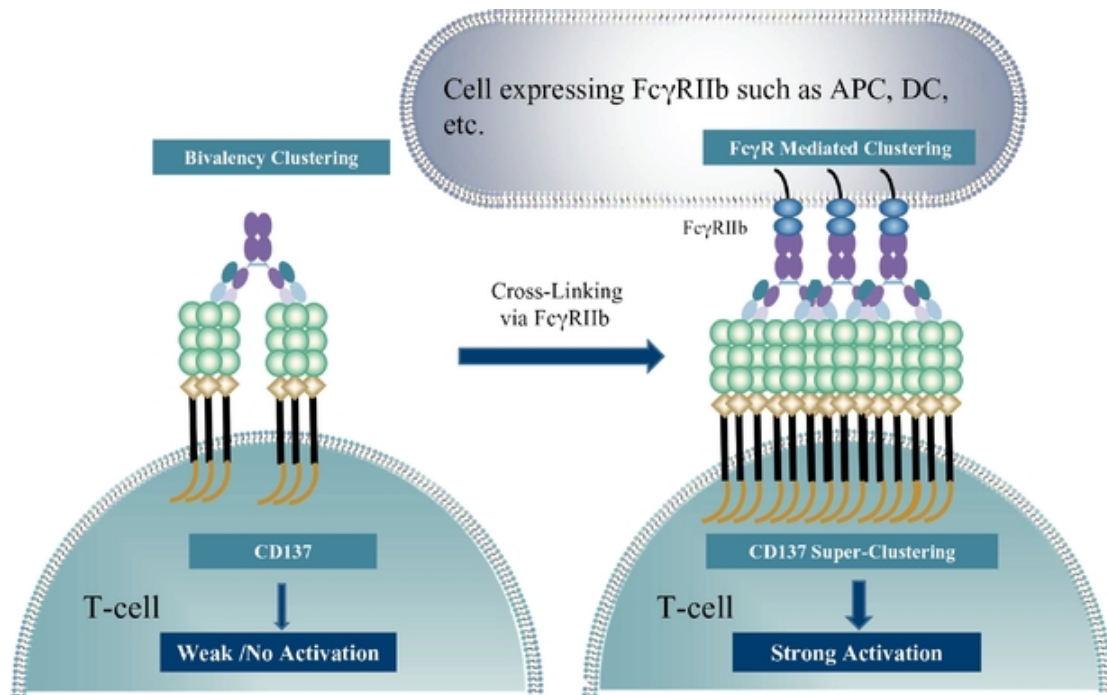
Our lead product candidate, ADG106, is a fully human ligand-blocking, agonistic anti-CD137 IgG4 mAb designed to target a unique conserved epitope of CD137. As of the Data Cut-off Date, we have completed the Phase Ia dose escalation in each of our Phase I studies of ADG106 as a monotherapy in patients with advanced or metastatic solid tumors and/or NHL in both the United States and China. ADG106 was well-tolerated in the completed Phase Ia dose escalation of the Phase I clinical trials. Early efficacy signals were also observed with tumor size reduction, significant target modulations and increased T-cell and NK proliferation, consistent with MOA.

Mechanism of Action

CD137 is a member of the tumor necrosis factor, or TNF, receptor superfamily. As illustrated in the figure below, the binding of an antibody to this receptor induces a co-stimulatory signal on activated enhanced cytotoxic T lymphocyte, or CD8⁺ T-cells, and natural killer, or NK cells, resulting in proliferation, and increased pro-inflammatory cytokine secretion and cytolytic function. CD137 co-stimulation is a clinically validated pathway for T-cell activation and its antitumor response is highlighted by the approval of a CD137-targeting CAR-T therapy by the FDA. Because most tumors are killed by cytotoxic T-cells in an antigen specific manner, we believe agents that mediate CD8⁺ T-cell activation can impart strong cytolytic activity. Therefore, we believe that CD137 agonists are promising candidates with potential to enhance and mediate long lasting antitumor immunity.

ADG106 is designed to bind to activated human CD4⁺ and CD8⁺ T-cells with low nanomolar affinity and block CD137 ligand binding in a concentration-dependent manner to disable its reverse signaling. As shown in the figure below, ADG106 is designed to target a unique conserved epitope of CD137 with a novel MOA for CD137 agonism by its natural ligand-like binding and cross-linking by Fcγ receptors. ADG106 has not been observed to bind to unstimulated naïve T-cells, which have no detectable level of CD137 expression. ADG106 is also designed to bind to activated NK cells to boost their cytotoxic and ADCC functions.



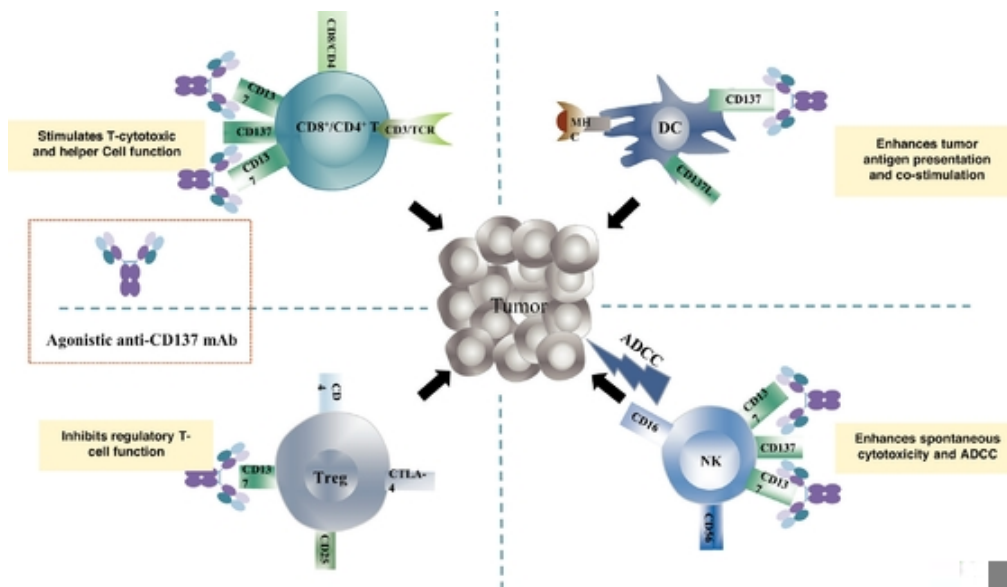


Notes:

"APC" refers to antigen-presenting cells;

"DC" refers to dendritic cells.

The MOA of ADG106 in tumor TME is illustrated below.



Notes:

"DC" refers to denritic cells;

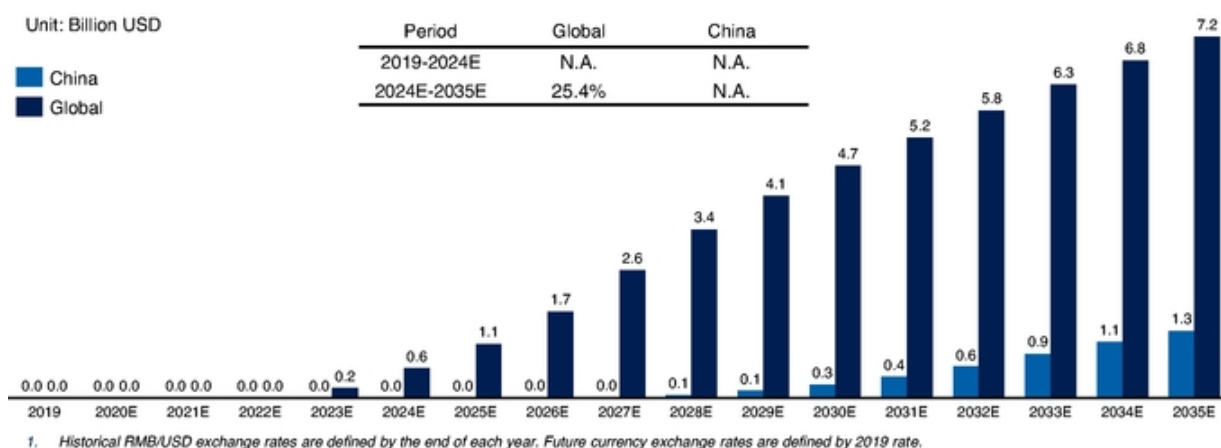
"Treg" refers to regulatory T-cell.

Market Opportunity and Competition

CD137 is an inducible costimulatory receptor expressed on activated T-cells in the TME. Agonistic mAbs targeting CD137 have been developed to harness CD137 signaling for cancer immunotherapy. An anti-CD137 agonist would possess the potential to target a wide spectrum of cancer types both as a monotherapy and in combination with various other therapies, especially in the chemotherapy-free setting, including anti-PD-1, anti-PD-L1 and anti-CTLA-4 antibodies, three proven immune checkpoint inhibitors. Although we believe there is compelling evidence that supports the therapeutic potential of a CD137 agonist, there are currently no marketed CD137 agonist drugs. According to Frost & Sullivan, there are two advanced CD137 agonist antibodies in clinical development. However, clinical development of one of these antibodies has been hampered by inflammatory liver toxicity, despite initial signs of efficacy. The other CD137 antibody under clinical development has demonstrated better safety features, but is a less potent CD137 agonist. One of the key challenges of CD137 agonist clinical development is CD137 agonist specific toxicity. Previous clinical trials have shown CD137 antibodies cause immune anomalies, notably polyclonal activation of CD8+ T-cells and secretion of inflammatory cytokines, which affect the function of the liver, spleen, and bone marrow. We believe that it is therefore critical to develop therapeutic candidates that are designed to maximize potency of CD137 agonism while minimizing CD137 agonist specific toxicity.

According to the Frost & Sullivan Report, the global CD137 monoclonal antibody market will emerge in 2023 and is estimated to reach US\$0.6 billion and US\$7.2 billion in 2024 and 2035 respectively, with a CAGR of 25.4% from 2024 to 2035. The China CD137 monoclonal antibody market is estimated to emerge in 2028 and is expected to grow to US\$1.3 billion in 2035.

Global and China CD137 Antibody Market, 2019-2035E



Source: Frost & Sullivan Report

Summary of Clinical Trial Data

Overview: We have completed Phase Ia dose escalation and are conducting cohort expansion in each of our two Phase I trials of ADG106 as a monotherapy in patients with advanced or metastatic solid tumors and/or NHL in the United States, or the ADG106-1001 clinical trial, and in China, or the ADG106-1002 clinical trial. As of the Data Cut-off Date, ADG106 was dose escalated up to 10 mg/kg and had been observed to be well-tolerated at dose expansion at 3 mg/kg, 5 mg/kg and at 300 mg and 400 mg flat doses for a total number of 92 patients dosed. Liver enzyme elevation of ³ Grade 3 was reported in three patients, with one determined to be not treatment related. We also observed tumor shrinkage in some patients.

Trial Design: Our Phase I clinical trials were designed to evaluate the safety and tolerability of ADG106, as well as its pharmacokinetics, or PK, immunogenicity, and preliminary clinical response. There were two parts to the dose escalation study: the accelerated titration phase at 0.03, 0.1, and 0.3 mg/kg and a conventional "3+3" dose escalation phase at 1, 3, 5 and 10 mg/kg. ADG106 was administered once every three weeks by intravenous infusion.

Dose-expansion cohorts started at dose levels that were observed to be well-tolerated in the dose escalation phase and showed evidence of clinical and biological activities. Patients with advanced or metastatic solid tumors or NHL, who were refractory or relapsed after exhausting almost all available therapies, were enrolled for a maximum of 24 months unless they withdrew earlier due to disease progression, intolerable toxicity, or withdrawal of consent. The protocol allowed for up to 52 patients in ADG106-1001 and 124 patients in ADG106-1002 to be enrolled in the accelerated titration, dose escalation, and dose expansion phases.

The primary objective of the Phase I clinical trials, including both the dose escalation phase and dose expansion phase, is to assess safety and tolerability at increasing dose levels of ADG106 as a monotherapy in patients with advanced or metastatic solid tumors and/or NHL. The secondary objectives of the Phase I clinical trial are to characterize the PK profile of ADG106 and to evaluate the immunogenicity of ADG106 and potential antitumor activity of ADG106. An exploratory objective of such Phase I clinical trial is to identify potential biomarkers of patients who respond to treatment with ADG106.

The endpoints of ADG106 Phase I trial are:

Primary endpoint:

- Dose-limiting toxicities, or DLT, in the first 2 cycles.

Secondary endpoints:

- Safety endpoints include AEs, clinical laboratory results, vital signs, physical examination findings, and ECG results.
- Objective response rate, or ORR, duration of response, or DOR, time to progression, or TTP, disease control rate, or DCR, progression-free survival, or PFS, and overall survival, or OS, as assessed by Response Evaluation Criteria in Solid Tumors, or RECIST, version 1.1, and/or immune-related RECIST, or irRECIST, for solid tumors, and the Lugano Classification for NHL/ Hodgkin's lymphoma.
- PK endpoints include peak serum concentration, or C_{max}, serum concentration at the end of a dosing interval, or C_{trough}, time to reach C_{max}, or T_{max}, area under the curve from time zero to the last timepoint, or AUC_{0-last}, AUC from time zero to infinity, or AUC_{0-inf}, AUC during a dosing interval, or AUC_{tau}, clearance, or CL, and volume of distribution at steady state, or V_{dss}, as data permit.
- Anti-drug antibody, or ADA, levels for ADG106.

Status: As of Data Cut-off Date, the Phase Ia dose escalation portion of the ADG106-1001 Phase I clinical trial in the United States and the ADG106-1002 Phase I clinical trial in China had been completed and the Phase Ib/II dose expansion phase for both trials was ongoing. We plan to complete the Phase I trials and initiate biomarker-driven global Phase II trials in 2021.

As of the Data Cut-off Date, 92 patients were enrolled, among whom 35 patients were enrolled and treated in the ADG106-1001 clinical trial and 57 patients were enrolled and treated in the ADG106-1002 clinical trial. Thirty-eight patients were enrolled in the dose escalation phase and

54 patients were enrolled in the dose expansion phase. Patients in the expansion cohort of both trials were still on the study treatment. The patient demographics and baseline parameters are listed below:

Patient Demographics	ADG106-1001 (n=35)	ADG106-1002 (n=57)
Age (Years)		
Median (Max, Min)	61(82, 25)	50(72, 21)
Sex		
Female	19	19
Male	16	38
Race		
African American	1	
Chinese	3	57
Caucasian	31	
Eastern Cooperative Oncology Group ("ECOG") Performance Status		
0	3	45
1	32	12
Number of Prior Systemic Regimens		
Median (Min, Max)	5 (0, 14)	3 (0, 10)
Cancer Types		
	Colorectal Cancer (n=7)	Nasopharyngeal Cancer (n=14)
	Renal Cell Carcinoma(n=4)	Non-small Cell Lung Cancer (n=13)
	Follicular Lymphoma (n=3)	Non-Hodgkin's lymphoma n=10)
	Head and Neck Squamous Cell Carcinoma (n=3)	Adenoid Cystic Cancer (n=5)
	Esophageal Carcinoma (n=2)	Cervical Cancer (n=4)
	Anal Cancer (n=2)	Others (n=11)
	Other (n=14)	

Safety Assessment: As of the Data Cut-off Date, 35 patients and 57 patients were enrolled in the two Phase I trials in the United States and China, respectively. ADG106 is generally well tolerated at 3mg/kg and 5mg/kg with dose escalation up to 10mg/kg. We observed limited TEAEs, liver toxicity or hematologic abnormalities in both trials.

As of the Data Cut-off Date, 35 patients enrolled in the ADG106-1001 clinical trial in United States had safety assessments. 24 of 35 (69%) patients discontinued ADG106 treatment (19 patients discontinued due to progression disease, 3 patients discontinued due to adverse events). Two DLTs were observed, including a Grade 2 hypoglossal nerve disorder at 5 mg/kg and a Grade 3 adrenal insufficiency at 10mg/kg cohorts. Across all dose cohorts, liver function tests showed two patients with >Grade 3 AST increase, including one patient who had abnormal baseline liver enzyme exhibited Grade 3 AST increase. Hematologic tests showed two patients with Grade 3 hemoglobin decreases, and five patients had Grade 3 lymphocyte decreases.

As of the Data Cut-off Date, 57 patients enrolled in the ADG106-1002 clinical trial in China had safety assessments. 45 of 57 (68.9%) patients discontinued ADG106 treatment (31 patients discontinued due to progression disease, 7 patients discontinued due to adverse events, 5 due to withdrawal of consent and 2 due to clinical deterioration). One DLT (Grade 4 neutrophil count decreased, 10mg/kg) was observed in one patient with mantle cell lymphoma. Across all dose cohorts of 57 patients, only one patient showed Grade 3 AST and ALT liver enzyme increases. Two patients dosed at 5mg/kg or 10mg/kg experienced Grade 3 and 4 hemolytic anemia, respectively. Both patients were treated and subsequently recovered.

Clinical Efficacy Assessment: As of the Data Cut-off Date, 71 of 92 patients enrolled in both trials received post-treatment scans and were evaluated for preliminary efficacy assessment. The responses observed in these patients included one partial response, 39 stable disease, and 31 progressive disease. The disease control rate was 56% (40 of 71 patients). Three patients enrolled in ADG106-1002 clinical trial achieved greater than 30% tumor shrinkage. One patient with a solid tumor had a partial response

with an approximately 40% tumor size reduction in the target lesions and two NHL patients showed tumor shrinkage of 32% and 33% at the end of the Cycle two. A third ADG106-1002 NHL patient showed a tumor size reduction of 17% and a patient with esophageal carcinoma enrolled in ADG106-1001 achieved 18% tumor shrinkage.

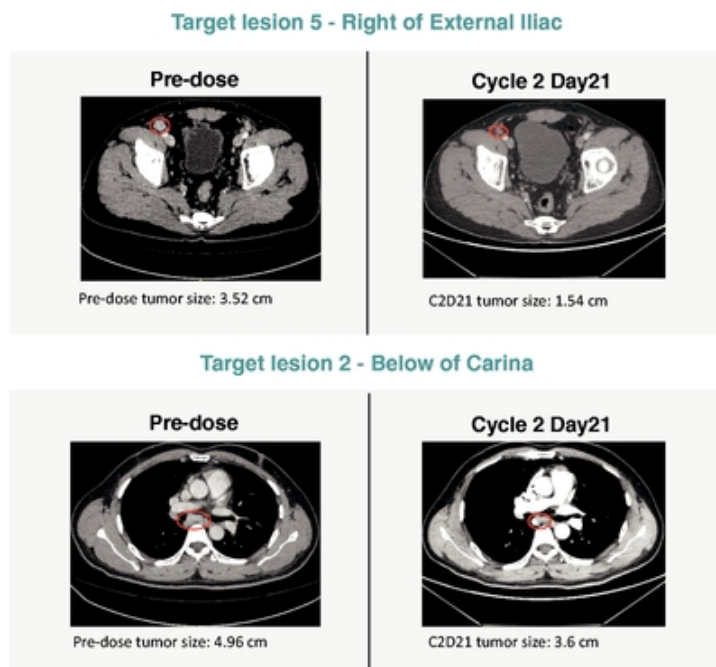
As of the Data Cut-off Date, of the 71 patients who received post-treatment scans, 23 were enrolled in ADG106-1001 clinical trial. The observed responses were 12 stable disease and 11 progressive disease and a disease control rate of 52% (12 of 23 patients). Tumor shrinkage and a decrease in standard uptake value scores on PET CT images were observed in three patients, including one patient with stage IV follicular lymphoma who was refractory to multiple prior therapies including rituximab, another patient with endometrial carcinoma, and a third patient with esophageal carcinoma.

As of the Data Cut-off Date, of the 71 patients who received post-treatment scans, 48 were enrolled in ADG106-1002 clinical trial. The observed responses were one partial response, 27 stable disease, and 20 progressive disease with a disease control rate of 58% (28 of 48 patients). One patient with nasopharyngeal carcinoma demonstrated partial response with an approximately 40% tumor size reduction in the target lesion.

All three patients with T cell lymphoma enrolled in ADG106-1002 responded to ADG106 treatment and showed tumor size reductions of 32%, 33%, and 17%. Given its unmet medical needs, we are exploring Orphan Drug Designation with the FDA. In addition, we are collaborating with principal investigators who specialize in T cell lymphoma to conduct both investigator-initiated trials and an Adagene-sponsored trial to further validate the anti-tumor activities of ADG106 in this indication. We believe these efforts may lead to expedited regulatory approval of ADG106 for T cell lymphoma.

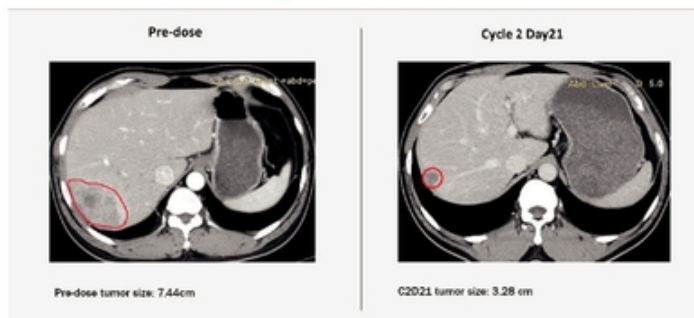
In conclusion, we observed a disease control rate of 56% among 92 patients including one patient with refractory nasopharyngeal carcinoma who had failed four prior therapies including anti-PD-L1 and four other patients with deep responses that exhibited 33%, 32%, 18%, and 17% tumor size reductions. Six patients remained on ADG106 treatment for over six months including one patient who received 3 mg/kg ADG106 treatment for over one year.

Case 1: Patient A with NHL, who relapsed after multiple chemotherapies and stem cell transplantation achieved 33% tumor size reduction after one dose of ADG106 at 3 mg/kg.

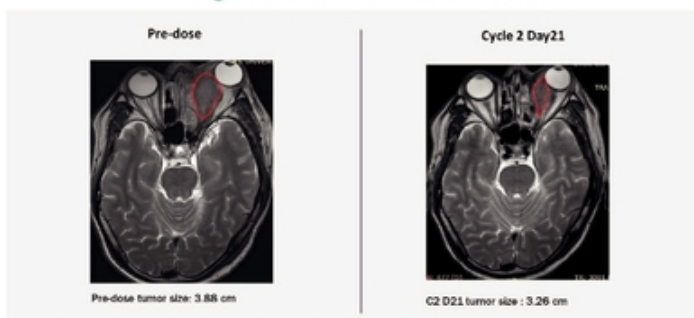


Case 2: Patient B with solid tumor (stage IV), who relapsed after multiple lines of treatment (chemotherapies, radiotherapy, target therapy, and immune checkpoint inhibition therapy), showed 40% tumor size reduction after two doses of ADG106 at 5 mg/kg.

Target Lesion 1 – Liver

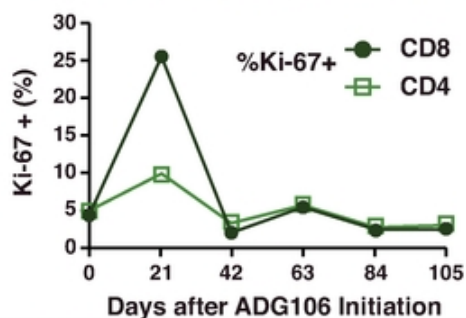


Target Lesion 3 – Left Ethmoid Sinus



Pharmacodynamic Biomarker Measure of CD137 Target Modulation: As shown below, we have observed stimulation of T-cell proliferation and expansion in one patient treated with ADG106, evidenced by an increase in percentage of Ki-67 protein (an indicator of proliferative activity).

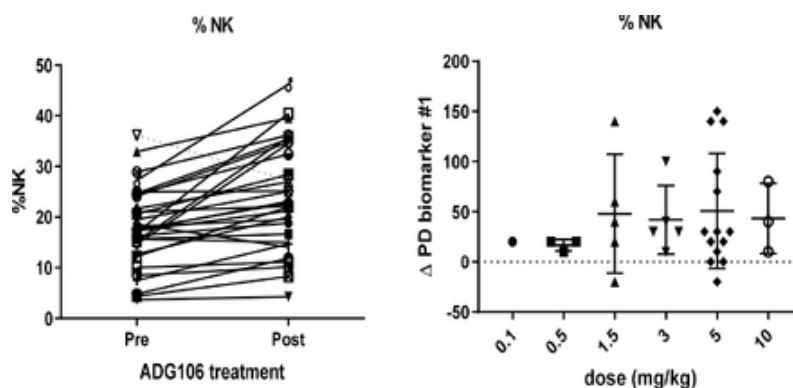
ADG106 (180mg) for Patient D



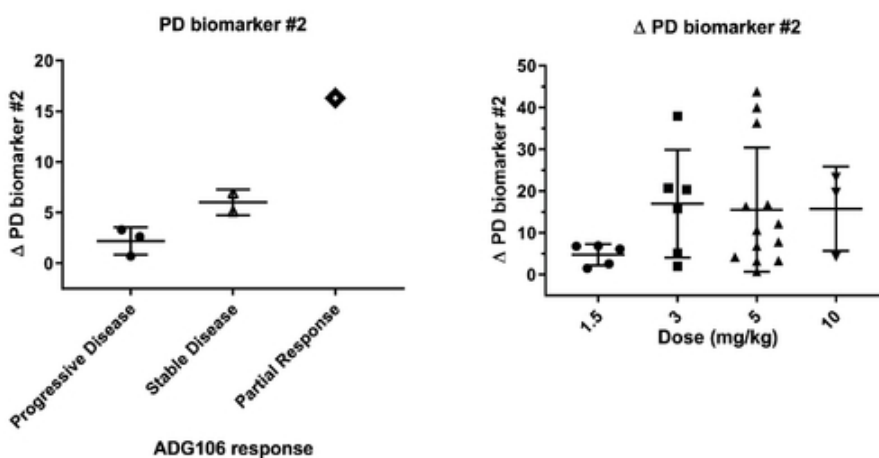
Stimulation of T-cell proliferation upon ADG106 administration

As illustrated below, we have also observed increased proliferation of NK cells in patients treated with ADG106 after the first cycle of treatment. Dose dependent stimulations of the identified immune cell subpopulation following ADG106 treatment at doses of 0.1, 0.5, and 1.5 mg/kg (see figure below on the right) were observed. No significant difference was observed from 1.5 to 10 mg/kg, suggesting that the drug may have saturated CD137 receptors on the target cells at the doses ≥ 1.5 mg/kg. This is the

first report of an anti-CD137 antibody drug showing stimulation of NK cell proliferation in human. This finding also clarifies the MOA of ADG106, suggesting involvement of diverse immune cells in ADG106 mediated anti-tumor activities in human.

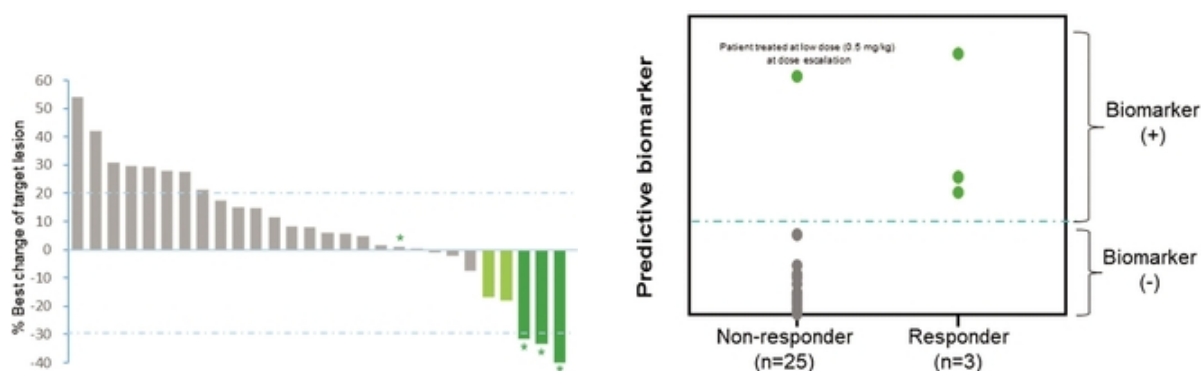


Our ADG106 Phase I clinical trials enrolled cancer patients without regard to cancer types. As shown in the figure below on the left, the effect of ADG106 on soluble CD137 was not tumor type specific, but rather a universal event observed for most of the patients. As seen in the figures below, we observed dose-dependent increases in the biomarker level in patient serum after ADG106 treatment and correlation of the change of the soluble CD137 with patient response.



Identification of a Potential Predictive Biomarker for ADG106 Treatment Response: We have identified a potential predictive biomarker that correlated with patient treatment responses based on a retrospective analysis of pretreatment biopsy specimens from 28 patients with clinical outcome data as shown in the diagram below on the right panel. As of the Data Cut-off Date, three patients who achieved greater than 30% tumor shrinkage after ADG106 treatment were found to be biomarker positive. One Phase Ia patient who was treated with a low dose of ADG106 was biomarker positive and

showed stable disease, as shown in the waterfall plot below on the left panel. Twenty-four biomarker negative patients did not show significant response as measured by tumor size reduction in CT images.



We have developed a high sensitivity multiplex IHC assay to measure the expression of the predictive biomarker. We measured expression of the predictive biomarker in 746 tumor microarray, or TMA, samples from patients with 14 different tumor types and identified biomarker positive expression of up to 42% in various types of solid tumors and hematological malignancies.

We will stratify and preselect patients using this biomarker in the upcoming Phase II clinical trials in 2021. In the early stage of the Phase II clinical trials, biomarker positive and negative patients will be identified using our multiplex IHC clinical trial assay in order to prospectively validate the predictive biomarker. After an interim analysis, we intend to collaborate with a diagnostic partner to identify patients for clinical trial enrollment using an analytically validated investigational diagnostic assay in the next stage of the Phase II clinical trials. This tumor biomarker test may be developed as companion diagnostic to facilitate AD106 registration according to FDA requirements.

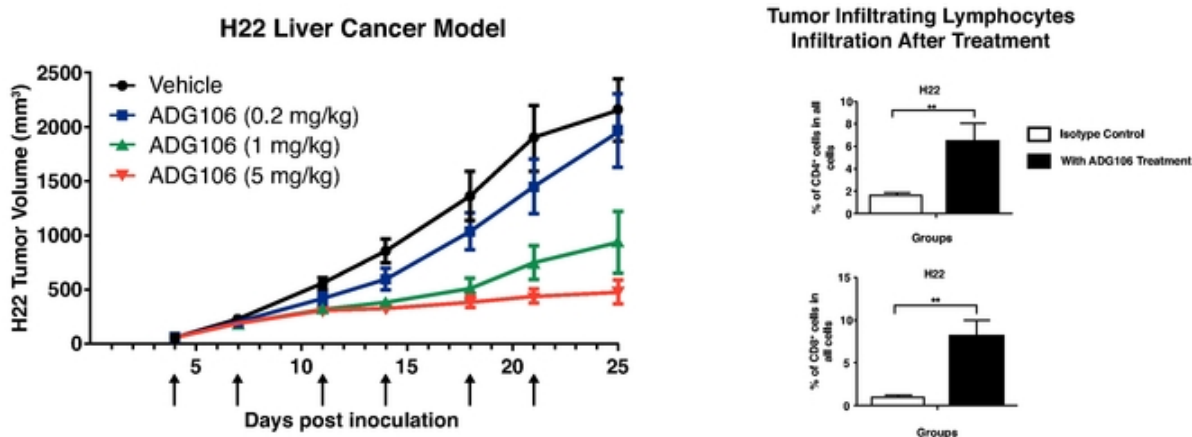
Summary of Preclinical Studies and Results

Overview: ADG106 was evaluated extensively in both *in vitro* and *in vivo* preclinical studies. ADG106 was observed in *in vitro* studies to enhance activation of, and cytokine release by, primed T-cells alone or in combination with other immunomodulatory agents. ADG106 was also observed to have antitumor activities *in vivo* as a monotherapy in a dose dependent manner in multiple tumor models. When combined with a variety of cancer therapeutics, ADG106 exhibited antitumor activity *in vivo* in animal models, including in immunotherapy resistant models. Our mechanistic analyses suggest that ADG106 promotes an antitumor response by stimulating infiltration and expansion of CD4⁺ and CD8⁺ T-cells in tumors. ADG106 was well tolerated in animals, with a no-observed-adverse-effect-level, or NOAEL, ³ 100 mg/kg/dose and ³ 200 mg/kg/dose in rats and cynomolgus monkeys, respectively. These findings support our belief that the ADG106-boosted immune response could offer an effective alternative solution for cancer immunotherapy as a monotherapy and in combination with other therapies, especially for PD-1/PD-L1 resistant patient populations.

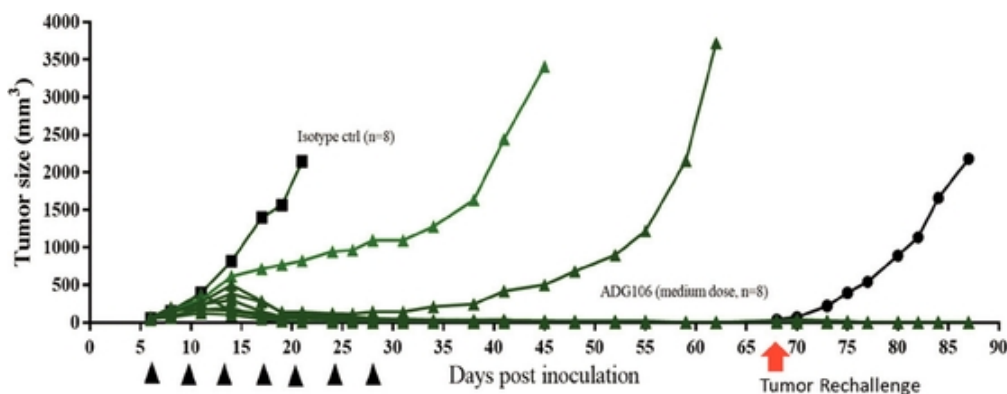
Preclinical Pharmacology: Our *in vitro* studies showed that ADG106 bound with high affinity and specificity to a unique epitope of CD137 that blocks the CD137L ligand binding with subsequent enhancement and proliferation of T-cells and pro-inflammatory interferon-g responses. Interferon-g is a master checkpoint regulator for many cytokines that targets a common component of many heterodimeric cytokine receptors.

ADG106 was species cross-reactive against human, mouse, and cynomolgus monkey CD137 and was active as a monotherapy in multiple preclinical tumor models. As shown below (left figure), in an H22 liver cancer syngeneic mouse model, ADG106 was observed to have *in vivo* antitumor activities as

a monotherapy in a dose dependent manner. As shown below (right figure), tumors treated with ADG106 showed increased CD4⁺ and CD8⁺ infiltration compared to an isotype control.



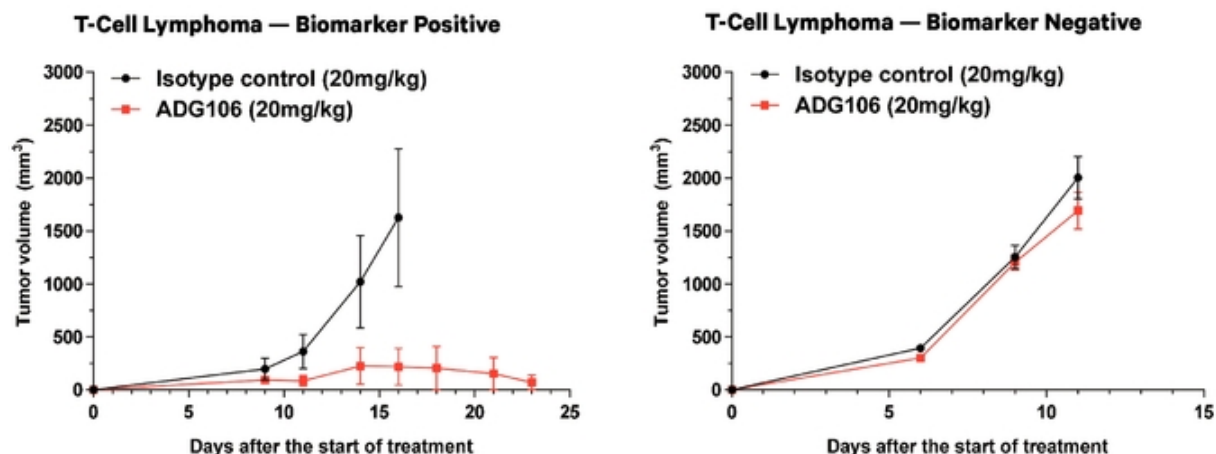
As illustrated in the figure below, ADG106 also showed a durable antitumor response in a CT26 colon cancer syngeneic mouse model. ADG106 treatment induced a complete response, or CR, in six out of eight mice. These CR mice received reinoculation of CT26 tumor cells, or rechallenge, and remained tumor-free without additional ADG106 treatment, indicating development of antitumor memory response elicited by ADG106. Tumors treated with ADG106 showed increased CD4⁺ and CD8⁺ infiltration compared to an isotype control, similar to the H22 liver cancer model (data not shown). These data are consistent with the mechanism of ADG106 as a CD137 agonist in stimulating T-cell proliferation, activation, and infiltration into the TME to induce an antitumor effect.



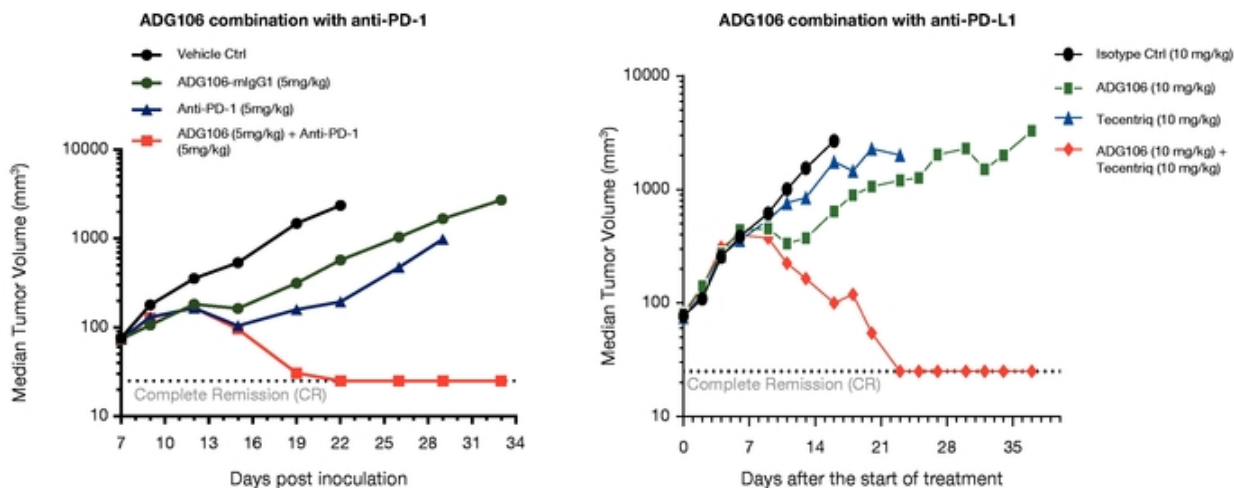
- durable response in syngeneic mouse colon cancer model
- CR previously dosed with ADG106 showed no tumor growth upon tumor re-challenge

Patient stratification: Understanding which patient populations may preferentially benefit from ADG106 treatment is important in designing clinical trials. Our preclinical studies have identified a biomarker that is associated with *in vivo* sensitivity of tumor models to ADG106 treatment. We determined biomarker level in ten syngeneic mouse tumor models and tested ADG106 *in vivo* in these ten models. In the seven models that were biomarker positive, four models were sensitive to ADG106

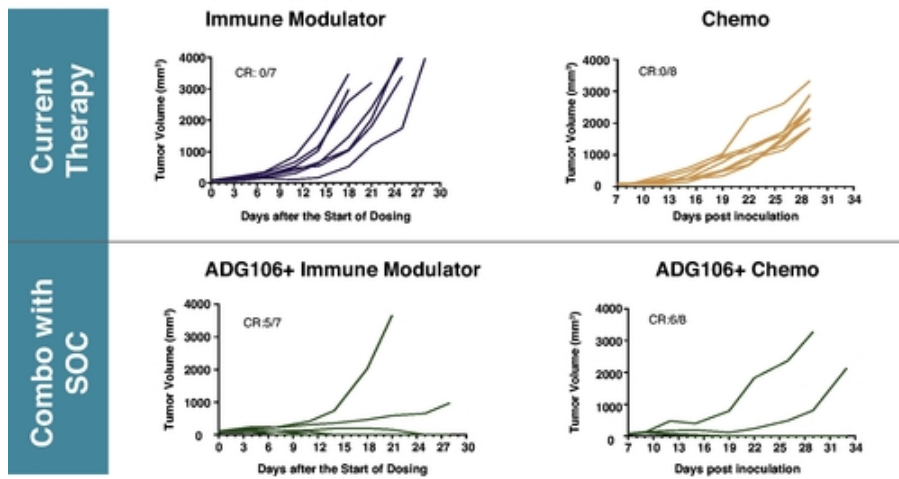
treatment while three were resistant or less sensitive to ADG106. The three biomarker negative models were resistant to ADG106 treatment. Additionally, the three resistant or less sensitive biomarker positive models showed synergistic effect to ADG106 treatments in combination with anti-CTLA-4 or anti-PD-L1. As illustrated in the figures below, among two T-cell lymphoma models, the biomarker positive model was observed to be sensitive to ADG106 treatment (see figure below on the left) and the biomarker negative model was observed to be resistant to ADG106 treatment (see figure below on the right). This biomarker provides a potential mechanism for stratifying patients that may benefit from ADG106 treatment in future clinical trials.



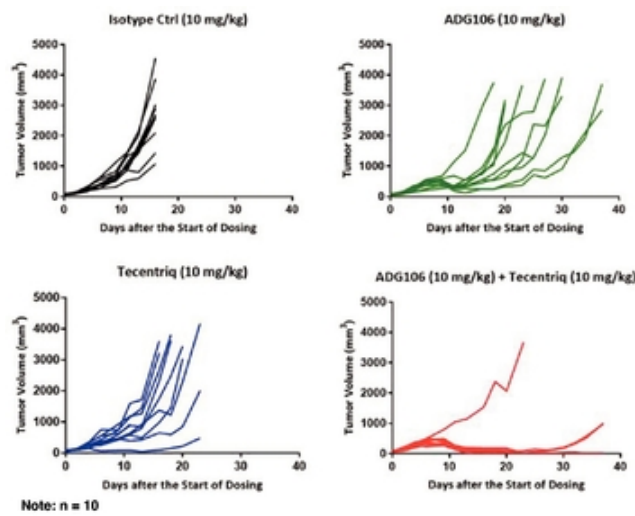
In addition to monotherapy antitumor activities, we observed that ADG106 had antitumor activities in combination with other therapies in preclinical studies. As shown below, ADG106 produced a synergistic effect with an anti-PD-1/L1 therapy, and SOC, in *in vivo* tumor models, including models that are resistant to current PD-1/PD-L1 and SOC therapies.



Combination with Standard of Care

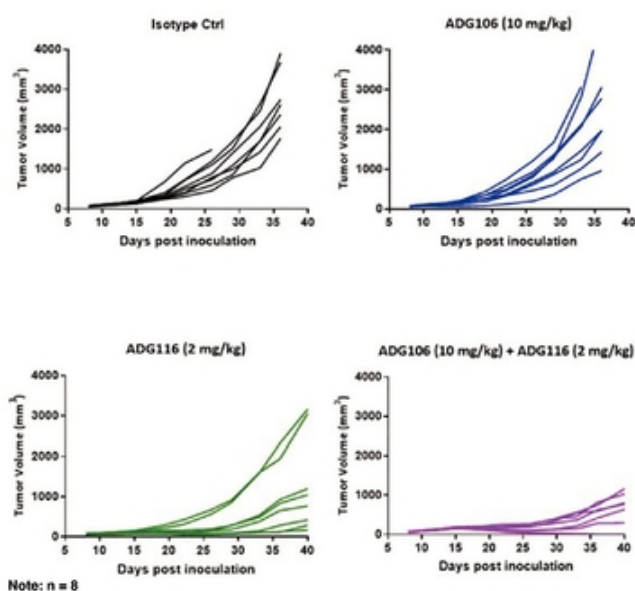


In a syngeneic tumor that is biomarker (mentioned above) positive, although ADG106 did not show anti-tumor activity in this model, combination of ADG106 with atezolizumab (Tecentriq), an approved anti-PD-L1 antibody that blocks PD-1/PD-L1 interaction, showed enhanced antitumor response compared to the corresponding monotherapies.



As demonstrated below, ADG106 was also observed to show antitumor activities when combined with our CTLA-4 antibody ADG116 in a mouse tumor model that is biomarker (mentioned above)

positive, while ADG106 alone did not show antitumor activities. We believe that this data provides a strong rationale for combining our pipeline programs including ADG106 with ADG116.



Clinical Development Plan

We have completed the Phase Ia dose escalation portion of each of our Phase I clinical trials and are conducting a Phase Ib cohort expansion phase using the selected doses in several indications. Our ADG106 Phase Ia clinical trials are designed to evaluate the safety, tolerability, and PK of repeat ascending doses of ADG106 in advanced or metastatic solid tumors and NHL. The secondary objective of these trials is to characterize the PK profile of ADG106, to evaluate and potentially identify a maximum tolerated dose or optimal dose of ADG106, assess relevant biomarkers, and evaluate preliminary signs of antitumor activity.

Our ongoing Phase Ib clinical trials are designed to evaluate the preliminary signs of antitumor activity. Based on Phase Ib data, we expect to choose the indications to be further explored in our future trials, for which plan to stratify and preselect patients based on their biomarker status to improve the response rate of ADG106. As a part of our strategies to ensure success of ADG106 clinical development, we may also explore niche indications for expedited approval. We may consider testing different dosages, different dosing schedules as well as ADG106 as a monotherapy or in combination with other immunotherapies in different disease subgroups and/or different disease stages. There are also investigator-initiated trials of ADG106, sponsored in Singapore by investigators, targeting various indications.

We are evaluating combinations of ADG106 with various therapies in clinical trials. We have received NMPA approval for the ADG106-1008 study in China which is a Phase Ib/II combination trial of ADG106 with anti-PD-1 in advanced solid tumors and hematological malignancies. We are also in the preparation for a submission of a CTN for ADG106-1003 study in Australia which is to combine ADG106 with anti-PD-1, anti-VEGF, and anti-CD20 antibodies in advanced solid tumors and hematological malignancies.

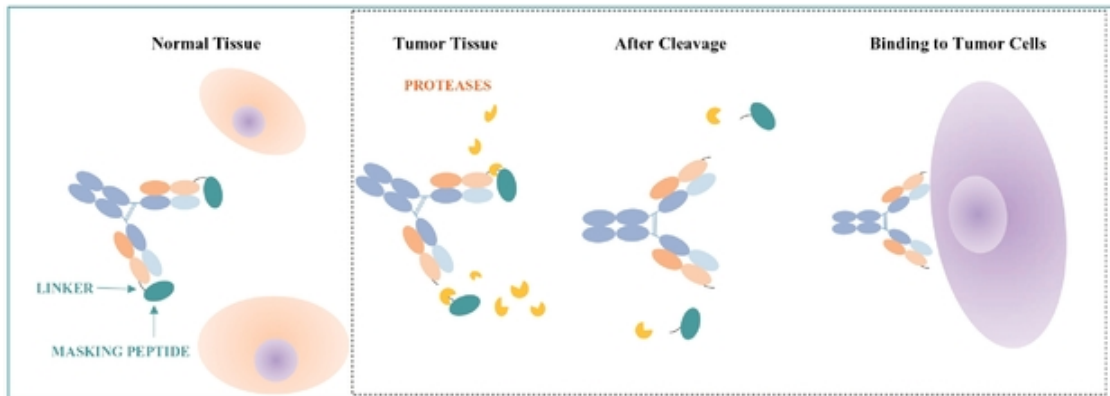
ADG126: Novel Anti-CTLA-4 SAFEbody Candidate

Summary

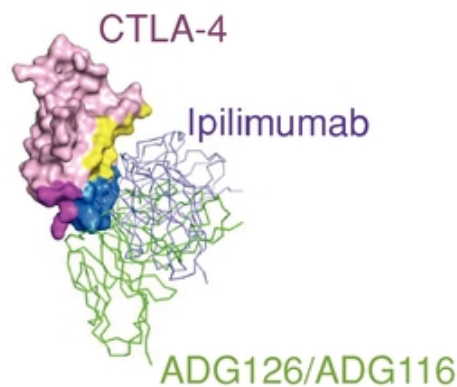
Our most advanced SAFEbody program, ADG126, is a fully human anti-CTLA-4 mAb generated using our SAFEbody technology designed to address the safety concerns with associated existing CTLA-4 therapeutics, while maintaining efficacy in the TME. Our proprietary SAFEbody technology is designed to enable ADG126 to be activated only in tumor tissues rather than healthy tissues. In preclinical studies, we observed that ADG126 had an enhanced therapeutic window and improved safety features. Furthermore, in PD studies of ADG126, it was observed that while the CTLA-4 binding affinity was masked in an intact ADG126 antibody, once the masking peptide was cleaved off of ADG126, its high binding affinity to CTLA-4 was restored.

Mechanism of Action

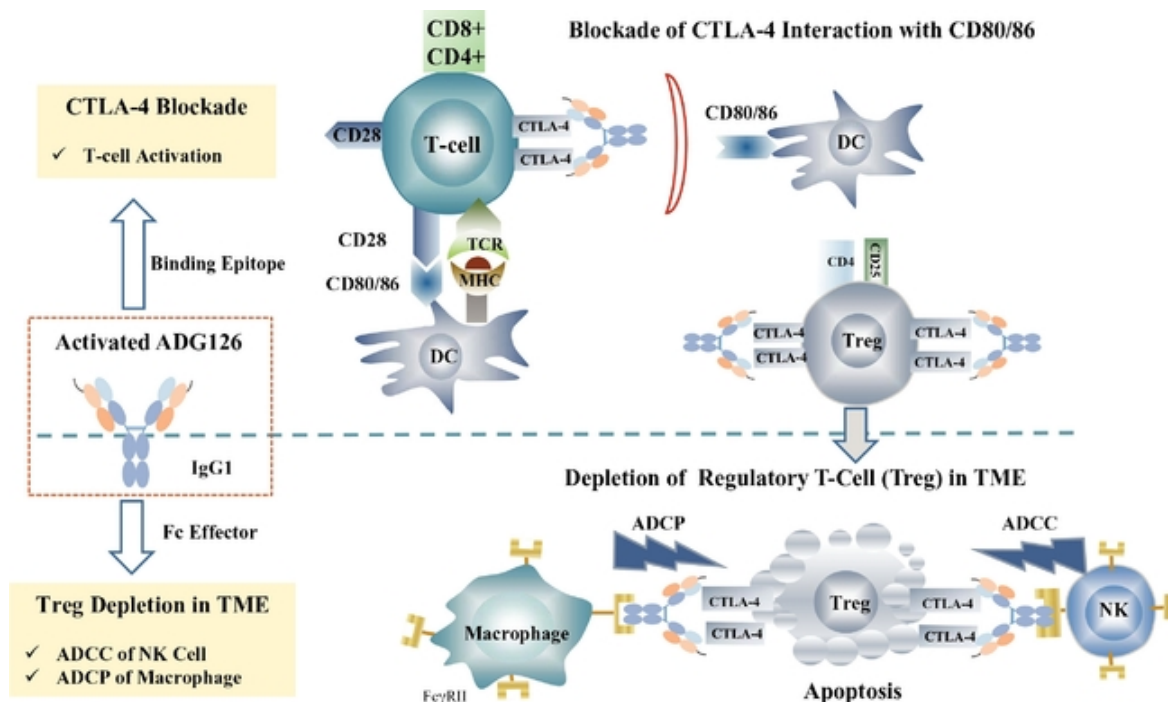
ADG126 is a fully human anti-CTLA-4 mAb. The masking moiety in ADG126 functions to block the interaction between ADG126 and its target CTLA-4 protein. Once ADG126 enters the TME, proteases overexpressed in the TME cleave off the masking moiety, and the antibody is then activated, binding to CTLA-4 and inhibiting its function. ADG126 is locally activated specifically in the TME, rather than systemically, to stimulate antitumor immune response. The following diagram illustrates the activation process of ADG126 in tumor tissue.



We believe that activated ADG126 potentiates T-cell immune response by blocking the inhibitory effect of CTLA-4. ADG126/ADG116 is designed to target CTLA-4 conserved epitope with species cross-reactivity for translational fidelity.



Moreover, ADG126 has been observed in preclinical animal studies to mediate effector functions to eliminate CTLA-4 expressing cells, particularly regulatory T-cells, primarily through ADCC. These actions of ADG126 could lead to enhanced activation and proliferation of tumor infiltrating T-effector cells and reduced T-regulatory cell function, which may contribute to a general increase in T-cell responsiveness, including an enhanced antitumor immune response. The below diagram illustrates the MOA of activated ADG126.



Notes:

"ADCP" refers to antibody-dependent cell-mediated phagocytosis;

"MHC" refers to major histocompatibility complex;

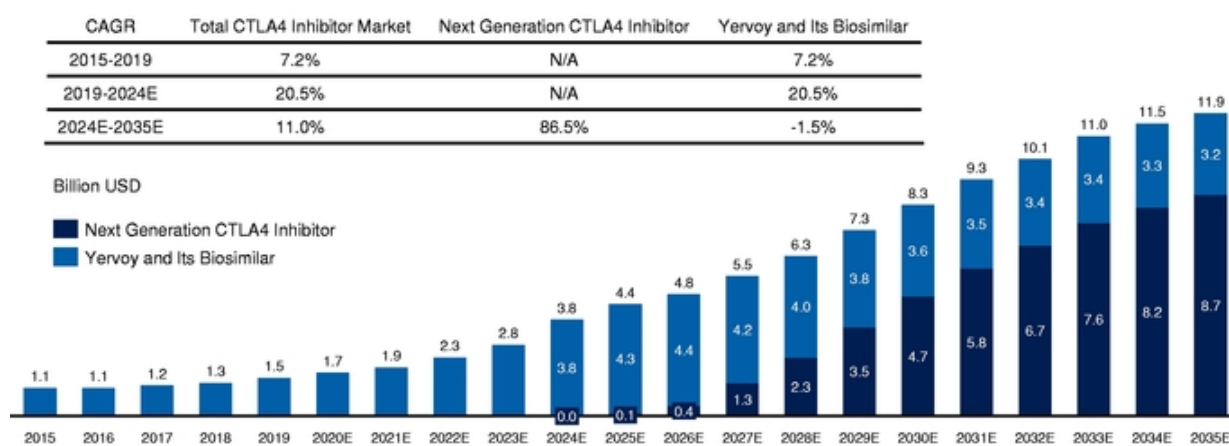
"TCR" refers to T-cell receptor;

Market Opportunity and Competition

According to Frost & Sullivan, as of December 31, 2020, ipilimumab is the only marketed CTLA-4 drug targeting cancer and it has been approved for six indications including monotherapy and combination therapies approved by the FDA. While ipilimumab is the only marketed CTLA-4 antibody drug, there are over ten CTLA-4 antibodies in clinical development globally, according to Frost & Sullivan. There are currently no marketed CTLA-4 antibody drugs in China, but at least seven CTLA-4 antibodies in clinical development, according to Frost & Sullivan.

The global total CTLA-4 inhibitors market reached US\$1.5 billion in 2019 and is estimated to reach US\$3.8 billion and US\$11.9 billion in 2024 and 2035, respectively, representing a CAGR of 11.0% from 2024 to 2035, according to the Frost & Sullivan Report.

Breakdown of Global CTLA-4 Inhibitors Market By Category, 2015-2035E



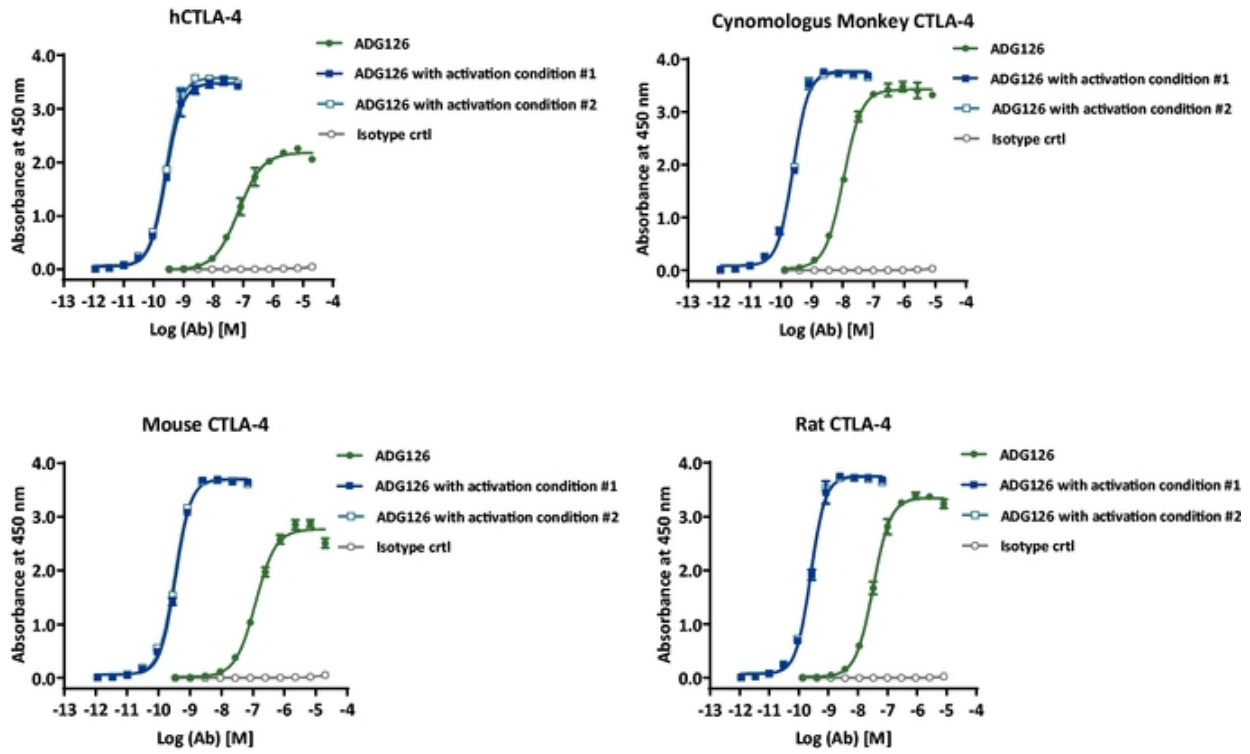
Source: Frost & Sullivan Report

Primary limitation of CTLA-4 antibodies is toxicity. We believe that the continued expansion of indications, and the launch of innovative novel CTLA-4 antibodies with potential for improved safety and better efficacy may increase the market for CTLA-4 antibodies significantly.

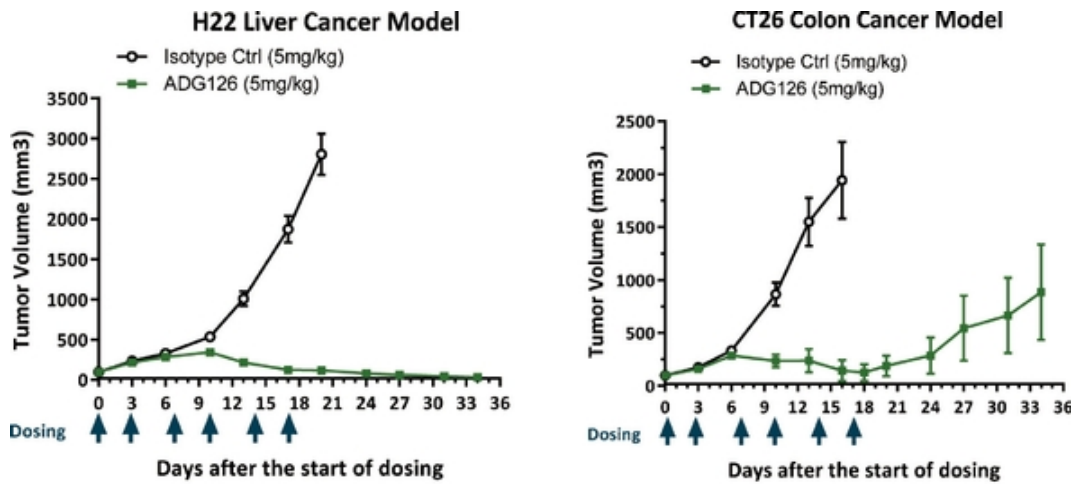
Summary of Preclinical Studies

Preclinical Pharmacology: We observed in PD studies that ADG126, in its intact SAFEbody form, bound weakly to CTLA-4. However, once the proteases cleaved off the masking peptide, ADG126 was activated and bound to a high affinity to human, cynomolgus monkey and mouse CTLA-4, as shown in the figures below. Activated ADG126 was observed to lead to the release of CD80/CD86 ligands from CTLA-4 sequestration, and stimulation of CD28 signaling to boost T-cell activity. It also targets

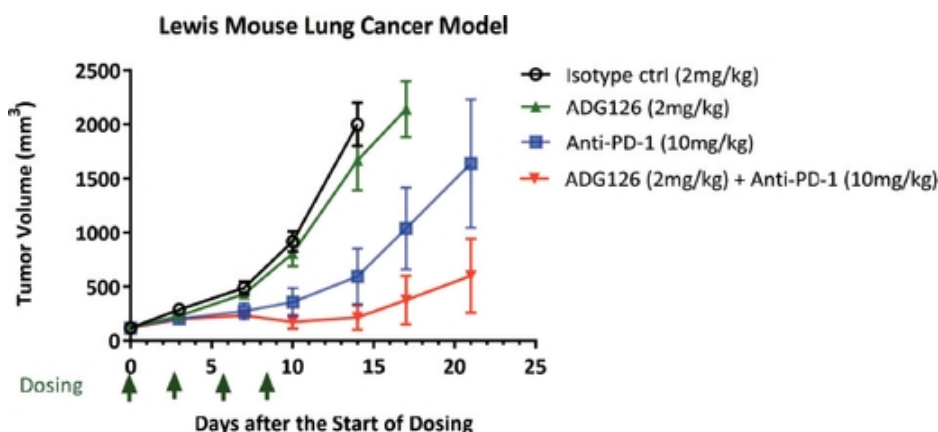
regulatory T-cells for depletion within the TME by means of ADCC, to mediate antitumor T-cell immunity.



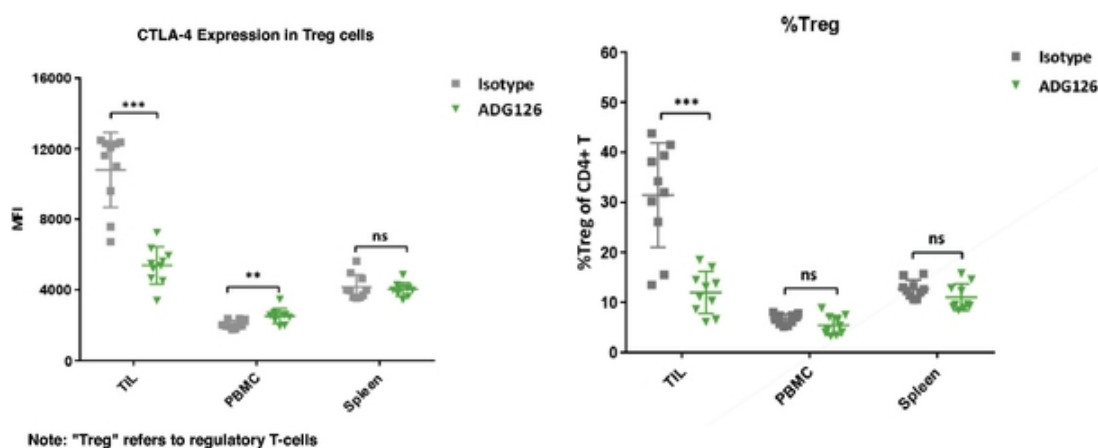
We evaluated the *in vivo* antitumor efficacy of ADG126 in syngeneic mouse tumor models. As shown in the figures below, in these studies, ADG126 was observed to inhibit tumor growth in different mouse tumor models as a monotherapy.



As shown in the figure below, ADG126 synergized with an anti-PD-1 antibody to elicit a stronger anti-tumor response than ADG126 or the anti-PD-1 antibody alone in a Lewis lung cancer syngeneic mouse model.



ADG126 treatment was observed to specifically deplete regulatory T-cells in the tumor, but not in peripheral tissues. The following figures illustrate the reduction of CTLA-4 expression in infiltrating lymphocytes, or TILs (see figure below on the left) and regulatory T-cell depletion (see figure below on the right) by ADG126 in CT26 tumor model.



Preclinical Toxicology: We performed preclinical toxicology studies designed to assess the toxicity features of ADG126. We selected cynomolgus monkeys and mice as toxicology species for animal toxicity evaluation. No abnormal findings attributable to ADG126 were observed. We utilized a nonobese diabetic mouse model to determine percentage of survival with AD126 treatment. All mice survived after six treatments of ADG126 at 50 mg/kg.

In a four-week GLP repeat-dose toxicology studies, intravenous infusion of ADG126 to cynomolgus monkeys at 5, 30, or 200 mg/kg/dose once weekly for five doses followed by a 28-day recovery period was well-tolerated. Adverse, but reversible, microscopic findings of minimal to moderate mixed perivascular infiltrates were observed at 200 mg/kg in both sexes in the kidney, liver, pancreas, epididymis, skin, and were observed in the ovaries in females, and the connective tissue associated with the mesenteric lymph node and thyroid gland. The NOAEL was considered to be 30 mg/kg/dose and the highest non-severely toxic dose was considered to be 200 mg/kg/dose.

Clinical Development Plan

The Australian Therapeutic Goods Administration authorized us to initiate a Phase I trial of ADG126-1001 in Australia. The first site of ADG126-1001 has been approved by the Australian authorities, investigational products have been released, and the site is ready to start patient screening. In addition, three potential patients have been identified for enrollment in the ADG126-1001 trial and we expect to commence patient enrollment by February. We also have received approval from the FDA to initiate the Phase I clinical trial of ADG126 in the United States in 2021. These two Phase I dose escalation trials will test five doses at 0.1, 0.3, 1, 3, and 10 mg/kg, based on our preclinical primate toxicity study in comparison with ipilimumab. We anticipate that these clinical trials will allow us to determine the ADG126 effective dose in late 2021. Meanwhile, we are preparing IND submissions to initiate ADG126 clinical trials in China.

ADG116: Novel Anti-CTLA-4 NEObody Candidate

Summary

ADG116 is a fully human anti-CTLA-4 antibody generated using our NEObody technology, designed to enhance the efficacy and to address toxicity concerns associated with existing CTLA-4 therapeutics. In our preclinical studies, ADG116 was observed to potentiate CD28 signaling and T-cell immune responses (cytokine production) in the presence of primary stimulatory signaling. Furthermore, ADG116 reduced immunosuppressive regulatory T-cell activity and enhanced CD8⁺ T-cells activity in the TME to induce antitumor responses. We believe these preclinical results support the further clinical evaluation of ADG116 both as a monotherapy and in combination with other therapies for a wide range of tumor types. In July 2020, we obtained authorization from the Australian Therapeutic Goods Administration under a CTN to start a Phase I clinical trial of ADG116. As of January 15, 2021, we have successfully completed dose escalation at the first three doses in our ADG116-1003 clinical trial in Australia, including a dose level at 0.03mg/kg, which was the dose level used for the first patient who suffered a fatal incident in the ADG116 Phase I trial in the United States. As of January 15, 2021, no dose limiting toxicity or treatment related serious adverse events have been observed in any of the patients treated. There was no treatment induced liver toxicity and all three patients treated at 0.03mg/kg showed normal liver enzyme level post ADG116 treatment. The ADG116-1001 Phase I trial in the United States was placed on clinical hold on September 30, 2019 by the FDA, after we reported to the FDA the death of the only patient dosed in the trial. The FDA removed the clinical hold on December 5, 2019 after we submitted an amendment to the study protocol.

Mechanism of Action

ADG116 is designed to target a unique conserved epitope of CTLA-4. In preclinical studies, ADG116 was observed to have softer CTLA-4 ligand blocking and stronger ADCC for regulatory T-cell depletion than ipilimumab. Based on the PD results observed in our preclinical studies, we believe that ADG116 has the potential to specifically bind to human CTLA-4, or hCTLA-4, without binding to other CD28 family receptors, which could potentially block the CTLA-4/CD80 and CTLA-4/CD86 ligand interactions. We believe that ADG116 potentiates CD28 signaling and T-cell immune response in the presence of a primary stimulatory signaling. Moreover, ADG116 has been observed in preclinical animal studies to mediate effector functions to eliminate CTLA-4 expressing cells, particularly regulatory T-cells, primarily through ADCC by NK cells. These actions of ADG116 could lead to enhanced activation and proliferation of tumor infiltrating T-effector cells and reduced T-regulatory cell function, which may contribute to a general increase in T-cell responsiveness, including an enhanced antitumor immune response.

Clinical Trial Design

The following summarizes the Australia clinical trial design of ADG116. During the accelerated titration phase, one patient per dose level will be treated. If the patient experiences a DLT or two Grade 2 or above drug-related toxicity, as agreed upon by the investigator and sponsor, the dose level will be expanded according to a "3+3" dose escalation design. During the traditional "3+3" dose escalation phase, from the dosage level at 0.03mg/kg onwards, dose escalation will follow a traditional "3+3" design with three or six patients treated at each dosage level, depending on the incidence of DLTs. Initially, three patients will be enrolled into the dosage level with the sentinel patient treated at least 24 hours before the subsequent patients. Duration of treatment is expected to continue up to two years, if the investigator considers that continued treatment could benefit the patient, or until disease progression and/or unacceptable toxicity, or withdrawal of informed consent.

The primary objectives of the dose escalation stage of the planned clinical trial will be to assess DLT and recommended Phase II dose, or RP2D. The safety profile of ADG116 will also be assessed in terms of severity, seriousness, and frequency of AEs, laboratory results, vital signs, physical examinations, and ECGs of patients enrolled in the study. The secondary objectives of the planned clinical trial are to (i) assess plasma PK parameters; (ii) evaluate anti-drug antibodies, or ADAs, in patients' blood before, during and after treatment with ADG116; (iii) evaluate objective response in accordance with RECIST v1.1 and/or iRECIST; and (iv) evaluate preliminary evidence of antitumor activity of ADG116, as characterized by ORR, DCR, duration of response, or DOR, duration of stable disease, PFS, and OS. The exploratory objective is to identify PD biomarkers for ADG116, including but not limited to cytokines, plasma proteins, immune cells, and tissue biomarkers.

The endpoints of ADG116 Phase I trial are:

Primary endpoints:

- DLT and RP2D. The safety profile of ADG116 will be assessed in terms of severity, seriousness, and frequency of adverse events, laboratory results, vital signs, physical examinations, and ECGs enrolled in the study.

Secondary endpoints:

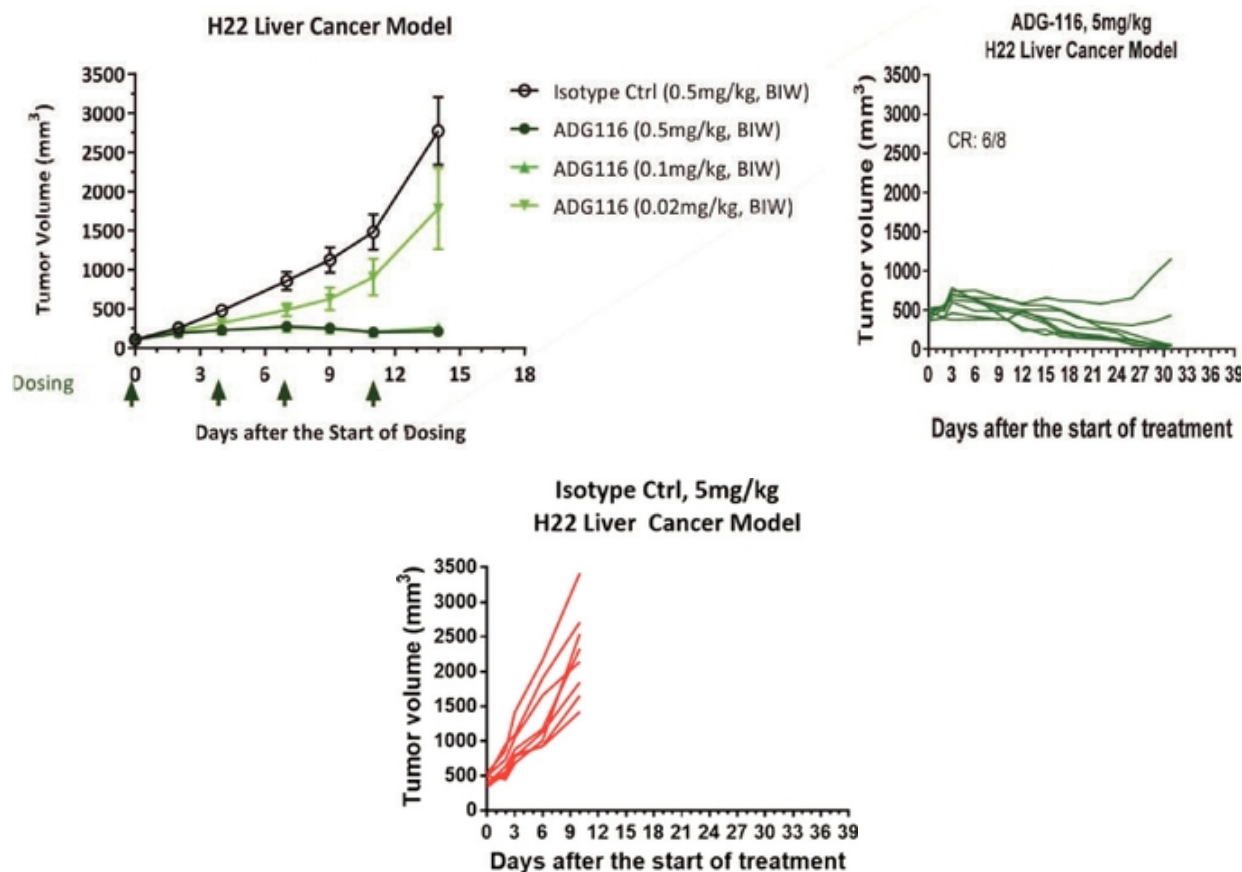
- Plasma PK parameters including but not limited to:
 - $AUC_{0-\infty}$
 - AUC from time zero to last measurable concentration
 - AUC from time zero to 21 days post-dose
 - Maximum (peak) plasma concentration
 - Time to maximum (peak) plasma concentration
 - Trough plasma concentration
 - Apparent half-life
 - Apparent total body clearance
 - Apparent volume of distribution
- Mean residence time
- Incidence of ADAs.
- Objective response in accordance with RECIST v1.1 and/or iRECIST.

- Preliminary evidence of antitumor activity as characterized by ORR, DCR, DOR, duration of stable disease, PFS, and OS.

Summary of Preclinical Studies and Results

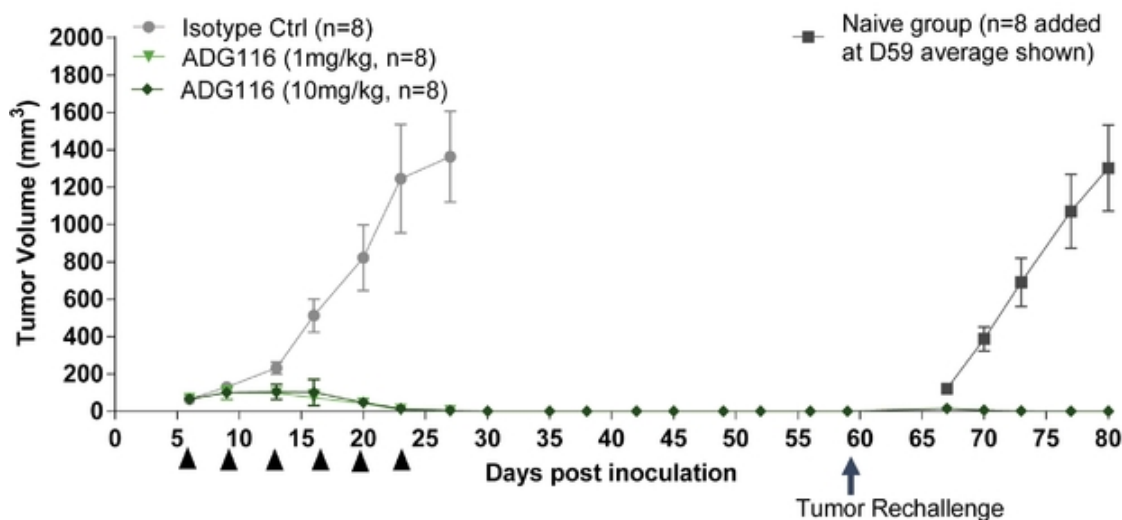
We extensively evaluated ADG116 in both *in vitro* and *in vivo* preclinical studies. Our *in vivo* efficacy studies were conducted in mice, and safety evaluations were conducted in both cynomolgus monkeys and rats. ADG116 was observed to show robust *in vivo* anti-tumor activity in multiple syngeneic mouse tumor models. Ipilimumab was included as a benchmark and was compared with ADG116 in a series of our preclinical studies. In these preclinical studies, we observed that ADG116 had a potential advantage over ipilimumab in potentiating T-cell activation. While ADG116 has softer CTLA-4 ligand blocking, it was observed to have superior ability in eliminating CTLA-4 positive regulatory T-cells via ADCC in tumors resulting in enhanced antitumor responses. ADG116 was well tolerated in rats and cynomolgus monkeys at doses up to 30 mg/kg in GLP-compliant four-week repeat-dose toxicology studies.

Preclinical Pharmacology: We evaluated ADG116 as a monotherapy *in vivo* in an H22 liver cancer syngeneic mouse model. As shown in the figure below, ADG116 was observed to induce an encouraging antitumor response at low doses in a dose-dependent manner (see figure below on the left). Additionally, ADG116 was observed to inhibit tumor growth of large tumors in the same model (see figure below on the right).

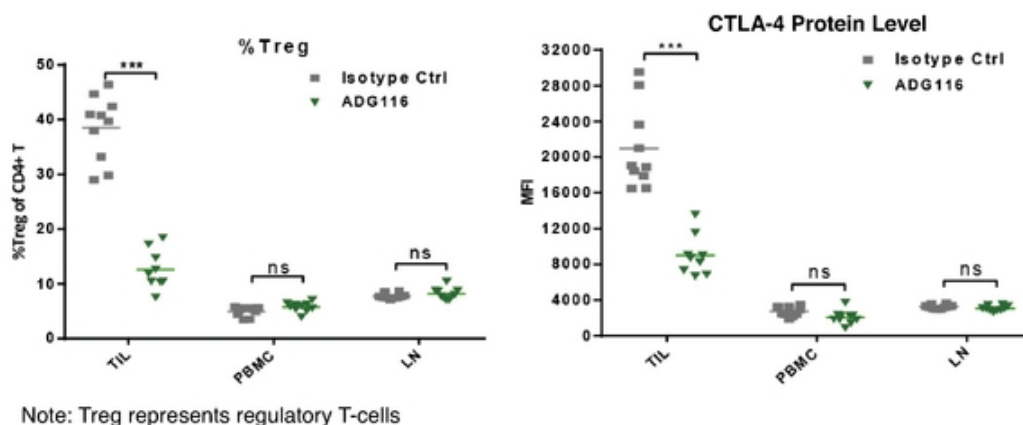


As illustrated in the below figure, in a liver cancer tumor rechallenge study, ADG116 was observed to induce significant antitumor response. Fifty-nine days after the initial tumor inoculation and more than 30 days after the last ADG116 treatment, mice were then rechallenged with the same tumor. We observed that mice that responded to the initial ADG116 treatment remained tumor-free even without additional ADG116 treatment while naïve mice developed tumors, indicating the development of antitumor memory response elicited by ADG116.

Liver Cancer Model (n=8 per group)



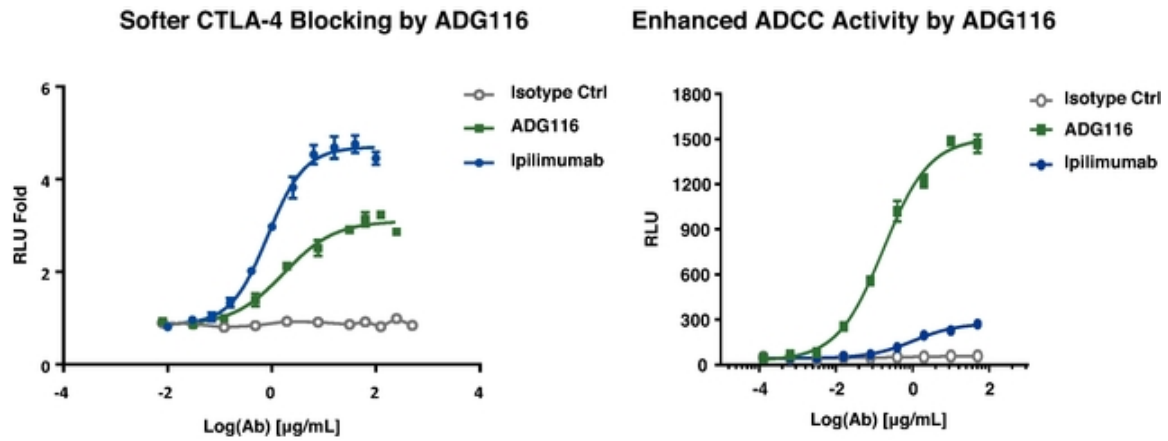
Since ADG116 was observed to exhibit strong ADCC activity *in vitro*, we evaluated the ability of ADG116 to deplete regulatory T-cells *in vivo* in a CT26 mouse colon cancer syngeneic model. ADG116 treatment was observed to specifically deplete regulatory T-cells in the tumor, but not in peripheral blood mononuclear cells, or PBMCs, or lymph nodes. The following figure shows significant regulatory T-cell depletion (left figure) and inhibition of CTLA-4 expression (right figure) in tumors by ADG116.



In a series of preclinical studies, we compared the effect of ADG116 to ipilimumab. We believe that these studies suggest that ADG116 has a unique MOA compared to ipilimumab and has a potential advantage over ipilimumab in eliciting antitumor responses. These preclinical trial results support the further clinical evaluation of ADG116 as a monotherapy and in combination with other therapies for a wide range of tumor types.

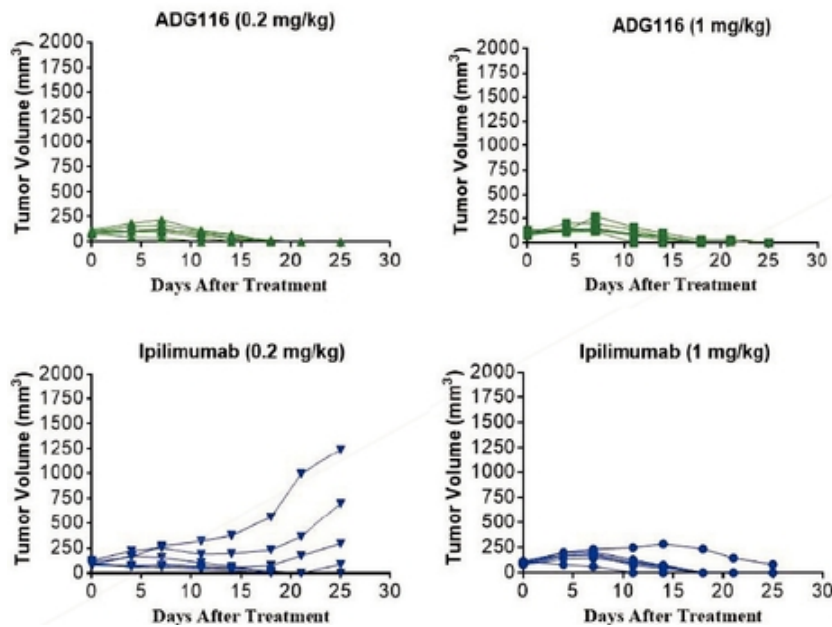
In the CTLA-4 blockade bioassay illustrated in the figure below on the left, ADG116 was observed to exhibit weaker blocking than ipilimumab of CTLA-4's ability to inhibit CD80- and CD86-induced IL-2 production. This result supports our belief that ADG116 can function as a CTLA-4 checkpoint inhibitor with weaker activity than ipilimumab, which may result in less systemic autoimmune side effects on normal tissues. On the other hand, ADG116 was observed to exhibit notably stronger ADCC activity than ipilimumab in the *in vitro* assay shown in the figure below on the right. Since CTLA-4 is expressed on regulatory T-cells, we believe that ADG116 offers a potential advantage over ipilimumab

in depleting regulatory T-cells by means of ADCC. We subsequently investigated this *in vivo* in a syngeneic mouse tumor model.



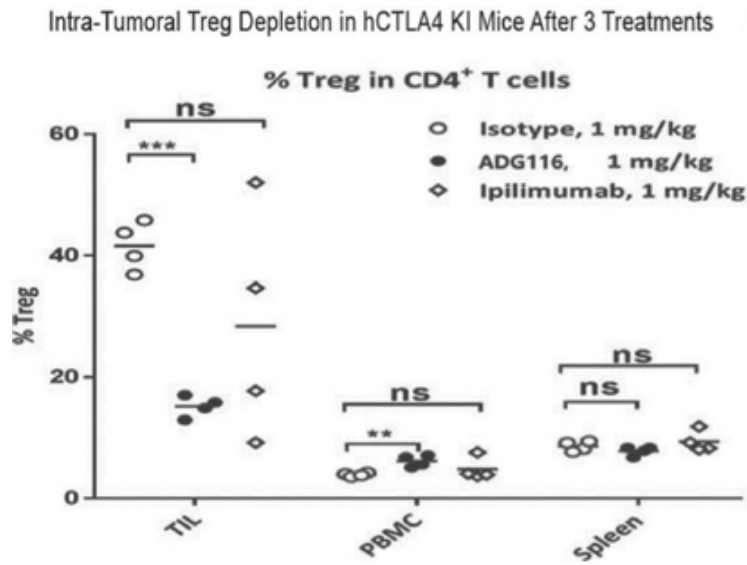
To compare ADG116's *in vivo* antitumor activity to ipilimumab, we utilized a subcutaneous MC38 mouse syngeneic colon cancer model in hCTLA-4, knock-in, or KI, C57BL/6 mice. We selected the hCTLA-4 KI mice as ipilimumab does not cross-react with mouse CTLA-4. As shown in the figure below, ADG116 was observed in this study to exhibit stronger antitumor activity than ipilimumab (ADG116 at 0.2 mg/kg induced equivalent antitumor response as 1 mg/kg of ipilimumab).

Antitumor Activity of ADG116 vs Ipilimumab in MC38 Colon Cancer Model in hCTLA-4 KI Mice



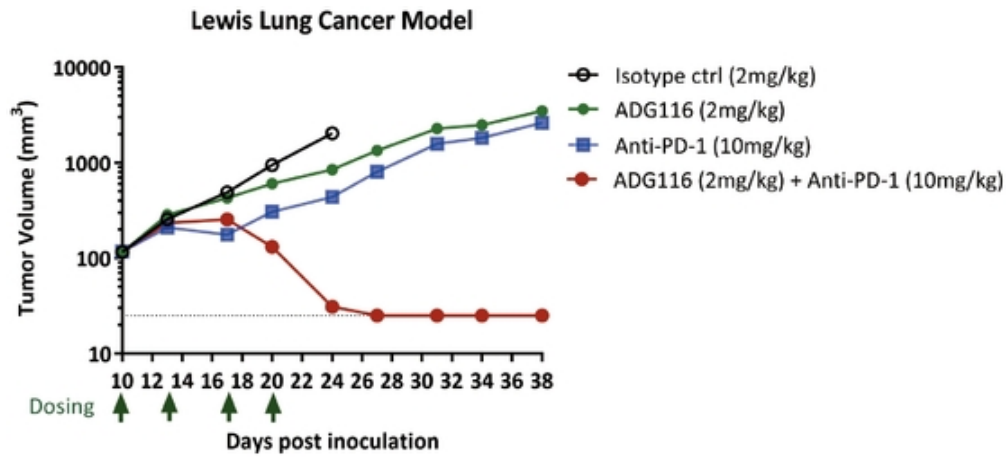
As shown in the figure below, we measured tumor infiltrating regulatory T-cells in the subcutaneous hCTLA-4 KI MC38 mouse colon cancer model after ADG116 or ipilimumab treatment. Among the TILs, the percentage of regulatory T-cells was observed to be significantly reduced after ADG116 treatment while the regulatory T-cell reduction after ipilimumab treatment was not significant. Notably, we observed that regulatory T-cell depletion by ADG116 occurred only in the TME, and not in PBMCs or the spleen. We

believe that these results provide a mechanistic rationale for the enhanced *in vivo* antitumor activity of ADG116 compared to ipilimumab. ADG116 may reduce the immunosuppressive regulatory T-cell activity specifically in the TME to enhance antitumor immune responses.

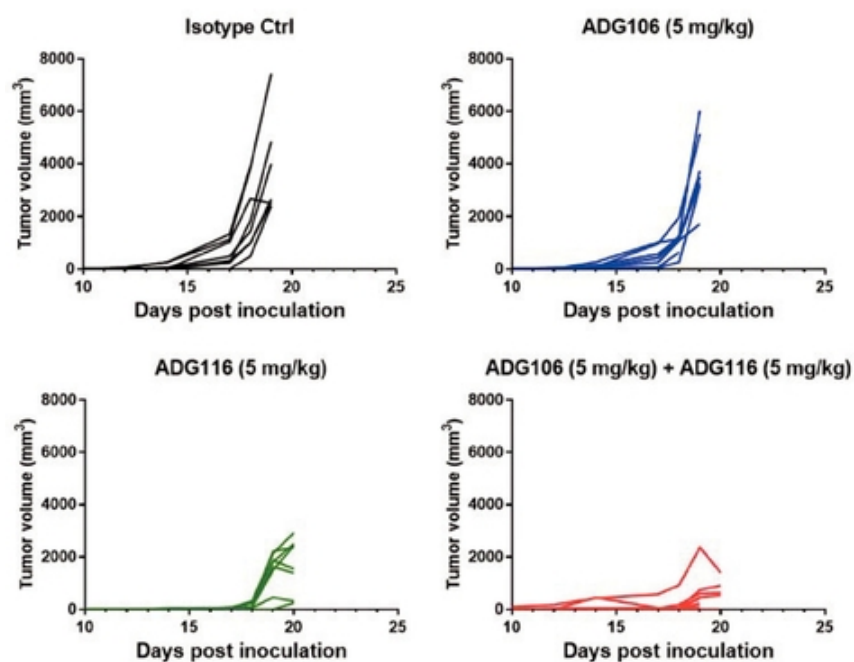


Note: "Treg" refers to regulatory T-cells.

As illustrated below, in addition to evaluating the efficacy of ADG116 as a monotherapy, we also evaluated ADG116 in combination therapies. ADG116 was observed to have an antitumor effect with an anti-PD-1 treatment in a Lewis lung cancer syngeneic mouse model.



We also examined the effects of ADG116 in combination with our CD137 agonistic antibody, ADG106, in a B16F10 melanoma syngeneic mouse model. As shown below, we observed that the combination of ADG116 with ADG106 enhanced the antitumor activity compared with ADG116 or ADG106 alone.



Preclinical Toxicology: We performed preclinical toxicology studies in cynomolgus monkeys and rats to evaluate the toxicity of ADG116. There were no abnormal findings in the single-dose toxicology studies. We observed that ADG116 was well tolerated in both cynomolgus monkeys and rats at up to 200 mg/kg. In a GLP-compliant, four-week repeat-dose toxicology study, ADG116 was tolerated at doses up to 30 mg/kg/dose (five doses per week). In this study, ADG116 related hematology parameter changes, serum chemistry changes, mononuclear infiltration of predominantly lymphocytes with fewer macrophages into the parenchyma of numerous organs were the primary test article-related effects evaluated. These changes were reversible at £ 30 mg/kg/dose, and consistent with the biological role of CTLA-4 in regulating and maintaining peripheral immune tolerance. The NOAEL was considered to be 30 mg/kg/dose in both rats and cynomolgus monkeys.

Clinical Development Plan

As of January 15, 2021, we have successfully completed the three dose escalation at 0.003 mg/kg, 0.01mg/kg and 0.03mg/kg in the Phase I clinical trial of ADG116 in Australia. 0.03mg/kg was the dose that we had initiated a Phase I trial of ADG116 in the United States. We intend to use clinical data collected from the Australia trial to facilitate clinical development in the United States. We also have a Phase I clinical trial open in the United States for ADG116 as a monotherapy in patients with advanced/metastatic solid tumors; however, we are not currently enrolling patients in this clinical trial.

DEEP, BROAD, DIFFERENTIATED PRECLINICAL DISCOVERY PIPELINE

In addition to our three clinical-stage product candidates, ADG106, ADG126 and ADG116, we are building a deep, broad and differentiated preclinical pipeline. Utilizing our DPL platform and three platform technologies NEObody, SAFEbody, and POWERbody, we have built a portfolio of programs that are at various stages of the drug discovery and development process. We have the ability to build multiple generations of drugs against the same targets, e.g., CD137 and CTLA-4, representing the depth of our preclinical pipeline, as shown in the illustration below.



We have also built a preclinical and discovery pipeline against multiple validated and novel targets, representing the breadth of our preclinical and discovery pipeline. Our SAFEbody programs include multiple SAFE NEObodies, such as SAFE CD47 NEObody, as well as a number of undisclosed SAFEbodies. The class of anti-CD47 therapies has attracted significant interests from leading biopharmaceutical companies given its significant potential. Our POWERbody programs include an immuno-oncology antagonist program, an immuno-oncology agonist program, multiple SAFE antibody drug conjugate, or ADC, programs and multiple SAFE bispecific T-cell engager programs, which include CD20xCD3 SAFE bispecific antibody, or bsAb, and HER2xCD3 SAFE bsAb.

Below is an example of our POWERbody preclinical discovery program, which is tumor-associated antigen, or TAA, and CD3 bispecific T-cell engager program.

SAFEbody TAAxCD3 BsAb T-cell Engager Program

Summary

Bispecific T-cell engager antibodies have been explored as a means to recruit cytolytic T-cells to kill tumor cells. This is based on the simultaneous recognition of an antigen on tumor cells and binding to the CD3 epsilon chain, or CD3e, within the T-cell receptor complex on T-cells that bridges malignant tumor cells directly to CD3⁺ T-cells. Blinatumomab, or Blincyto, the first bispecific T-cell engager reactive with the B-cell antigen CD19, was approved by the FDA in 2014 for the treatment of neoplasms. While early studies showed promising clinical efficacy, bispecific T-cell engagers were also hampered by severe dose-limiting toxicities primarily manifesting as cytokine release syndrome which resulted in a prohibitively narrow therapeutic window.

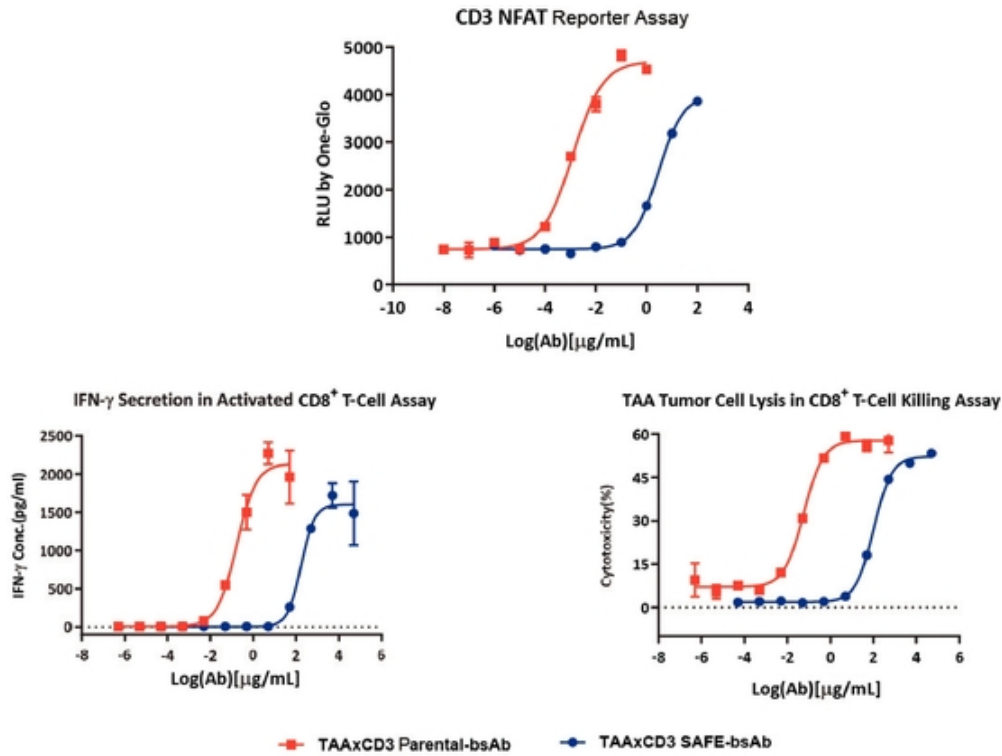
Through our DPL platform, we have developed a deep pipeline of bispecific T-cell engager antibodies for both liquid and solid tumors. To address the safety issues of current CD3 T-cell engager antibodies, we leveraged our proprietary SAFEbody technology to develop differentiated bispecific T-cell engagers.

Our TAAxCD3 SAFE-bsAbs comprise anti-human TAA and CD3 SAFEbodies and are intended for the treatment of TAA⁺ malignancies. Our TAAxCD3 SAFE-bsAb in their activated forms have been observed to stimulate T-cell activation and TAA⁺ tumor cell killing. We did not observe our TAAxCD3 SAFE-bsAbs to have visible cytokine release syndrome and other adverse events in exploratory toxicity studies in cynomolgus monkeys. Our TAAxCD3 SAFE-bsAbs highlight a potential new approach in tumor immunotherapy and provide a rationale for safe and effective treatment of TAA⁺ tumors. Based on our preclinical studies, we are moving forward with generating stable cell lines and CMC development of our two TAAxCD3 SAFE-bsAb programs.

Preclinical Studies

To evaluate the functional activity of parental and SAFE-bsAbs, we tested our bsAbs in multiple cell-based assays. As shown below, the TAAxCD3 parental-bsAb significantly stimulated CD3 signaling in a Jurkat cell-based NFAT reporter assay. Furthermore, the TAAxCD3 parental-bsAb was observed to significantly increase IFN γ secretion from CD8⁺ T-cells and induce TAA overexpressing tumor cell

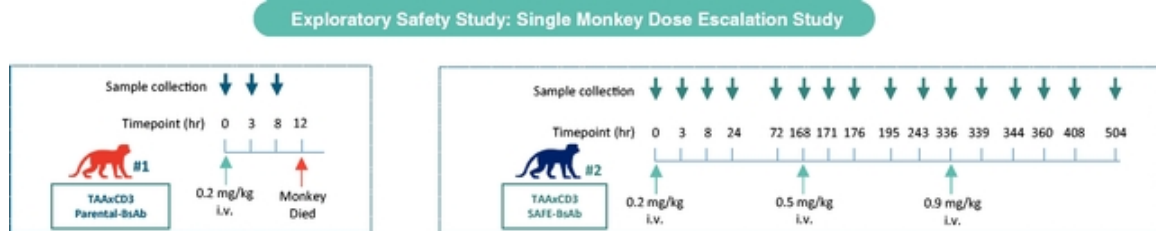
death by activating CD8⁺ T-cells in the T-cell-dependent cellular cytotoxicity assay. These results suggest that our CD3 bsAb potentially stimulated T-cells to kill tumor cells. Consistent with our expectation, the masked SAFE-bsAb was observed to have lower activity in these functional assays.



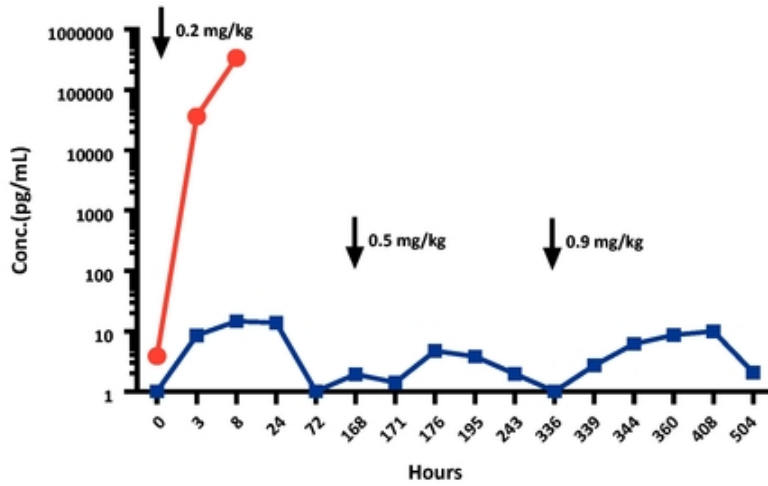
Exploratory Toxicity Study in Cynomolgus Monkeys

As current efforts in developing therapeutic CD3-based T-cell engagers are impeded by severe dose-limiting toxicities, primarily manifesting as non-specific cytokine release syndrome, we evaluated the safety results while developing CD3 bsAbs with our SAFEbody technology. Particularly, we evaluated the cytokine and immune-associated events, for TAAxCD3 bsAbs in cynomolgus monkeys.

Cynomolgus monkeys were dosed intravenously with our TAAxCD3 parental- or SAFE-bsAbs. We observed significant IL-6 release with the parental-bsAb; while we did not observe SAFE-bsAb induced IL-6 release at any of the three-dose levels during the testing period, which indicates a noticeable improvement in safety results by SAFEbody masking.

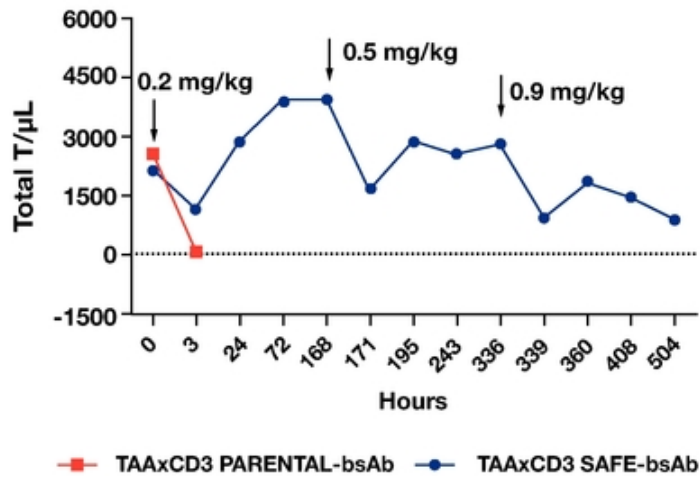


IL-6 release in cynomolgus monkeys treated with BsAbs



The cynomolgus monkey treated with the parental-bsAb died approximately 12 hours after first dosing at 0.2 mg/kg, likely due to acute cytokine release syndrome, which is consistent with severe dose-limiting toxicities observed in other bispecific T-cell engager antibodies. Moreover, total T-cells were nearly undetectable at three hours following the initial parental-bsAb treatment at 0.2 mg/kg. The SAFE-bsAb treated cynomolgus monkey also exhibited reduction of total T-cell counts, but to a lesser degree, and absolute lymphocyte counts rebounded at 24 hours after each dosing of SAFE-bsAb.

Absolute Lymphocyte Count After Treatment with bsAbs



OUR PLATFORM

Overview

Our proprietary DPL platform is built upon our insights into precise and dynamic antibody-antigen interaction. As such, our DPL platform has been designed to enable the discovery of antibodies with better developability properties. By addressing the challenges in traditional antibody design and engineering, we believe that our DPL platform will enable us to improve the efficacy and safety profile of antibody therapeutics. Our DPL platform is empowered by our computational platform, artificial intelligence and three innovative technologies: NEObody, SAFEbody, and POWERbody.

AI-Powered Antibody Discovery

We believe that efficient drug discovery requires significant commitment to resolve three key challenges: 1) the selection of targets with preclinical and/or clinical validation; 2) the availability of abundant hits at discovery stage for the identification of Pre-Clinical Candidates, or PCC, that usually takes a highly iterative process from target-to-hits, lead identification and lead optimization; and 3) deep understanding of target biology and MOA from detailed molecular, cellular, *in vivo* preclinical and human disease models. To address these challenges and to improve antibody R&D productivity, we have designed and built our AI-powered DPL platform to provide sufficient discovery capacity, capable of generating abundant high quality hits against targets, and then selecting the right leads for extensive preclinical testing in order to understand their MOAs, their efficacy and safety potential before going for clinical proof of concept testing. Our current focus on oncology portfolio is to establish early clinical proof of concept, preferably in Phase I trials.

We believe that it is critical to pair the right PCC with the specific epitope of a chosen target. The affinity and precise orientation by which the specific epitope of a given target is engaged by the PCC can have a profound impact on the pharmacology and toxicity of drug candidate. Therefore, we believe precision design and engineering of antibody-based drugs make a fundamental difference on the success of a drug discovery program. As demonstrated by extensive preclinical data of our ADG106, ADG126 and ADG116 programs and early clinical data of our ADG106 program, targeting the conserved epitope of CD137 by ADG106 and that of CTLA-4 by ADG126/ADG116, respectively, across humans, monkeys and mice can give rise to novel biological and pharmacological activities. The broad species cross-reactivity of these NEObody drug candidates above can facilitate seamless translational studies from different immune-competent syngeneic tumor models to monkeys, and then subsequent clinical trials in humans. The novel MOA and biology concerning CD137 and CTLA-4 targets include their regulation in peripheral and tumor microenvironments, the involvement of different immune cells upon target engagement such as NK cells by anti-CD137 ADG106 agonist and T-regulatory cells by anti-CTLA-4 specific antibodies ADG116 and ADG126. We believe these examples provide strong support for our proposal to search for "The Right PCC" that is paired with the specific epitope of a given target in order to unlock the biology of such target and its pathway and to facilitate a deep understanding of its MOA in triggering a cascade of biological and pharmacological events leading to disease control.

In order to search for "The Right PCC", our proprietary DPL platform uses a deep learning approach to explore the conformational diversity of antibody binding sites. Our DPL platform combines AI algorithms and ever-increasing big data in antibody sequence, structure, and binding epitope and affinity from public and our own proprietary databases to design, construct and screen high-quality proprietary antibody libraries with well-defined sequence, scaffold and biophysical attributes for antibody drug discovery. Our computational engine combines software, AI and visualization tools to mine, organize, compute, and interactively explore these immense multidimensional big data sets. We iteratively inform wetlab experimentation with *in silico* computation, and vice versa. By exploiting conformational diversity through the combination of our proprietary computational algorithms with AI,

we have designed and precisely constructed approximately one trillion (10^{12}) antibody sequences in our DPL. These antibodies feature broad epitope (the portion of an antigen that are recognized by an antibody) coverage and robust CMC attributes. The antibodies discovered can be further engineered into NEObody, SAFEbody and POWERbody antibodies depending on the targeted product profile of a given therapeutic target.

Our ADG106, ADG126 and ADG116 clinical programs illustrate how our AI-powered DPL platform has improved our R&D productivity. Our DPL library is a large physical phage library ($\sim 10^{12}$) that contain our AI-powered synthetic designer antibodies with flexible antibody binding interface. The combination of binding interface flexibility with the variation of amino acid sequences in the antigen binding interface exponentially increases the conformational diversity of our AI-powered DPL libraries, far beyond conventional natural or synthetic antibody libraries. Our AI-Powered DPL library screening against CD137 or CTLA4 antigens have yielded a large number of high affinity primary hits. For example, our anti-CD137 agonist antibody ADG106 was identified from 60 high affinity primary hits with a wide range of epitope diversity in terms of their species cross-reactivity, which compared favorably with the number and epitope diversity of high affinity anti-CD137 antibody hits reported by our global peers on their publicly filed documents. Our anti-CTLA-4 antibody ADG116 was identified from 74 high affinity primary hits with a wide range of epitope diversity in terms of their species cross-reactivities: 19 clones are human, monkey, canine and mouse cross-reactive; 12 clones are human and monkey cross reactive but weak mouse cross-reactive; 32 clones are human and monkey cross-reactive; 11 clones only recognize human CTLA-4. The abundant discovery hits with diversified binding epitopes for varying degree of species cross-reactivity show the power of our AI-powered DPL platform not only in creating novel ADG106 and ADG116 NEObodies in targeting different epitopes of a given antigen, but also in targeting the conserved epitope across different species of a given antigen for both CD137 and CTLA-4 to enable translational studies from immune-competent animal tumor models to human patients due to their broad species cross-reactivities.

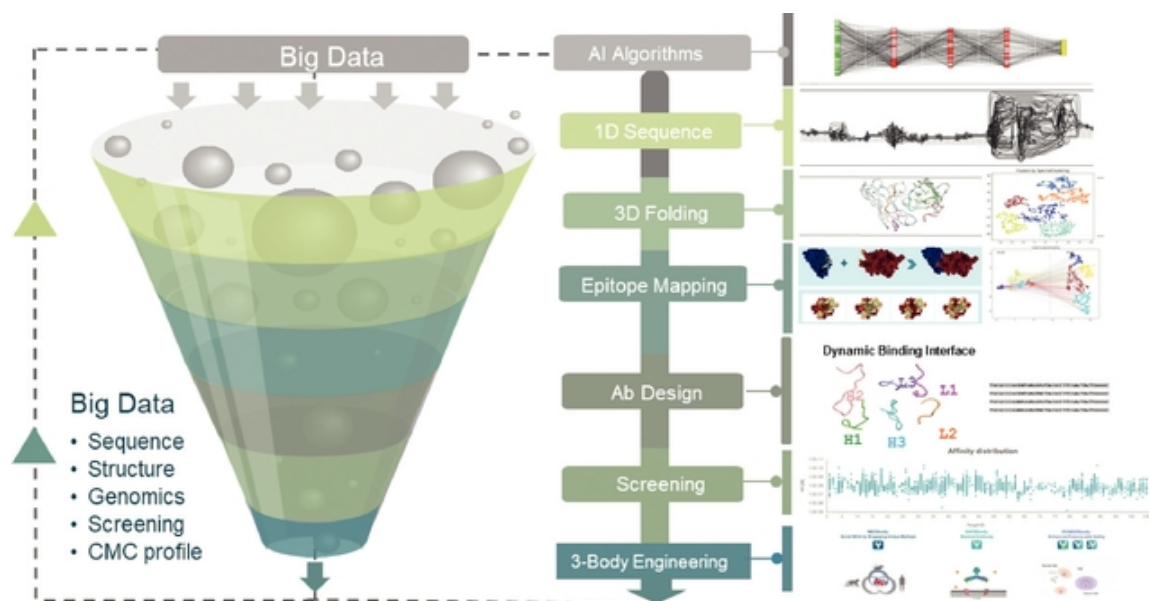
We believe that the high affinity and species cross-reactive primary hits from our DPL library screening saves time and cost from primary hits to straight PCC. The broad species cross-reactivity of the primary hits also streamlines lead identification through simple syngeneic tumor models for safety and efficacy testing. The robust CMC attributes of our DPL library are designed to also eliminate the requirement to optimize antibody sequences before starting CMC process for IND enabling and formulation studies. Furthermore, we believe our AI-powered dynamic precision library of approximately one trillion (10^{12}) antibody sequences expands the diversity at the start of discovery to maximize the chance that suitable leads are found on the first pass. In conclusion, we believe our AI-powered antibody discovery and engineering DPL platform significantly increases R&D productivity for antibody drug discovery. It has enabled us to generate highly differentiated PCC molecules that we believe can target the conserved epitope across different species for novel MOAs and also facilitate seamless translational studies from animals to human via their broad species cross-reactivity.

Briefly, our AI-powered antibody discovery engine includes six steps:

1. First principle of antibody folding and dynamics. We fold the antibody primary amino acid sequence into three-dimensional structures based on machine learning or homology modeling in combination with the Molecular Dynamic Simulation in the presence and absence of dynamic interaction with their antigen targets.
2. AI-powered deep learning of antibody sequence and structure and binding epitopes. AI algorithm and machine learning have been widely used to map protein sequence and structure variation and connectivity. Deep learning has been used to map the antibody and antigen epitope in order to learn the landscape of the epitope space and their correlation with antibody binding interface.

3. Antibody design that combines the conformational diversity with primary amino acid sequence variation. We have designed antibody libraries that contain dynamic binding units and their combination in the context of different CDRs and framework scaffold to capture their conformational diversity of antibody binding interface in combination with variation in primary amino acid sequences far beyond conventional natural or synthetic antibody libraries.
4. Synthetic dynamic precision library by AI-powered computational design. The amino acid sequence library is converted into synthetic DNAs that can be assembled and constructed for phage and yeast library screening.
5. Directed evolution by phage and yeast libraries. The synthetic phage and yeast libraries can be screened semi-automatically for more than 100 antigens to yield high affinity primary hits.
6. PCC identification by targeting conserved epitope with broad species cross-reactivity to unlock the deep biology of target and its MOA upon target engagement by the PCC. PCC is designed and selected to target the conserved epitope of a given target across different species. Traditionally, affinity of primary hits from synthetic antibody libraries have to be improved before subjecting them to extensive in vitro and in vivo assays for lead identification and optimization through an iterative process between functional assays and antibody engineering. Our DPL platform can yield high affinity primary hits, which can then become product candidate leads subject to functional characterization without intensive iterative lead optimization process.

The following diagram illustrates how AI algorithms and big data empower rapid antibody discovery with improved productivity:



NEObody™

NEObody technology is a fully synthetic phage display and yeast display-based antibody discovery technology, which we believe is differentiated from other synthetic antibody technologies through its innovative designs and precise constructions. Our designs are based on critical insights gained from extensive studies of antibody structural variability made possible by our proprietary in-house developed computational tools.

Innovative antibody library design: We believe that diversity of an antibody library should be defined at the quaternary structural level, instead of the traditional primary amino acid sequence level. Comparison of structural and sequence diversity revealed that variability as assessed by structural alignments was generally lower than the variability observed with sequence alignments. Based on our deep understanding of chemical principles governing antibody folding and extensive analysis of a vast number of antibody structural variability, through novel statistical tools developed in-house, we have redefined the antibody hyper-variable regions, or HVRs, that are critical for antigen recognition. These HVRs, as defined based on structural variability, are distinct from but complementary with complementarity-determining regions that are traditionally defined based on amino acid sequence variability. In addition, we have defined and designed dynamic motifs which adopt multiple conformations and incorporated them into our DPL antibody library design to enable the coverage of a wide range of structural diversity with a limited number of amino acid sequences.

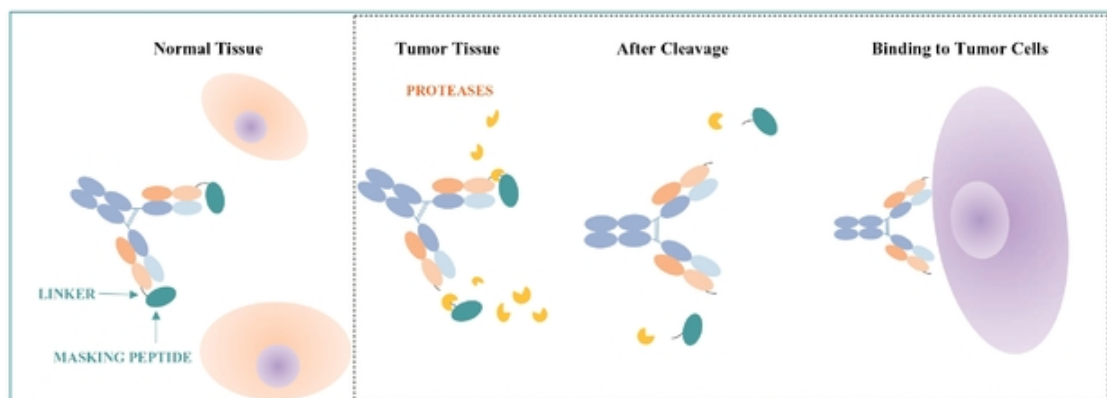
Broad epitope coverage, particularly of evolutionarily conserved epitopes: NEObody technology enables us to discover antibodies targeting numerous epitopes against a broad range of antigens. In particular, we focus on the antibodies targeting the conserved epitopes of divergent antigens. These species cross-reactive antibodies not only have the potential to reveal new biological functions of the targets, but also facilitate preclinical studies using various immune intact animal models, resulting in high fidelity translation from preclinical to clinical studies. Our NEObody technology has been evaluated in preclinical studies with numerous antigens, including some difficult antigens such as membrane or oligomeric proteins. The success criteria in our preclinical assays include high affinity of primary hits with diverse antibody sequences, broad epitope coverage, as well as favorable CMC development properties. For example, ADG106 was discovered using our NEObody technology to target a conserved epitope of CD137 that mostly overlaps with its ligand binding site, which we believe enables ADG106 to activate CD137 in its natural ligand-like fashion, with complete blocking of CD137L, which may differentiate it from two other investigational leading CD137 agonistic antibodies.

Enhanced developability profiles: NEObody technology is designed to preemptively eliminate the chemically unstable sites, or combinations of "problematic" sites that may pose risk for downstream manufacturing processes. This precision design strategy coupled with precision oligonucleotides and library construction have resulted in our high-quality antibody discovery libraries. The primary hits from these libraries generally lack severe "problematic" and spurious amino acid sites, and therefore, may offer promising characteristics for further development.

SAFEbodyTM

SAFEbody is our proprietary differentiated precision antibody masking technology designed to enable an antibody to bind its target specifically only after conditional activation of the antibody in target tissues. By engineering our antibody therapeutic candidates to selectively activate in the TME, our SAFEbody technology is designed to improve safety and tolerability of antibody therapeutics while simultaneously maintaining clinical activity. Through this technology, we believe we can provide a

solution to on-target off-tumor toxicity, one of the long-lasting challenges with many approved antibody therapeutics.



Activation in the TME: SAFEbody technology is designed to mask an antibody binding interface with a masking motif, which then prevents an antibody from binding to its target in healthy tissues. The masking motif is designed to activate or unmask the antibody binding in the TME where certain activation conditions such as a protease is upregulated as compared to healthy tissues, allowing the antibody to bind to and attack the tumor. Our SAFEbody enabled therapeutic candidates are therefore designed to be activated predominantly in the TME while remaining largely in an inactive state in healthy tissues.

Innovative masking moiety library design: Leveraging computational biology, we have designed multiple masking libraries with structured scaffolds and balanced chemistry of amino acids with favorable attributes for masking, easier manufacturing processes, and lower immunogenicity.

Precision masking without self-inhibition: To differentiate from other masking technologies, our antibody masking moieties have been designed and discovered in the right context in an effort to provide greater expression and specificity with no self-inhibition upon activation. We employ sophisticated screening processes for rapid discovery of highly masked SAFEbody product candidates subject to systematic tuning of the masking efficiency to match the target biology.

Improved pharmacology and safety features: We believe our SAFEbody technology has the potential to also reduce the dose-limiting toxicities observed in combination therapies and thus potentially enable new combinations with other cancer therapies that were previously difficult to use. Our SAFEbody technology may also provide favorable for PK and PD profiles for antibodies and ADCs to reduce the drug clearance from circulation in healthy tissues.

Our SAFEbody technology has been applied to multiple target antibodies that have been either discovered with NEObody technology or supplied by our partners. Our ADG126 was discovered by combining SAFEbody technology with its parental antibody discovered through our NEObody technology.

POWERbody™

In recent years, multiple modalities have been developed to enhance the potency of traditional antibody drugs. These include Fc engineering for enhanced ADCC or cross-linking efficiency, ADCs and TCEs. Some of them, such as TCEs, can be so potent that only micrograms of the active ingredient are needed for a single dose, in contrast to regular antibody where hundreds of milligrams or even grams are required for a single dose. NEObodies can be converted into SAFEbodies which can be further reformatted into SAFEbody ADCs, bispecific TCEs, and Fc engineered antibodies. We refer to these to as POWERbodies to indicate their potentially enhanced potency and safety profile.

We believe SAFEbodies can be applied to these modalities. Since an unmasked ADC, TCE, or Fc engineered antibody can be so potent, there is the risk that it will cause severe toxicity due to damage to healthy tissues or fast lysis of tumor cells, resulting in a narrow therapeutic window, as is frequently observed with some ADC drugs or TCEs that have been FDA approved or are in clinical development. We believe our POWERbodies enabled by our SAFEbody technology with ADC, TCE, etc., due to their inactivity in circulation and specific local activation in TME, could significantly reduce toxicity and at the same time retain efficacy.

POWERbody technology aims to boost the efficacy of antibody candidates with safety profiles enhanced by our SAFEbody technology. We believe our POWERbody technology will unleash the full power of antibody-based therapeutics to kill cancer cells with enhanced safety, achieving full potential in antibody based therapies such as bispecific TCEs and ADCs.

COLLABORATIONS WITH BIOPHARMACEUTICAL COMPANIES

We enter into collaborations with biotechnology and pharmaceutical companies to advance the development and commercialization of our technology platforms and product candidates. We seek collaborations that will allow us to retain significant future participation in product sales through royalties paid on net sales. For example, we have entered into agreements with ADC Therapeutics to develop antibody drug conjugates against tumor targets using our SAFEbody technology. In addition, we have also out-licensed the Greater China rights to an anti-PD-L1 antibody and second antibody candidate to Dragon Boat Pharmaceuticals and its affiliates. Our collaborations empower our growth in many ways, including generating cash flow and revenues that partially offset expenditures on our internal research and development programs, expanding our knowledge base regarding antibody technology across multiple targets and antibodies provided by our partners, and providing us with future pipeline opportunities through opt-in licensing rights to new product candidates created using our technology. Below are the highlights of collaborations we formed with our business partners:

Exelixis Collaboration and License Agreement

In February 2021, we entered into a collaboration and license agreement (the "Exelixis Agreement") with Exelixis, Inc., pursuant to which we agreed to generate masked antibodies with our SAFEbody technology against an initial target and a second target to be selected by Exelixis. Pursuant to the Exelixis Agreement, we granted Exelixis an exclusive, worldwide, sublicensable license to research, develop, make, have made, sell, offer for sale, import and commercialize products containing the masked antibodies to be generated by us with respect to both targets.

We are subject to certain exclusivity provisions limiting our right to research, develop, make, and commercialize any other antibodies directed to the selected targets during the term of the Exelixis Agreement, subject to exceptions for antibodies with sequence identities that differ by a specified amount. Exelixis is subject to certain exclusivity provisions limiting its rights to research, develop, make, and commercialize any other masked antibodies directed to the selected targets during the term of the Exelixis Agreement, subject to exceptions for antibodies with sequence identities that differ by a specified amount or use different masking technology.

We are required to use commercially reasonable efforts to generate masked antibodies in accordance the program plan for each target at our cost. Exelixis is required to use commercially reasonable efforts to further develop, obtain regulatory approval and commercialize a product for each target.

Under the Exelixis Agreement, we are entitled to receive an \$11,000,000 upfront payment. In addition, we could be eligible to receive, on a target-by-target basis, up to \$27,500,000 in development milestone payments and up to \$100,000,000 in regulatory milestone payments upon the achievement of specified development and regulatory milestones, respectively, and up to \$262,500,000 in sales milestone

payments upon the achievement of specified commercial milestones. In the aggregate, we could be eligible to receive up to \$55,000,000 in development milestone payments, \$200,000,000 in regulatory milestone payments, and up to \$525,000,000 in sales milestone payments for both targets under the Exelixis Agreement. In addition, we are entitled to receive mid-single-digit percentage net sales-based royalties on products developed under the Exelixis Agreement, subject to certain reductions. Royalties, if any, will be payable on a country-by-country and product-by-product basis, until the latest of (i) the tenth anniversary of the first commercial sale of such product in such country, (ii) the expiration of the last-to-expire of certain specified patents that cover such product's composition of matter or method of use as sold in such country or (iii) the expiration of regulatory or data exclusivity for such product in such country.

The Exelixis Agreement will expire upon the termination of all royalty obligations. Exelixis will own the inventions relating to the compounds arising in connection with the Exelixis Agreement and we will own the inventions relating to our platform. Exelixis may terminate the Exelixis Agreement without cause, in its entirety or on a target-by-target basis, upon a specified notice period, and may terminate for safety reasons on a target-by-target basis immediately upon notice. Either party has the right to terminate the Exelixis Agreement for cause upon the other party's uncured material breach or insolvency, provided that, upon our material uncured breach, Exelixis may elect to instead continue the Exelixis Agreement, and any subsequent milestones or royalty amounts payable to us after such time will be reduced by a specified percentage. Upon termination of the Exelixis Agreement for any reason, the license granted to Exelixis will terminate.

ADC Therapeutics Agreements

In April 2019, we entered into a material transfer and collaboration agreement (the "ADCT Collaboration Agreement") and a license agreement (the "ADCT License Agreement") with ADC Therapeutics, a late clinical-stage oncology-focused biotechnology company pioneering the development and commercialization of targeted ADCs.

ADCT Collaboration Agreement

Pursuant to the ADCT Collaboration Agreement, we agreed to generate masked antibodies with our SAFEbody technology that could be combined with the pyrrolobenzodiazepine-based cytotoxic payload used in ADC Therapeutics' ADCs to create novel ADCs. ADC Therapeutics entered into the ADCT Collaboration Agreement with us to evaluate the use of our SAFEbody technology with respect to up to two exclusive targets selected by ADC Therapeutics. Upon our delivery of certain initial results, ADC Therapeutics has the option to license our SAFEbody™ technology with respect to one or both targets as further detailed below. ADC Therapeutics has not yet exercised such options as of September 15, 2020.

Both parties are required to use commercially reasonable efforts to perform certain development obligations under the ADCT Collaboration Agreement. Additionally, we are subject to exclusivity obligations to ADC Therapeutics under the ADCT Collaboration Agreement with respect to (i) the targets for which ADC Therapeutics has a license or an option to license and (ii) the use or licensing of our intellectual property that is necessary or useful to the development plan under the ADCT Collaboration Agreement or that would preclude us from granting to ADC Therapeutics the licenses under the ADCT License Agreement. ADC Therapeutics owns intellectual property that are specific to the SAFEbodies that we develop under the ADCT Collaboration Agreement with respect to the two elected targets, and we will own all intellectual property developed under the ADCT Collaboration Agreement that relates generally to our SAFEbody platform.

Under the ADCT Collaboration Agreement, we are eligible to receive up to US\$1,600,000 in consideration for our exclusivity obligations, upon achievement of certain development milestones and

upon ADC Therapeutics' election to proceed with development for the two elected targets. As of the date of this prospectus, we have received US\$437,500 in aggregate payments under the ADCT Collaboration Agreement. ADC Therapeutics has the right to terminate the ADCT Collaboration Agreement at any time and for any reason in its entirety or on a target-by-target basis upon thirty days' prior written notice to us. Either party may terminate the ADCT Collaboration Agreement, in its entirety or on a target-by-target basis, upon the other party's uncured material breach of the agreement or the other party's insolvency-related events.

ADCT License Agreement

Subject to the exercise of the options contained in the ADCT Collaboration Agreement, we have granted ADC Therapeutics, with respect to each elected target, an exclusive, worldwide, perpetual and irrevocable (subject only to the termination provisions) license (with the right to grant sublicenses) to develop, make, use, commercialize and import the antibody drug conjugates that comprise masked antibodies generated by us under these programs. Subject to certain conditions, including the exercise by ADC Therapeutics of its first option to license our SAFEbody™ technology, ADC Therapeutics will grant us the option to negotiate a license to develop, manufacture and commercialize ADCs containing our SAFEbody™ technology in Greater China.

Under the ADCT License Agreement, if ADC Therapeutics exercises both of its options granted thereunder, we could be eligible to receive up to US\$166,090,000 in aggregate milestone payments, in addition to mid-single-digit percentage net sales-based tiered royalties on products licensed under the ADCT License Agreement, subject to certain reductions. Royalties, if any, will be payable on a country-by-country and product-by-product basis, until the earlier of (i) the tenth anniversary of the first commercial sale of such product or (ii) the expiration of the last-to-expire patent licensed under the agreement in such country, unless earlier terminated by the parties, following which any licenses granted to ADC Therapeutics under the ADCT License Agreement shall become fully paid up, perpetual and irrevocable. In addition to the contingent milestone and royalty payments, if ADC Therapeutics exercises both of its options granted under the ADCT Collaboration Agreement, we are also entitled to a mid-six-figure dollar amount annual maintenance fee. ADC Therapeutics has not yet exercised the options contained in the ADCT Collaboration Agreement as of the date of this prospectus. Therefore, we have not received any payment under the ADCT License Agreement.

ADC Therapeutics has the right to terminate the ADCT License Agreement before the expiration of the royalty term on a product-by-product basis or in its entirety (i) for any reason or no reason upon thirty days' written notice to us, or (ii) if ADC Therapeutics chooses to discontinue the development or sale of the applicable licensed product worldwide. Each party has certain rights to terminate the ADCT License Agreement with prior written notice upon the other party's uncured material breach or insolvency.

Sanjin Collaboration/ Out-Licensing Agreements

2018 Collaboration Agreements

In December 2018, we entered into (i) a collaboration agreement (the "Sanjin Greater China Agreement") that covers Greater China with Guilin Sanjin Pharmaceutical Co., Ltd. ("Sanjin") and certain of its subsidiaries (collectively, "Sanjin Parties") and (ii) a collaboration agreement (the "Sanjin ROW Agreement", together with the Sanjin Greater China Agreement, the "2018 Sanjin Agreements") that covers the regions other than Greater China with Sanjin. Pursuant to the Sanjin Greater China Agreement, we transferred the Chinese intellectual property directly related to a monospecific antibody molecule that binds to the PD-L1 target (the "PD-L1 Project"), including patent rights, patent application rights and technologies based on the core sequence of the molecule, to Sanjin Parties. Sanjin Parties will own all the Chinese intellectual property developed in the exercise of Sanjin Parties'

rights under the agreement, including but not limited to improvements (including combination products), clinical trials, regulatory filings, and commercialization rights relating thereto. We also granted Sanjin Parties a royalty-free license to use our other existing intellectual property and improvements thereto which are related to the PD-L1 Project for the purposes of exploiting its rights and performing its obligations under the agreement. Sanjin Parties will enjoy all the economic benefits deriving from the PD-L1 Project in Greater China, including but not limited to patent transfer fee, licensing fee, sales revenue and sales commission, etc. Sanjin Parties will pay us (i) within the validity period of the patent of PD-L1 molecule, single-digit percentage of net sales of the products that use the licensed antibody after such products enter the market and (ii) a low to mid-low double-digit percentage of the profits resulting from any transfer of the license to any third parties depending on the timing of the transfer relative to the development stage of the product. We also received a low-seven figure dollar upfront fee upon the effectiveness of the agreement from Sanjin Parties.

Pursuant to the Sanjin ROW Agreement, we granted Sanjin a royalty-free license to use all intellectual property relating to (i) the collaboration under the agreement that we controlled before we entered into the agreement or acquired independently of the agreement and (ii) improvements thereto for the purposes of exploiting its rights and performing its obligations under the agreement. Any intellectual property generated independently by a party under the agreement will be solely owned by that party who generated such intellectual property, and any intellectual property generated from cooperation between us and Sanjin's affiliates in connection with the collaboration will be jointly owned. We have the right to apply for the patents derived from our core and key technologies pertaining to the PD-L1 molecule in the rest of the world and we retain a majority of the economic benefits derived from the Sanjin ROW Agreement, including but not limited to any patent transfer fee, licensing fee and gains realized under such transfer. In case we intend to transfer to a third party our share of economic interests in any country outside of Greater China, we must notify Sanjin and Sanjin will receive a right of first refusal if it pays us a deposit equal to a low double-digit percentage of the consideration that we expect to receive from such third party. If Sanjin waives the right of first refusal, we can proceed with the transfer, provided that the final transaction price with the third party is not lower than the amount of the offering price that was included in our notice to Sanjin.

Under the 2018 Sanjin Agreements, we agreed not to (i) independently develop any monospecific antibodies that bind to the PD-L1 target or (ii) grant any rights associated with such antibodies to any third parties during the three-year period from the effective date of the agreements. The exclusivity obligation does not prevent us from (i) developing or granting any licenses to third parties for intellectual property that covers bispecific antibodies, ADCs, diagnostic antibodies, nano-particles and probody against PD-L1 target and (ii) continuing to provide antibody screening service that were commenced before the execution of the Sanjin Greater China Agreement and either party has the independent right to conduct combination therapy studies outside of the Greater China. Either non-breaching party may terminate the 2018 Sanjin Agreements if the other party's ability to comply with its respective obligations under the agreements is negatively affected by contingencies such as failure to maintain operation or changes in core project management and the other party fails to take effective remedial measures. Each agreement automatically terminates upon the termination of the other agreement. Upon the rescission or termination, Sanjin Parties will return to us all the intellectual property transferred by us to the Sanjin Parties as well as documents and data provided by us under the 2018 Sanjin Agreements. The 2018 Sanjin Agreements are governed by PRC law.

2019 Collaboration Agreements

In May 2019, we entered into (i) a collaboration agreement that covers Greater China (the "Dragon Boat Greater China Agreement") and (ii) a collaboration agreement that covers the regions other than Greater China (the "Dragon Boat ROW Agreement," together with the Dragon Boat Greater China Agreement, the "2019 Dragon Boat Agreements"), with Dragon Boat Biopharmaceutical

(Shanghai) Limited. ("Dragon Boat"), a subsidiary of Sanjin. Pursuant to the Dragon Boat Greater China Agreement, we will transfer the Chinese intellectual property directly related to a certain monospecific antibody molecule (the "Specified Molecule") that binds to a specified target (the "Specified Project"), including the patent rights, patent application rights and technologies based on the core sequence of the molecule, to Dragon Boat. Dragon Boat will own all the Chinese intellectual property developed in the exercise of Dragon Boat's rights under the agreement, including but not limited to improvements (including combination products), clinical trials, regulatory filings, and commercialization rights relating thereto. We also granted Dragon Boat a royalty-free license to use our other existing intellectual property and improvements thereto which are related to the Specified Project for the purposes of exploiting its rights and performing its obligations under the agreement. Dragon Boat will enjoy all the economic benefits deriving from the Specified Project in Greater China, including but not limited to patent transfer fee, licensing fee, sales revenue and sales commission, etc. and will pay us (i) RMB6,000,000 (equivalent to approximately US\$0.9 million) milestone payments and (ii) a single-digit percentage of net sales of the products that within the validity period of the patent of the Specified Molecule use the licensed antibody after such products enter the market. Dragon Boat also paid us a mid-six figure dollar upfront fee upon the signing of the agreement.

Pursuant to the Dragon Boat ROW Agreement, we granted Dragon Boat a royalty-free license to use all intellectual property relating to (i) the collaboration under the agreement that we controlled before we entered into the agreement or acquired independently of the agreement and (ii) improvements thereto for the purposes of exploiting its rights and performing its obligations under the agreement. Any intellectual property generated independently by a party under the agreement will be solely owned by that party who generated such intellectual property, and any intellectual property generated from cooperation between us and Dragon Boat in connection with the collaboration will be jointly owned. We have the right to apply for the patents derived from our core and key technologies pertaining to the Specified Molecule in the rest of the world and we retain a majority of the economic benefits derived from the Dragon Boat ROW Agreement, including but not limited to any patent transfer fee, licensing fee and gains realized under such transfer. In case we intend to transfer to a third party our share of economic interests in any country outside of Greater China, we must notify Dragon Boat and Dragon Boat will receive a right of first refusal if it pays us a deposit equal to a low double-digit percentage of the consideration that we expect to receive from such third party. If Dragon Boat waives the right of first refusal, we can proceed with the transfer, provided that the final transaction price with the third party is not lower than the amount of the offering price that was included in our notice to Dragon Boat.

Under the 2019 Dragon Boat Agreements, we agreed not to (i) independently develop any monospecific antibodies that bind to the specified target or (ii) grant any rights associated with such antibodies to any third parties during the three-year period from the effective date of the agreements. The exclusivity obligation does not prevent us from (i) developing or granting any licenses to third parties for intellectual property that covers bispecific antibodies, ADCs, diagnostic antibodies, nano-particles and probody against the specific target and (ii) continuing to provide antibody screening service that were commenced before the execution of the Dragon Boat Greater China Agreement and either party has the independent right to conduct combination therapy studies outside of the Greater China. Either nonbreaching party may terminate the 2019 Dragon Boat Agreements if the other party's ability to comply with its obligations under the agreements is negatively affected by contingencies such as failure to maintain operation or changes in core project management and the other party fails to take effective remedial measures. Each agreement automatically terminates upon the termination of the other agreement. Upon the rescission or termination, Dragon Boat will return to us all the intellectual property transferred by us to Dragon Boat as well as documents and data provided by us under the 2019 Dragon Boat Agreements. The 2019 Dragon Boat Agreements are governed by PRC law.

As of the date of this prospectus, we have received approximately US\$1.5 million and US\$0.6 million in aggregate payments under the 2018 Sanjin Agreements and the 2019 Dragon Boat Agreements, respectively.

INTELLECTUAL PROPERTY

Protection of our intellectual property is fundamental to the long-term success of our business. Specifically, our success is dependent on our ability to obtain and maintain protection for our technology and the know-how related to our business, defend and enforce our intellectual property rights, and operate our business without infringing, misappropriating, or otherwise violating valid and enforceable intellectual property rights of others. Our patent strategy is focused on seeking coverage for our core technologies and products, such as the DPL platform, ADG106, ADG126, and ADG116. We also rely on trade secret protection of our confidential information and know-how relating to our proprietary technology, platforms and product candidates, and continuing innovation to develop, strengthen, and maintain our proprietary position in our technology, platforms and product candidates. We expect to rely on data exclusivity, market exclusivity, patent term adjustment and patent term extensions when available.

As of January 15, 2021, we own two issued patents and nine pending patent applications in China and we also own two issued patents and seven pending applications in Europe filed with the European Patent Office, thirteen pending patent applications in the United States and 67 pending patent applications in other jurisdictions. Our patents and patent applications cover our key technologies and products, including the DPL platform, ADG106, ADG126, and ADG116. Excluding any patent term adjustment and patent term extension, our currently issued patents are expected to expire from 2033 to 2034. If any patents issue from our pending patent applications, excluding any patent term adjustments and patent term extension, such patents will be expected to expire from 2033 to 2041. The following table summarizes material pending patent applications in the United States, China, Europe and under Patent Cooperation Treaty, or PCT, covering our product candidates, including ADG106, ADG126, ADG116 and ADG104.

<u>Product Candidates</u>	<u>Title of Patent Application</u>	<u>Type of Patent Applications⁽¹⁾</u>	<u>Jurisdiction</u>
ADG106	Anti-CD137 molecules and uses thereof Combination therapy comprising anti-CD137 antibodies	Composition of matter/ method of use/ method of making method of treatment/ method of use	United States of America China and European Patent Office United States of America
ADG126	Anti-CTLA4 antibodies and methods of making and using the same	Composition of matter/ method of use/ method of making	United States of America, China and European Patent Office
ADG116	Anti-CTLA4 antibodies and methods of making and using the same	Composition of matter/ method of use/ method of making	United States of America, China and European Patent Office
ADG104	Anti-PD-L1 antibodies and use thereof	Composition of matter/ method of use/ method of making	United States of America and European Patent Office

(1) You should read the Risk Factors included elsewhere in this prospectus for important information about risks posed by the loss of patent protection, in particular the risks described under "Risk Factors—Risks Related to Our Intellectual Property".

The following table summarizes material issued patents in Europe and China covering our proprietary technologies.

<u>Application No.</u>	<u>Title of Patent</u>	<u>Type of Patent⁽¹⁾</u>	<u>Jurisdiction</u>
13877452.6	An Integrated System for Library Construction, Affinity Binder Screening and Expression Thereof	Method	European Patent Office
201380074656.1	An Integrated System for Library Construction, Affinity Binder Screening and Expression Thereof	Method	China
14908246.3	Methods and Systems for Autoinduction of Protein Expression	Method	European Patent Office
201410789857.6	Filter Vector System And Its Applications	Method	China

(1) You should read the Risk Factors included elsewhere in this prospectus for important information about risks posed by the loss of patent protection, in particular the risks described under "Risk Factors—Risks Related to Our Intellectual Property".

We continually assess and refine our intellectual property strategy as we develop new platform technologies and product candidates. To that end, we are prepared to file additional patent applications relating to the new technologies that we develop if our intellectual property strategy requires such filings, or where we seek to adapt to competition or seize business opportunities. In addition to filing and prosecuting patent applications in China and the United States, we may elect to file counterpart patent applications in additional countries and regions where we believe such foreign filing is likely to be beneficial.

As with other biopharmaceutical companies, our ability to maintain and solidify our proprietary and intellectual property position for our platform technologies and product candidates will depend on our success in obtaining effective patent claims and enforcing those claims if granted. However, our owned pending patent applications, and any patent applications that we may in the future file or license from third parties may not result in the issuance of patents. In addition, the term of individual issued patents depends upon the legal term for patents in the countries in which they are obtained. In most countries in which we have filed or intend to file in the future, including the United States, the patent term is 20 years from the earliest filing date of a nonprovisional patent application. We intend to seek patent term extensions to any of our issued patents in any jurisdiction where these are available; however, there is no guarantee that the applicable authorities will agree with our assessment of whether such extensions should be granted, and even if granted, the length of such extensions. The life of a patent, and the protection it affords, is therefore limited and once the patent life of our issued patents has expired, we may face competition, including from other competing technologies. As a result, our patent portfolio may not provide us with sufficient rights to exclude others from commercializing products similar or identical to ours.

We seek to ensure that investments made into the development of our technology are protected by relying on a combination of patents, trademarks, copyrights, trade secrets and contractual rights, including license agreements. In addition to our patent portfolio described above, as of January 15, 2021, our PRC subsidiary owns 25 registered trademarks relating to various aspects of our operations, and two registered domain names in China. To protect our rights, we seek to enter into confidentiality

agreements, nondisclosure agreements and employee disclosure and invention assignment agreements with our employees, contractors and other third parties who may have or need access to our confidential information. We have also employed internal policies, encryptions and data security measures to protect our proprietary rights. However, there can be no assurance that our efforts will be successful. If our employees, contractors or other third parties violate these agreements or otherwise infringe upon, misappropriate or otherwise violate our intellectual property rights, we may seek to enforce our rights against such parties. In addition, from time to time, third parties may initiate litigation against us alleging infringement, misappropriation or other violation of their proprietary rights or declaring their noninfringement of our intellectual property rights. An adverse result in any such proceeding could enjoin the commercialization of our technology platform and product candidates, result in significant damages, and have a material adverse effect on our business. Even if we are successful in any such litigation, we may be required to incur significant costs and dedicate significant personnel time in defending such litigation. For more information on these and other risks related to intellectual property, see "Risk Factors—Risks Related to Our Intellectual Property".

MANUFACTURING AND SUPPLY

We do not currently operate a cGMP-compliant manufacturing facility. We currently outsource the GMP manufacturing and quality control testing and cGMP quality assurance release of clinical trial material to WuXi Biologics. We have entered into a framework agreement with Wuxi Biologics, under which it provides services to us on a project-by-project basis. We are also working with other qualified manufacturers to provide diversified manufacturing and supply services. We also monitor the manufacturing activities of clinical trial material to ensure the compliance with local and international cGMP and applicable regulations. We have assembled a seasoned internal team with rich experience to drive and monitor the manufacturing process. Currently, Wuxi Biologics obtains raw materials and supplies for manufacturing activities from multiple suppliers who we believe have sufficient capacity to meet our demands. We expect to continue our relationships with WuXi Biologics but are continuously evaluating multiple vendors globally to ensure continuous supply of products for global clinical trials.

COMPETITION

The biotechnology and pharmaceutical industries are highly competitive and characterized by continuing technological advancement, significant competition and an emphasis on intellectual property. While we believe that our management's research, development and commercialization experience provide us with competitive advantages, we face potential competition from many different sources, including global biopharmaceutical companies, major multinational pharmaceutical companies, established biotechnology companies, specialty pharmaceutical companies, universities, academic institutions, government agencies and other public and private research organizations that conduct research, seek patent protection and establish collaborative arrangements for the research, development, manufacturing and commercialization of cancer immunotherapies. Any product candidates that we successfully develop and commercialize will compete with new immunotherapies and other drug products that may become available in the future. Our commercial opportunity could be reduced or eliminated if our competitors develop and commercialize products that are safer and more effective, have fewer or less severe side effects or are more convenient than any products that we may develop. Our competitors also may obtain FDA or other regulatory approval for their products more rapidly than we do.

We compete in the segments of the pharmaceutical, biotechnology and other related markets that develop cancer treatments. There are many other companies that have commercialized and/or are developing immuno-oncology treatments for cancer, including large pharmaceutical and biotechnology companies. Many of our competitors have significantly greater financial, technical and human resources than we have, and mergers and acquisitions in the biopharmaceutical industry may result in even more

resources being concentrated among a smaller number of our competitors. Smaller or early-stage companies may also prove to be significant competitors, particularly through collaborative arrangements with large and established companies. Our commercial opportunities could be reduced or eliminated if our competitors develop or market products or other novel therapies that are more effective, safer or less costly than our current or future product candidates, or obtain regulatory approval for their products more rapidly than we may obtain approval for our product candidates.

Our lead product candidate, ADG106, is an investigational fully human ligand-blocking, agonistic anti-CD137 IgG4 mAb that is designed to target a unique conserved epitope of CD137. While potentially unique for this type of product candidate, we expect our primary competition to be with other clinical-stage CD137 agonist product candidates. According to Frost & Sullivan, there are no marketed CD137 agonist drugs. The two leading molecules in clinical trials are utomilumab (PF-05082566) from Pfizer and urelumab (BMS-663513) from BMS. The following chart reflects CD137 agonist mAb programs in the global pipeline:

Global Pipelines of CD137 Agonist Mab Programs		
Drug	Company	Phase
		I/II
		I
		I
PF-05082566	Pfizer	II
		I/II
		I
		I/II
BMS-663513	BMS	II
		I/II
ATOR-1017	Alligator	I
AGEN2373	Agenus/Gilead	I
CTX-471	Compass	I
LVGN6051	Lyvgen	I

ADG126 and ADG116 are investigational, fully human anti-CTLA-4 mAbs generated through our NEObody and SAFEbody technologies, respectively. We expect our primary competition to be within the CTLA-4 antibody market, especially with ipilimumab. According to Frost & Sullivan, as of July 31, 2020, Yervoy (ipilimumab) from BMS is the only market CTLA-4 drug targeting cancer with six indications including monotherapy and combination therapies approved by FDA.

In January 2020, BMS submitted a biologics license application, or BLA, with the FDA for Opdivo in combination with Yervoy for the first-line treatment of patients with metastatic or recurrent NSCLC with no EGFR or ALK genomic tumor aberrations. In April 2020, the FDA approved Opdivo plus Yervoy, administered concomitantly with a limited course of chemotherapy, for the first-line treatment of patients with metastatic or recurrent NSCLC with no EGFR or ALK genomic tumor aberrations.

While Yervoy is the only marketed CTLA-4 antibody drug, there are over ten CTLA-4 antibody monotherapies and combination therapies in clinical development globally, according to Frost & Sullivan. Among the pipeline candidates, tremelimumab by AstraZeneca, as part of a combination

therapy with durvalumab, is undergoing Phase III clinical trial for six different indications. Four other drug candidates are undergoing Phase II clinical trial, according to Frost & Sullivan.

There are no marketed CTLA-4 antibody drugs in China, but at least seven CTLA-4 antibodies pipelines are in clinical development, according to the Frost & Sullivan Report. For example, BMS has submitted a new drug application for ipilimumab; tremelimumab by AstraZeneca has entered Phase III clinical trials for indications including NSCLC, small cell lung cancer and melanoma. Major limitations of CTLA-4 mAbs include toxicity. According to the Frost & Sullivan Report, nivolumab and ipilimumab combination therapy has shown relatively higher toxicity in clinical studies even at lower dosages, which was observed from the published clinical data. We believe that the continued expansion of indications, and the launch of innovative novel CTLA-4 antibodies with potential for higher safety and better efficacy may increase the market for CTLA-4 antibodies significantly.

EMPLOYEES

We had a total of 198 employees as of December 31, 2020. The following table sets forth the numbers of our employees categorized by function as of December 31, 2020. We also engage consultants and part-time staff as and when appropriate.

<u>Function</u>	<u>Number of Employees</u>
Research and Development	159
Computational Biology and Informatics	25
Technology Development	71
Drug Discovery	40
Clinical Development	23
General Administration	36
HR	6
Finance	9
IT	5
Administration	16
Business Development and Marketing	3
Total	<u>198</u>

Our success depends on our ability to attract, motivate, train and retain qualified personnel. We believe we offer our employees competitive compensation packages and an environment that encourages self-development. We regularly recruit new talents through campus events and colleague referral to build and develop our own talent pool. Through employee succession planning, we help employees understand their career path within Adagene, motivate them to remain in the organization and to achieve their personal career goals. Other initiatives for talent retention include executive coaching, employee surveys or engagement, training and development, compensation and rewards. As a result of these efforts, we have generally been able to attract and retain qualified personnel and maintain a stable core management team.

As required by regulations in China, we participate in various employee social security plans that are organized by municipal and provincial governments, including pension insurance, unemployment insurance, maternity insurance, work-related injury insurance, medical insurance and housing funds. We are required under PRC law to make contributions to employee benefit plans at specified percentages of the salaries, bonuses and certain allowances of our employees, up to a maximum amount specified by the local government from time to time. We have granted, and plan to continue to grant, share-based incentive awards to our employees in the future to incentivize their contributions to our growth and development.

We believe that we maintain a good working relationship with our employees, and we have not experienced any material labor disputes. None of our employees is subject to a collective bargaining agreement.

FACILITIES

Our headquarter is based in Suzhou, China, where we have our main administrative and laboratory offices, which are approximately 2,946 square meters in size. The expiration dates of the lease agreements for this facility range from March 31, 2021 to October 7, 2023. We also have a 2,673 square feet facility in San Diego, California for laboratory, research and development functions, the lease for which expires on August 31, 2023. We believe our current facilities are sufficient to meet our near-term needs and we do not expect difficulties in renewing lease agreements for our facilities.

INSURANCE

We provide social security insurance including pension insurance, unemployment insurance, work-related injury insurance and medical insurance for our employees. We maintain property insurance, general liability insurance, products/completed operations insurance, auto and international auto liability insurance, workers compensation insurance, international workers compensation insurance, accident and health insurance and director and officer liabilities insurance. We consider our insurance coverage sufficient and in line with market practice for our business operations in the industry.

LEGAL PROCEEDINGS

From time to time, we may become involved in legal proceedings or be subject to claims arising in the ordinary course of our business. We are not presently a party to any legal proceedings that, if determined adversely to us, would individually or taken together have a material adverse effect on our business, results of operations, financial condition or cash flows. Regardless of the outcome, litigation can have an adverse impact on us because of defense and settlement costs, diversion of management resources and other factors.

REGULATION

United States Regulation

The FDA and other regulatory authorities at federal, state, and local levels, as well as in foreign countries, extensively regulate, among other things, the research, development, testing, manufacture, quality control, import, export, safety, effectiveness, labeling, packaging, storage, distribution, record keeping, approval, advertising, promotion, marketing, post-approval monitoring and post-approval reporting of biologics such as those we are developing. We, along with third-party contractors, will be required to navigate the various preclinical, clinical and commercial approval requirements of the governing regulatory agencies of the countries in which we wish to conduct studies or seek approval or licensure of our product candidates.

In the United States, the FDA regulates biologic products under the Federal Food, Drug and Cosmetic Act, its implementing regulations and other laws, including, in the case of biologics, the Public Health Service Act. Our product candidates are subject to regulation by the FDA as biologics. Biologics require the submission of a BLA and licensure, which constitutes approval, by the FDA before being marketed in the United States. None of our product candidates has been approved by the FDA for marketing in the United States, and we currently have no BLAs pending. Failure to comply with applicable FDA or other requirements at any time during product development, clinical testing, the approval process or after approval may result in administrative or judicial sanctions. These sanctions could include the FDA's refusal to approve pending applications, suspension or revocation of approved applications, warning letters, product recalls, product seizures, total or partial suspensions of manufacturing or distribution, injunctions, fines, civil penalties or criminal prosecution.

The process required by the FDA before biologic product candidates may be marketed in the United States generally involves the following:

- completion of preclinical laboratory tests and animal studies performed in accordance with the FDA's GLP regulations;
- submission to the FDA of an IND, which must become effective before clinical trials may begin and must be updated annually or when significant changes are made;
- approval by an independent Institutional Review Board, or IRB, or ethics committee at each clinical site before the trial is commenced;
- performance of adequate and well-controlled human clinical trials to establish the safety and effectiveness of the proposed biologic product candidate for its intended indications;
- preparation of and submission to the FDA of a BLA when adequate data are obtained from pivotal clinical trials;
- a determination by the FDA within 60 days of its receipt of a BLA to accept the application for review;
- satisfactory completion of an FDA Advisory Committee review, if applicable;
- satisfactory completion of an FDA pre-approval inspection of the manufacturing facility or facilities at which the proposed product is produced to assess compliance with cGMP and to assure that the facilities, methods and controls are adequate to preserve the biological product's continued safety, purity and potency, and of selected clinical investigation sites to assess compliance with Good Clinical Practices, or GCP regulations; and
- FDA review and approval of the BLA to permit commercial marketing of the product for particular indications for use in the United States.

Preclinical and Clinical Development

Prior to beginning the first clinical trial with a product candidate in the United States, we must submit an IND application to the FDA. An IND application is a request for authorization from the FDA to administer an investigational new drug product to humans. The central focus of an IND application is on the general investigational plan and the protocol(s) for clinical trials. The IND application also includes results of animal and in vitro studies assessing the toxicology, pharmacokinetics, pharmacology, and pharmacodynamic characteristics of the product; chemistry, manufacturing, and controls information; and any available human data or literature to support the use of the investigational product. An IND must become effective before human clinical trials may begin. The IND automatically becomes effective 30 days after receipt by the FDA, unless the FDA, within the 30-day time period, raises safety concerns or questions about the proposed clinical trial. If the IND sponsor is not able to address FDA's concerns satisfactorily within the 30-day time frame, the IND may be placed on clinical hold. The IND sponsor and the FDA must resolve any outstanding concerns or questions before the IND is cleared by the FDA and the clinical trial can begin. Submission of an IND therefore may or may not result in FDA authorization to begin a clinical trial.

Clinical trials involve the administration of the investigational product to human subjects under the supervision of qualified investigators in accordance with GCPs, which include the requirement that all research subjects provide their informed consent for their participation in any clinical trial. Clinical trials are conducted under protocols detailing, among other things, the objectives of the study, the parameters to be used in monitoring safety and the effectiveness criteria to be evaluated. Generally, a separate submission to the existing IND must be made for each successive clinical trial conducted during product development and for any subsequent protocol amendments. Furthermore, an independent IRB for each site proposing to conduct the clinical trial must review and approve the plan for any clinical trial and its informed consent form before the clinical trial begins at that site, and must monitor the study until completed. Regulatory authorities, the IRB or the sponsor may suspend a clinical trial at any time on various grounds, including a finding that the subjects are being exposed to an unacceptable health risk or that the trial is unlikely to meet its stated objectives. Some studies also include oversight by an independent group of qualified experts organized by the clinical trial sponsor, known as a data safety monitoring board, or DSMB, which provides recommendation on whether or not a study should move forward at designated check points based on access to certain data from the study. The DSMB may recommend halting of the clinical trial if it determines that there is an unacceptable safety risk for subjects or on other grounds, such as no demonstration of efficacy. There are also requirements governing the reporting of ongoing clinical trials and clinical trial results to public registries.

For purposes of BLA approval, human clinical trials are typically conducted in three sequential phases that may overlap.

- Phase I—The investigational product is initially introduced into healthy human subjects or patients with the target disease or condition. These studies are designed to test the safety, dosage tolerance, absorption, metabolism and distribution of the investigational product in humans, the side effects associated with increasing doses, and, if possible, to gain early evidence on effectiveness. For investigational products developed for oncology indications, the Phase I trials are normally conducted in patients with serious or life-threatening diseases without other treatment alternatives.
- Phase II—The investigational product is administered to a limited patient population with a specified disease or condition to evaluate the preliminary efficacy, optimal dosages and dosing schedule and to identify possible adverse side effects and safety risks. Multiple Phase II clinical trials may be conducted to obtain information prior to beginning larger and more expensive Phase III clinical trials. For certain indications in patients with serious or life-threatening

diseases and with no available therapies, it may be possible to obtain BLA approval based on data from Phase II trials if a positive benefit risk profile is demonstrated.

- Phase III—The investigational product is administered to an expanded patient population to further evaluate dosage, to provide statistically significant evidence of clinical efficacy and to further test for safety, generally at multiple geographically dispersed clinical trial sites. These clinical trials are intended to establish the overall risk/benefit ratio of the investigational product and to provide an adequate basis for product approval.

In some cases, the FDA may require, or companies may voluntarily pursue, additional clinical trials after a product is approved to gain more information about the product. These so-called Phase 4 studies may be made a condition to approval of the BLA. Concurrent with clinical trials, companies may complete additional animal studies and develop additional information about the biological characteristics of the product candidate, and must finalize a process for manufacturing the product in commercial quantities in accordance with cGMP requirements. The manufacturing process must be capable of consistently producing quality batches of the product candidate and, among other things, must develop methods for testing the identity, strength, quality and purity of the final product, or for biologics, the safety, purity and potency. Additionally, appropriate packaging must be selected and tested and stability studies must be conducted to demonstrate that the product candidate does not undergo unacceptable deterioration over its shelf life.

BLA Submission and Review

Assuming successful completion of all required testing in accordance with all applicable regulatory requirements, the results of product development, preclinical studies and clinical trials are submitted to the FDA as part of a BLA requesting approval to market the product for one or more indications. The BLA must include all relevant data available from pertinent preclinical and clinical trials, including negative or ambiguous results as well as positive findings, together with detailed information relating to the product's chemistry, manufacturing, controls and proposed labeling, among other things. The submission of a BLA requires payment of a substantial application user fee to the FDA unless a waiver or exemption applies.

Once an original BLA has been submitted, FDA has 60 days to determine whether the application can be filed. If FDA determines that an application to be deficient, on its face, in a way that precludes a complete review, FDA may not accept the application for review and may issue a refuse-to-file letter to the sponsor. If FDA determines the application is filable, the FDA's goal is to review standard applications within ten months after it accepts the application for filing, or, if the application qualifies for priority review, six months after the FDA accepts the application for filing. In both standard and priority reviews, the review process is often significantly extended by FDA requests for additional information or clarification. The FDA reviews a BLA to determine, among other things, whether a product is safe, pure and potent and the facilities in which it is manufactured, processed, packed, or held meets standards designed to assure the product's continued safety, purity and potency. The FDA may convene an advisory committee to provide clinical insight on application review questions. Before approving a BLA, the FDA will typically inspect the facility or facilities where the product is manufactured. The FDA will not approve an application unless it determines that the manufacturing processes and facilities are in compliance with cGMP requirements and adequate to assure consistent production of the product within required specifications. Additionally, before approving a BLA, the FDA will typically inspect one or more clinical sites to assure compliance with GCP. If the FDA determines that the application, manufacturing process or manufacturing facilities are not acceptable, it will outline the deficiencies in the submission and often will request additional testing or information. Notwithstanding the submission of any requested additional information, the FDA ultimately may decide that the application does not satisfy the regulatory criteria for approval.

After the FDA evaluates a BLA and conducts inspections of manufacturing facilities where the commercial product and/or its drug substance will be produced, the FDA may issue an approval letter or a Complete Response letter. An approval letter authorizes commercial marketing of the product with specific prescribing information for specific indications. A Complete Response letter will describe all of the deficiencies that the FDA has identified in the BLA, except that where the FDA determines that the data supporting the application are inadequate to support approval, the FDA may issue the Complete Response letter without first conducting required inspections, testing submitted product lots, and/or reviewing proposed labeling. In issuing the Complete Response letter, the FDA may recommend actions that the applicant might take to place the BLA in condition for approval, including requests for additional information or clarification. The FDA may delay or refuse approval of a BLA if applicable regulatory criteria are not satisfied, require additional testing or information and/or require post-marketing testing and surveillance to monitor safety or efficacy of a product.

If regulatory approval of a product is granted, such approval will be granted for particular indications and may entail limitations on the indicated uses for which such product may be marketed. For example, the FDA may approve the BLA with a Risk Evaluation and Mitigation Strategy, or REMS, to ensure the benefits of the product outweigh its risks. A REMS is a safety strategy to manage a known or potential serious risk associated with a product and to enable patients to have continued access to such medicines by managing their safe use, and could include medication guides, physician communication plans, or elements to assure safe use, such as restricted distribution methods, patient registries and other risk minimization tools. The FDA also may condition approval on, among other things, changes to proposed labeling or the development of adequate controls and specifications. Once approved, the FDA may withdraw the product approval if compliance with pre- and post-marketing requirements is not maintained or if problems occur after the product reaches the marketplace. The FDA may require one or more Phase 4 post-market studies and surveillance to further assess and monitor the product's safety and effectiveness after commercialization, and may limit further marketing of the product based on the results of these post-marketing studies.

Expedited Development and Review Programs

The FDA offers a number of expedited development and review programs for qualifying product candidates. The fast track program is intended to expedite or facilitate the process for reviewing new products that meet certain criteria. Specifically, new products are eligible for fast track designation if they are intended to treat a serious or life-threatening disease or condition and demonstrate the potential to address unmet medical needs for the disease or condition. Fast track designation applies to the combination of the product and the specific indication for which it is being studied. The sponsor of a fast track product has opportunities for frequent interactions with the review team during product development and, once a BLA is submitted, the product may be eligible for priority review. A fast track product may also be eligible for rolling review, in which case the FDA may consider for review sections of the BLA on a rolling basis before the complete application is submitted, if the sponsor provides a schedule for the submission of the sections of the BLA, the FDA agrees to accept sections of the BLA and determines that the schedule is acceptable, and the sponsor pays any required user fees upon submission of the first section of the BLA.

A product intended to treat a serious or life-threatening disease or condition may also be eligible for breakthrough therapy designation to expedite its development and review. A product can receive breakthrough therapy designation if preliminary clinical evidence indicates that the product may demonstrate substantial improvement over existing therapies on one or more clinically significant endpoints, such as substantial treatment effects observed early in clinical development. The designation includes all of the fast track program features, as well as more intensive FDA interaction and guidance beginning as early as Phase I and an organizational commitment to expedite the development and review of the product, including involvement of senior managers.

Any marketing application for a biologic submitted to the FDA for approval, including a product with a fast track designation and/or breakthrough therapy designation, may be eligible for other types of FDA programs intended to expedite the FDA review and approval process, such as priority review and accelerated approval. A product is eligible for priority review if it has the potential to provide a significant improvement in the treatment, diagnosis or prevention of a serious disease or condition compared to marketed products. For products containing new molecular entities, priority review designation means the FDA's goal is to take action on the marketing application within six months of the 60-day filing date (compared with ten months under standard review).

Additionally, products studied for their safety and effectiveness in treating serious or life-threatening diseases or conditions may receive accelerated approval upon a determination that the product has an effect on a surrogate endpoint that is reasonably likely to predict clinical benefit, or on a clinical endpoint that can be measured earlier than irreversible morbidity or mortality, that is reasonably likely to predict an effect on irreversible morbidity or mortality or other clinical benefit, taking into account the severity, rarity, or prevalence of the condition and the availability or lack of alternative treatments. As a condition of accelerated approval, the FDA will generally require the sponsor to perform adequate and well-controlled post-marketing clinical trials to verify and describe the anticipated effect on irreversible morbidity or mortality or other clinical benefit. In addition, the FDA currently requires as a condition for accelerated approval pre-approval of promotional materials, which could adversely impact the timing of the commercial launch of the product.

In 2017, FDA established a new regenerative medicine advanced therapy, or RMAT, designation as part of its implementation of the 21st Century Cures Act, which was signed into law in December 2016. The RMAT designation program is intended to fulfill the 21st Century Cures Act requirement that FDA facilitate an efficient development program for, and expedite review of, any drug that meets the following criteria: (1) it qualifies as a RMAT, which is defined as a cell therapy, therapeutic tissue engineering product, human cell and tissue product, or any combination product using such therapies or products, with limited exceptions; (2) it is intended to treat, modify, reverse, or cure a serious or life-threatening disease or condition; and (3) preliminary clinical evidence indicates that the drug has the potential to address unmet medical needs for such a disease or condition. Like fast track and breakthrough therapy designation, RMAT designation provides potential benefits that include more frequent meetings with the FDA to discuss the development plan for the product candidate and eligibility for rolling review and priority review. Products granted RMAT designation may also be eligible for accelerated approval on the basis of a surrogate or intermediate endpoint reasonably likely to predict long-term clinical benefit, or reliance upon data obtained from a meaningful number of sites, including through expansion to additional sites. Once approved, when appropriate, the FDA can permit fulfillment of post-approval requirements under accelerated approval through the submission of clinical evidence, clinical trials, patient registries, or other sources of real-world evidence such as electronic health records; through the collection of larger confirmatory datasets; or through post-approval monitoring of all patients treated with the therapy prior to approval.

Fast track designation, breakthrough therapy designation, priority review, accelerated approval, and RMAT designation do not change the standards for approval but may expedite the development or approval process.

Orphan Drug Designation

Under the Orphan Drug Act, the FDA may grant orphan designation to a drug or biologic intended to treat a rare disease or condition, which is a disease or condition that affects fewer than 200,000 individuals in the United States, or if it affects more than 200,000 individuals in the United States, there is no reasonable expectation that the cost of developing and making available a drug or biologic for this type of disease or condition will be recovered from sales in the United States for that drug or biologic. Orphan drug designation must be requested before submitting a BLA. After the FDA

grants orphan drug designation, the generic identity of the therapeutic agent and its potential orphan use are disclosed publicly by the FDA. The orphan drug designation does not convey any advantage in, or shorten the duration of, the regulatory review or approval process.

If a product that has orphan drug designation subsequently receives the first FDA approval for the disease for which it has such designation, the product is entitled to orphan drug exclusive approval (or exclusivity), which means that the FDA may not approve any other applications, including a full BLA, to market the same biologic for the same indication for seven years, except in limited circumstances, such as a showing of clinical superiority to the product with orphan drug exclusivity. Orphan drug exclusivity does not prevent the FDA from approving a different drug or biologic for the same disease or condition, or the same drug or biologic for a different disease or condition. Among the other benefits of orphan drug designation are tax credits for certain research and a waiver of the BLA application fee.

A designated orphan drug may not receive orphan drug exclusivity if it is approved for a use that is broader than the indication for which it received orphan designation. In addition, exclusive marketing rights in the United States may be lost if the FDA later determines that the request for designation was materially defective or if the manufacturer is unable to assure sufficient quantities of the product to meet the needs of patients with the rare disease or condition.

Post-Approval Requirements

Any products manufactured or distributed by us pursuant to FDA approvals are subject to pervasive and continuing regulation by the FDA, including, among other things, requirements relating to record keeping, reporting of adverse experiences, periodic reporting, product sampling and distribution, and advertising and promotion of the product. After approval, most changes to the approved product, such as adding new indications or other labeling claims, are subject to prior FDA review and approval. There also are continuing user fee requirements, under which FDA assesses an annual program fee for each product identified in an approved BLA. Biologic manufacturers and their subcontractors are required to register their establishments with the FDA and certain state agencies, and are subject to periodic unannounced inspections by the FDA and certain state agencies for compliance with cGMP, which impose certain procedural and documentation requirements upon us and our third-party manufacturers. Changes to the manufacturing process are strictly regulated, and, depending on the significance of the change, may require prior FDA approval before being implemented. FDA regulations also require investigation and correction of any deviations from cGMP and impose reporting requirements upon us and any third-party manufacturers that we may decide to use. Accordingly, manufacturers must continue to expend time, money and effort in the area of production and quality control to maintain compliance with cGMP and other aspects of regulatory compliance.

The FDA may withdraw approval if compliance with regulatory requirements and standards is not maintained or if problems occur after the product reaches the market. Later discovery of previously unknown problems with a product, including adverse events of unanticipated severity or frequency, or with manufacturing processes, or failure to comply with regulatory requirements, may result in revisions to the approved labeling to add new safety information; imposition of post-market studies or clinical trials to assess new safety risks; or imposition of distribution restrictions or other restrictions under a REMS program. Other potential consequences include, among other things:

- restrictions on the marketing or manufacturing of a product, complete withdrawal of the product from the market or product recalls;

- fines, warning letters or holds on post-approval clinical trials;
- refusal of the FDA to approve pending applications or supplements to approved applications, or suspension or revocation of existing product approvals;
- product seizure or detention, or refusal of the FDA to permit the import or export of products; or
- injunctions or the imposition of civil or criminal penalties.

The FDA closely regulates the marketing, labeling, advertising and promotion of biologics. A company can make only those claims relating to safety and efficacy, purity and potency that are approved by the FDA and in accordance with the provisions of the approved label. The FDA and other agencies actively enforce the laws and regulations prohibiting the promotion of off-label uses. Failure to comply with these requirements can result in, among other things, adverse publicity, warning letters, corrective advertising and potential civil and criminal penalties. Physicians may prescribe legally available products for uses that are not described in the product's labeling and that differ from those tested by us and approved by the FDA. Such off-label uses are common across medical specialties. Physicians may believe that such off-label uses are the best treatment for many patients in varied circumstances. The FDA does not regulate the behavior of physicians in their choice of treatments. The FDA does, however, restrict manufacturer's communications on the subject of off-label use of their products.

Biosimilars and Reference Product Exclusivity

The Patient Protection and Affordable Care Act, as amended by the Health Care and Education Reconciliation Act, or collectively, the ACA, signed into law in 2010, includes a subtitle called the Biologics Price Competition and Innovation Act of 2009, or BPCIA, which created an abbreviated approval pathway for biological products that are biosimilar to or interchangeable with an FDA-approved reference biological product.

Biosimilarity, which requires that there be no clinically meaningful differences between the biological product and the reference product in terms of safety, purity, and potency, can be shown through analytical studies, animal studies, and a clinical trial or studies. Interchangeability requires that a product be biosimilar to the reference product and the product must demonstrate that it can be expected to produce the same clinical results as the reference product in any given patient and, for products that are administered to a patient more than once, the biologic and the reference biologic may be alternated or switched after one has been previously administered without increasing safety risks or risks of diminished efficacy relative to exclusive use of the reference biologic. Complexities associated with the larger, and often more complex, structures of biological products, as well as the processes by which such products are manufactured, pose significant hurdles to implementation of the abbreviated approval pathway that are still being worked out by the FDA.

Under the BPCIA, an application for a biosimilar product may not be submitted to the FDA until four years following the date that the reference product was first licensed by the FDA. In addition, the FDA may not approve a biosimilar product until 12 years from the date on which the reference product was first licensed. During this 12-year period of exclusivity, another company may still market a competing version of the reference product if the FDA approves a full BLA containing that applicant's own preclinical data and data from adequate and well-controlled clinical trials to demonstrate the safety, purity and potency of the competing product. The BPCIA also created certain exclusivity periods for biosimilars approved as interchangeable products. At this juncture, it is unclear whether products deemed "interchangeable" by the FDA will, in fact, be readily substituted by pharmacies, which are governed by state pharmacy law.

The BPCIA is complex and continues to be interpreted and implemented by the FDA. In addition, government proposals have sought to reduce the 12-year reference product exclusivity period. Other aspects of the BPCIA, some of which may impact the BPCIA exclusivity provisions, have also been the subject of recent litigation. As a result, the ultimate implementation and impact of the BPCIA is subject to significant uncertainty.

Other Healthcare Laws and Compliance Requirements

Pharmaceutical companies are subject to additional healthcare regulation and enforcement by the federal government and by authorities in the states and foreign jurisdictions in which they conduct their business. Such laws include, without limitation:

- the U.S. federal Anti-Kickback Statute, which prohibits, among other things, persons and entities from knowingly and willfully soliciting, receiving, offering or paying remuneration, to induce, or in return for, either the referral of an individual, or the purchase or recommendation of an item or service for which payment may be made under any federal healthcare program;
- federal civil and criminal false claims laws, including the civil False Claims Act, and civil monetary penalty laws, which prohibit, among other things, individuals or entities from knowingly presenting, or causing to be presented, claims for payment to the federal government, including federal healthcare programs, that are false or fraudulent;
- the federal Health Insurance Portability and Accountability Act of 1996, or HIPAA, which created additional federal criminal statutes which prohibit, among other things, executing a scheme to defraud any healthcare benefit program and making false statements relating to healthcare matters, and which, as amended by the Health Information Technology for Economic and Clinical Health Act of 2009, or HITECH, also imposes certain requirements on HIPAA covered entities and their business associates relating to the privacy, security and transmission of individually identifiable health information;
- the U.S. federal Physician Payments Sunshine Act, which requires certain manufacturers of drugs, devices, biologics and medical supplies for which payment is available under Medicare, Medicaid or the Children's Health Insurance Program, with specific exceptions, to annually report to the federal government, information related to payments or other transfers of value made to physicians, as defined by such law, certain other health care providers beginning in 2022, and teaching hospitals, as well as ownership and investment interests held by physicians and their immediate family members; and
- United States state and foreign law equivalents of each of the above federal laws, which, in some cases, differ from each other in significant ways, and may not have the same effect, thus complicating compliance efforts, including laws governing the privacy and security of personal data, such as the GDPR, which imposes obligations and restrictions on the collection and use of personal data relating to individuals located in the EU and EEA (including with regard to health data).

If their operations are found to be in violation of any of such laws or any other governmental regulations that apply, they may be subject to significant penalties, including, without limitation, civil, criminal and administrative penalties, damages, fines, exclusion from government-funded healthcare programs, such as Medicare and Medicaid or similar programs in other countries or jurisdictions, integrity oversight and reporting obligations to resolve allegations of non-compliance, disgorgement, imprisonment, contractual damages, reputational harm, diminished profits and the curtailment or restructuring of our operations.

Coverage and Reimbursement

Significant uncertainty exists as to the coverage and reimbursement status of any pharmaceutical or biological product for which we obtain regulatory approval. Sales of any product depend, in part, on the extent to which such product will be covered by third-party payers, such as federal, state, and foreign government healthcare programs, commercial insurance and managed healthcare organizations, and the level of reimbursement for such product by third-party payers. Decisions regarding the extent of coverage and amount of reimbursement to be provided are made on a plan-by-plan basis. As there is no uniform policy of coverage and reimbursement for drug products among third-party payers in the United States, coverage and reimbursement policies for drug products can differ significantly from payer to payer. There may be significant delays in obtaining coverage and reimbursement as the process of determining coverage and reimbursement is often time-consuming and costly which will require us to provide scientific and clinical support for the use of our products to each payer separately, with no assurance that coverage or adequate reimbursement will be obtained. It is difficult to predict at this time what government authorities and third-party payers will decide with respect to coverage and reimbursement for our drug products. For products administered under the supervision of a physician, obtaining coverage and adequate reimbursement may be particularly difficult because of the higher prices often associated with such drugs. Additionally, separate reimbursement for the product itself or the treatment or procedure in which the product is used may not be available, which may impact physician utilization.

In addition, the U.S. government, state legislatures and foreign governments have continued implementing cost-containment programs, including price controls, restrictions on coverage and reimbursement and requirements for substitution of generic products. Third-party payers are increasingly challenging the prices charged for medical products and services, examining the medical necessity and reviewing the cost effectiveness of pharmaceutical or biological products, medical devices and medical services, in addition to questioning safety and efficacy. Adoption of price controls and cost-containment measures, and adoption of more restrictive policies in jurisdictions with existing controls and measures, could further limit sales of any product. Decreases in third-party reimbursement for any product or a decision by a third-party payer not to cover a product could reduce physician usage and patient demand for the product.

Healthcare Reform

The United States and some foreign jurisdictions are considering or have enacted a number of reform proposals to change the healthcare system. There is significant interest in promoting changes in healthcare systems with the stated goals of containing healthcare costs, improving quality or expanding access. In the United States, the pharmaceutical industry has been a particular focus of these efforts and has been significantly affected by federal and state legislative initiatives, including those designed to limit the pricing, coverage, and reimbursement of pharmaceutical and biopharmaceutical products, especially under government-funded healthcare programs, and increased governmental control of drug pricing.

In March 2010, the ACA was signed into law, which substantially changed the way healthcare is financed by both governmental and private insurers in the United States, and significantly affected the pharmaceutical industry. The ACA contains a number of provisions of particular import to the pharmaceutical and biotechnology industries, including, but not limited to, those governing enrollment in federal healthcare programs, a new methodology by which rebates owed by manufacturers under the Medicaid Drug Rebate Program are calculated for drugs that are inhaled, infused, instilled, implanted or injected, and annual fees based on pharmaceutical companies' share of sales to federal healthcare programs.

Since its enactment, there have been judicial, Congressional, and executive branch challenges to certain aspects of the ACA, and we expect there will be additional challenges and amendments to the ACA in the future. For example, the 2020 federal spending package permanently eliminates, effective January 1, 2020, the ACA-mandated "Cadillac" tax on high-cost employer-sponsored health coverage and medical device tax and, effective January 1, 2021, also eliminates the health insurer tax. In addition, the Tax Act was enacted, which, among other things, removes penalties for not complying with ACA's individual mandate to carry health insurance. On December 14, 2018, a U.S. District Court Judge in the Northern District of Texas ruled that the individual mandate is a critical and inseparable feature of the ACA, and therefore, because it was repealed as part of the Tax Act, the remaining provisions of the ACA are invalid as well. Additionally, on December 18, 2019, the U.S. Court of Appeals for the 5th Circuit upheld the District Court ruling that the individual mandate was unconstitutional and remanded the case back to the District Court to determine whether the remaining provisions of the ACA are invalid as well. On March 2, 2020, the U.S. Supreme Court granted the petitions for writs of certiorari to review this case, although it is unclear when or how the Supreme Court will rule. It is also unclear how other efforts to challenge, repeal or replace the ACA will impact the ACA.

Other legislative changes have been proposed and adopted since the ACA was enacted, including aggregate reductions of Medicare payments to providers of 2% per fiscal year and reduced payments to several types of Medicare providers, which will remain in effect through 2030, with the exception of a temporary suspension from May 1, 2020 through December 31, 2020, unless additional Congressional action is taken. Moreover, there has recently been heightened governmental scrutiny over the manner in which manufacturers set prices for their marketed products, which has resulted in several Congressional inquiries and proposed and enacted federal and state legislation designed to, among other things, bring more transparency to product pricing, review the relationship between pricing and manufacturer patient programs, and reform government program reimbursement methodologies for drug products.

At the federal level, the Trump administration's budget proposal for fiscal year 2021 includes a US\$135 billion allowance to support legislative proposals seeking to reduce drug prices, increase competition, lower out-of-pocket drug costs for patients, and increase patient access to lower-cost generic and biosimilar drugs. Further, the Trump administration released a "Blueprint," or plan, to lower drug prices and reduce out of pocket costs of drugs that contains additional proposals to increase drug manufacturer competition, increase the negotiating power of certain federal healthcare programs, incentivize manufacturers to lower the list price of their products, and reduce the out-of-pocket costs of drug products paid by consumers. The Department of Health and Human Services, or HHS, has solicited feedback on some of these measures and has implemented others under its existing authority. For example, in May 2019, the Centers for Medicare & Medicaid Services, or CMS, issued a final rule to allow Medicare Advantage plans the option to use step therapy for Part B drugs beginning January 1, 2020. This final rule codified CMS's policy change that was effective January 1, 2019. Most recently, in July 2020, President Trump also signed a number of executive orders that attempt to implement several of the Administration's proposals. While some of measures may require additional authorization to become effective, Congress and the Trump administration have each indicated that it will continue to seek new legislative and/or administrative measures to control drug costs.

At the state level, legislatures have increasingly passed legislation and implemented regulations designed to control pharmaceutical product pricing, including price or patient reimbursement constraints, discounts, restrictions on certain product access and marketing cost disclosure and transparency measures, and, in some cases, designed to encourage importation from other countries and bulk purchasing. We expect that these and other healthcare reform measures that may be adopted in the future, may result in more rigorous coverage criteria and lower reimbursement, and in additional downward pressure on the price that we receive for any approved product.

PRC Regulation

We are subject to a variety of PRC laws, rules and regulations affecting many aspects of our business. This section sets out a summary of the major relevant laws, regulations, rules and policies which may have material impact on our business and operations.

Regulations on Company Establishment and Foreign Investment

The establishment, operation and management of corporate entities in China are governed by the Company Law of PRC, or the PRC Company Law, which was promulgated by the Standing Committee of the National People's Congress, or the NPC, in December 1993 and further amended in December 1999, August 2004, October 2005, December 2013 and October 2018, respectively. According to the PRC Company Law, companies are generally classified into two categories: limited liability companies and companies limited by shares. The PRC Company Law also applies to foreign-invested limited liability companies. According to the PRC Company Law, where laws on foreign investment have other stipulations, such stipulations shall prevail.

Investment activities in the PRC by foreign investors are governed by the Guiding Foreign Investment Direction, which was promulgated by the State Council in February 2002 and came into effect in April 2002, and the Special Administrative Measures for the Access of Foreign Investment (Negative List), or the Negative List, which was promulgated by the Ministry of Commerce of the PRC, or the MOFCOM, and the National Development and Reform Commission, or the NDRC, in June 2020 and came into effect in July 2020. The Negative List sets out the restrictive measures in a unified manner, such as the requirements on shareholding percentages and management, for the access of foreign investments, and the industries that are prohibited from receiving foreign investment. The Negative List covers 12 industries, and any field not falling under the Negative List shall be administered under the principle of equal treatment to domestic and foreign investment.

Foreign Investment Law of the PRC, or the Foreign Investment Law, was promulgated by the NPC in March 2019 and came into effect in January 2020. When the Foreign Investment Law came into effect, the Law on Wholly Foreign-owned Enterprises of the PRC, the Law on Sino-foreign Equity Joint Ventures of the PRC and the Law on Sino-foreign Cooperative Joint Ventures of the PRC were repealed simultaneously. The investment activities of foreign natural persons, enterprises or other organizations (collectively, the "foreign investors") directly or indirectly within the territory of China shall comply with and be governed by the Foreign Investment Law. Such activities include: 1) establishing by foreign investors of foreign-invested enterprises in China alone or jointly with other investors; 2) acquiring by foreign investors of shares, equity, property shares, or other similar interests of Chinese domestic enterprises; 3) investing by foreign investors in new projects in China alone or jointly with other investors; and 4) other forms of investment prescribed by laws, administrative regulations or the State Council.

In December 2019, the State Council promulgated the Regulations on Implementing the Foreign Investment Law of the PRC, which came into effect in January 2020. When the Regulations on Implementing the Foreign Investment Law of the PRC came into effect, the Regulation on Implementing the Sino-Foreign Equity Joint Venture Enterprise Law of the PRC, Provisional Regulations on the Duration of Sino-Foreign Equity Joint Venture Enterprise, the Regulations on Implementing the Wholly Foreign-Invested Enterprise Law of the PRC and the Regulations on Implementing the Sino-Foreign Cooperative Joint Venture Enterprise Law of the PRC were repealed simultaneously.

In December 2019, the MOFCOM and the State Administration for Market Regulation, or the SAMR promulgated the Measures on Reporting of Foreign Investment Information, which came into effect in January 2020. When the Measures on Reporting of Foreign Investment Information came into effect, the Interim Measures for the Administration of Filing for Establishment and Changes in Foreign

Investment Enterprises were repealed simultaneously. Since January 1, 2020, for foreign investors carrying out investment activities directly or indirectly in China, the foreign investors or foreign-invested enterprises shall submit investment information to the relevant commerce administrative authorities according to the Measure on Reporting of Foreign Investment Information.

Regulation on Pharmaceutical Product Development, Approval and Registration

Drug Regulatory Regime

The Drug Administration Law of the PRC, or the Drug Administration Law, was promulgated by the Standing Committee of the NPC, in September 1984. The two latest amendments to the Drug Administration Law were the amendment promulgated in April 2015 and in August 2019. The Regulations for the Implementation of the Drug Administration Law was promulgated by the State Council in August 2002, and was last amended in March 2019. The Drug Administration Law and the Regulations for the Implementation of the Drug Administration Law have jointly established the legal framework for the administration of pharmaceutical products in China, including the research, development and manufacturing of new drugs. The Drug Administration Law applies to entities and individuals engaged in the development, production, trade, application, supervision and administration of pharmaceutical products. It regulates and provides for a framework for the administration of pharmaceutical manufacturers, pharmaceutical trading companies and medicinal preparations of medical institutions, and the development, research, manufacturing, distribution, packaging, pricing and advertisements of pharmaceutical products. The Regulations for the Implementation of the Drug Administration Law, at the same time, provide the detailed implementation regulations for the Drug Administration Law.

In 2017, the drug regulatory system entered a new and significant period of reform. The General Office of the State Council and the General Committee of China Communist Party jointly issued Opinions on Deepening the Reform of the Evaluation and Approval Systems and Encouraging Innovation on Drugs and Medical Devices, or the Innovation Opinions. The expedited programs, the record-filing system, the prioritized review mechanism, the acceptance of foreign clinical data under the Innovation Opinions and other recent reforms encourage drug manufacturers to seek marketing approval in China first for the development of drugs in highly prioritized therapeutic areas, such as oncology or rare diseases.

To implement the regulatory reform introduced by Innovation Opinions, the Standing Committee of the NPC, the National Medical Products Administration, or the NMPA, a newly formed government authority as well as other authorities, are currently responsible for revising the laws, regulations and rules governing the pharmaceutical products and the industry.

In August 2019, the Standing Committee of the NPC promulgated the new Drug Administration Law, or the 2019 Amendment, which came into effect in December 2019. The 2019 Amendment contains many of the major reform initiatives implemented by the Chinese government since 2015, including but not limited to the Marketing Authorization Holder, or the MAH, system, conditional approvals of drugs, traceability system of drugs, and the cancellation of relevant certification according to the Good Manufacturing Practice, or the GMP, and the Good Supply Practice, or the GSP.

Regulatory Authorities

Pharmaceutical products in China are monitored and supervised on a national scale by the NMPA. The local provincial medical products administrative authorities are responsible for supervision and administration of drugs within their respective administrative regions. The NMPA was newly formed under the SAMR. The NMPA's predecessor, the State Drug Administration, or the SDA, was replaced by the State Food and Drug Administration, or the SFDA, which was later reorganized into the China

Food and Drug Administration, or the CFDA, as part of the institutional reforms implemented by the State Council.

The primary responsibilities of the NMPA include:

- monitoring and supervising the administration of pharmaceutical products, medical appliances and equipment as well as cosmetics in the PRC;
- formulating administrative rules and policies concerning the supervision and administration of pharmaceutical, medical devices, and cosmetics industry;
- evaluating, registering and approving new drugs, generic drugs, imported drugs and traditional Chinese medicine;
- approving and issuing permits for the manufacture and export/import of pharmaceutical products, medical appliances and equipment;
- approving the establishment of enterprises to be engaged in the manufacture and distribution of pharmaceutical products;
- examining and evaluating the safety of pharmaceutical products, medical devices, and cosmetics; and
- managing significant accidents involving pharmaceutical products, medical devices and cosmetics.

In 2013, the Ministry of Health, or the MOH, and the National Population and Family Planning Commission were integrated into the National Health and Family Planning Commission of the PRC, or the NHFPC. In March 2018, the First Session of the Thirteenth NPC approved the State Council Institutional Reform Proposal, according to which, NHFPC and certain other governmental authorities were consolidated into the National Health Commission, or the NHC. The responsibilities of the NHC include coordinating the formulation of national drug policies, the national essential medicine system and the National Essential Medicines List and drafting the administrative rules for the procurement, distribution and use of national essential medicines.

According to the Decision of the CFDA on Adjusting the Approval Procedures under the Administrative Approval Items for Certain Drugs, which was promulgated by the CFDA in March 2017 and came into effect in May 2017, the Investigational New Drug Application, or the IND, approval should be issued by the Center for Drug Evaluation, or the CDE, on behalf of the CFDA.

Regulations on Clinical Trials and Registration of Drugs

Administrative Measures for Drug Registration

In July 2007, the SFDA promulgated the amended version of the Administrative Measures for Drug Registration, or the Registration Measures, which became effective in October 2007. The Registration Measures mainly cover: (1) definitions of drug registration applications and regulatory responsibilities of drug administration; (2) general requirements for drug registration, including application for registration of new drugs, generic drugs, imported drugs and supplemental application, as well as application for re-registration; (3) clinical trials; (4) application, examination and approval of new drugs, generic drugs and imported drugs; (5) supplemental applications and re-registrations of drugs; (6) inspections; (7) registration standards and specifications; (8) time limit; (9) re-examination; and (10) liabilities and other supplementary provisions.

According to the Registration Measures, drug registration applications are divided into three different types, namely Domestic New Drug Application, Domestic Generic Drug Application and Imported Drug Application. Drugs which fall into one of three general types are divided according to the drug's working mechanism, namely whether the drug is classified as a chemical medicine, a

biological product, a traditional Chinese medicine or a natural medicine. New Drug Application, or NDA, refers to an application for registration of a drug that has not yet been marketed for sale in China. In addition, the registration of drugs that change the dosage form of the marketed drugs, change the route of administration and increase the new indications shall be reported in accordance with the application procedures for new drugs. Under the Registration Measures, a Category 1 drug refers to a new drug that has never been marketed in any country, and such drug is eligible for special review or fast track approval by the NMPA.

In January 2020, the SAMR released the amended Administrative Measures for Drug Registration, or the Amended Registration Measures, which came into effect in July 2020. The Amended Registration Measures provide detailed procedural and substantive requirements for the key regulatory concepts established by the Drug Administration Law, and confirms a number of reform actions that have been taken in the past years, including but not limited to: (i) the full implementation of the MAH system and implied approval of the commencement of clinical trial; (ii) the implementation of associated review of drugs, excipients and packaging materials; and (iii) the introduction of four procedures for expedited registration of drugs, which are procedures for groundbreaking therapeutic drugs, procedures for conditional approval, procedures for prioritized reviews and approval, and procedures for special examination and approval. Detailed implementation rules for drug classification and requirements for corresponding application materials will be promulgated by the NMPA.

In March 2016, the CFDA issued the Reform Plan for Registration Category of Chemical Medicine, which outlined the reclassifications of drug applications under the Registration Measures. According to the Reform Plan for Registration Category of Chemical Medicine, Category 1 drugs refer to innovative new drugs that have not been marketed anywhere in the world. Improved new drugs that are not marketed anywhere in the world fall into Category 2 drugs. Generic drugs, that have equivalent quality and efficacy to the originator's drugs and have been marketed abroad but not yet in China, can be classified as Category 3 drugs. Generic drugs, that have equivalent quality and efficacy to the originator's drugs and have been marketed in China, fall into Category 4 drugs. Category 5 drugs are drugs which have already been marketed abroad, but are not yet approved in China. Category 1 drugs and Category 5 drugs can be registered through the Domestic New Drug Application and the Imported Drug Application Procedures under the Registration Measures, respectively.

The SFDA promulgated the Administrative Provisions on Special Examination and Approval of Registration of New Drugs in January 2009, according to which, the SFDA conducts special examination and approval for new drug registration applications when: (1) the effective constituent of drug extracted from plants, animals, minerals, etc., as well as the preparations thereof have never been marketed in China, and the material medicines and the preparations thereof are newly discovered; (2) the chemical raw material medicines as well as the preparations thereof and the biological product have not been approved for marketing at home and abroad; (3) the new drugs have obvious clinical treatment advantages for such diseases as AIDS, malignant tumors and orphan diseases, etc. or (4) the new drugs treat diseases currently with no effective methods of treatment.

The Special Examination and Approval of Registration of New Drugs provides that the applicant may file for special examination and approval at the clinical trial application stage if the product candidate falls within items (1) or (2). The provisions provide that for product candidates that fall within items (3) or (4), the application for special examination and approval cannot be made until filing for production.

Accelerated Approval for Clinical Trial and Registration

The Innovation Opinions established a framework for reforming the evaluation and approval system for drugs, medical devices and equipment. The Innovation Opinions enhanced the standard of

approval for drug registration and accelerated the evaluation and approval process for innovative drugs as well as drug clinical trials.

The CFDA released the Circular Concerning Several Policies on Drug Registration Review and Approval in November 2015, which further clarified the measures and policies for simplifying and accelerating the approval process of clinical trials, including:

- a one-time umbrella approval procedure allowing the overall approval of all phases of a new drug's clinical trials, replacing the current phase-by-phase application and approval procedure; and
- a fast track drug registration or clinical trial approval pathway for the following applications: (1) registration of innovative new drugs for treating HIV, cancer, serious infectious diseases and orphan diseases, etc.; (2) registration of pediatric drugs; (3) registration of geriatric drugs and drugs treating PRC-prevalent diseases in elders; (4) registration of drugs listed in national major science and technology projects or national key research and development plan; (5) registration of clinical urgently needed drugs using advanced technology, using innovative treatment methods, or having distinctive clinical benefits; (6) registration of foreign innovative drugs to be manufactured locally in China; (7) concurrent applications for new drug clinical trials which are already approved in the United States or EU or concurrent drug registration applications for drugs which have applied to the competent drug approval authorities for marketing authorization and passed such authorities' onsite inspections in the United States or EU and are manufactured using the same production line in China; and (8) clinical trial applications for drugs with urgent clinical need and patent expiry within three years, and manufacturing authorization applications for drugs with urgent clinical need and patent expiry within one year.

The NMPA released the Circular on Adjusting Evaluation and Approval procedures for Clinical Trials for Drugs in July 2018, according to which, within 60 days after the acceptance of and the fees paid for the IND application, the applicant may conduct the clinical trials for the drug in accordance with the clinical trial protocol submitted, if the applicant has not received any negative or questioning opinion from the CDE. Such approval process has been further enacted into the 2019 Amendment.

Trial Exemptions and Acceptance of Foreign Data

The NMPA issued the Technical Guidance Principles on Accepting Foreign Drug Clinical Trial Data in July 2018, as one of the implementing rules for the Innovation Opinions, which provides that overseas clinical data can be submitted for the drug registration applications in China. Such applications can be in the form of waivers to China-based clinical trials, bridging trials and direct NDAs. According to the Technical Guidance Principles on Accepting Foreign Drug Clinical Trial Data, sponsors may use the data of foreign clinical trials to support drug registration in China, provided that sponsors must ensure the authenticity, completeness, accuracy and traceability of foreign clinical trial data and such data must be obtained consistent with the relevant requirements under the Good Clinical Trial Practice of the International Conference on Harmonization of Technical Requirements for Registration of Pharmaceuticals for Human Use, or the ICH. Sponsors must also comply with other relevant sections of the Registration Measures when applying for drug registrations in China using foreign clinical trial data.

The NMPA now officially permits, and its predecessor agencies have permitted on a case-by-case basis in the past, drugs approved outside of China to be approved in China on a conditional basis without pre-approval clinical trials being conducted in China. Specifically, the NMPA and the NHC released the Procedures for Reviewing and Approval of Clinical Urgently Needed Overseas New Drugs in October 2018, permitting drugs that have been approved within the last ten years in the United States, the EU or Japan and that prevent or treat orphan diseases, or prevent or treat serious life-threatening illnesses for which there is either no effective therapy in China, or for which the

foreign-approved drug would have clear clinical advantages. Applicants will be required to establish a risk mitigation plan and may be required to complete trials in China after the drug has been marketed. The CDE has developed a list of qualifying drugs that meet the foregoing criteria.

Clinical Trial Process and Good Clinical Practices

According to the Registration Measures, a clinical trial consists of Phases I, II, III and IV. Phase I refers to the initial clinical pharmacology and safety evaluation studies in humans. Phase II refers to the preliminary evaluation of a product candidate's therapeutic effectiveness and safety for particular indications in patients, to provide evidence and support for the design of Phase III clinical trials and to settle the administrative dose regimen. Phase III refers to clinical trials undertaken to confirm the therapeutic effectiveness and safety on patients with target indications, to evaluate the overall benefit-risk relationships of the drug, and ultimately to provide sufficient evidence for the review of drug registration application. Phase IV refers to a new drug's post-marketing study to assess therapeutic effectiveness and adverse reactions when the drug is widely used, to evaluate the overall benefit-risk relationships of the drug when used among the general population or specific groups and to adjust the administration dose.

To improve the quality of clinical trials, the SFDA promulgated the Good Clinical Trial Practice for Drugs in August 2003, or the GCP Rules, which was replaced by the revised Good Clinical Trial Practice for Drugs, the Revised GCP Rules, promulgated by the NMPA and the NHC in April 2020 and coming into effect in July 2020. According to the Administration of Quality of Drug Clinical Practice, clinical trial means systematical investigation of drugs conducted on human subjects (patients or healthy volunteers) to prove or reveal the function, adverse reactions and/or absorption, distribution, metabolism and excretion of the drug being investigated. The purpose of a clinical trial is to determine the therapeutic efficacy and safety of the drug. The Revised GCP Rules provide comprehensive and substantive requirements on the design and conduct of clinical trials in China. In particular, the Revised GCP Rules enhance the protection for study subjects and tighten the control over bio-samples collected under clinical trials.

The Revised GCP Rules also set out the qualifications and requirements for the investigators and centers participating in clinical trial, who must: (i) have professional certification at a clinical trial center, professional knowledge, training experience and capability of clinical trial, and be able to provide the latest resume and relevant qualification documents per request; (ii) be familiar with the trial protocol, investigator's brochure and relevant information of the trial drug provided by the applicant; (iii) be familiar with and comply with the Revised GCP Rules and relevant laws and regulations relating to clinical trials; (iv) keep a copy of the authorization form on work allocation signed by investigators; (v) accept supervision and inspection organized by the applicant and inspection by the drug regulatory authorities; and (vi) in the case of investigators and clinical trial centers authorizing other individual or institution to undertake certain responsibilities and functions relating to clinical trial, they shall ensure such individual or institution are qualified and establish complete procedures to ensure the responsibilities and functions are fully performed and generate reliable data.

Communication with the CDE

According to the Circular on Adjusting Evaluation and Approval Procedures for Clinical Trials for Drugs, where the application for clinical trial of new investigational drug has been approved, upon the completion of Phases I and II clinical trials and prior to Phase III clinical trial, the applicant shall submit the application for Communication Session to the CDE to discuss the key technical questions including the design of Phase III clinical trial protocol. Within 60 days after the acceptance of and the fees paid for the IND application, the applicant may conduct the clinical trials for the drug in accordance with the clinical trial protocol submitted, if the applicant has not received any negative or questioning opinion from the CDE.

The NMPA promulgated the Administrative Measures for Communication on the Research, Development and Technical Evaluation of Drugs in September 2018, according to which, during the research and development periods and in the registration applications of, among others, the innovative new drugs, the applicants may propose to conduct communication meetings with the CDE. The communication meetings can be classified into three types. Type I meetings are convened to address key safety issues in clinical trials of drugs and key technical issues in the research and development of breakthrough therapeutic drugs. Type II meetings are held during the key research and development periods of drugs, mainly including meetings before the IND application, meetings upon the completion of Phase II trials and before the commencement of Phase III trials, meetings before submitting a marketing application for a new drug, and meetings for risk evaluation and control. Type III meetings refer to meetings not classified as Type I or Type II.

Drug Clinical Trial Registration

According to the Registration Measures, upon obtaining the approval of its IND applications and before conducting a clinical trial, an applicant shall file a registration form with the SFDA containing various details, including the clinical trial protocol, the name of the principal researcher of the leading institution, the names of participating institutions and researchers, an approval letter from the ethics committee, and a sample of the informed consent form, with a copy sent to the competent provincial administration departments where the trial institutions will be located. The CFDA released the Announcement on Drug Clinical Trial Information Platform in September 2013, according to which, instead of the aforementioned registration field with the CFDA, all clinical trials approved by the CFDA and conducted in China shall complete a clinical trial registration and publish trial information through the Drug Clinical Trial Information Platform. The applicant shall complete the trial pre-registration within one month after obtaining the approval of the clinical trial approval in order to obtain the trial's unique registration number and complete registration of certain follow-up information before the first subject's enrollment in the trial. If the registration is not completed within one year after the approval of the IND applications, the applicant shall submit an explanation, and if the first submission is not completed within three years, the approval of the IND applications shall automatically expire.

New Drug Application

According to the Registration Measures, drug registration applications include domestic NDA, domestic generic drug application and imported drug application. Drugs are classified into chemical drugs, biological products and traditional Chinese medicine or natural drugs. When Phases I, II and III clinical trials have been completed, the applicant may apply to the SFDA for approval of the NDA. The SFDA then determines whether to approve the application according to the comprehensive evaluation opinion provided by the CDE.

Pilot Plan for the MAH System

The Innovation Opinions provide a pilot plan for the MAH system.

Under the authorization of the Standing Committee of the NPC, the General Office of the State Council issued the Pilot Plan for the Drug Marketing Authorization Holder Mechanism in May 2016, which provides a detailed pilot plan for the MAH system in 10 Chinese provinces. Under the MAH system, domestic drug research and development institutions and individuals in the pilot regions are eligible to be holders of drug registrations without having to become drug manufacturers. The marketing authorization holders may engage contract manufacturers for manufacturing, provided that the contract manufacturers are licensed and located within the pilot regions. Drugs that qualify for the MAH system are: (1) new drugs (including but not limited to drugs under category I to category IV of chemical drugs, and targeted preparation, sustained release preparation, controlled release preparation

under category V of chemical drugs, biological products approved as category I and VII drugs and biosimilars under the Registration Measures) approved after the implementation of the MAH system; (2) generic drugs approved as category III or IV drugs under the Reform Plan for Registration Category of Chemical Medicine; (3) previously approved generics that have passed equivalence assessments against original drugs; and (4) previously approved drugs whose licenses were held by drug manufacturers originally located within the pilot regions, but which have moved out of the pilot regions due to corporate mergers or other reasons.

The CFDA promulgated the Circular on the Matters Relating to Promotion of the Pilot Program for the Drug Marketing Authorization Holder System in August 2017. It clarified the legal liability of the MAH, who is responsible for managing the whole manufacturing and marketing chain and the whole life cycle of drugs and legally liable for preclinical drug study, clinical trials, manufacturing, marketing, distribution and adverse drug reaction monitoring. According to the Circular on the Matters Relating to Promotion of the Pilot Program for the Drug Marketing Authorization Holder System, the MAH shall submit a report of drug manufacturing, marketing, prescription, techniques, pharmacovigilance, quality control measures and other situations to the CFDA within 20 working days after the end of each year.

According to the Pilot Plan for the Drug Marketing Authorization Holder Mechanism, the pilot plan was originally set for a three-year period and was scheduled to expire in November 2018. The Standing Committee of the NPC promulgated the Decision of Extending the Pilot Period of Authorizing the State Council to Carry Out the Pilot Plan for the Drug Marketing Authorization Holder Mechanism in Certain Places in October 2018, which extended the term of the MAH system to November 4, 2019.

According to the 2019 Amendment, which came into effect on December 1, 2019, the MAH system will be applicable throughout the country and the legal representative and the key person-in-charge of a drug MAH shall be fully responsible for the quality of drugs.

International Multi-Center Clinical Trials

The International Multi-Center Clinical Trial Guidelines (Trial), or the Multi-Center Clinical Trial Guidelines, which was promulgated by the CFDA in January 2015 and came into effect in March 2015, provided guidance on the implementation of Multi-Regional Clinical Trials, or the MRCT, in China. According to the Multi-Center Clinical Trial Guidelines, international multi-center clinical trial applicants may simultaneously perform clinical trials in different centers using the same clinical trial protocol. Where the applicants plan to implement the international multi-center clinical trials in the PRC, the applicants shall comply with relevant laws and regulations, such as the Drug Administration Law, the Implementing Regulations of the Drug Administration Law and the Registration Measures, execute the GCP Rules, make reference to universal international principles such as the ICH-GCP and comply with the laws and regulations of the countries involved in the international multi-center clinical trials. Where the applicants plan to use the data derived from the international multi-center clinical trials for approval of a drug registration in the PRC, it shall involve at least two countries, including China, and shall satisfy the requirements for clinical trials set forth in the Multi-Center Clinical Trial Guidelines, Registration Measures and other related laws and regulations.

In April 2020, the NMPA and the NHC promulgated the Revised GCP Rules, which came into effect in July 2020. The Revised GCP Rules summarize the requirements for initiating an MRCT, that is, before initiating an MRCT: (i) the applicant shall ensure that all the centers participating in the clinical trial comply with the trial protocol; (ii) the applicant shall provide each center with the same trial protocol, and each center shall comply with the same unified evaluation criterion for clinical trial and laboratory data and the same guidance for case report form; (iii) each center shall use the same case report form to record the data of each human subject obtained during the trial; (iv) before

initiating a clinical trial, a written document is required to specify the responsibilities of the investigators of each center; and (v) the applicant shall ensure the communication among the investigators of each center.

Data derived from international multi-center clinical trials can be used for the new drug applications with the NMPA. When using international multi-center clinical trial data to support new drug applications in China, applicants shall submit the completed global clinical trial report, statistical analysis report and database, along with relevant supporting data in accordance with the content and format requirements under the International Conference on Harmonization-Common Technical Document; subgroup research results summary and comparative analysis shall also be conducted concurrently. Leveraging the clinical trial data derived from international multi-center clinical trials conducted by our partners, we may avoid unnecessarily repetitive clinical trials and thus further accelerate the NDA process in China.

The CFDA released the Decision on Adjusting Items concerning the Administration of Imported Drug Registration in October 2017, which includes the following key points:

- If the International Multicenter Clinical Trial, or the IMCCT, of a drug is conducted in China, Phase I clinical trial of the drug is allowed simultaneously. The IMCCT drug does not need to be approved or to enter into either a Phase II or III clinical trial in a foreign country, except for preventive biological products;
- If the IMCCT is conducted in China, the application for drug marketing authorization can be submitted directly after the completion of the IMCCT. The Registration Measures and relevant laws and regulations shall be complied with for registration application;
- With respect to applications for clinical trial and marketing of the imported innovative chemical drugs and therapeutic biological products, the marketing authorization in the country or region where the foreign drug manufacturer is located will not be required; and
- With respect to drug applications that have been accepted before the release of this Decision, if relevant requirements are met, importation permission can be granted if such applications request exemption of clinical trials for the imported drugs based on the data generated from the IMCCT.

Approval of Human Genetic Resources

The Interim Administrative Measures on Human Genetic Resources, promulgated by the Ministry of Science and Technology and the MOH in June 1998, aimed at protecting and fairly utilizing human genetic resources in the PRC. The Ministry of Science and Technology promulgated the Service Guide for Administrative Licensing Items concerning Examination and Approval of Sampling, Collecting, Trading or Exporting Human Genetic Resources, or Taking Such Resources out of the PRC in July 2015, according to which, the sampling, collection or research activities of human genetic resources by a foreign-invested sponsor fall within the scope of international cooperation, and the cooperating Chinese organization shall apply for approval of the China Human Genetic Resources Management Office through an online system. The Ministry of Science and Technology further promulgated the Circular on Optimizing the Administrative Examination and Approval of Human Genetic Resources in October 2017 and October 2020, which became effective in December 2017 and October 2020, respectively and simplified the approval of sampling and collecting human genetic resources for listing a drug in the PRC.

The Regulations of the PRC on the Administration of Human Genetic Resources, which was promulgated by the State Council in May 2019 and came into effect in July 2019, further stipulates that, in order to obtain marketing authorization for relevant drugs and medical devices in China, no approval is required in international clinical trial cooperation using China's human genetic resources at

clinical institutions without export of human genetic resource materials. However, the type, quantity and usage of the human genetic resource to be used shall be filed with the administrative department of science and technology under the State Council before clinical trials.

Regulations on Drug Manufacturing and Distribution

Drug Manufacturing

According to the Drug Administration Law and the Implementing Regulations of the Drug Administration Law, a drug manufacturing enterprise is required to obtain a drug manufacturing license from the relevant provincial drug administration authority of the PRC. The grant of such license is subject to an inspection of the manufacturing facilities, and an inspection to determine whether the sanitary condition, quality assurance systems, management structure and equipment meet the required standards. According to the Implementing Regulations of the Drug Administration Law and the Measures on the Supervision and Administration of the Manufacture of Drugs, which was promulgated in August 2004, amended in November 2017 and January 2020 and came into effect in July 2020, the drug manufacturing license is valid for five years and shall be renewed at least six months prior to its expiration date upon a re-examination by the relevant authority. In addition, the name, legal representative, registered address and type of the enterprise specified in the drug manufacturing certificate shall be identical to that set forth in the business license as approved and issued by the industrial and commercial administrative department. To the extent the MAH does not manufacture the drug internally but through a contract manufacturing organization, the MAH shall apply for drug manufacturing license with the provincial counterpart of the NMPA, subject itself to inspections and other regulatory oversight by the agency.

The Good Manufacturing Practice for Drugs was promulgated in March 1988 and was amended in December 1992 and June 1999 and January 2011. The latest amendment was in June 2020 and will come into effect in October 2020. The Good Manufacturing Practice for Drugs comprises a set of detailed standard guidelines governing the manufacture of drugs, which include institution and staff qualifications, production premises and facilities, equipment, hygiene conditions, production management, quality controls, product operation, raw material management, maintenance of sales records, management of customer complaints and adverse event reports.

Drug Distribution

According to the Drug Administration Law, its implementing regulations and the Measures for the Supervision and Administration of Circulation of Pharmaceuticals, which was promulgated by the SFDA in December 2006 and came into effect in May 2007, pharmaceutical enterprises shall be responsible for the quality of the pharmaceuticals that they manufacture, operate, use, purchase, sell, transport, or store.

According to the Measures for the Administration of Pharmaceutical Operation Certificate, which was promulgated in February 2004 and amended in November 2017 by the CFDA, a Medicine Operation Certificate is valid for five years. Each holder of the Medicine Operation Certificate must apply for an extension of its permit six months prior to expiration. The establishment of a wholesale pharmaceutical distribution company requires the approval of provincial medicine administrative authorities. Upon approval, the authority will grant a Medicine Operation Certificate in respect of the wholesale pharmaceutical product distribution company. The establishment of a retail pharmacy store requires the approval of the local medicine administrative authorities at or above the county level. Upon approval, the authority will grant a Medicine Operation Certificate in respect of the retail pharmacy store.

Other PRC Government Regulations

Regulations on Intellectual Property Rights

In terms of international conventions, China has entered into (including but not limited to) the Agreement on Trade-Related Aspects of Intellectual Property Rights, the Paris Convention for the Protection of Industrial Property, the Madrid Agreement Concerning the International Registration of Marks and the Patent Cooperation Treaty.

Patents

According to the Patent Law of the PRC, which was promulgated by the Standing Committee of the NPC in March 1984, amended in September 1992, August 2000 and December 2008, and came into effect in October 2009, and the Implementation Rules of the Patent Law of the PRC, which was promulgated by the State Council in June 2001 and amended in December 2002 and January 2010, there are three types of patents in the PRC: invention patents, utility model patents and design patents. The protection period is 20 years for an invention patent and 10 years for a utility model patent and a design patent, commencing from their respective application dates. Any individual or entity that utilizes a patent or conducts any other activities that infringe a patent without prior authorization of the patent holder shall pay compensation to the patent holder and is subject to a fine imposed by relevant administrative authorities and, if constituting a crime, shall be held criminally liable in accordance with the law. According to the Patent Law of the PRC, any organization or individual that applies for a patent in a foreign country for an invention or utility model patent established in China is required to report to the NIPA for confidentiality examination.

Trade Secrets

According to the PRC Anti-Unfair Competition Law, which was promulgated by the Standing Committee of the NPC in September 1993 and amended in November 2017 and April 2019, respectively, the term "trade secrets" refers to technical and business information that is unknown to the public, has utility, may create business interests or profits for its legal owners or holders, and is maintained as a secret by its legal owners or holders. Under the PRC Anti-Unfair Competition Law, business persons are prohibited from infringing others' trade secrets by: (1) obtaining the trade secrets from the legal owners or holders by any unfair methods such as theft, bribery, fraud, coercion, electronic intrusion, or any other illicit means; (2) disclosing, using or permitting others to use the trade secrets obtained illegally under item (1) above; (3) disclosing, using or permitting others to use the trade secrets, in violation of any contractual agreements or any requirements of the legal owners or holders to keep such trade secrets in confidence; or (4) instigating, inducing or assisting others to violate a confidentiality obligation or to violate a rights holder's requirements on keeping confidentiality of trade secrets, disclosing, using or permitting others to use the trade secrets of the rights holder. If a third party knows or should have known of the above-mentioned illegal conduct but nevertheless obtains, uses or discloses trade secrets of others, the third party may be deemed to have committed a misappropriation of the others' trade secrets. The parties whose trade secrets are being misappropriated may petition for administrative corrections, and regulatory authorities may stop any illegal activities and fine infringing parties.

Trademarks

According to the Trademark Law of the PRC promulgated by the Standing Committee of the NPC in August 1982, and amended in February 1993, October 2001, August 2013 and April 2019, respectively, the period of validity for a registered trademark is ten years, commencing on the date of registration. The registrant shall go through the formalities for renewal within twelve months prior to the expiry date of the trademark if continued use is intended. Where the registrant fails to do so, a

grace period of six months may be granted. The validity period for each renewal of registration is ten years, commencing on the day immediately after the expiry of the preceding period of validity for the trademark. In the absence of a renewal upon expiry, the registered trademark shall be cancelled. Industrial and commercial administrative authorities have the authority to investigate any behavior that infringes the exclusive right under a registered trademark in accordance with the law. In case of a suspected criminal offense, the case shall be timely referred to a judicial authority and decided according to the law.

Domain Names

Domain names are protected under the Administrative Measures on the Internet Domain Names, which was promulgated by the Ministry of Industry and Information Technology in August 2017, and the Implementing Rules on Registration of National Top-level Domain Names, which was promulgated by China Internet Network Information Center in and came into effect in June 2019. The Ministry of Industry and Information Technology is the main regulatory body responsible for the administration of PRC internet domain names. Domain name registrations are handled through domain name service agencies established under the relevant regulations, and the applicants become domain name holders upon successful registration.

Regulations on Product Liability

In addition to the strict new drug approval process, certain PRC laws have been promulgated to protect the rights of consumers and to strengthen the control of medical products in the PRC. Under current PRC laws, manufacturers and vendors of defective products in the PRC may incur liability for loss and injury caused by such products. According to the General Principles of the Civil Law of the PRC promulgated in April 1986 and amended in August 2009 and General Rules of the Civil Law of the People's Republic of China promulgated in March 2017 and took effect in October 2017 (collectively, the "PRC Civil Law"), the manufacturer or vendor of a defective product which causes property damage or physical injury to any person may be subject to civil liability for such damage or injury.

In February 1993, the Product Quality Law of the PRC, or the Product Quality Law, was promulgated to supplement the PRC Civil Law, aiming to protect the legitimate rights and interests of the end-users and consumers and to strengthen the supervision and control of the quality of products. The Product Quality Law was last revised in December 2018. According to the revised Product Quality Law, manufacturers who produce defective products may be subject to civil or criminal liability and have their business licenses revoked.

The Law of the PRC on the Protection of the Rights and Interests of Consumers was promulgated in October 1993 and amended in October 2013 to protect consumer rights when they purchase or use goods and services. According to which, all business operators must comply with this law when they manufacture or sell goods and/or provide services to customers. Under the latest amendment, all business operators shall protect the customers' privacy and keep any consumer information they obtain during the business operation strictly confidential. In addition, in extreme situations, pharmaceutical product manufacturers and operators may be subject to criminal liability if their goods or services lead to the death or injuries of customers or other third parties.

Regulations on Tort

According to the Tort Law of the PRC promulgated by the Standing Committee of the NPC in December 2009, if damages to other persons are caused by defective products due to the fault of third parties, such as the parties providing transportation or warehousing, the producers and the sellers of the products have the right to recover their respective losses from such third parties. If defective

products are identified after they have been put into circulation, the producers or the sellers shall take remedial measures such as issuance of a warning, recall of products, etc., in a timely manner. The producers or the sellers shall be liable under tort if they fail to take remedial measures in a timely manner or have not made efforts to take remedial measures, thus causing damages. If the products are produced or sold with known defects, causing deaths or severe adverse health issues, the infringing party has the right to claim punitive damages in addition to compensatory damages.

Regulations on Environment Protection

Pursuant to the Environmental Protection Law of the PRC promulgated by the Standing Committee of the NPC, in December 1989, amended in April 2014 and effective in January 2015, any entity which discharges or will discharge pollutants during its course of operations or other activities must implement effective environmental protection safeguards and procedures to control and properly treat waste gas, waste water, waste residue, dust, malodorous gases, radioactive substances, noise vibrations, electromagnetic radiation and other hazards produced during such activities. According to the provisions of the Environmental Protection Law, in addition to other relevant laws and regulations of the PRC, the Ministry of Environmental Protection and its local counterparts take charge of administering and supervising said environmental protection matters.

Pursuant to the Environmental Protection Law, the environmental impact statement on any construction project must assess the pollution that the project is likely to produce and its impact on the environment, and stipulate preventive and curative measures; the statement shall be submitted to competent administrative department of environmental protection for approval. Installations for the prevention and control of pollution in construction projects must be designed, built and commissioned together with the principal part of the project.

Pursuant to the Law of the People's Republic of China on Environment Impact Assessment, which was promulgated in October 2002 and most recently amended in December 2018, the State implements a classification-based management on the environmental impact assessment of construction projects according to the impact of the construction projects on the environment. Construction units shall prepare an Environmental Impact Report or an Environmental Impact Statement, or fill out the Environmental Impact Registration Form.

Pursuant to the Regulations on Urban Drainage and Sewage Disposal, which was promulgated in October 2013 and came into effect in January 2014, and the Measures for the Administration of Permits for the Discharge of Urban Sewage into the Drainage Network, which was promulgated in January 2015 and came into effect in March 2015, drainage entities covered by urban drainage facilities shall discharge sewage into urban drainage facilities in accordance with the relevant provisions of the state. Where a drainage entity needs to discharge sewage into urban drainage facilities, it shall apply for a drainage license in accordance with the provisions of these Measures. The drainage entity that has not obtained the drainage license shall not discharge sewage into urban drainage facilities.

Regulations on Fire Protection

The Fire Prevention Law of the PRC, or the Fire Prevention Law, was adopted in April 1998 and last amended in April 2019. The Fire Prevention Law provides that fire control design and construction of a construction project shall comply with the State's fire control technical standards. Developers, designers, builders and project supervisors shall be responsible for the quality of the fire control design and construction of the construction project pursuant to the law. Development project fire safety design examinations and acceptance systems shall be implemented for development projects which are required to have fire safety design in accordance with the national fire protection technical standards.

According to the Eight Measures for the Public Security Fire Department to Deepen Reform and Serve Economic and Social Development promulgated by the Ministry of Public Security of the PRC in

August 2015, the fire protection design and completion acceptance fire protection record of construction projects with an investment of less than RMB300,000 or a building area of less than 300 square meters (or below the limit set by the housing and urban construction department of the provincial people's government) was no longer required.

Regulations on Foreign Exchange and Dividend Distribution

Foreign Exchange Control

According to the PRC Regulation for the Foreign Exchange promulgated by the State Council in January 1996, which was amended in January 1997 and August 2008, and the Regulation on the Administration of the Foreign Exchange Settlement, Sales and Payment promulgated by the People's Bank of China in June 1996, foreign exchanges required for distribution of profits and payment of dividends may be purchased from designated foreign exchange banks in the PRC upon presentation of a board resolution authorizing distribution of profits or payment of dividends.

According to the Circular of the State Administration of Foreign Exchange, or the SAFE, on Further Improving and Adjusting the Foreign Exchange Policies on Direct Investment and its appendix promulgated in November 2012 and amended in May 2015, October 2018 and December 2019 by the SAFE, (1) the opening of and payment into foreign exchange accounts under direct investment accounts are no longer subject to approval by the SAFE; (2) reinvestment with legal income of foreign investors in China is no longer subject to approval by SAFE; (3) the procedures for capital verification and confirmation that foreign-funded enterprises need to go through are simplified; (4) purchase and external payment of foreign exchange under direct investment accounts are no longer subject to approval by SAFE; (5) domestic transfer of foreign exchange under direct investment account is no longer subject to approval by SAFE; and (6) the administration over the conversion of foreign exchange capital of foreign-invested enterprises is improved. Later, the SAFE promulgated the Circular on Further Simplifying and Improving Foreign Exchange Administration Policies in Respect of Direct Investment in February 2015, which was further amended in December 2019 and prescribed that the bank instead of the SAFE can directly handle the foreign exchange registration and approval under foreign direct investment while the SAFE and its branches indirectly supervise the foreign exchange registration and approval under foreign direct investment through the bank.

The Provisions on the Administration of Foreign Exchange in Foreign Direct Investments by Foreign Investors, which were promulgated by the SAFE in May 2013 and amended in October 2018 and December 2019, regulate and clarify the administration over foreign exchange administration in foreign direct investments.

According to the Circular on the Reform of the Management Method for the Settlement of Foreign Exchange Capital of Foreign-invested Enterprises promulgated by the SAFE in March 2015 and amended in December 2019, and the Circular on the Reform and Standardization of the Management Policy of the Settlement of Capital Projects promulgated by the SAFE in June 2016, the settlement of foreign exchange by foreign invested enterprises shall be governed by the policy of foreign exchange settlement on a discretionary basis. However, the settlement of foreign exchange shall only be used for their own operational purposes within the business scope of the foreign invested enterprises and follow the principles of authenticity.

Dividend Distribution

The SAFE promulgated the Notice on Improving the Check of Authenticity and Compliance to Further Promote Foreign Exchange Control in January 2017, which stipulates several capital control measures with respect to outbound remittance of profits from domestic entities to offshore entities, including the following: (1) under the principle of genuine transaction, banks shall check board resolutions regarding profit distribution, the original version of tax filing records and audited financial

statements; and (2) domestic entities shall hold income to account for previous years' losses before remitting the profits. Moreover, domestic entities shall make detailed explanations of sources of capital and utilization arrangements, and provide board resolutions, contracts and other proof when completing the registration procedures in connection with an outbound investment.

Foreign Exchange Registration of Offshore Investment by PRC Residents

The SAFE promulgated the SAFE Circular 37 in July 2014. The SAFE Circular 37 requires PRC residents (including PRC institutions and individuals) to register with local branches of SAFE in connection with their direct or indirect offshore investment in an overseas special purpose vehicle, or the SPV, directly established or indirectly controlled by PRC residents for offshore investment and financing with their legally owned assets or interests in domestic enterprises, or their legally owned offshore assets or interests. Such PRC residents are also required to amend their registrations with the SAFE when there is a change to the basic information of the SPV, such as changes of a PRC resident individual shareholder, the name or operating period of the SPV, or when there is a significant change to the SPV, such as changes of the PRC individual resident's increase or decrease of its capital contribution in the SPV, or any share transfer or exchange, merger, division of the SPV.

The Circular on Further Simplifying and Improving Foreign Exchange Administration Policies in Respect of Direct Investment, which was promulgated in February 2015 and effective in June 2015 and further amended in December 2019, provides that PRC residents may register with qualified banks instead of the SAFE in connection with their establishment or control of an offshore entity established for the purpose of overseas direct investment. The SAFE and its branches shall implement indirect supervision over foreign exchange registration of direct investment via the banks.

Failure to comply with the registration procedures set forth in the SAFE Circular 37 may result in restrictions on the foreign exchange activities of the relevant onshore company, including the payment of dividends and other distributions to its offshore parent or affiliate, the capital inflow from the offshore entities and settlement of foreign exchange capital, and may also subject relevant onshore company or PRC residents to penalties under PRC foreign exchange administration regulations.

Regulations on Labor

Labor Law and Labor Contract Law

According to the PRC Labor Law, which was promulgated by the Standing Committee of the NPC in July 1994 and amended in August 2009 and December 2018, respectively, the PRC Labor Contract Law, which was promulgated by the Standing Committee of the NPC in June 2007 and amended in December 2012 and came into effect July 2013, and the Implementing Regulations of the Employment Contracts Law of the PRC, which was promulgated by the State Council in September 2008, labor contracts in written form shall be executed to establish labor relationships between employers and employees. In addition, wages cannot be lower than the local minimum wage. The employers must establish a system for labor safety and sanitation, strictly abide by State rules and standards, provide education regarding labor safety and sanitation to its employees, provide employees with labor safety and sanitation conditions and necessary protection materials in compliance with State rules, and carry out regular health examinations for employees engaged in work involving occupational hazards.

Social Insurance and Housing Provident Funds

According to the Social Insurance Law of PRC, which was promulgated by the Standing Committee of the NPC in October 2010 and came into effect in July 2011, and further amended in December 2018, and the Interim Regulations on the Collection and Payment of Social Security Funds, which was promulgated by the State Council in January 1999 and amended in March 2019, and the Regulations on the Administration of Housing Provident Funds, which was promulgated by the State

Council in April 1999 and amended in March 2002 and March 2019, employers are required to contribute, on behalf of their employees, to a number of social security funds, including funds for basic pension insurance, unemployment insurance, basic medical insurance, occupational injury insurance, maternity insurance and to housing provident funds. Any employer who fails to contribute may be fined and ordered to make good the deficit within a stipulated time limit.

Regulations on Taxation

Enterprise Income Tax

According to the Enterprise Income Tax Law promulgated by the NPC in March 2007 and amended in February 2017 and December 2018, and the Implementation Rules of the Enterprise Income Tax Law of the PRC promulgated by the State Council in December 2007 and amended in April 2019, other than a few exceptions, the income tax rate for both domestic enterprises and foreign-invested enterprises is 25%. Enterprises are classified as either "resident enterprises" or "non-resident enterprises". Besides enterprises established within the PRC, enterprises established outside China whose "de facto management bodies" are located in China are considered "resident enterprises" and subject to the uniform 25% enterprise income tax rate for their global income. A non-resident enterprise refers to an entity established under foreign law whose "de facto management bodies" are not within the PRC but which have an establishment or place of business in the PRC, or which do not have an establishment or place of business in the PRC but have income sourced within the PRC. An income tax rate of 10% will normally be applicable to dividends declared to non-PRC resident enterprise investors that do not have an establishment or place of business in the PRC, or that have such establishment or place of business but the relevant income is not effectively connected with the establishment or place of business, to the extent such dividends are derived from sources within the PRC.

According to the Notice on Promoting the Implementation of Corporate Income Tax Policies for Advanced Technology Service Enterprises Nationwide, or the Notice, since January 2017, an enterprise which is recognized as an "Advanced Technology Service Enterprises" under the Notice enjoys a reduced enterprise income tax rate of 15%.

According to the Arrangement Between the Mainland of China and the Hong Kong Special Administrative Region for the Avoidance of Double Taxation and Prevention of Fiscal Evasion with Respect to Taxes on Income, or the Double Tax Avoidance Arrangement, which was promulgated and came into effect in August 2006, and other applicable PRC laws, if a Hong Kong resident enterprise is determined by the competent PRC tax authority to have satisfied the relevant conditions and requirements under such Double Tax Avoidance Arrangement and other applicable laws, the 10% withholding tax on the dividends the Hong Kong resident enterprise receives from a PRC resident enterprise may be reduced to 5%. However, based on the Circular on Certain Issues with Respect to the Enforcement of Dividend Provisions in Tax Treaties which was promulgated by the State Administration of Taxation, the STA, in February 2009, if the relevant PRC tax authorities determine, in their discretion, that a company benefits from such reduced income tax rate due to a structure or arrangement that is primarily tax-driven, such PRC tax authorities may adjust the preferential tax treatment. Based on the Announcement on Certain Issues with Respect to the "Beneficial Owner" in Tax Treaties, which was promulgated by the STA in February 2018 and came into effect in April 2018, if an applicant's business activities do not constitute substantive business activities, it could result in the negative determination of the applicant's status as a "beneficial owner", and consequently, the applicant could be precluded from enjoying the above-mentioned reduced income tax rate of 5% under the Double Tax Avoidance Arrangement.

Value Added Tax

According to the Provisional Regulations of the PRC on Value-Added Tax, effective in January 1994 and further amended in November 2008, February 2016, and November 2017, and its implementation rules effected in January 1994 and amended in December 2008 and October 2011, except stipulated otherwise, taxpayers who sell goods, labor services or tangible personal property leasing services or import goods shall be subject to a 17% tax rate; taxpayers who sell transport services, postal services, basic telecommunications services, construction services, or real property leasing services, sell real property, transfer the land use right shall be subject to an 11% tax rate, and taxpayers who sell services or intangible assets shall be subject to a 6% tax rate.

According to the Circular of the Ministry of Finance and the State Administration of Taxation on Adjusting Value-added Tax Rates adopted in April 2018, as of May 2018, where a taxpayer engages in a taxable sales activity for the value-added tax purpose or imports goods, the previous applicable 17% and 11% rates are adjusted to 16% and 10%.

According to the Announcement on Relevant Policies for Deepening Value-Added Tax Reform, effective in April 2019, the 16% VAT tax rate, which applies to the sales or imported goods of a VAT general taxpayer, will be lowered to 13%; and the 10% VAT tax rate will be lowered to 9%.

According to the Measures for the Exemption of Value-Added Tax from Cross-Border Taxable Activities in the Collection of Value-Added Tax in Lieu of Business Tax (for Trial Implementation) revised in June 2018, if domestic enterprises provide cross-border taxable activities such as professional technical services, technology transfer, software services, the above-mentioned cross-border taxable activities are exempt from VAT.

MANAGEMENT**Executive Officers, Key Employees and Directors**

The following table sets forth information regarding our directors and executive officers as of the date of this prospectus.

<u>Executive Officers</u>	<u>Age</u>	<u>Position/Title</u>
Peter (Peizhi) Luo, Ph.D.	55	Co-Founder, Chief Executive Officer and Chairman of the Board
Fangyong (Felix) Du, Ph.D.	51	Chief Technology Officer
Hua Gong, M.D., Ph.D.	53	Chief Operating Officer & Head of Clinical Development and Precision Medicine
JC Xu, M.D., Ph.D.	56	Chief Scientific Officer
Qinghai Zhao, Ph.D.	60	Chief Manufacturing Officer
Man Kin (Raymond) Tam, M.B.A., B. Eng.	43	Chief Financial Officer and Director Nominee [#]
Yu (Albert) Ren, Ph.D.	46	Chief Strategy and Corporate Development Officer

<u>Key Employees</u>	<u>Age</u>	<u>Position/Title</u>
Yan Li, M.B.A.	46	Senior Vice President, Bioinformatics and Information Technology
Xiaohong (Kristine) She	54	Senior Vice President, Head of Clinical Operations
Guizhong Liu, Ph.D.	50	Head of Biology and Pharmacology
Alexander Goergen	34	Head of Business Development
Yuren (Ron) Wang, Ph.D.	59	Head of Portfolio Management

<u>Non-Employee Directors</u>	<u>Age</u>	<u>Position/Title</u>
Daniel Auerbach, M.B.A.*	62	Director
Chong Xu, Ph.D.*	39	Director
Yu Miao	32	Director
Yunxia Yang*	47	Director
Lefei Sun	41	Director
Andy (Yiu Leung) Cheung	61	Independent Director Nominee [#]
Min Li	58	Independent Director Nominee [#]

* Each of Mr. Daniel Auerbach, Mr. Chong Xu and Yunxia Yang will resign from our board of directors immediately prior to the SEC's declaration of effectiveness of our registration statement on Form F-1, of which this prospectus is a part.

Each of Mr. Man Kin (Raymond) Tam, Mr. Andy (Yiu Leung) Cheung and Mr. Min Li has accepted appointment as a director, which will be immediately effective upon the declaration of effectiveness of our registration statement on Form F-1 by the SEC, of which this prospectus is a part.

Executive Officers

Peter (Peizhi) Luo, Ph.D. is our Co-Founder and has served as our Chief Executive Officer since November 2011 and Chairman of the Board of the Directors since February 2018. Dr. Luo served as the first lead scientist in computational protein design and protein laboratory at Xencor (Nasdaq: XNCR) from July 1998 to August 2000. In September 2000, Dr. Luo founded Abmaxis Inc. and served as its Co-Founder, Chief Technology Officer, president, and director. In May 2006, Dr. Luo led the acquisition of Abmaxis Inc. by Merck & Co. (NYSE: MRK), after which Dr. Luo served as a director of Biologics Technology at Merck, and Chief Technology Officer of Abmaxis, the subsidiary of Merck & Co. Throughout his career, Dr. Luo also led the business development efforts in connection with collaborations and strategic partnerships with multiple global partners. Dr. Luo received his

bachelor's degree in applied chemistry in technical physics from Peking University in 1986, master's degree in applied physics from The Institute of High Energy Physics of the Chinese Academy of Sciences in 1989, and Ph.D. degree in chemistry from The University of Chicago in 1995. Dr. Luo also completed his postdoctoral research in protein folding at Stanford University in 1998. Dr. Luo's spouse is Xiaohong (Kristine) She who is our Senior Vice President, Head of Clinical Operations.

Fangyong (Felix) Du, Ph.D. joined us as Vice President of Technology Development in January 2012 and has served as our Chief Technology Officer since May 2019. Dr. Du has over 20 years of experience in biological research and discovery industry, and has published numerous peer-reviewed articles in world-renowned scientific journals such as Nature and Science. Dr. Du worked at Affomix from October 2009 to July 2010 and Illumina (Nasdaq: ILMN) from July 2010 to January 2012, and then joined the Company in January 2012 as the Vice President of Technology Development. Dr. Du received his bachelor's degree and master's degree in physiology and biophysics in 1991 and 1994, respectively, from Peking University, and his Ph.D. degree in biology from the California Institute of Technology in 2001. Dr. Du also completed his postdoctoral research from Yale University in 2007.

Hua Gong, M.D., Ph.D. has served as our Chief Operating Officer and Head of Clinical Development and Precision Medicine since July 2020. Dr. Gong served as the principal scientist of Pfizer (NYSE: PFE) from January 1999 to March 2009. From April 2009 to October 2011, she served as the associate director of Prometheus Diagnostics & Therapeutics. From November 2011 to January 2013, Dr. Gong served as an executive director of Premier Research. From January 2013 to June 2020, she served as the senior director and head of genomics biomarker of Novartis. Dr. Gong received her M.D. degree from Anhui Medical University in 1989, master's degree in biostatistics from Sun Yat-Sen University of Medical Sciences in 1992 and Ph.D. degree in cancer biology from Wayne State University in 1999.

JC Xu, M.D., Ph.D. has served as our Chief Scientific Officer since August 2020. Dr. Xu has more than 20 years of experience in oncology drug discovery and development, and more than four years of experience in business development, strategy, and operations in the biopharmaceutical industry in the United States. Prior to joining Adagene, Dr. Xu was head of R&ED China Strategy at Celgene now BMS from 2017 to 2020. Prior to that, Dr. Xu was Director of Strategy & Operations at Celgene Quantical Research and Director of Biology at Quantical Pharmaceuticals from 2012 to 2017. Prior to Quantical, Dr. Xu worked in leadership roles at a number of biopharmaceutical companies, including Pfizer, Amgen, and Corixa.

Dr. Xu received her M.D. degree from Beijing Medical University (now Peking University Health Science Center) in 1987 and her Ph.D. degree in Immunology from University of Alabama at Birmingham in 1993. She completed her post-doctoral training at DNAX Research Institute (now Merck Palo Alto) in 1996. She is an inventor of more than 120 issued and pending patents and has published more than 50 articles in peer-reviewed journals.

Qinghai Zhao, Ph.D. has served as our Chief Manufacturing Officer since October 2020. Dr. Zhao has more than 25 years of experience in protein therapeutics development and defining product development strategy from early stage IND through commercial filing. Prior to joining Adagene, Dr. Zhao was Vice President of Technical Development and Manufacturing at Forty Seven Inc. (NASDAQ:FTSV) from 2017 to October 2020. Prior to that, Dr. Zhao was vice president of CMC/Manufacturing at AnaptysBio (NASDAQ:ANAB) from 2016 to 2017. Prior to AnaptysBio, Dr. Zhao worked as an Executive Director and the Head of CMC at NGM Biopharmaceuticals from 2014 to 2016. Dr. Zhao also held various positions at Human Genome Sciences, Inc., Cogenesys and Teva Biopharmaceuticals USA from 2001 to 2014. Dr. Zhao received his bachelor's degree from Shanghai Chemical Engineering Institute (now East China University of Sciences and Technology) in 1982, master's degree from Shanghai Medical University (now merged with Fudan University) in 1986 and Ph.D. degree from University of Uppsala in 1998.

Man Kin (Raymond) Tam has served as our Chief Financial Officer since September 2019 and will serve as our director immediately upon the effectiveness of our registration statement on Form F-1, of which this prospectus is a part. Mr. Tam has over 20 years of management experience in finance and banking across the Asia-Pacific region. Mr. Tam worked in HSBC and J.P. Morgan Chase Bank, N.A. from 1999 to 2010. Mr. Tam served as project director of Mineralogy Pty Limited and Chief Financial Officer of Resourcehouse Limited from April 2010 to October 2015. From October 2015 to August 2019, Mr. Tam consecutively served as the Chief Financial Officer of China Regenerative Medicine International Limited (HKEx: 8158), Beijing Gas Blue Sky Holdings Limited (HKEx: 6828), and AgenTus Therapeutics, Inc. Mr. Tam is a fellow of CPA Australia, a member of the American Institute of Certified Public Accountants and the Hong Kong Institute of Certified Public Accountants. He is also a CFA and FRM charter-holder. He received his bachelor's degree in civil & resources engineering from the University of Auckland in 1997, master's degree in practising accounting from Monash University in 2001 and an Executive Master of Business Administration degree from the University of Western Ontario in 2005.

Yu (Albert) Ren, Ph.D. has served as our Chief Strategy and Corporate Development Officer since January 2021. Prior to joining us, Dr. Ren held various positions at Merck & Co., Inc. (NYSE: MRK). He had previously worked at Portfolio Management & Global Clinical Development Finance responsible for financial / strategic planning on KEYTRUDA® development and facilitation of KEYTRUDA® clinical trials prioritization. He also worked at Merck Corporate Strategy Office on enterprise strategy and at Merck Corporate Business Development on a variety of business development transactions. Before returning to Merck, Dr. Ren served as a principal consultant at Quintiles/IQVIA Consulting from 2009 to 2010. Dr. Ren gained extensive consulting experience at McKinsey & Company from 2007 to 2009. Dr. Ren started his professional career at Merck Research Lab working on Pharmaceutical Research & Development from discovery to early clinical development from 2001 to 2006. Dr. Ren received his Ph.D. in Chemistry from University of Minnesota in 2001, and bachelor's degree in Chemistry from Peking University in 1996.

Key Employees

Yan Li, M.B.A. has served as our Senior Vice President of Bioinformatics and Information Technology since November 2011. Ms. Li has over 25 years of experience in software development, with about 20 years focusing on the development of informatics software tools for antibody library design and analysis. Ms. Li served as the senior software engineer and applications scientist at Abmaxis from November 2001 to May 2006. Following the acquisition of Abmaxis by Merck & Co, Ms. Li worked at Merck & Co from May 2006 to December 2010. She received the Merck Award for Excellence in "Innovative Technologies" with the team and a Special Award for her contribution. Ms. Li received her bachelor's degree in information science from East China University of Science and Technology in 1995 and a Master of Business Administration degree in 2010 from Santa Clara University.

Xiaohong (Kristine) She has served as our Senior Vice President, Head of Clinical Operations since November 2011. Ms. She has over 20 years of laboratory and laboratory management experience in the renowned laboratories in the United States and has completed multiple projects in molecular biology and immunology at the labs of University of Chicago, the Genome Center at Stanford University, neurological animal studies at Stanford Palo Alto Veteran's Hospital, and cloning and functional screening of cDNA libraries at Caltech. Her work has been published in Nature, Science, Biotechnology, among many other notable publications. Ms. She received her bachelor's degree in microbiology from Wuhan University in 1986 and master's degree in biochemistry and microbiology from the Institute of Microbiology of the Chinese Academy of Sciences in 1989. Ms. She's spouse is Peter (Peizhi) Luo, Ph.D., our Co-Founder, Chief Executive Officer and Chairman of the Board.

Guizhong Liu, Ph.D. has served as our Head of Biology and Pharmacology since October 2015. Dr. Liu has over 15 years of experience in drug discovery and development, both in small molecule

kinase inhibitors and large molecule antibodies in oncology and immunology field. He has published over 40 peer-reviewed papers in high-profile journals involving key signalling pathways and targets in cancer biology. From July 2007 to August 2011, Dr. Liu served as an assistant professor of the department of oncological science at Mount Sinai School of Medicine. Prior to joining us, Dr. Liu served as head of molecular cancer biology in CrownBio from October 2011 to September 2015. Dr. Liu received his bachelor's degree in biology in 1992 and master's degree in cell biology in 1995 from Beijing Normal University and Ph.D. degree in cell biology from Peking Union Medical College in 1998. He also completed his postdoctoral training in cancer biology at Mount Sinai School of Medicine in 2004.

Alexander Goergen has served as our Head of Business Development since October 2017 when he joined the Company. Mr. Goergen has worked in various roles at the Covance, TRC and International AIDS Vaccine Initiative from October 2008 to October 2012. Prior to joining us, Alexander worked in business development for Catalent Pharma Solutions Biologics Division since October 2012. Mr. Goergen completed many licensing, manufacturing, and cell line development programs both domestically and internationally during his previous employment. Mr. Goergen received his bachelor's degree in Chemistry from Lafayette College in 2008 and master's degree in Biotechnology from the University of Wisconsin-Madison in 2011.

Yuren (Ron) Wang, Ph.D. has served as our Head of Portfolio Management since September 2019. Dr. Wang served as the senior director of business development in Reaction Biology Corporation from March 2013 to December 2018. Prior to joining us, he served as the vice president of R&D research for Jemincare Therapeutics (USA), responsible for the scientific evaluation of projects, in-licensing and portfolio management from December 2018 to August 2019. Dr. Wang received his bachelor's degree in plant science from Shandong Agricultural University in 1982, master's degree in biochemistry from the Graduate School of Chinese Academy of Agricultural Sciences in 1986 and Ph.D. degree in cell and molecular biology from the University of Pennsylvania in 1996. He also completed his postdoctoral training in molecular pharmacology at the University of Pennsylvania in 1999.

Non-Employee Directors

Daniel Auerbach has served as our director since July 2013. He joined Fidelity in 1994 and has been serving as the senior managing partner of Eight Roads, a proprietary investment arm backed by Fidelity. Mr. Auerbach held key board roles with a multitude of companies, including the former director of Alibaba Group, Wuxi Pharmatech during 2005 to 2007, Innovent (HKEx: 1801) during 2011 to 2018, Hua Medicine (HKEx: 2552) during 2010 to 2018 and others. Mr. Auerbach received his bachelor's degree in Economics and Foreign Languages from Dartmouth College in 1980 and Master of Business Administration degree from The Harvard Business School in 1987.

Chong Xu, Ph.D. has served as our director since May 2020. Dr. Xu joined F-Prime Capital in 2015 and is currently serving as its principal. Prior to joining F-Prime Capital, Dr. Xu worked at McKinsey & Company's Boston office as a consulting professional from September 2014 to October 2015. Dr. Xu also worked at Massif Partners from April 2011 to May 2012 as an investment professional. From December 2009 to April 2011, Dr. Xu served as an investment professional at Affirmed Healthcare. Dr. Xu received his bachelor's degree in biology from Zhejiang University in 2001, a Ph.D. degree in cell biology from University of Virginia in 2009, and an MBA degree from Darden School of Business in 2014.

Yu Miao has served as our director since May 2020. He also serves as the executive director of GP Healthcare Capital Co., Ltd, mainly responsible for equity investment focusing on healthcare industry. Prior to joining GP Healthcare Capital in April 2015, Mr. Yu worked at Eli Lilly & Co. focusing on quality assurance from July 2011 to June 2012 and Oriza Seed Fund Management Co., Ltd., focusing on equity investment from July 2014 to April 2015. Mr. Miao received

his bachelor's degree in pharmacy from China Pharmaceutical University in 2011, and master's degree in pharmaceutical science from Northeastern University in 2014.

Yunxia Yang has served as our director since June 2019. She has also served as a managing director of Sequoia Capital China since May 2015 where she focuses on healthcare investment. Ms. Yang has also served as a director of Burning Rock Biotech Ltd since January 2017. Before starting in venture capital, Ms. Yang worked as a product manager at GE Healthcare from June 2006 to July 2007 and a business development manager at Johnson & Johnson from July 2009 to April 2011. She then worked at Legend Capital, where she led investment in areas covering gene diagnostics, medical devices and healthcare service, as a member of the healthcare team from April 2011 to May 2015. Ms. Yang received her master's degree in clinical science from Huazhong Technology University in 1997 and Master of Business Administration degree from Duke University in 2009.

Lefei Sun has served as our director since December 2019. Mr Sun has been a non-executive director of Hong Kong Asia Medical Holding Limited, a leading hospital management group in Asia with hospital assets such as Wuhan Asia Heart Hospital, from November 2018. He is also a non-executive director of various biotech companies such as Ocumension Therapeutics (HKEx: 1477) and CANbridge Pharmaceuticals Inc. Mr. Sun has served as head of China healthcare at General Atlantic since May 2018, and has been a managing director since January 2020, in charge of private equity investment and portfolio management in healthcare and life sciences sectors. From December 2014 to April 2018, Mr. Sun was a founding partner and a member of investment committee of Beijing HuaTai Ruihe Investment Fund Management Company (LLP), also known as Huatai Healthcare Investment Fund. Prior to joining General Atlantic, Mr. Sun served as various investment roles at Hony Capital, Credit Suisse, OrbiMed, Orchid Asia, and as a management consultant at McKinsey & Company, all in the healthcare sector. Mr. Sun obtained his master's degree in neurosciences from Johns Hopkins University School of Medicine in 2006, and his bachelor's degree in mathematics and physics from Tsinghua University in 2002.

Andy (Yiu Leung) Cheung will serve as our independent director immediately upon the effectiveness of our registration statement on Form F-1, of which this prospectus is a part. Mr. Cheung has many years of auditing and accounting professional experience. Mr. Cheung was deputy area managing partner of Ernst & Young ("EY") in Asia Pacific overseeing the business operations, finance, information technology and risk management functions from July 2018 to June 2020. Mr. Cheung was the assurance leader for EY in Greater China from July 2013 to June 2018. Mr. Cheung has served as an independent non-executive director of JW (Cayman) Therapeutics Co. Ltd (HKEX:2126) since October 2020. Mr. Cheung received his bachelor's degree in accounting and finance from the University of Lancaster in the United Kingdom in June 1982. He obtained a master's degree in accounting and finance from London School of Economics in the United Kingdom in August 1983. Mr. Cheung is a member of Hong Kong Institute of Certified Public Accountants.

Min Li will serve as our independent director immediately upon the effectiveness of our registration statement on Form F-1, of which this prospectus is a part. Dr. Min Li is the founder, CEO and director of SciNeuro Pharmaceuticals, a biotechnology company headquartered in Shanghai focused on developing innovative therapeutics for CNS diseases. Dr. Li was formerly Senior Vice President, Global Head of Neuroscience R&D at GSK from 2013 to 2019. During his employment with GSK, he also served as General Manager of GSK R&D China, the highest-ranking officer at the country level. Before moving into industry, Dr. Li had served as a tenured Professor of Neuroscience at Johns Hopkins University School of Medicine for over 20 years. Dr. Li served on scientific advisory boards, non-profit and corporate boards that include multinational corporations (Corning, Pfizer and Merck), disease foundations (RYR-foundation, AHA) and biotech companies (Xencor, Affymax, Datavant, Icagen, and Coda Biotechnologies). He is also an Adjunct Professor of Neuroscience at Johns Hopkins School of Medicine. Dr. Li received his B.S. degree in biochemistry from Wuhan University in 1984 and his Ph.D. in molecular immunology from Johns Hopkins University in 1990. As

a Helen Hay Whitney Fellow, he performed postdoctoral research at University of California San Francisco. With a Sloan Award, in 1994 he returned to Johns Hopkins to join the faculty as an Esther A & Joseph Klingenstein Neuroscience Fellow. Dr. Li has published more than 120 scientific articles and holds patents in biotechnologies. His research and scholarship were recognized internationally with awards including being a fellow of American Association for Advancement of Science and an established fellow of American Heart Association.

Employment Agreements and Indemnification Agreements

We have entered into employment agreements with each of our executive officers. Each of our executive officers is employed for a period of twelve months, which will be renewed automatically and extended automatically for a period of twelve months unless the executive or us gives prior written notice. We may terminate an executive officer's employment for cause at any time without advance notice in certain events. We may terminate an executive officer's employment by giving a prior written notice or by paying certain compensation. An executive officer may terminate his or her employment at any time by giving a prior written notice.

Each executive officer has agreed to hold, unless expressly consented to by us, at all times during and after the termination of his or her employment agreement, in strict confidence and not to use, any of our confidential information or the confidential information of our customers and suppliers. In addition, each executive officer has agreed to be bound by certain non-competition and non-solicitation restrictions during the term of his or her employment and for a maximum of two years following the last date of employment.

We plan to enter into indemnification agreements with each of our directors and executive officers. Under these agreements, we agree to indemnify our directors and executive officers against certain liabilities and expenses incurred by such persons in connection with claims made by reason of their being a director or officer of our company.

Board of Directors

Our board of directors will consist of six directors, including two independent directors, namely Andy (Yiu Leung) Cheung and Min Li, upon the SEC's declaration of effectiveness of our registration statement on Form F-1 to which this prospectus forms a part. A director is not required to hold any shares in our company to qualify to serve as a director. The Corporate Governance Rules of the Nasdaq generally require that a majority of an issuer's board of directors must consist of independent directors. However, the Corporate Governance Rules of the Nasdaq permit foreign private issuers like us to follow "home country practice" in certain corporate governance matters. We rely on this "home country practice" exception and do not have a majority of independent directors serving on our board of directors.

A director who is in any way, whether directly or indirectly, interested in a contract or proposed contract with our company is required to declare the nature of his or her interest at a meeting of our directors. A general notice given to the directors by any director to the effect that he or she is a member, shareholder, director, partner, officer or employee of any specified company or firm and is to be regarded as interested in any contract or transaction with that company or firm shall be deemed a sufficient declaration of interest for the purposes of voting on a resolution in respect to a contract or transaction in which he/she has an interest, and after such general notice it shall not be necessary to give special notice relating to any particular transaction. A director may vote in respect of any contract or proposed contract or arrangement notwithstanding that he/she may be interested therein and if he/she does so, his/her vote shall be counted and he/she may be counted in the quorum at any meeting of the directors at which any such contract or proposed contract or arrangement is considered, subject to any separate requirement for Audit Committee approval under applicable law or the Listing Rules of

the Nasdaq. Our board of directors may exercise all of the powers of our company to borrow money, to mortgage or charge its undertaking, property and uncalled capital, or any part thereof, and to issue debentures, debenture stock or other securities whenever money is borrowed or as security for any debt, liability or obligation of our company or of any third party. None of our directors has a service contract with us that provides for benefits upon termination of service as a director.

Committees of the Board of Directors

Prior to the completion of this offering, we intend to establish an audit committee, a compensation committee and a nominating and corporate governance committee under our board of directors. We intend to adopt a charter for each of the three committees prior to the completion of this offering. Each committee's members and functions are described below.

Audit Committee. Our audit committee will consist of Andy (Yiu Leung) Cheung and Min Li, and is chaired by Andy (Yiu Leung) Cheung. We have determined that Andy (Yiu Leung) Cheung and Min Li satisfy the requirements of Rule 5605(a)(2) of the Listing Rules of the Nasdaq and meet the independence standards under Rule 10A-3 under the Securities Exchange Act of 1934, as amended. We have determined that Andy (Yiu Leung) Cheung qualifies as an "audit committee financial expert." The audit committee oversees our accounting and financial reporting processes and the audits of the financial statements of our company. The audit committee is responsible for, among other things:

- reviewing and recommending to our board for approval, the appointment, re-appointment or removal of the independent auditor, after considering its annual performance evaluation of the independent auditor;
- approving the remuneration and terms of engagement of the independent auditor and pre-approving all auditing and non-auditing services permitted to be performed by our independent auditors at least annually;
- obtaining a written report from our independent auditor describing matters relating to its independence and quality control procedures;
- reviewing with the independent registered public accounting firm any audit problems or difficulties and management's response;
- discussing with our independent auditor, among other things, the audits of the financial statements, including whether any material information should be disclosed, issues regarding accounting and auditing principles and practices;
- reviewing and approving all proposed related party transactions, as defined in Item 404 of Regulation S-K under the Securities Act;
- reviewing and recommending the financial statements for inclusion within our quarterly earnings releases and to our board for inclusion in our annual reports;
- discussing the annual audited financial statements with management and the independent registered public accounting firm;
- reviewing the adequacy and effectiveness of our accounting and internal control policies and procedures and any special steps taken to monitor and control major financial risk exposures;
- at least annually, reviewing and reassessing the adequacy of the committee charter;
- approving annual audit plans, and undertaking an annual performance evaluation of the internal audit function;
- establishing and overseeing procedures for the handling of complaints and whistleblowing;

- meeting separately and periodically with management and the independent registered public accounting firm;
- monitoring compliance with our code of business conduct and ethics, including reviewing the adequacy and effectiveness of our procedures to ensure proper compliance; and
- reporting regularly to the board.

Compensation Committee. Our compensation committee will consist of Peter Luo, Andy (Yiu Leung) Cheung and Min Li and is chaired by Peter Luo. We have determined that Andy (Yiu Leung) Cheung and Min Li satisfy the "independence" requirements of Rule 5605(a)(2) of the Listing Rules of the Nasdaq. The compensation committee assists the board in reviewing and approving the compensation structure, including all forms of compensation, relating to our directors and executive officers. Our Chief Executive Officer may not be present at any committee meeting during which their compensation is deliberated upon. The compensation committee is responsible for, among other things:

- overseeing the development and implementation of compensation programs in consultation with our management;
- at least annually, reviewing and approving, or recommending to the board for its approval, the compensation for our executive officers;
- at least annually, reviewing and recommending to the board for determination with respect to the compensation of our non-executive directors;
- at least annually, reviewing periodically and approving any incentive compensation or equity plans, programs or other similar arrangements;
- reviewing executive officer and director indemnification and insurance matters;
- overseeing our regulatory compliance with respect to compensation matters, including our policies on restrictions on compensation plans and loans to directors and executive officers;
- at least annually, reviewing and reassessing the adequacy of the committee charter;
- selecting compensation consultant, legal counsel or other adviser only after taking into consideration all factors relevant to that person's independence from management; and
- reporting regularly to the board.

Nominating and Corporate Governance Committee. Our nominating and corporate governance committee will consist of Peter Luo, Andy (Yiu Leung) Cheung and Min Li, and is chaired by Peter Luo. We have determined that Andy (Yiu Leung) Cheung and Min Li satisfy the "independence" requirements of Rule 5605(a)(2) of the Listing Rules of the Nasdaq. The nominating and corporate governance committee assists the board in selecting individuals qualified to become our directors and in determining the composition of the board and its committees. The nominating and corporate governance committee is responsible for, among other things:

- recommending nominees to the board for election or re-election to the board, or for appointment to fill any vacancy on the board;
- reviewing annually with the board the current composition of the board with regards to characteristics such as independence, knowledge, skills, experience, expertise, diversity and availability of service to us;
- developing and recommending to our board such policies and procedures with respect to nomination or appointment of members of our board and chairs and members of its committees or other corporate governance matters as may be required pursuant to any SEC or Nasdaq rules, or otherwise considered desirable and appropriate;

- selecting and recommending to the board the names of directors to serve as members of the audit committee and the compensation committee, as well as of the nominating and corporate governance committee itself;
- at least annually, reviewing and reassessing the adequacy of the committee charter;
- developing and reviewing at least annually the corporate governance principles adopted by the board and advising the board with respect to significant developments in the law and practice of corporate governance and our compliance with such laws and practices; and
- evaluating the performance and effectiveness of the board as a whole.

Duties and Functions of Directors

Under Cayman Islands law, our directors owe fiduciary duties to our company, including a duty of loyalty, a duty to act honestly and a duty to act in what they consider in good faith to be in our best interests. Our directors must also exercise their powers only for a proper purpose. Our directors also owe to our company a duty to exercise the skill they actually possess and such care and diligence that a reasonable prudent person would exercise in comparable circumstances. It was previously considered that a director need not exhibit in the performance of his duties a greater degree of skill than may reasonably be expected from a person of his knowledge and experience. However, English and Commonwealth courts have moved towards an objective standard with regard to the required skill and care and these authorities are likely to be followed in the Cayman Islands. In fulfilling their duty of care to us, our directors must ensure compliance with our memorandum and articles of association, as amended and restated from time to time. Our company has the right to seek damages if a duty owed by our directors is breached. In limited exceptional circumstances, a shareholder may have the right to seek damages in our name if a duty owed by our directors is breached. The functions and powers of our board of directors include, among others, (i) convening shareholders' annual general meetings and reporting its work to shareholders at such meetings, (ii) declaring dividends, (iii) appointing officers and determining their terms of offices and responsibilities, (iv) exercising the borrowing powers of our company, and (v) approving the transfer of shares of our company, including the registering of such shares in our share register.

Terms of Directors and Officers

Our officers are elected by and serve at the discretion of the board of directors. Other than Andy (Yiu Leung) Cheung, Min Li and Man Kin (Raymond) Tam, each director is not subject to a term of office and holds office until such time as his successor takes office or until the earlier of his death, resignation or removal from office pursuant to the applicable provisions of our memorandum and articles of association. The initial term of our independent director nominees, namely Andy (Yiu Leung) Cheung and Min Li, is two-year from the effectiveness of the appointment. The initial term of our non-independent director nominee, Man Kin (Raymond) Tam, is one-year from the effectiveness of the appointment. The initial term for each of Andy (Yiu Leung) Cheung, Min Li and Man Kin (Raymond) Tam is subject to re-appointment by the board of directors and affirmative vote of Chairman of the board of directors. A director will be removed from office automatically if, among other things, the director (i) becomes bankrupt or makes any arrangement or composition with his creditors; (ii) dies or is found by our company to be of unsound mind; (iii) resigns by notice in writing to our company; (iv) without special leave of absence from our board of directors, is absent from three consecutive meetings of the board and the board resolves that his office be vacated; (v) is prohibited by law from being a director; or (vi) is removed from office pursuant to any other provisions of our post-offering amended and restated memorandum and articles of association.

Interested Transactions

A director may, subject to any separate requirement for audit and risk committee approval under applicable law or applicable Nasdaq listing rules, vote in respect of any contract or transaction in which he or she is interested, provided that the nature of the interest of any directors in such contract or transaction is disclosed by him or her at or prior to its consideration and any vote in that matter.

Compensation of Directors and Executive Officers

For the fiscal year ended December 31, 2020, we paid an aggregate of US\$2.6 million in cash to our executive officers, and we paid US\$54,000 cash compensation to our then non-executive director, Mr. Tom Beck, for the fiscal year ended December 31, 2020. We did not pay any cash compensation to other non-executive directors. For the fiscal year ended December 31, 2020, we did not set aside or accrue expenses related to pension, retirement or other similar benefits to our executive officers and directors. Our PRC subsidiary is required by law to make contributions equal to certain percentages of each employee's salary for his or her pension insurance, medical insurance, unemployment insurance and other statutory benefits and a housing provident fund. For share incentive grants to our directors and executive officers, see "—Share Incentive Plan."

Share Incentive Plan

Adagene Inc. Second Amended and Restated Share Incentive Plan

In November 2015, we adopted Adagene Inc. Share Incentive Plan, or the 2015 Plan, which was later superseded and replaced by Adagene Inc. Amended and Restated Share Incentive Plan, or the 2017 Plan, in September 2017. In December 2019, we adopted the Second Amended and Restated Share Incentive Plan, or the 2019 Plan, to supersede and replace the 2017 Plan. The terms of the 2015 Plan, the 2017 Plan and the 2019 Plan are substantially the same other than the maximum aggregate number of shares we may issue under the respective plan.

The purpose of the 2019 Plan is to attract, motivate, retain and reward certain officers, employees, directors and other eligible persons and to further link the interests of award recipients with those of our shareholders generally. The 2019 Plan provides for the issuance of up to an aggregate of 11,391,131 of our ordinary shares. Upon the effectiveness of our registration statement on Form F-1, we will terminate the authority to grant additional awards under the 2019 Plan and all future awards will be granted under the 2021 Plan. Therefore, the effective maximum number of shares issuable under the 2019 Plan will be 10,728,501. As of the date of this prospectus, the aggregate number of our ordinary shares underlying our outstanding awards under the 2019 Plan is 3,601,522, excluding awards that were forfeited, cancelled or exercised after the relevant grant dates. The term of the awards will expire not more than ten years after the date of grant.

The following paragraphs summarize the principal terms of the 2019 Plan.

Types of Awards. The 2019 Plan permits the awards of options, share appreciation rights, ordinary shares or restricted shares.

Plan Administration. The 2019 Plan shall be administered by our board of directors or one or more committees appointed by the board of directors or another committee (within its delegated authority), the Plan Administrator.

Promissory Notes. The promissory notes with respect to the 2019 Plan, or the 2019 Promissory Notes are full recourse, repayable within a period of time determined by the Plan Administrator, which should not exceed five years (subject to certain early repayment events), and bear interest at the interest rate determined by the Plan Administrator but not less than the interest rate necessary to avoid the imputation of interest under United States Internal Revenue Code of 1986, as amended or other

applicable tax law. Certain plan participants previously purchased our shares with such promissory notes. The current amount outstanding under these notes (which is also the largest aggregate amount outstanding since the first issuance of the promissory notes) was US\$1.8 million immediately.

Repayment of the Promissory Notes. The terms, repayment provisions, and collateral release provisions of the note and the pledge securing the note shall conform with all applicable rules and regulations, including those of the Federal Reserve Board of the United States and any applicable law, as then in effect.

Eligibility. The plan administrators may decide that an award under the 2019 Plan be granted to any employee, officer or director of the Company or its affiliates, or that it be granted to any consultant or adviser who provides services to the Company or its affiliates.

Award Agreements. Each award under the 2019 Plan shall be evidenced by an award agreement in the form approved by the plan administrators. The terms of the award agreements will be determined by the plan administrators and consistent with the terms of the 2019 Plan.

Conditions of Award. The plan administrators shall determine the participants, types of awards, numbers of shares to be covered by awards, terms and conditions of each award, including, but not limited to, the price and number of securities to be offered or awarded, the installments (if applicable) in which such awards will become exercisable or will vest, performance targets (if applicable), the events of termination or reversion of such awards.

Transfer Restrictions. With a few exceptions, no right of interest of a participant in any award may be subject in any manner to sale, transfer, anticipation, alienation, assignment, pledge, encumbrance or charge. This restriction does not apply to (i) transfers to our company, (ii) transfers by gift or domestic relations order to one or more family members, (iii) the designation of a beneficiary to receive benefits if a participant dies or transfers by will, (iv) permitted transfers or exercises on behalf of a participant by the participant's duly authorized legal representative if the participant has suffered a disability.

Reduction or Clawback of Awards. The awards granted under the 2019 Plan are subject to the terms of our recoupment, clawback or similar policy as it may be in effect from time to time, as well as any similar provisions of applicable law, any of which could in certain circumstances require repayment or forfeiture of awards or any ordinary shares or other cash or property received with respect to the awards (including any value received from a disposition of the shares acquired upon payment of the Awards).

Amendment and Termination of the 2019 Plan. The board of directors may, at any time, terminate or, from time to time, amend, modify or suspend the 2019 Plan, in whole or in part. No awards may be granted during any period that the board of directors suspends the 2019 Plan. To the extent then required by applicable law or listing agency, any amendment to the 2019 Plan may be subject to shareholder approval. Unless earlier terminated by the board of directors, the 2019 Plan will terminate at the close of business on the day before the 10th anniversary of the date the board of directors approved the 2019 Plan.

Adagene Inc. 2021 Performance Incentive Plan

In January 2021, our board of directors adopted, and our shareholders approved, the 2021 Performance Incentive Plan, or the 2021 Plan, to provide an additional means through the grant of awards to attract, motivate, retain and reward selected employees and other eligible persons. The terms of the 2021 Plan are summarized below. Employees, officers, directors and consultants that provide services to us or one of our subsidiaries may be selected to receive awards under the 2021 Plan.

Our board of directors or a committee appointed by the board administers the 2021 Plan. The plan administrator has broad authority to:

- select participants and determine the types of awards that they are to receive;
- determine the number of shares that are to be subject to awards and the terms and conditions of awards, including the price (if any) to be paid for the shares or the award and establish the vesting conditions (if applicable) of such shares or awards;
- cancel, modify or waive our rights with respect to, or modify, discontinue, suspend or terminate any or all outstanding awards, subject to any required consents;
- construe and interpret the terms of the 2021 Plan and any agreements relating to the plan;
- accelerate or extend the vesting or exercisability or extend the term of any or all outstanding awards subject to any required consent;
- subject to the other provisions of the 2021 Plan, make certain adjustments to one or more outstanding awards (including a repricing of the exercise or base price of any outstanding option or share appreciation right without shareholder approval) and authorize the termination, conversion, substitution or succession of awards; and
- allow the purchase price of an award or our ordinary shares to be paid in the form of cash, check or electronic funds transfer, by the delivery of previously-owned ordinary shares or by a reduction of the number of shares deliverable pursuant to the award, by services rendered by the recipient of the award, by notice and third party payment or cashless exercise on such terms as the plan administrator may authorize or any other form permitted by law.

A total of 2,994,000 of our ordinary shares is authorized for issuance with respect to awards granted under the 2021 Plan. The share limit will automatically increase on the first trading day in January of each year (commencing with 2022) by an amount equal to (1) 5% of the total number of our outstanding ordinary shares on December 31 of the prior year, or (2) such lesser number as determined by our board of directors. Any shares subject to awards granted under the 2021 Plan or our Second Amended and Restated Share Incentive Plan that are not paid, delivered or exercised before they expire or are canceled or terminated, or otherwise fail to vest, as well as shares used to pay the purchase or exercise price of such awards or related tax withholding obligations, will become available for new award grants under the 2021 Plan. As of the date of this prospectus, no awards have been granted under the 2021 Plan, and the full number of shares authorized under the 2021 Plan is available for award purposes.

Awards under the 2021 Plan may be in the form of incentive or nonqualified stock options, share appreciation rights, share bonuses, restricted shares, share units, restricted share units and other forms of awards including cash awards. Awards under the plan generally will not be transferable other than by will or the laws of descent and distribution, except that the plan administrator may authorize certain transfers.

Options and share appreciation rights may not be granted under the 2021 Plan at exercise prices below the fair market value of the underlying ordinary shares on the date of grant, except that the plan administrator may grant such awards with exercise prices below the fair market value of the underlying shares to service-providers who are not subject to U.S. taxes. The maximum term of options and share appreciation rights granted under the plan is ten years. Incentive stock options granted to any 10% owner of our ordinary shares must have an exercise price that is at least 110% of the fair market value of our ordinary shares on the grant date and a maximum term of five years. These and other awards may also be issued solely or in part for services. Awards are generally paid in cash or our ordinary shares. The plan administrator may provide for the deferred payment of awards and may determine the terms applicable to deferrals.

As is customary in incentive plans of this nature, the number and type of shares available under the 2021 Plan and any outstanding awards, as well as the exercise or purchase prices of awards, will be subject to adjustment in the event of certain reorganizations, mergers, combinations, recapitalizations, share splits, share dividends or other similar events that change the number or kind of shares outstanding, and extraordinary dividends or distributions of property to the shareholders.

Generally, and subject to limited exceptions set forth in the 2021 Plan, if we dissolve or undergo certain corporate transactions such as a merger, business combination or other reorganization, or a sale of all or substantially all of our assets, all awards then-outstanding under the 2021 Plan will become fully vested or paid, as applicable, and will terminate or be terminated in such circumstances, unless the plan administrator provides for the assumption, substitution or other continuation of the award. The plan administrator also has the discretion to establish other change in control provisions with respect to awards granted under the 2021 Plan. For example, the administrator could provide for the acceleration of vesting or payment of an award in connection with a corporate event that is not described above and provide that any such acceleration shall be automatic upon the occurrence of any such event.

Our board of directors may amend or terminate the 2021 Plan at any time, but no such action will affect any outstanding award in any manner materially adverse to a participant without the consent of the participant. Plan amendments will be submitted to shareholders for their approval as required by applicable law or deemed necessary or advisable by the board. The 2021 Plan is not exclusive; our board of directors and compensation committee may grant shares and performance incentives or other compensation, in shares or cash, under other plans or authority.

The plan will terminate on January 16, 2031. However, the plan administrator will retain its authority until all outstanding awards are exercised or terminated.

The following table summarizes, as of the date of this prospectus, the number of ordinary shares under outstanding awards that we granted to our directors and executive officers under the 2019 Plan,

which replaced the 2015 Plan, excluding awards that were exercised, forfeited or canceled after the relevant grant dates.

Name	Ordinary Shares Underlying Equity Awards Granted	Exercise Price (US\$/Share)	Date of Grant	Date of Expiration
Executive Officers				
Peter (Peizhi) Luo, Ph.D.	287,415	\$13.85	January 16, 2021	January 16, 2031
Fangyong (Felix) Du, Ph.D.	111,744	\$13.85	January 16, 2021	January 16, 2031
Hua Gong, M.D., Ph.D.	*	\$2.26	August 2020	August 2030
JC Xu, M.D., Ph.D.	*	\$2.26	August 2020	August 2030
Qinghai Zhao, Ph.D.	*	\$2.26	October 2020	October 2030
Man Kin (Raymond) Tam, M.B.A., B. Eng.	*	\$1.83	August 2020	August 2030
	*	\$1.48	March 2020	March 2030
Yu (Albert) Ren, Ph.D.	—	—	—	—
Non-Employee Directors				
Daniel Auerbach, M.B.A.	—	—	—	—
Chong Xu, Ph.D.	—	—	—	—
Yu Miao	—	—	—	—
Yunxia Yang, M.B.A.	—	—	—	—
Lefei Sun	—	—	—	—
Andy (Yiu Leung) Cheung (independent director nominee)	—	—	—	—
Min Li (independent director nominee)	—	—	—	—
All directors and executive officers as a group	2,112,986		Various dates from March 2020 to January 2021	Various dates from March 2030 to January 2031

Note:

* The shares held by each of these directors and executive officers represent less than 1% of our total outstanding shares.

As of the date of this prospectus, our award holders other than our directors and executive officers as a group held outstanding awards to purchase 1,488,536 ordinary shares. For discussions of our accounting policies and estimates for awards granted pursuant to the 2019 Plan, see "Management's Discussion and Analysis of Financial Condition and Results of Operations—Critical Accounting Policies, Judgments and Estimates—Share-based compensation."

PRINCIPAL SHAREHOLDERS

The following table sets forth information concerning the beneficial ownership of our ordinary shares as of the date of this prospectus, assuming conversion of all of our outstanding series A-1 preferred shares, series A-2 preferred shares, series B preferred shares, series C-1 preferred shares, series C-2 preferred shares and series C-3 preferred shares into ordinary shares, on a one-to-one basis by:

- each of our directors and executive officers; and
- each person known to us to beneficially own more than 5% of our ordinary shares.

The calculations in the table below are based on 43,532,275 ordinary shares on an as-converted basis outstanding as of the date of this prospectus and ordinary shares outstanding immediately after the completion of this offering, including (i) ordinary shares to be sold by us in this offering in the form of ADSs, and (ii) 27,249,824 ordinary shares converted from our outstanding preferred shares, assuming that the underwriters do not exercise their option to purchase additional ADSs and excluding the 2,262,500 ordinary shares, issued but deemed to be not outstanding as of the date of this prospectus, held by Great Han Fortune LP.

Beneficial ownership is determined in accordance with the rules and regulations of the SEC. In computing the number of shares beneficially owned by a person and the percentage ownership of that person, we have included shares that the person has the right to acquire within 60 days, including through the exercise of any option, warrant, or other right or the conversion of any other security.

These shares, however, are not included in the computation of the percentage ownership of any other person.

	Ordinary Shares Beneficially Owned Prior to this Offering		Ordinary Shares Beneficially Owned After this Offering		
	Number	%**	Number	Percentage of total ordinary shares on an as-converted basis***	Percentage of aggregate voting power****
Directors and Executive Officers:†					
Peter (Peizhi) Luo ⁽¹⁾	10,449,636	23.5%			
Fangyong (Felix) Du ⁽²⁾	1,052,188	2.4%			
Hua Gong	*	*			
JC Xu	*	*			
Qinghai Zhao	*	*			
Man Kin (Raymond) Tam	*	*			
Yu (Albert) Ren	—	—			
Daniel Auerbach	—	—			
Chong Xu	—	—			
Yunxia Yang	—	—			
Yu Miao	—	—			
Lefei Sun	—	—			
Andy (Yiu Leung) Cheung (independent director nominee)	—	—			
Min Li (independent director nominee)					
All directors and executive officers as a group	10,449,636	23.5%			
Principal Shareholders:					
Peter Luo act-in-concert group ⁽³⁾	10,449,636	23.5%			
JSR Limited ⁽⁴⁾	5,353,242	12.3%			
Asia Ventures II L.P. ⁽⁵⁾	4,826,037	11.1%			
F-Prime Capital Partners Healthcare Fund III LP ⁽⁶⁾	4,826,037	11.1%			
Wuxi Pharmatech Healthcare Fund I L.P. ⁽⁷⁾	4,706,946	10.8%			
General Atlantic Singapore AI Pte. Ltd. ⁽⁸⁾	4,452,441	10.2%			

Notes:

* Less than 1% of our total outstanding shares on an as-converted basis.

** For each person and group included in this table, percentage ownership is calculated by dividing the number of shares beneficially owned by such person or group by the sum of (i) 43,532,275 being the number of ordinary shares on an as-converted basis as of the date of this prospectus and (ii) the number of ordinary shares underlying share options held by such person or group that are exercisable within 60 days after the completion of this offering.

*** For each person and group included in this table, percentage ownership is calculated by dividing the number of shares beneficially owned by such person or group by the sum of (i) , being the number of ordinary shares on an as-converted basis outstanding immediately after this offering, (ii) , being the number of ordinary shares to be sold by us in this offering in the form of ADSs, and (iii) the number of ordinary shares underlying share options held by such person or group that are exercisable within 60 days after this offering.

**** For each person and group included in this column, percentage of voting power is calculated by dividing the voting power beneficially owned by such person or group by the voting power of all of our ordinary shares as a single class.

- † The business address of our directors and executive officers, except for Fangyong (Felix) Du, Man Kin (Raymond) Tam, Daniel Auerbach, Chong Xu, Yunxia Yang, Yu Miao and Lefei Sun, is Center 2150, 315 Montgomery Street, 9th Floor, San Francisco CA 94104, United States of America. The business address of Fangyong (Felix) Du and Man Kin (Raymond) Tam is 4F, Building C14, No. 218, Xinghu Street, Suzhou Industrial Park, China 215123; the business address of Daniel Auerbach is Suite 2201, Level 22, Pacific Place Two, 88 Queensway, Admiralty, Hong Kong, the business address of Chong Xu is One Main Street, 13th Floor, Cambridge, MA 02142; the business address of Yunxia Yang is Room 3606, China Central Place Tower 3, 77 Jianguo Road, Beijing 100025, PRC, the business address of Yu Miao is Unit 4901, One Lujiazui, No.68, Yin Cheng(C) Rd., Shanghai 200120, PRC; and the business address of Lefei Sun is Suite 5704-5706, 57F, Two IFC, 8 Finance Street, Central, Hong Kong; the business address of Andy (Yiu Leung) Cheung is Queen's Terrace, Tower 1 51/F Unit D, 1 Queen Street, Sheungwan, Hong Kong.
- (1) Represents 10,449,636 ordinary shares on an as-converted basis held by Peter Luo act-in-concert group, as set forth in note (3) below.
 - (2) Represents (i) 810,944 ordinary shares held by Ping Ren, who is the spouse of Fangyong (Felix) Du, (ii) 228,744 ordinary shares held by Great Han Fortune LP for the benefit of Dr. Du that are vested or will be vested within 60 days after this offering, and (iii) 12,500 ordinary shares held by Great Han Fortune LP. Great Han Fortune LP is controlled by Peter Luo and is formed to hold ordinary shares in connection with the share awards issued under the 2019 Plan.
 - (3) Represents (i) 7,187,314 ordinary shares held by HAN 2020 GRAT, for which Peter Luo is a Trustee, (ii) 596,174 ordinary shares issuable immediately prior to the completion of this offering upon the conversion of 596,174 Series A-1 preferred shares held by HAN 2020 GRAT, for which Peter Luo is a Trustee, (iii) 287,415 ordinary shares underlying 287,415 share options granted to Peter Luo that are vested or will be vested within 60 days after this offering, (iv) 83,333 ordinary shares held by Great Han Fortune LP for the benefit of Peter Luo and (v) 83,333 ordinary shares held by Great Han Fortune LP for the benefit of Peter Luo that are vested or will be vested within 60 days after this offering; (vi) 45,980 ordinary shares underlying 45,980 share options granted to Xiaohong She that are vested or will be vested within 60 days after this offering, (vii) 10,417 ordinary shares held by Great Han Fortune LP for the benefit of Xiaohong She, and (viii) 59,667 ordinary shares held by Great Han Fortune LP for the benefit of Xiaohong She that are vested or will be vested within 60 days after this offering; (ix) 1,052,188 ordinary shares and share options beneficially owned by Fangyong (Felix) Du, as set forth in note (2) above; (x) total of 827,407 ordinary shares held by Raymond Tam and several key employees of the Company, and (xi) total of 216,408 share options granted to Raymond Tam, Hua Gong, JC Xu and several key employees that are vested or will be vested within 60 days after this offering. On December 14, 2020, Peter Luo, Fangyong (Felix) Du, Ping Ren, Hua Gong, JC Xu, Qinghai Zhao, Man Kin (Raymond) Tam, Xiaohong (Kristine) She, Yan Li, Guizhong Liu and Alexander Goergen entered into a concert party agreement, pursuant to which the parties agree to (i) always be acting in concert in respect of their respective direct or indirect voting rights at our shareholders' general meetings, (ii) recognize the controlling position of Peter Luo; and (iii) act in concert in accordance with Peter Luo's opinions in respect of the daily operations and management and the major decision-making of us.
 - (4) Represents 5,353,242 ordinary shares issuable upon the conversion of 5,353,242 Series B preferred shares held by JSR Limited, a British Virgin Islands company. JSR Limited is controlled by GP Healthcare Capital Co., Ltd. The registered address of JSR Limited is Vistra Corporate Services Centre, Wickhams Cay II, Road Town, Tortola, VG1110, British Virgin Islands.
 - (5) Represents (i) 2,000,000 ordinary shares and (ii) 2,826,037 ordinary shares issuable upon the conversion of 1,589,796 Series A-1 preferred shares, 790,138 Series A-2 preferred shares and 446,103 Series B preferred shares held by Asia Ventures II L.P., a limited partnership incorporated in the Bermuda. The general partner of Asia Ventures II L.P. is Asia Partners II L.P., a Bermuda exempt limited partnership. The general partner of Asia Partners II L.P. is Eight Roads GP, who is ultimately controlled by Eight Roads Holdings Limited. The registered address of Asia Ventures II L.P. is Pembroke Hall, 42 Crow Lane, Pembroke, Bermuda HM 19.
 - (6) Represents (i) 2,000,000 ordinary shares and (ii) 2,826,037 ordinary shares issuable upon the conversion of 1,589,796 Series A-1 preferred shares, 790,138 Series A-2 preferred shares and 446,103 Series B preferred shares held by F-Prime Capital Partners Healthcare Fund III LP. F-Prime Capital Partners Healthcare Advisors Fund III LP is the general partner of F-Prime Capital Partners Healthcare Fund III LP. F-Prime Capital Partners Healthcare Advisors Fund III LP is solely managed by Impresa Management LLC, its general partner and investment manager. Each of the entities listed above expressly disclaims beneficial ownership of the securities listed above except to the extent of any pecuniary interest therein. The address of these entities is 245 Summer Street, Boston, MA 02210.
 - (7) Represents (i) 1,880,909 ordinary shares and (ii) 2,826,037 ordinary shares issuable upon the conversion of 1,589,796 Series A-1 preferred shares, 790,138 Series A-2 preferred shares and 446,103 Series B preferred shares held by Wuxi Pharmatech Healthcare Fund I L.P., a limited partnership incorporated in the Cayman Islands. Wuxi Pharmatech Healthcare Fund I L.P. is an indirect wholly owned subsidiary of WuXi AppTec Co., Ltd (SSE: 603259; SEHK: 2359). WuXi AppTec Co., Ltd. is a listed company on the Shanghai Stock Exchange and the Main Board of the Hong Kong Stock

Exchange. The registered address of WuXi Pharmatech Healthcare Fund I L.P. is P.O. Box 309, Ugland House, Grand Cayman, KY1-1104, Cayman Islands.

- (8) Represents 4,452,441 ordinary shares issuable upon the conversion of 4,452,441 Series C-3 preferred shares held by General Atlantic Singapore AI Pte. Ltd., a company incorporated under the laws of Singapore. General Atlantic Singapore AI Pte. Ltd. is wholly-owned by General Atlantic Singapore Fund Pte. Ltd., which is controlled by General Atlantic Singapore Interholdco Ltd. The registered address of General Atlantic Singapore AI Pte. Ltd. is 80 Robinson Road, #02-00 Singapore 068898.

As of the date of this prospectus, 35.9% of our outstanding ordinary shares or outstanding preferred shares are held by record holders in the United States.

We are not aware of any arrangement that may, at a subsequent date, result in a change of control of our company. See "Description of Share Capital—History of Securities Issuances" for a description of issuances of our ordinary shares and preferred shares that have resulted in significant changes in ownership held by our major shareholders.

RELATED PARTY TRANSACTIONS

The following is a summary of transactions since January 1, 2017 to which we have been a participant in which any of our then directors, executive officers or holders of more than 5% of any class of our voting securities at the time of such transaction, or any members of their immediate family, had or will have a direct or indirect material interest.

Employment Agreements and Indemnification Agreements

See "Management—Employment Agreements and Indemnification Agreements."

Private Placements

See "Description of Share Capital—History of Securities Issuances."

Share Incentives

See "Management—Share Incentive Plan."

Other Related Party Transactions

Transactions with Peter Luo

In May 2014, we received a subsidy of US\$84.6 thousand from the local government for attracting highly skilled personnel on behalf of Peter Luo, our Chairman, CEO and ordinary and preferred shareholder. We settled this transaction with Peter Luo in May 2020.

Transactions with WuXi AppTec Group

We received research and development services from WuXi AppTec Group, the parent company of one of our principal shareholders. The amounts for the purchase of the services were US\$1.0 million and US\$2.1 million in 2018 and 2019, respectively. As of December 31, 2018 and 2019, the amounts due to WuXi AppTec Group were US\$0.2 million and US\$0.4 million, respectively.

The amounts for the purchase of the services were US\$1.3 million and US\$2.3 million in the nine months ended September 30, 2019 and 2020, respectively. As of September 30, 2020, the amount due to WuXi AppTec Group was US\$354.3 thousand.

Transactions with WuXi Biologics (Shanghai) Co., Ltd. or WuXi Biologics

We received research and development services, including provision of manufacturing and quality control testing services, from WuXi Biologics, an entity controlled by the ultimate controlling party of one of our principal shareholders. The amounts for the purchase of the services were US\$6.5 million and US\$3.6 million in 2018 and 2019, respectively. As of December 31, 2018 and 2019, the amounts due to WuXi Biologics were US\$3.4 million and US\$1.4 million, respectively.

The amounts for the purchase of the services were US\$2.8 million and US\$5.1 million in the nine months ended September 30, 2019 and 2020, respectively. As of September 30, 2020, the amount due to WuXi Biologics was US\$2.0 million.

Director and Executive Officers

Exercise of Share Options

In October and November 2017, Dr. Peter (Peizhi) Luo, Xiaohong (Kristine) She and Dr. Fangyong (Felix) Du exercised their respective vested share options under the 2015 Plan. The payment for exercise of such share options is repaid in installments with nil interest. The aggregate

exercise price for Dr. Luo, Ms. She and Dr. Du was US\$365.0 thousand, US\$67.4 thousand and US\$173.4 thousand, respectively. Each of Dr. Luo, Ms. She and Dr. Du has settled the payments for their respective exercise of such share options.

Promissory Notes

The following table sets forth the material terms of the promissory notes issued in connection with the share purchase plans between us and our directors and executive officers:

<u>Name of Director or Executive Officer</u>	<u>Latest Amount Previously Outstanding</u>
Peter (Peizhi) Luo	US\$4.2 million
Xiaohong (Kristine) She*	US\$0.6 million
Fangyong (Felix) Du	US\$1.5 million

Note:

Xiaohong (Kristine) She is the spouse of Dr. Peter (Peizhi) Luo

All promissory notes were made pursuant to our 2019 Plan. All amounts outstanding under the promissory notes have been settled. For further information, see "Management—Share Incentive Plan—Adagene Inc. Second Amended and Restated Share Incentive Plans—Repayment of the Promissory Notes."

DESCRIPTION OF SHARE CAPITAL

We are a Cayman Islands exempted company and our affairs are governed by our memorandum and articles of association, as amended and restated from time to time, and Companies Act (as amended) of the Cayman Islands, which we refer to as the "Companies Act" below, and the common law of the Cayman Islands.

As of the date hereof, our authorized share capital consists of US\$50,000 divided into 500,000,000 shares with a par value of US\$0.0001, including: (i) 472,750,176 ordinary shares of par value US\$0.0001 each, (ii) 7,844,371 series A preferred shares of par value US\$0.0001 each (the "Series A Preferred Shares"), which are further divided into 5,473,957 series A-1 preferred shares of par value of US\$0.0001 each (the "Series A-1 Preferred Shares"), 2,370,414 series A-2 preferred shares of par value of US\$0.0001 each (the "Series A-2 Preferred Shares"), (iii) 7,494,537 series B preferred shares of par value US\$0.0001 each (the "Series B Preferred Shares"), and (iv) 11,910,916 authorized series C preferred shares with par value of US\$0.0001 each (the "Series C Preferred Shares"), which are further divided into 5,597,354 series C-1 preferred shares with par value of US\$0.0001 each (the "Series C-1 Preferred Shares"), 1,861,121 series C-2 preferred shares with par value of US\$0.0001 each (the "Series C-2 Preferred Shares") and 4,452,441 series C-3 preferred shares with par value of US\$0.0001 each (the "Series C-3 Preferred Shares").

As of the date of this prospectus, there are 16,282,451 ordinary shares, excluding 2,262,500 ordinary shares, issued but deemed to be not outstanding as of the date of this prospectus, held by Great Han Fortune LP, 5,473,957 Series A-1 Preferred Shares, 2,370,414 Series A-2 Preferred Shares, 7,494,537 Series B Preferred Shares, 5,597,354 Series C-1 Preferred Shares, 1,861,121 Series C-2 Preferred Shares and 4,452,441 Series C-3 Preferred Shares issued and outstanding. All of our issued and outstanding shares are fully paid. Immediately prior to the completion of this offering, all of our issued and outstanding preferred shares will be converted into ordinary shares on a one-for-one basis ordinary shares.

We plan to adopt an amended and restated memorandum and articles of association, which will become effective and replace the current sixth amended and restated memorandum and articles of association in its entirety immediately prior to completion of this offering. Our authorized share capital immediately prior to completion of the offering will be US\$80,000 divided into 800,000,000 shares comprising (i) 640,000,000 ordinary shares of a par value of US\$0.0001 each, and (ii) 160,000,000 shares of a par value of US\$0.0001 each of such class or classes (however designated) as our board of directors may determine in accordance with our amended and restated memorandum and articles of association. We will issue _____ ordinary shares represented by ADSs in this offering, assuming the underwriters do not exercise their over-allotment option. All awards under the 2019 Plan, regardless of grant dates, will entitle holders to an equivalent number of ordinary shares once the vesting and exercising conditions are met.

The following are summaries of material provisions of our post-offering amended and restated memorandum and articles of association and the Companies Act insofar as they relate to the material terms of our ordinary shares that we expect will become effective immediately prior to the completion of this offering.

Ordinary shares

General. Immediately prior to the completion of this offering, our authorized share capital is US\$80,000 divided into 800,000,000 shares comprising (i) 640,000,000 ordinary shares of a par value of US\$0.0001 each, and (ii) 160,000,000 shares of a par value of US\$0.0001 each of such class or classes (however designated) as our board of directors may determine in accordance with our amended and restated memorandum and articles of association. Holders of ordinary shares will have the same rights. All of our issued and outstanding ordinary shares are fully paid and non-assessable. Certificates

representing the ordinary shares are issued in registered form. We may not issue share to bearer. Our shareholders who are nonresidents of the Cayman Islands may freely hold, transfer and vote their ordinary shares.

Dividends. The holders of our ordinary shares are entitled to such dividends as may be declared by our board of directors subject to our post-offering memorandum and articles of association and the Companies Act. In addition, our shareholders may, subject to the provisions of our articles of association, by ordinary resolution declare a dividend, but no dividend may exceed the amount recommended by our directors. Our post-offering memorandum and articles of association provide that dividends may be declared and paid out of our profits, realized or unrealized, or from any reserve set aside from profits which our board of directors determine is no longer needed. Dividends may also be declared and paid out of the share premium account or any other fund or account which can be authorized for this purpose in accordance with the Companies Act. No dividend may be declared and paid unless our directors determine that, immediately after the payment, we will be able to pay our debts as they become due in the ordinary course of business and we have funds lawfully available for such purpose.

Voting Rights. In respect of all matters subject to a shareholders' vote, holder of an each Ordinary Share is entitled to one vote for each Ordinary Share registered in his or her name on our register of members. Voting at any meeting of shareholders is by way of a poll unless a show of hands is permitted. A poll may be demanded by the chairman of such meeting or any one shareholder.

A quorum required for a meeting of shareholders consists of two or more shareholders holding not less than one-half of the votes attaching to the issued and outstanding shares entitled to vote at general meetings present in person or by proxy or, if a corporation or other non-natural person, by its duly authorized representative. As a Cayman Islands exempted company, we are not obliged by the Companies Act to call shareholders' annual general meetings. Our post-offering memorandum and articles of association provide that we may (but are not obliged to) in each year hold a general meeting as our annual general meeting in which case we will specify the meeting as such in the notices calling it, and the annual general meeting will be held at such time and place as may be determined by our board of directors. We, however, will hold an annual shareholders' meeting during each fiscal year, as required by the Listing Rules of the Nasdaq Global Market. Each general meeting, other than an annual general meeting, shall be an extraordinary general meeting. Shareholders' annual general meetings and any other general meetings of our shareholders may be called by a majority of our board of directors or our chairman or upon a requisition of shareholders holding at the date of deposit of the requisition not less than one-third of the votes attaching to the issued and outstanding shares entitled to vote at general meetings, in which case the directors are obliged to call such meeting and to put the resolutions so requisitioned to a vote at such meeting; however, our post-offering memorandum and articles of association do not provide our shareholders with any right to put any proposals before annual general meetings or extraordinary general meetings not called by such shareholders. Advance notice of at least seven (7) days is required for the convening of our annual general meeting and other general meetings in accordance with our post-offering memorandum and articles of association.

An ordinary resolution to be passed at a meeting by the shareholders requires the affirmative vote of a simple majority of the votes attaching to the ordinary shares cast by those shareholders entitled to vote who are present in person or by proxy at a general meeting, while a special resolution also requires the affirmative vote of no less than two-thirds of the votes attaching to the ordinary shares cast by those shareholders entitled to vote who are present in person or by proxy at a general meeting. A special resolution will be required for important matters such as a change of name or making changes to our post-offering memorandum and articles of association.

Transfer of Ordinary Shares. Subject to the restrictions in our post-offering memorandum and articles of association as set out below, any of our shareholders may transfer all or any of his or her

ordinary shares by an instrument of transfer in the usual or common form or any other form approved by our board of directors.

Our board of directors may, in its absolute discretion, decline to register any transfer of any Ordinary Share. Our board of directors may also decline to register any transfer of any Ordinary Share unless:

- the instrument of transfer is lodged with us, accompanied by the certificate for the ordinary shares to which it relates and such other evidence as our board of directors may reasonably require to show the right of the transferor to make the transfer;
- the instrument of transfer is in respect of only one class of shares;
- the instrument of transfer is properly stamped, if required;
- in the case of a transfer to joint holders, the number of joint holders to whom the Ordinary Share is to be transferred does not exceed four;
- the shares are free from any lien in favor of us; and
- a fee of such maximum sum as the Nasdaq may determine to be payable or such lesser sum as our directors may from time to time require is paid to us in respect thereof.

If our directors refuse to register a transfer, they shall, within three months after the date on which the instrument of transfer was lodged, send to each of the transferor and the transferee notice of such refusal.

The registration of transfers may, after compliance with any notice requirement of the Nasdaq, be suspended and the register closed at such times and for such periods as our board of directors may from time to time determine, *provided, however*, that the registration of transfers shall not be suspended nor the register closed for more than 30 days in any year as our board may determine.

Liquidation. On a return of capital on winding up or otherwise (other than on conversion, redemption or purchase of ordinary shares), if the assets available for distribution amongst our shareholders shall be more than sufficient to repay the whole of the share capital at the commencement of the winding up, the surplus shall be distributed amongst our shareholders in proportion to the par value of the shares held by them at the commencement of the winding up, subject to a deduction from those shares in respect of which there are monies due, of all monies payable to our company for unpaid calls or otherwise. If our assets available for distribution are insufficient to repay all of the paid-up capital, the assets will be distributed so that the losses are borne by our shareholders in proportion to the par value of the shares held by them. Any distribution of assets or capital to a holder of ordinary shares will be the same in any liquidation event.

Calls on Ordinary Shares and Forfeiture of Ordinary Shares. Our board of directors may from time to time make calls upon shareholders for any amounts unpaid on their ordinary shares in a notice served to such shareholders at least 14 clear days prior to the specified time of payment. The ordinary shares that have been called upon and remain unpaid are subject to forfeiture.

Redemption, Repurchase and Surrender of Ordinary Shares. We may issue shares on terms that such shares are subject to redemption, at our option or at the option of the holders thereof, on such terms and in such manner as may be determined, before the issue of such shares, by our board of directors or by an ordinary resolution of our shareholders. Our company may also repurchase any of our shares provided that the manner and terms of such purchase have been approved by our board of directors or by an ordinary resolution of our shareholders, or are otherwise authorized by our post-offering memorandum and articles of association. Under the Companies Act, the redemption or repurchase of any share may be paid out of our company's profits or out of the proceeds of a fresh issue of shares

made for the purpose of such redemption or repurchase, or out of capital (including share premium account and capital redemption reserve) if the Company can, immediately following such payment, pay its debts as they fall due in the ordinary course of business. In addition, under the Companies Act no such share may be redeemed or repurchased (a) unless it is fully paid up, (b) if such redemption or repurchase would result in there being no shares outstanding, or (c) if the company has commenced liquidation. In addition, our company may accept the surrender of any fully paid share for no consideration.

Variations of Rights of Shares. If at any time our share capital is divided into different classes or series of shares, the rights attached to any class or series of shares (unless otherwise provided by the terms of issue of the shares of that class or series), whether or not our company is being wound-up, may be varied with the consent in writing of holders of not less than a majority of the issued shares of that class or series or with the sanction of a special resolution at a separate meeting of the holders of the shares of the class or series. The rights conferred upon the holders of the shares of any class issued shall not, unless otherwise expressly provided by the terms of issue of the shares of that class, be deemed to be varied by the creation or issue of further shares ranking *pari passu* with such existing class of shares.

Inspection of Books and Records. Holders of our ordinary shares have no general right under Cayman Islands law to inspect or obtain copies of our list of shareholders or our corporate records. However, we will provide our shareholders with annual audited financial statements. See "Where You Can Find Additional Information."

Issuance of Additional Shares. Our post-offering memorandum and articles of association authorizes our board of directors to issue additional ordinary shares from time to time as our board of directors shall determine, to the extent of available authorized but unissued shares.

Our post-offering memorandum and articles of association also authorizes our board of directors to establish from time to time one or more series of preferred shares and to determine, with respect to any series of preferred shares, the terms and rights of that series, including:

- the designation of the series;
- the number of shares of the series;
- the dividend rights, dividend rates, conversion rights, voting rights; and
- the rights and terms of redemption and liquidation preferences.

Our board of directors may issue preferred shares without action by our shareholders to the extent authorized but unissued. Issuance of these shares may dilute the voting power of holders of ordinary shares.

Anti-Takeover Provisions. Some provisions of our post-offering memorandum and articles of association may discourage, delay or prevent a change of control of our company or management that shareholders may consider favorable, including provisions that authorize our board of directors to issue preferred shares in one or more series and to designate the price, rights, preferences, privileges and restrictions of such preferred shares without any further vote or action by our shareholders.

Changes in Capital.

We may from time to time by ordinary resolution:

- increase the share capital by such sum, to be divided into shares of such classes and amount, as the resolution shall prescribe;

- consolidate and divide all or any of our share capital into shares of a larger amount than our existing shares;
- sub-divide our existing shares, or any of them into shares of a smaller amount; or
- cancel any shares which, at the date of the passing of the resolution, have not been taken or agreed to be taken by any person and diminish the amount of our share capital by the amount of the shares so canceled.

We may by special resolution, subject to any confirmation or consent required by the Companies Act, reduce our share capital or any capital redemption reserve in any manner permitted by law.

Exempted Company. We are an exempted company with limited liability under the Companies Act. The Companies Act distinguishes between ordinary resident companies and exempted companies. Any company that is registered in the Cayman Islands but conducts business mainly outside of the Cayman Islands may apply to be registered as an exempted company. The requirements for an exempted company are essentially the same as for an ordinary company except that an exempted company:

- does not have to file an annual return of its shareholders with the Registrar of Companies;
- is not required to open its register of members for inspection;
- does not have to hold an annual general meeting;
- may issue shares with no par value;
- may obtain an undertaking against the imposition of any future taxation (such undertakings are usually given for 30 years in the first instance);
- may register by way of continuation in another jurisdiction and be deregistered in the Cayman Islands;
- may register as a limited duration company; and
- may register as a segregated portfolio company.

"Limited liability" means that the liability of each shareholder is limited to the amount unpaid by the shareholder on that shareholder's shares of the company.

Register of Members

Under the Companies Act, we must keep a register of members and there should be entered therein:

- the names and addresses of our members, a statement of the shares held by each member, and of the amount paid or agreed to be considered as paid, on the shares of each member;
- the date on which the name of any person was entered on the register as a member; and
- the date on which any person ceased to be a member.

Under the Companies Act, the register of members of our company is prima facie evidence of the matters set out therein (that is, the register of members will raise a presumption of fact on the matters referred to above unless rebutted) and a member registered in the register of members is deemed as a matter of the Companies Act to have legal title to the shares as set against its name in the register of members. Upon completion of this offering, we will perform the procedure necessary to immediately update the register of members to record and give effect to the issuance of shares by us to the Depositary (or its nominee) as the depositary. Once our register of members has been updated, the shareholders recorded in the register of members will be deemed to have legal title to the shares set against their name.

If the name of any person is incorrectly entered in or omitted from our register of members, or if there is any default or unnecessary delay in entering on the register the fact of any person having ceased to be a member of our company, the person or member aggrieved (or any member of our company or our company itself) may apply to the Grand Court of the Cayman Islands for an order that the register be rectified, and the Court may either refuse such application or it may, if satisfied of the justice of the case, make an order for the rectification of the register.

Differences in Corporate Law

The Companies Act is derived, to a large extent, from the older Companies Acts of England, but does not follow many recent English law statutory enactments. In addition, the Companies Act differs from laws applicable to United States corporations and their shareholders. Set forth below is a summary of the significant differences between the provisions of the Companies Law applicable to us and the laws applicable to companies incorporated in the State of Delaware.

Mergers and Similar Arrangements. The Companies Act permits mergers and consolidations between Cayman Islands companies and between Cayman Islands companies and non-Cayman Islands companies. For these purposes, (a) "merger" means the merging of two or more constituent companies and the vesting of their undertaking, property and liabilities in one of such companies as the surviving company, and (b) a "consolidation" means the combination of two or more constituent companies into a consolidated company and the vesting of the undertaking, property and liabilities of such companies to the consolidated company. In order to effect such a merger or consolidation, the directors of each constituent company must approve a written plan of merger or consolidation, which must then be authorized by (a) a special resolution of the shareholders of each constituent company, and (b) such other authorization, if any, as may be specified in such constituent company's articles of association. The written plan of merger or consolidation must be filed with the Registrar of Companies of the Cayman Islands together with a declaration as to the solvency of the consolidated or surviving company, a declaration as to the assets and liabilities of each constituent company and an undertaking that a copy of the certificate of merger or consolidation will be given to the members and creditors of each constituent company and that notification of the merger or consolidation will be published in the Cayman Islands Gazette. Court approval is not required for a merger or consolidation which is effected in compliance with these statutory procedures.

A merger between a Cayman parent company and its Cayman subsidiary or subsidiaries does not require authorization by a resolution of shareholders of that Cayman subsidiary if a copy of the plan of merger is given to every member of that Cayman subsidiary to be merged unless that member agrees otherwise. For this purpose, a company is a "parent" of a subsidiary if it holds issued shares that together represent at least ninety percent (90%) of the votes at a general meeting of the subsidiary.

The consent of each holder of a fixed or floating security interest over a constituent company is required unless this requirement is waived by a court in the Cayman Islands.

Save in certain limited circumstances, a shareholder of a Cayman constituent company who dissents from the merger or consolidation is entitled to payment of the fair value of his shares (which, if not agreed between the parties, will be determined by the Cayman Islands court) upon dissenting to the merger or consolidation, *provided* that the dissenting shareholder complies strictly with the procedures set out in the Companies Act. The exercise of dissenter rights will preclude the exercise by the dissenting shareholder of any other rights to which he or she might otherwise be entitled by virtue of holding shares, save for the right to seek relief on the grounds that the merger or consolidation is void or unlawful.

Separate from the statutory provisions relating to mergers and consolidations, the Companies Act also contains statutory provisions that facilitate the reconstruction and amalgamation of companies by way of schemes of arrangement, *provided* that the arrangement is approved by a majority in number of

each class of shareholders and creditors with whom the arrangement is to be made, and who must in addition represent three- fourths in value of each such class of shareholders or creditors, as the case may be, that are present and voting either in person or by proxy at a meeting, or meetings, convened for that purpose. The convening of the meetings and subsequently the arrangement must be sanctioned by the Grand Court of the Cayman Islands. While a dissenting shareholder has the right to express to the court the view that the transaction ought not to be approved, the court can be expected to approve the arrangement if it determines that:

- the statutory provisions as to the required majority vote have been met;
- the shareholders have been fairly represented at the meeting in question and the statutory majority are acting bona fide without coercion of the minority to promote interests adverse to those of the class;
- the arrangement is such that may be reasonably approved by an intelligent and honest man of that class acting in respect of his interest; and
- the arrangement is not one that would more properly be sanctioned under some other provision of the Companies Act.

The Companies Act also contains a statutory power of compulsory acquisition which may facilitate the "squeeze out" of a dissenting minority shareholder upon a tender offer. When a tender offer is made and accepted by holders of 90.0% of the shares affected within four months, the offeror may, within a two-month period commencing on the expiration of such four-month period, require the holders of the remaining shares to transfer such shares to the offeror on the terms of the offer. An objection can be made to the Grand Court of the Cayman Islands but this is unlikely to succeed in the case of an offer which has been so approved unless there is evidence of fraud, bad faith or collusion.

If an arrangement and reconstruction is thus approved, or if a tender offer is made and accepted, a dissenting shareholder would have no rights comparable to appraisal rights, which would otherwise ordinarily be available to dissenting shareholders of Delaware corporations, providing rights to receive payment in cash for the judicially determined value of the shares.

Shareholders' Suits. In principle, we will normally be the proper plaintiff to sue for a wrong done to us as a company, and as a general rule a derivative action may not be brought by a minority shareholder. However, based on English authorities, which would in all likelihood be of persuasive authority in the Cayman Islands, the Cayman Islands court can be expected to follow and apply the common law principles (namely the rule in *Foss v. Harbottle* and the exceptions thereto) which permit a minority shareholder to commence a class action against or derivative actions in the name of the company to challenge actions where:

- a company acts or proposes to act illegally or ultra vires;
- the act complained of, although not ultra vires, could only be effected duly if authorized by more than a simple majority vote that has not been obtained; and
- those who control the company are perpetrating a "fraud on the minority."

Indemnification of Directors and Executive Officers and Limitation of Liability. Cayman Islands law does not limit the extent to which a company's memorandum and articles of association may provide for indemnification of officers and directors, except to the extent any such provision may be held by the Cayman Islands courts to be contrary to public policy, such as to provide indemnification against civil fraud or the consequences of committing a crime. Our post-offering memorandum and articles of association provide that that we shall indemnify our officers and directors against all actions, proceedings, costs, charges, expenses, losses, damages or liabilities incurred or sustained by such directors or officer, other than by reason of such person's dishonesty, willful default or fraud, in or

about the conduct of our company's business or affairs (including as a result of any mistake of judgment) or in the execution or discharge of his duties, powers, authorities or discretions, including without prejudice to the generality of the foregoing, any costs, expenses, losses or liabilities incurred by such director or officer in defending (whether successfully or otherwise) any civil proceedings concerning our company or its affairs in any court whether in the Cayman Islands or elsewhere. This standard of conduct is generally the same as permitted under the Delaware General Corporation Law for a Delaware corporation.

In addition, we have entered into indemnification agreements with our directors and executive officers that provide such persons with additional indemnification beyond that provided in our post-offering memorandum and articles of association.

Insofar as indemnification for liabilities arising under the Securities Act may be permitted to our directors, officers or persons controlling us under the foregoing provisions, we have been informed that in the opinion of the SEC, such indemnification is against public policy as expressed in the Securities Act and is therefore unenforceable.

Directors' Fiduciary Duties. Under Delaware corporate law, a director of a Delaware corporation has a fiduciary duty to the corporation and its shareholders. This duty has two components: the duty of care and the duty of loyalty. The duty of care requires that a director act in good faith, with the care that an ordinarily prudent person would exercise under similar circumstances. Under this duty, a director must inform himself of, and disclose to shareholders, all material information reasonably available regarding a significant transaction. The duty of loyalty requires that a director acts in a manner he reasonably believes to be in the best interests of the corporation. He must not use his corporate position for personal gain or advantage. This duty prohibits self-dealing by a director and mandates that the best interest of the corporation and its shareholders take precedence over any interest possessed by a director, officer or controlling shareholder and not shared by the shareholders generally. In general, actions of a director are presumed to have been made on an informed basis, in good faith and in the honest belief that the action taken was in the best interests of the corporation. However, this presumption may be rebutted by evidence of a breach of one of the fiduciary duties. Should such evidence be presented concerning a transaction by a director, the director must prove the procedural fairness of the transaction, and that the transaction was of fair value to the corporation.

As a matter of Cayman Islands law, a director of a Cayman Islands company is in the position of a fiduciary with respect to the company and therefore it is considered that he owes the following duties to the company—a duty to act bona fide in the best interests of the company, a duty not to make a profit based on his position as director (unless the company permits him to do so), a duty not to put himself in a position where the interests of the company conflict with his personal interest or his duty to a third party, and a duty to exercise powers for the purpose for which such powers were intended. A director of a Cayman Islands company owes to the company a duty to act with skill and care. It was previously considered that a director need not exhibit in the performance of his duties a greater degree of skill than may reasonably be expected from a person of his knowledge and experience. However, English and Commonwealth courts have moved towards an objective standard with regard to the required skill and care and these authorities are likely to be followed in the Cayman Islands.

Under our post-offering memorandum and articles of association, directors who are in any way, whether directly or indirectly, interested in a contract or proposed contract with our company must declare the nature of their interest at a meeting of the board of directors.

Shareholder Action by Written Consent. Under the Delaware General Corporation Law, a corporation may eliminate the right of shareholders to act by written consent by amendment to its certificate of incorporation. The Companies Act and our post-offering memorandum and articles of association provide that our shareholders may approve corporate matters by way of a unanimous

written resolution signed by or on behalf of each shareholder who would have been entitled to vote on such matter at a general meeting without a meeting being held.

Shareholder Proposals. Under the Delaware General Corporation Law, a shareholder has the right to put any proposal before the annual meeting of shareholders, provided it complies with the notice provisions in the governing documents. A special meeting may be called by the board of directors or any other person authorized to do so in the governing documents, but shareholders may be precluded from calling special meetings.

The Companies Act provides shareholders with only limited rights to requisition a general meeting, and does not provide shareholders with any right to put any proposal before a general meeting. However, these rights may be provided in a company's articles of association. Our post-offering memorandum and articles of association allow our shareholders holding in aggregate not less than twenty percent of all votes attaching to the issued and outstanding shares of our company entitled to vote at general meetings to requisition an extraordinary general meeting of our shareholders, in which case our board is obliged to convene an extraordinary general meeting and to put the resolutions so requisitioned to a vote at such meeting. Other than this right to requisition a shareholders' meeting, our post-offering memorandum and articles of association do not provide our shareholders with any other right to put proposals before annual general meetings or extraordinary general meetings not called by such shareholders. As an exempted Cayman Islands company, we are not obliged by law to call shareholders' annual general meetings.

Cumulative Voting. Under the Delaware General Corporation Law, cumulative voting for elections of directors is not permitted unless the corporation's certificate of incorporation specifically provides for it. Cumulative voting potentially facilitates the representation of minority shareholders on a board of directors since it permits the minority shareholder to cast all the votes to which the shareholder is entitled on a single director, which increases the shareholder's voting power with respect to electing such director. There are no prohibitions in relation to cumulative voting under the laws of the Cayman Islands but our post-offering memorandum and articles of association do not provide for cumulative voting. As a result, our shareholders are not afforded any less protections or rights on this issue than shareholders of a Delaware corporation.

Removal of Directors. Under the Delaware General Corporation Law, a director of a corporation with a classified board may be removed only for cause with the approval of a majority of the outstanding shares entitled to vote, unless the certificate of incorporation provides otherwise. Under our post-offering memorandum and articles of association, directors not appointed by Peter Peizhi Luo, JSR Limited, and General Atlantic Singapore AI Pte. Ltd. may be removed with or without cause, by an ordinary resolution of our shareholders. A director shall hold office until the expiration of his or her term or his or her successor shall have been elected and qualified, or until his or her office is otherwise vacated. In addition, a director's office shall be vacated if the director (i) becomes bankrupt or makes any arrangement or composition with his creditors; (ii) is found to be or becomes of unsound mind or dies; (iii) resigns his office by notice in writing to the company; (iv) without special leave of absence from our board of directors, is absent from three consecutive meetings of the board and the board resolves that his office be vacated; (v) is prohibited by law from being a director; or (vi) is removed from office pursuant to any other provisions of our post-offering memorandum and articles of association.

Transactions with Interested Shareholders. The Delaware General Corporation Law contains a business combination statute applicable to Delaware corporations whereby, unless the corporation has specifically elected not to be governed by such statute by amendment to its certificate of incorporation, it is prohibited from engaging in certain business combinations with an "interested shareholder" for three years following the date that such person becomes an interested shareholder. An interested shareholder generally is a person or a group who or which owns or owned 15% or more of the target's

outstanding voting share within the past three years. This has the effect of limiting the ability of a potential acquirer to make a two-tiered bid for the target in which all shareholders would not be treated equally. The statute does not apply if, among other things, prior to the date on which such shareholder becomes an interested shareholder, the board of directors approves either the business combination or the transaction which resulted in the person becoming an interested shareholder. This encourages any potential acquirer of a Delaware corporation to negotiate the terms of any acquisition transaction with the target's board of directors.

Cayman Islands law has no comparable statute. As a result, we cannot avail ourselves of the types of protections afforded by the Delaware business combination statute. However, although Cayman Islands law does not regulate transactions between a company and its significant shareholders, the directors of the Company are required to comply with fiduciary duties which they owe to the Company under Cayman Islands laws, including the duty to ensure that, in their opinion, any such transactions must be entered into bona fide in the best interests of the company, and are entered into for a proper corporate purpose and not with the effect of constituting a fraud on the minority shareholders.

Dissolution; Winding up. Under the Delaware General Corporation Law, unless the board of directors approves the proposal to dissolve, dissolution must be approved by shareholders holding 100% of the total voting power of the corporation. Only if the dissolution is initiated by the board of directors may it be approved by a simple majority of the corporation's outstanding shares. Delaware law allows a Delaware corporation to include in its certificate of incorporation a supermajority voting requirement in connection with dissolutions initiated by the board.

Under Cayman Islands law, a company may be wound up by either an order of the courts of the Cayman Islands or by a special resolution of its members or, if the company is unable to pay its debts as they fall due, by an ordinary resolution of its members. The court has authority to order winding up in a number of specified circumstances including where it is, in the opinion of the court, just and equitable to do so.

Variation of Rights of Shares. Under the Delaware General Corporation Law, a corporation may vary the rights of a class of shares with the approval of a majority of the outstanding shares of such class, unless the certificate of incorporation provides otherwise. Under Cayman Islands law and our post-offering memorandum and articles of association, if our share capital is divided into more than one class of shares, we may vary the rights attached to any class with the written consent of the holders of not less than a majority of the issued shares of that class or with the sanction of a special resolution passed at a general meeting of the holders of the shares of that class.

Amendment of Governing Documents. Under the Delaware General Corporation Law, a corporation's governing documents may be amended with the approval of a majority of the outstanding shares entitled to vote, unless the certificate of incorporation provides otherwise. Under the Companies Act and our post-offering memorandum and articles of association, our memorandum and articles of association may only be amended by a special resolution of our shareholders.

Rights of Nonresident or Foreign Shareholders. There are no limitations imposed by our post-offering memorandum and articles of association on the rights of nonresident or foreign shareholders to hold or exercise voting rights on our shares. In addition, there are no provisions in our post-offering amended and restated memorandum and articles of association governing the ownership threshold above which shareholder ownership must be disclosed.

History of Securities Issuances

The following is a summary of securities issuances by Adagene Inc. in the past three years.

Ordinary Shares

From August 17, 2017 to January 19, 2021, we issued a total of 6,761,534 ordinary shares to employees for an aggregate consideration of US\$7.9 million upon the exercise of certain share incentive awards. For the purpose of this paragraph only, such number of shares includes the 2,262,500 ordinary shares, issued but deemed to be not outstanding as of the date of this prospectus, held by Great Han Fortune LP.

Preferred Shares

On February 2, 2018, we issued a total of 3,358,412 Series C-1 Preferred Shares to SCC Venture VI Holdco, Ltd., Gopher Harvest Co-Investment Fund LP and AVICT Global Holdings Limited for an aggregate consideration of US\$30.0 million.

On March 19, 2018, we issued a total of 1,679,206 Series C-1 Preferred Shares to King Star Med LP and WEALTHY TECHNOLOGIES LIMITED for an aggregate consideration of US\$15.0 million.

On May 16, 2018, we issued 559,736 Series C-1 Preferred Shares to Chief Strategic International Limited for a consideration of US\$5.0 million.

On June 13, 2019, we issued a total of 1,567,260 Series C-2 Preferred Shares to Mega Prime Development Limited, Poly Platinum Enterprises Limited and Chief Strategic International Limited for an aggregate consideration of US\$16.0 million.

On November 21, 2019, we issued 293,861 Series C-2 Preferred Shares to MODEST CHAMPION LIMITED for a consideration of US\$3.0 million.

On December 19, 2019, we issued 4,452,441 Series C-3 Preferred Shares to General Atlantic Singapore AI Pte. Ltd. for a consideration of US\$50.0 million.

Warrant

On February 2, 2018, we granted warrants to SCC Venture VI Holdco, Ltd. and Gopher Harvest Co-Investment Fund LP to purchase up to a total of US\$7.5 million worth of Series C-2 Preferred Shares at the exercise price of US\$10.2089 per share (as may be adjusted from time to time). The warrants were not exercised and have expired.

Award Grants

We have granted awards to purchase our ordinary shares to certain of our directors, executive officers and employees pursuant to the 2015 Plan and the 2019 Plan. As of the date of this prospectus, the aggregate number of our ordinary shares underlying our outstanding awards under the 2019 Plan is 3,601,522. We have not granted any awards under the 2021 Plan. See "Management—Share Incentive Plan."

Share Cancellation

On January 16, 2021, certain management members surrendered a total of 491,119 ordinary shares for cancellation to settle the balance due under their respective promissory notes issued in connection with the share purchase plans.

Shareholders' Agreements

We entered into a Fifth Amended and Restated Shareholders Agreement and a Fourth Amended and Restated Right of First Refusal and Co-Sale Agreement (collectively, "Shareholders Agreements")

on December 19, 2019 with our shareholders, which consist of holders of our ordinary shares, Series A-1 Preferred Shares, Series A-2 Preferred Shares, Series B Preferred Shares, Series C-1 Preferred Shares, Series C-2 Preferred Shares and Series C-3 Preferred Shares.

The Shareholders Agreements provide for certain special rights, including information and inspection rights, right of participation, right of first refusal, co-sale right, drag-along right, redemption, liquidation and protective provisions. Except for board representation right and registration right, all preferred shareholders' rights will automatically terminate upon the completion of this offering.

Board Representation

Each of Asia Ventures II L.P., F-Prime Capital Partners Healthcare Fund III LP and JSR Limited shall have the right to designate, appoint, remove and replace and reappoint one director so long as they each hold at least five percent of the shares outstanding on a fully-diluted basis and an as-converted basis, respectively. Notwithstanding of the foregoing, each of Asia Ventures II L.P. and F-Prime Capital Partners Healthcare Fund III LP will cease to have such board representation right upon the effectiveness of our post-offering memorandum and articles of association.

As long as Wuxi Pharmatech Healthcare Fund I L.P. holds at least five percent of the shares outstanding on a fully-diluted basis, it shall have the right to nominate one independent non-executive director and such one independent non-executive director shall be appointed and agreed by the board.

As long as Peter Luo holds any shares or is employed by us or any of our controlled affiliates, he shall have the right to designate, appoint, remove and replace and reappoint one director. Notwithstanding of the foregoing, our post-offering memorandum and articles of association will provide that for as long as Pete Luo holds or beneficially owns any shares or is employed by us or any of our subsidiaries, he will serve as one of our directors and the Chairman of the Board of Directors. In addition, during the period commencing upon the effectiveness of our registration statement on Form F-1, of which this prospectus is a part, and ending on the earlier of (i) the date upon which Peter Luo beneficially owns less than five percent of the shares outstanding on a fully diluted basis, (ii) the death or legal determination of Peter Luo's incapacity or (iii) the termination of Peter Luo as an executive officer or principal scientific advisor to us or any of our subsidiaries or for cause (as determined under his related employment or consulting arrangements and subject to all related cure provisions), Peter Luo shall have the right to designate, appoint, remove and replace and reappoint one additional director.

As long as General Atlantic Singapore AI Pte. Ltd. and its affiliates hold at least five percent of the shares outstanding on a fully-diluted basis, they shall have the right to designate, appoint, remove and replace and reappoint one director.

Registration Rights

Pursuant to our Fifth Amended and Restated Shareholders Agreement dated December 19, 2019, we have granted certain registration rights to our shareholders. Set forth below is a description of the registration rights granted under the agreement.

Demand Registration Rights. At any time or from time to time after the date that is six months after the closing of the IPO, holders holding thirty percent or more of the voting power of the then outstanding registrable securities held by all holders may request in writing that we effect a registration on any internationally recognized exchange that is reasonably acceptable to such requesting holders. Upon receipt of such a request, we shall (x) promptly give written notice of the proposed registration to all other holders and (y) as soon as practicable, use its good faith commercially reasonable efforts to cause the registrable securities specified in the request, together with any registrable securities of any holder who requests in writing to join such registration within fifteen (15) days after our delivery of

written notice, to be registered and/or qualified for sale and distribution in such jurisdiction as the initiating holders may request. We shall be obligated to consummate no more than two registrations that have been declared and ordered effective; provided that if the registrable securities sought to be included in the registration are not fully included in the registration for any reason other than solely due to the action or inaction of the holders including registrable securities in such registration, such registration shall not be deemed to constitute one of the registration rights granted. We shall not be obligated to take any action to effect any registration unless the aggregate proceeds from the offering that is the subject of the registration exceeds US\$5,000,000 and at least 40% of the registrable securities then outstanding shall participate in such registration. Furthermore, we have the right to defer filing of a registration statement if, after receiving a request from holders, we furnish to the holders a certificate signed by our chief executive officer stating that, in the good faith judgment of the board, it would be detrimental to us or our members for a registration statement to be filed in the near future, but we may not (i) utilize this right for more than ninety days on any one occasion or more than once during any twelve-month period, or (ii) register any other of our securities during such period.

Registration on Form F-3 or Form S-3. We shall use our good faith commercially reasonable efforts to qualify for registration on Form F-3 or Form S-3. If we qualify for registration on Form F-3 or Form S-3 (or any comparable form for registration in a jurisdiction other than the United States), holders holding ten percent or more of the voting power of the then outstanding registrable securities held by all holders may request us in writing to file, in any jurisdiction in which we have had a registered underwritten public offering, a registration statement on Form F-3 or Form S-3 (or any comparable form for registration in a jurisdiction other than the United States). We shall be obligated to consummate no more than two registrations that have been declared and ordered effective within any twelve-month period; provided that if the registrable securities sought to be included in the registration are not fully included in such registration for any reason other than solely due to the action or inaction of the holders including registrable securities in such registration, such registration shall not be deemed to constitute one of the registration rights granted. We shall not be obligated to take any action to effect any registration unless the aggregate proceeds from the offering that is the subject of the registration exceeds US\$5,000,000. Furthermore, we have the right to defer filing of a registration statement if, after receiving a request from holders, we furnish to the holders a certificate signed by our chief executive officer stating that, in the good faith judgment of the board, it would be detrimental to us or our members for a registration statement to be filed in the near future, but we may not (i) utilize this right for more than ninety days on any one occasion or more than once during any twelve-month period, or (ii) register any other of our securities during such period.

Piggyback Registration Rights. If we propose to register for our own account any of our equity securities, or for the account of any holder (other than a holder of registrable securities who is a party to the shareholder agreement) of equity securities any of such holder's equity securities, in connection with the public offering of such securities, we shall promptly give each holder written notice of such registration and, upon the written request of any holder given within fifteen days after delivery of such notice, we shall use our good faith commercially reasonable efforts to include in such registration any registrable securities thereby requested to be registered by such holder. If a holder decides not to include all or any of its registrable securities in such registration by us, such holder shall nevertheless continue to have the right to include any registrable securities in any subsequent registration statement or registration statements as may be filed by us.

Expenses of Registration. We will bear all registration expenses. Each holder, however, should bear its proportionate share of all of the underwriting discounts and selling commissions applicable to the sale of registrable securities.

DESCRIPTION OF AMERICAN DEPOSITARY SHARES

American Depositary Receipts

JPMorgan Chase Bank, N.A. ("JPMorgan"), as depositary, will issue the ADSs which you will be entitled to receive in this offering. Each ADS will represent an ownership interest in a designated number of shares which we will deposit with the custodian, as agent of the depositary, under the deposit agreement among ourselves, the depositary, yourself as an ADR holder and all other ADR holders, and all beneficial owners of an interest in the ADSs evidenced by ADRs from time to time.

The depositary's office is located at 383 Madison Avenue, Floor 11, New York, NY 10179.

The ADS to share ratio is subject to amendment as provided in the form of ADR (which may give rise to fees contemplated by the form of ADR). In the future, each ADS will also represent any securities, cash or other property deposited with the depositary but which they have not distributed directly to you.

A beneficial owner is any person or entity having a beneficial ownership interest ADSs. A beneficial owner need not be the holder of the ADR evidencing such ADS. If a beneficial owner of ADSs is not an ADR holder, it must rely on the holder of the ADR(s) evidencing such ADSs in order to assert any rights or receive any benefits under the deposit agreement. A beneficial owner shall only be able to exercise any right or receive any benefit under the deposit agreement solely through the holder of the ADR(s) evidencing the ADSs owned by such beneficial owner. The arrangements between a beneficial owner of ADSs and the holder of the corresponding ADRs may affect the beneficial owner's ability to exercise any rights it may have.

An ADR holder shall be deemed to have all requisite authority to act on behalf of any and all beneficial owners of the ADSs evidenced by the ADRs registered in such ADR holder's name for all purposes under the deposit agreement and ADRs. The depositary's only notification obligations under the deposit agreement and the ADRs is to registered ADR holders. Notice to an ADR holder shall be deemed, for all purposes of the deposit agreement and the ADRs, to constitute notice to any and all beneficial owners of the ADSs evidenced by such ADR holder's ADRs.

Unless certificated ADRs are specifically requested, all ADSs will be issued on the books of our depositary in book-entry form and periodic statements will be mailed to you which reflect your ownership interest in such ADSs. In our description, references to American depositary receipts or ADRs shall include the statements you will receive which reflect your ownership of ADSs.

You may hold ADSs either directly or indirectly through your broker or other financial institution. If you hold ADSs directly, by having an ADS registered in your name on the books of the depositary, you are an ADR holder. This description assumes you hold your ADSs directly. If you hold the ADSs through your broker or financial institution nominee, you must rely on the procedures of such broker or financial institution to assert the rights of an ADR holder described in this section. You should consult with your broker or financial institution to find out what those procedures are.

As an ADR holder or beneficial owner, we will not treat you as a shareholder of ours and you will not have any shareholder rights. Cayman Island law governs shareholder rights. Because the depositary or its nominee will be the shareholder of record for the shares represented by all outstanding ADSs, shareholder rights rest with such record holder. Your rights are those of an ADR holder or of a beneficial owner. Such rights derive from the terms of the deposit agreement to be entered into among us, the depositary and all holders and beneficial owners from time to time of ADRs issued under the deposit agreement and, in the case of a beneficial owner, from the arrangements between the beneficial owner and the holder of the corresponding ADRs. The obligations of the depositary and its agents are also set out in the deposit agreement. Because the depositary or its nominee will actually be the

registered owner of the shares, you must rely on it to exercise the rights of a shareholder on your behalf.

The following is a summary of what we believe to be the material terms of the deposit agreement. Notwithstanding this, because it is a summary, it may not contain all the information that you may otherwise deem important. For more complete information, you should read the entire deposit agreement and the form of ADR which contains the terms of your ADSs. You can read a copy of the deposit agreement which is filed as an exhibit to the registration statement of which this prospectus forms a part. You may also obtain a copy of the deposit agreement at the SEC's Public Reference Room which is located at 100 F Street, NE, Washington, DC 20549. You may obtain information on the operation of the Public Reference Room by calling the SEC at 1-800-732-0330. You may also find the registration statement and the attached deposit agreement on the SEC's website at <http://www.sec.gov>.

Share Dividends and Other Distributions

How will I receive dividends and other distributions on the shares underlying my ADSs?

We may make various types of distributions with respect to our securities. The depositary has agreed that, to the extent practicable, it will pay to you the cash dividends or other distributions it or the custodian receives on shares or other deposited securities, after converting any cash received into U.S. dollars (if it determines such conversion may be made on a reasonable basis) and, in all cases, making any necessary deductions provided for in the deposit agreement. The depositary may utilize a division, branch or affiliate of JPMorgan to direct, manage and/or execute any public and/or private sale of securities under the deposit agreement. Such division, branch and/or affiliate may charge the depositary a fee in connection with such sales, which fee is considered an expense of the depositary. You will receive these distributions in proportion to the number of underlying securities that your ADSs represent.

Except as stated below, the depositary will deliver such distributions to ADR holders in proportion to their interests in the following manner:

- *Cash.* The depositary will distribute any U.S. dollars available to it resulting from a cash dividend or other cash distribution or the net proceeds of sales of any other distribution or portion thereof (to the extent applicable), on an averaged or other practicable basis, subject to (i) appropriate adjustments for taxes withheld, (ii) such distribution being impermissible or impracticable with respect to certain registered ADR holders, and (iii) deduction of the depositary's and/or its agents' expenses in (1) converting any foreign currency to U.S. dollars to the extent that it determines that such conversion may be made on a reasonable basis, (2) transferring foreign currency or U.S. dollars to the United States by such means as the depositary may determine to the extent that it determines that such transfer may be made on a reasonable basis, (3) obtaining any approval or license of any governmental authority required for such conversion or transfer, which is obtainable at a reasonable cost and within a reasonable time and (4) making any sale by public or private means in any commercially reasonable manner. *If exchange rates fluctuate during a time when the depositary cannot convert a foreign currency, you may lose some or all of the value of the distribution.*
- *Shares.* In the case of a distribution in shares, the depositary will issue additional ADRs to evidence the number of ADSs representing such shares. Only whole ADSs will be issued. Any shares which would result in fractional ADSs will be sold and the net proceeds will be distributed in the same manner as cash to the ADR holders entitled thereto.
- *Rights to receive additional shares.* In the case of a distribution of rights to subscribe for additional shares or other rights, if we timely provide evidence satisfactory to the depositary that

it may lawfully distribute such rights, the depositary will distribute warrants or other instruments in the discretion of the depositary representing such rights. However, if we do not timely furnish such evidence, the depositary may:

- (i) sell such rights if practicable and distribute the net proceeds in the same manner as cash to the ADR holders entitled thereto; or
 - (ii) if it is not practicable to sell such rights by reason of the non-transferability of the rights, limited markets therefor, their short duration or otherwise, do nothing and allow such rights to lapse, in which case ADR holders will receive nothing and the rights may lapse.
- *Other Distributions.* In the case of a distribution of securities or property other than those described above, the depositary may either (i) distribute such securities or property in any manner it deems equitable and practicable or (ii) to the extent the depositary deems distribution of such securities or property not to be equitable and practicable, sell such securities or property and distribute any net proceeds in the same way it distributes cash.

If the depositary determines in its discretion that any distribution described above is not practicable with respect to any specific registered ADR holder, the depositary may choose any method of distribution that it deems practicable for such ADR holder, including the distribution of foreign currency, securities or property, or it may retain such items, without paying interest on or investing them, on behalf of the ADR holder as deposited securities, in which case the ADSs will also represent the retained items.

Any U.S. dollars will be distributed by checks drawn on a bank in the United States for whole dollars and cents. Fractional cents will be withheld without liability and dealt with by the depositary in accordance with its then current practices.

The depositary is not responsible if it fails to determine that any distribution or action is lawful or reasonably practicable.

There can be no assurance that the depositary will be able to convert any currency at a specified exchange rate or sell any property, rights, shares or other securities at a specified price, nor that any of such transactions can be completed within a specified time period. All purchases and sales of securities will be handled by the depositary in accordance with its then current policies, which are currently set forth on the "Disclosures" page (or successor page) of www.adr.com (as updated by the depositary from time to time, "ADR.com").

Deposit, Withdrawal and Cancellation

How does the depositary issue ADSs?

The depositary will issue ADSs if you or your broker deposit shares or evidence of rights to receive shares with the custodian and pay the fees and expenses owing to the depositary in connection with such issuance. In the case of the ADSs to be issued under this prospectus, we will arrange with the underwriters named herein to deposit such shares.

Shares deposited in the future with the custodian must be accompanied by certain delivery documentation and shall, at the time of such deposit, be registered in the name of JPMorgan Chase Bank, N.A., as depositary for the benefit of holders of ADRs or in such other name as the depositary shall direct.

The custodian will hold all deposited shares (including those being deposited by or on our behalf in connection with the offering to which this prospectus relates) for the account and to the order of the depositary, in each case for the benefit of ADR holders. ADR holders and beneficial owners thus have no direct ownership interest in the shares and only have such rights as are contained in the deposit

agreement. The custodian will also hold any additional securities, property and cash received on or in substitution for the deposited shares. The deposited shares and any such additional items are referred to as "deposited securities".

Deposited securities are not intended to, and shall not, constitute proprietary assets of the depositary, the custodian or their nominees. Beneficial ownership in deposited securities is intended to be, and shall at all times during the term of the deposit agreement continue to be, vested in the beneficial owners of the ADSs representing such deposited securities. Notwithstanding anything else contained herein, in the deposit agreement, in the form of ADR and/or in any outstanding ADSs, the depositary, the custodian and their respective nominees are intended to be, and shall at all times during the term of the deposit agreement be, the record holder(s) only of the deposited securities represented by the ADSs for the benefit of the ADR holders. The depositary, on its own behalf and on behalf of the custodian and their respective nominees, disclaims any beneficial ownership interest in the deposited securities held on behalf of the ADR holders.

Upon each deposit of shares, receipt of related delivery documentation and compliance with the other provisions of the deposit agreement, including the payment of the fees and charges of the depositary and any taxes or other fees or charges owing, the depositary will issue an ADR or ADRs in the name or upon the order of the person entitled thereto evidencing the number of ADSs to which such person is entitled. All of the ADSs issued will, unless specifically requested to the contrary, be part of the depositary's direct registration system, and a registered holder will receive periodic statements from the depositary which will show the number of ADSs registered in such holder's name. An ADR holder can request that the ADSs not be held through the depositary's direct registration system and that a certificated ADR be issued.

How do ADR holders cancel an ADS and obtain deposited securities?

When you turn in your ADR certificate at the depositary's office, or when you provide proper instructions and documentation in the case of direct registration ADSs, the depositary will, upon payment of certain applicable fees, charges and taxes, deliver the underlying shares to you or upon your written order. Delivery of deposited securities in certificated form will be made at the custodian's office. At your risk, expense and request, the depositary may deliver deposited securities at such other place as you may request.

The depositary may only restrict the withdrawal of deposited securities in connection with:

- temporary delays caused by closing our transfer books or those of the depositary or the deposit of shares in connection with voting at a shareholders' meeting, or the payment of dividends;
- the payment of fees, taxes and similar charges; or
- compliance with any U.S. or foreign laws or governmental regulations relating to the ADRs or to the withdrawal of deposited securities.

This right of withdrawal may not be limited by any other provision of the deposit agreement.

Record Dates

The depositary may, after consultation with us if practicable, fix record dates (which, to the extent applicable, shall be as near as practicable to any corresponding record dates set by us) for the determination of the registered ADR holders who will be entitled (or obligated, as the case may be):

- to receive any distribution on or in respect of deposited securities,
- to give instructions for the exercise of voting rights at a meeting of holders of shares, or

- to pay the fee assessed by the depositary for administration of the ADR program and for any expenses as provided for in the ADR,
- to receive any notice or to act in respect of other matters,

all subject to the provisions of the deposit agreement.

Voting Rights

How do I vote?

If you are an ADR holder and the depositary asks you to provide it with voting instructions, you may instruct the depositary how to exercise the voting rights for the shares which underlie your ADSs. As soon as practicable after receipt from us of notice of any meeting at which the holders of shares are entitled to vote, or of our solicitation of consents or proxies from holders of shares, the depositary shall fix the ADS record date in accordance with the provisions of the deposit agreement, provided that if the depositary receives a written request from us in a timely manner and at least 30 days prior to the date of such vote or meeting, the depositary shall, at our expense, distribute to the registered ADR holders a "voting notice" stating (i) final information particular to such vote and meeting and any solicitation materials, (ii) that each ADR holder on the record date set by the depositary will, subject to any applicable provisions of Cayman Islands law, be entitled to instruct the depositary as to the exercise of the voting rights, if any, pertaining to the deposited securities represented by the ADSs evidenced by such ADR holder's ADRs and (iii) the manner in which such instructions may be given or deemed given in accordance with the deposit agreement, including instructions for giving a discretionary proxy to a person designated by us. Each ADR holder shall be solely responsible for the forwarding of voting notices to the beneficial owners of ADSs registered in such ADR holder's name. There is no guarantee that ADR holders and beneficial owners generally or any holder or beneficial owner in particular will receive the notice described above with sufficient time to enable such ADR holder or beneficial owner to return any voting instructions to the depositary in a timely manner.

Following actual receipt by the ADR department responsible for proxies and voting of ADR holders' instructions (including, without limitation, instructions of any entity or entities acting on behalf of the nominee for DTC), the depositary shall, in the manner and on or before the time established by the depositary for such purpose, endeavor to vote or cause to be voted the deposited securities represented by the ADSs evidenced by such ADR holders' ADRs in accordance with such instructions insofar as practicable and permitted under the provisions of or governing deposited securities.

To the extent that (A) we have provided the depositary with at least 35 days' notice of the proposed meeting, (B) the voting notice will be received by all ADR holders and beneficial owners no less than 10 days prior to the date of the meeting and/or the cut-off date for the solicitation of consents, and (C) the depositary does not receive instructions on a particular agenda item from an ADR holder (including, without limitation, any entity or entities acting on behalf of the nominee for DTC) in a timely manner, such ADR holder shall be deemed, and in the deposit agreement the depositary is instructed to deem such ADR holder, to have instructed the depositary to give a discretionary proxy for such agenda item(s) to a person designated by us to vote the deposited securities represented by the ADSs for which actual instructions were not so given by all such ADR holders on such agenda item(s), provided that no such instruction shall be deemed given and no discretionary proxy shall be given unless (1) we inform the depositary in writing (and we agree to provide the depositary with such instruction promptly in writing) that (a) we wish such proxy to be given with respect to such agenda item(s), (b) there is no substantial opposition existing with respect to such agenda item(s) and (c) such agenda item(s), if approved, would not materially or adversely affect the rights of holders of shares, and (2) the depositary has obtained an opinion of counsel, in form and substance satisfactory to the depositary, confirming that (i) the granting of such discretionary proxy does not subject the depositary to any reporting obligations in the Cayman Islands, (ii) the granting of

such proxy will not result in a violation of the laws, rules, regulations or permits of the Cayman Islands, (iii) the voting arrangement and deemed instruction as contemplated herein will be given effect under the laws, rules and regulations of the Cayman Islands, and (iv) the granting of such discretionary proxy will not under any circumstances result in the shares represented by the ADSs being treated as assets of the depositary under the laws, rules or regulations of the Cayman Islands.

The depositary may from time to time access information available to it to consider whether any of the circumstances described above exist, or request additional information from us in respect thereto. By taking any such action, the depositary shall not in any way be deemed or inferred to have been required, or have had any duty or responsibility (contractual or otherwise), to monitor or inquire whether any of the circumstances described above existed. In addition to the limitations provided for in the deposit agreement, ADR holders and beneficial owners are advised and agree that (a) the depositary will rely fully and exclusively on us to inform it of any of the circumstances set forth above, and (b) neither the depositary, the custodian nor any of their respective agents shall be obliged to inquire or investigate whether any of the circumstances described above exist and/or whether we complied with our obligation to timely inform the depositary of such circumstances. Neither the depositary, the custodian nor any of their respective agents shall incur any liability to ADR holders or beneficial owners (i) as a result of our failure to determine that any of the circumstances described above exist or our failure to timely notify the depositary of any such circumstances or (ii) if any agenda item which is approved at a meeting has, or is claimed to have, a material or adverse effect on the rights of holders of shares. Because there is no guarantee that ADR holders and beneficial owners will receive the notices described above with sufficient time to enable such ADR holders or beneficial owners to return any voting instructions to the depositary in a timely manner, ADR holders and beneficial owners may be deemed to have instructed the depositary to give a discretionary proxy to a person designated by us in such circumstances, and neither the depositary, the custodian nor any of their respective agents shall incur any liability to ADR holders or beneficial owners in such circumstances.

ADR holders are strongly encouraged to forward their voting instructions to the depositary as soon as possible. For instructions to be valid, the ADR department of the depositary that is responsible for proxies and voting must receive them in the manner and on or before the time specified, notwithstanding that such instructions may have been physically received by the depositary prior to such time. The depositary will not itself exercise any voting discretion in respect of deposited securities. The depositary and its agents will not be responsible for any failure to carry out any instructions to vote any of the deposited securities, for the manner in which any voting instructions are given or deemed to be given in accordance with the deposit agreement, including instructions to give a discretionary proxy to a person designated by us, for the manner in which any vote is cast, including, without limitation, any vote cast by a person to whom the depositary is instructed to grant a discretionary proxy (or deemed to have been instructed to grant a discretionary proxy in accordance with the deposit agreement), or for the effect of any such vote. Notwithstanding anything contained in the deposit agreement or any ADR, the depositary may, to the extent not prohibited by any law, regulation, or requirement of the stock exchange on which the ADSs are listed, in lieu of distribution of the materials provided to the depositary in connection with any meeting of or solicitation of consents or proxies from holders of deposited securities, distribute to the registered holders of ADRs a notice that provides such ADR holders with or otherwise publicizes to such ADR holders instructions on how to retrieve such materials or receive such materials upon request (*i.e.*, by reference to a website containing the materials for retrieval or a contact for requesting copies of the materials).

We have advised the depositary that under Cayman Islands law and our constituent documents, each as in effect as of the date of the deposit agreement, voting at any meeting of shareholders is by show of hands unless a poll is (before or on the declaration of the results of the show of hands) demanded. In the event that voting on any resolution or matter is conducted on a show of hands basis

in accordance with our constituent documents, the depositary will refrain from voting and the voting instructions received by the depositary from ADR holders shall lapse. The depositary will not demand a poll or join in demanding a poll, whether or not requested to do so by ADR holders or beneficial owners.

There is no guarantee that you will receive voting materials in time to instruct the depositary to vote and it is possible that you, or persons who hold their ADSs through brokers, dealers or other third parties, will not have the opportunity to exercise a right to vote.

Reports and Other Communications

Will ADR holders be able to view our reports?

The depositary will make available for inspection by ADR holders at the offices of the depositary and the custodian the deposit agreement, the provisions of or governing deposited securities, and any written communications from us which are both received by the custodian or its nominee as a holder of deposited securities and made generally available to the holders of deposited securities.

Additionally, if we make any written communications generally available to holders of our shares, and we furnish copies thereof (or English translations or summaries) to the depositary, it will distribute the same to registered ADR holders.

Fees and Expenses

What fees and expenses will I be responsible for paying?

The depositary may charge each person to whom ADSs are issued, including, without limitation, issuances against deposits of shares, issuances in respect of share distributions, rights and other distributions, issuances pursuant to a stock dividend or stock split declared by us or issuances pursuant to a merger, exchange of securities or any other transaction or event affecting the ADSs or deposited securities, and each person surrendering ADSs for withdrawal of deposited securities or whose ADRs are cancelled or reduced for any other reason, \$5.00 for each 100 ADSs (or any portion thereof) issued, delivered, reduced, cancelled or surrendered, or upon which a share distribution or elective distribution is made or offered, as the case may be. The depositary may sell (by public or private sale) sufficient securities and property received in respect of a share distribution, rights and/or other distribution prior to such deposit to pay such charge.

The following additional charges shall also be incurred by the ADR holders, the beneficial owners, by any party depositing or withdrawing shares or by any party surrendering ADSs and/or to whom ADSs are issued (including, without limitation, issuance pursuant to a stock dividend or stock split declared by us or an exchange of stock regarding the ADSs or the deposited securities or a distribution of ADSs), whichever is applicable:

- a fee of U.S.\$1.50 per ADR or ADRs for transfers of certificated or direct registration ADRs;
- a fee of U.S.\$0.05 or less per ADS held for any cash distribution made, or for any elective cash/stock dividend offered, pursuant to the deposit agreement;
- an aggregate fee of U.S.\$0.05 or less per ADS per calendar year (or portion thereof) for services performed by the depositary in administering the ADRs (which fee may be charged on a periodic basis during each calendar year and shall be assessed against holders of ADRs as of the record date or record dates set by the depositary during each calendar year and shall be payable in the manner described in the next succeeding provision);
- a fee for the reimbursement of such fees, charges and expenses as are incurred by the depositary and/or any of its agents (including, without limitation, the custodian and expenses incurred on

behalf of ADR holders in connection with compliance with foreign exchange control regulations or any law or regulation relating to foreign investment) in connection with the servicing of the shares or other deposited securities, the sale of securities (including, without limitation, deposited securities), the delivery of deposited securities or otherwise in connection with the depository's or its custodian's compliance with applicable law, rule or regulation (which fees and charges shall be assessed on a proportionate basis against ADR holders as of the record date or dates set by the depository and shall be payable at the sole discretion of the depository by billing such ADR holders or by deducting such charge from one or more cash dividends or other cash distributions);

- a fee for the distribution of securities (or the sale of securities in connection with a distribution), such fee being in an amount equal to the \$0.05 per ADS issuance fee for the execution and delivery of ADSs which would have been charged as a result of the deposit of such securities (treating all such securities as if they were shares) but which securities or the net cash proceeds from the sale thereof are instead distributed by the depository to those ADR holders entitled thereto;
- stock transfer or other taxes and other governmental charges;
- cable, telex and facsimile transmission and delivery charges incurred at your request in connection with the deposit or delivery of shares, ADRs or deposited securities;
- transfer or registration fees for the registration of transfer of deposited securities on any applicable register in connection with the deposit or withdrawal of deposited securities; and
- fees of any division, branch or affiliate of the depository utilized by the depository to direct, manage and/or execute any public and/or private sale of securities under the deposit agreement.

To facilitate the administration of various depository receipt transactions, including disbursement of dividends or other cash distributions and other corporate actions, the depository may engage the foreign exchange desk within JPMorgan Chase Bank, N.A. (the "**Bank**") and/or its affiliates in order to enter into spot foreign exchange transactions to convert foreign currency into U.S. dollars. For certain currencies, foreign exchange transactions are entered into with the Bank or an affiliate, as the case may be, acting in a principal capacity. For other currencies, foreign exchange transactions are routed directly to and managed by an unaffiliated local custodian (or other third party local liquidity provider), and neither the Bank nor any of its affiliates is a party to such foreign exchange transactions.

The foreign exchange rate applied to an foreign exchange transaction will be either (a) a published benchmark rate, or (b) a rate determined by a third party local liquidity provider, in each case plus or minus a spread, as applicable. The depository will disclose which foreign exchange rate and spread, if any, apply to such currency on the "Disclosures" page (or successor page) of ADR.com. Such applicable foreign exchange rate and spread may (and neither the depository, the Bank nor any of their affiliates is under any obligation to ensure that such rate does not) differ from rates and spreads at which comparable transactions are entered into with other customers or the range of foreign exchange rates and spreads at which the Bank or any of its affiliates enters into foreign exchange transactions in the relevant currency pair on the date of the foreign exchange transaction. Additionally, the timing of execution of an foreign exchange transaction varies according to local market dynamics, which may include regulatory requirements, market hours and liquidity in the foreign exchange market or other factors. Furthermore, the Bank and its affiliates may manage the associated risks of their position in the market in a manner they deem appropriate without regard to the impact of such activities on the depository, us, holders or beneficial owners. *The spread applied does not reflect any gains or losses that may be earned or incurred by the Bank and its affiliates as a result of risk management or other hedging related activity.*

Notwithstanding the foregoing, to the extent we provide U.S. dollars to the depository, neither the Bank nor any of its affiliates will execute a foreign exchange transaction as set forth herein. In such case, the depository will distribute the U.S. dollars received from us.

Further details relating to the applicable foreign exchange rate, the applicable spread and the execution of foreign exchange transactions will be provided by the depository on ADR.com. Each holder and beneficial owner by holding or owning an ADR or ADS or an interest therein, and we, each acknowledge and agree that the terms applicable to foreign exchange transactions disclosed from time to time on ADR.com will apply to any foreign exchange transaction executed pursuant to the deposit agreement.

We will pay all other charges and expenses of the depository and any agent of the depository (except the custodian) pursuant to agreements from time to time between us and the depository.

The right of the depository to receive payment of fees, charges and expenses survives the termination of the deposit agreement, and shall extend for those fees, charges and expenses incurred prior to the effectiveness of any resignation or removal of the depository.

The fees and charges described above may be amended from time to time by agreement between us and the depository.

The depository may make available to us a set amount or a portion of the depository fees charged in respect of the ADR program or otherwise upon such terms and conditions as we and the depository may agree from time to time. The depository collects its fees for issuance and cancellation of ADSs directly from investors depositing shares or surrendering ADSs for the purpose of withdrawal or from intermediaries acting for them. The depository collects fees for making distributions to investors by deducting those fees from the amounts distributed or by selling a portion of distributable property to pay the fees. The depository may collect its annual fee for depository services by deduction from cash distributions, or by directly billing investors, or by charging the book-entry system accounts of participants acting for them. The depository will generally set off the amounts owing from distributions made to holders of ADSs. If, however, no distribution exists and payment owing is not timely received by the depository, the depository may refuse to provide any further services to ADR holders that have not paid those fees and expenses owing until such fees and expenses have been paid. At the discretion of the depository, all fees and charges owing under the deposit agreement are due in advance and/or when declared owing by the depository.

Payment of Taxes

ADR holders or beneficial owners must pay any tax or other governmental charge payable by the custodian or the depository on any ADS or ADR, deposited security or distribution. If any taxes or other governmental charges (including any penalties and/or interest) shall become payable by or on behalf of the custodian or the depository with respect to any ADR, any deposited securities represented by the ADSs evidenced thereby or any distribution thereon, including, without limitation, any Chinese Enterprise Income Tax owing if the Circular Guoshuifa [2009] No. 82 issued by the Chinese State Administration of Taxation (SAT) or any other circular, edict, order or ruling, as issued and as from time to time amended, is applied or otherwise, such tax or other governmental charge shall be paid by the ADR holder thereof to the depository and by holding or owning, or having held or owned, an ADR or any ADSs evidenced thereby, the ADR holder and all beneficial owners thereof, and all prior ADR holders and beneficial owners thereof, jointly and severally, agree to indemnify, defend and save harmless each of the depository, us and its and our agents in respect of such tax or other governmental charge. Notwithstanding the depository's right to seek payment from current and former beneficial owners, by holding or owning, or having held or owned, an ADR, the ADR holder thereof (and prior ADR holder thereof) acknowledges and agrees that the depository has no obligation to seek payment of amounts owing from any current or former beneficial owner. If an ADR holder owes any tax or other governmental charge, the depository may (i) deduct the amount thereof from any cash

distributions, or (ii) sell deposited securities (by public or private sale) and deduct the amount owing from the net proceeds of such sale. In either case the ADR holder remains liable for any shortfall. If any tax or governmental charge is unpaid, the depository may also refuse to effect any registration, registration of transfer, split-up or combination of deposited securities or withdrawal of deposited securities until such payment is made. If any tax or governmental charge is required to be withheld on any cash distribution, the depository may deduct the amount required to be withheld from any cash distribution or, in the case of a non-cash distribution, sell the distributed property or securities (by public or private sale) in such amounts and in such manner as the depository deems necessary and practicable to pay such taxes and distribute any remaining net proceeds or the balance of any such property after deduction of such taxes to the ADR holders entitled thereto.

As an ADR holder or beneficial owner, you will be agreeing to indemnify us, the depository, its custodian and any of our or their respective officers, directors, employees, agents and affiliates against, and hold each of them harmless from, any claims by any governmental authority with respect to taxes, additions to tax, penalties or interest arising out of any refund of taxes, reduced rate of withholding at source or other tax benefit obtained.

Reclassifications, Recapitalizations and Mergers

If we take certain actions that affect the deposited securities, including (i) any change in par value, split-up, consolidation, cancellation or other reclassification of deposited securities or (ii) any distributions of shares or other property not made to holders of ADRs or (iii) any recapitalization, reorganization, merger, consolidation, liquidation, receivership, bankruptcy or sale of all or substantially all of our assets, then the depository may choose to, and shall if reasonably requested by us:

- amend the form of ADR;
- distribute additional or amended ADRs;
- distribute cash, securities or other property it has received in connection with such actions;
- sell any securities or property received and distribute the proceeds as cash; or
- none of the above.

If the depository does not choose any of the above options, any of the cash, securities or other property it receives will constitute part of the deposited securities and each ADS will then represent a proportionate interest in such property.

Amendment and Termination

How may the deposit agreement be amended?

We may agree with the depository to amend the deposit agreement and the ADSs without your consent for any reason. ADR holders must be given at least 30 days' notice of any amendment that imposes or increases any fees or charges (other than stock transfer or other taxes and other governmental charges, transfer or registration fees, SWIFT, cable, telex or facsimile transmission costs, delivery costs or other such expenses), or otherwise prejudices any substantial existing right of ADR holders or beneficial owners. Such notice need not describe in detail the specific amendments effectuated thereby, but must identify to ADR holders and beneficial owners a means to access the text of such amendment. If an ADR holder continues to hold an ADR or ADRs after being so notified, such ADR holder and any beneficial owner are deemed to agree to such amendment and to be bound by the deposit agreement as so amended. No amendment, however, will impair your right to surrender your ADSs and receive the underlying securities, except in order to comply with mandatory provisions of applicable law.

Any amendments or supplements which (i) are reasonably necessary (as agreed by us and the depositary) in order for (a) the ADSs to be registered on Form F-6 under the Securities Act of 1933 or (b) the ADSs or shares to be traded solely in electronic book-entry form and (ii) do not in either such case impose or increase any fees or charges to be borne by ADR holders, shall be deemed not to prejudice any substantial rights of ADR holders or beneficial owners. Notwithstanding the foregoing, if any governmental body or regulatory body should adopt new laws, rules or regulations which would require amendment or supplement of the deposit agreement or the form of ADR to ensure compliance therewith, we and the depositary may amend or supplement the deposit agreement and the ADR at any time in accordance with such changed laws, rules or regulations. Such amendment or supplement to the deposit agreement in such circumstances may become effective before a notice of such amendment or supplement is given to ADR holders or within any other period of time as required for compliance.

Notice of any amendment to the deposit agreement or form of ADRs shall not need to describe in detail the specific amendments effectuated thereby, and failure to describe the specific amendments in any such notice shall not render such notice invalid, provided, however, that, in each such case, the notice given to the ADR holders identifies a means for ADR holders and beneficial owners to retrieve or receive the text of such amendment (*i.e.*, upon retrieval from the SEC's, the depositary's or our website or upon request from the depositary).

How may the deposit agreement be terminated?

The depositary may, and shall at our written direction, terminate the deposit agreement and the ADRs by mailing notice of such termination to the registered holders of ADRs at least 30 days prior to the date fixed in such notice for such termination; provided, however, if the depositary shall have (i) resigned as depositary under the deposit agreement, notice of such termination by the depositary shall not be provided to registered ADR holders unless a successor depositary shall not be operating under the deposit agreement within 60 days of the date of such resignation, and (ii) been removed as depositary under the deposit agreement, notice of such termination by the depositary shall not be provided to registered holders of ADRs unless a successor depositary shall not be operating under the deposit agreement on the 60th day after our notice of removal was first provided to the depositary.

If the shares are not listed or quoted for trading on a stock exchange or in a securities market as of the date so fixed for termination, then after such date fixed for termination (a) all direct registration ADRs shall cease to be eligible for the direct registration system and shall be considered ADRs issued on the ADR register maintained by the depositary and (b) the depositary shall use its reasonable efforts to ensure that the ADSs cease to be DTC eligible so that neither DTC nor any of its nominees shall thereafter be a holder of ADRs. At such time as the ADSs cease to be DTC eligible and/or neither DTC nor any of its nominees is a holder of ADRs, the depositary shall (a) instruct its custodian to deliver all shares and/or deposited securities to us along with a general stock power that refers to the names set forth on the ADR register maintained by the depositary and (b) provide us with a copy of the ADR register maintained by the depositary. Upon receipt of such shares and/or deposited securities and the ADR register maintained by the depositary, we have agreed to use our best efforts to issue to each registered ADR holder a share certificate representing the shares represented by the ADSs reflected on the ADR register maintained by the depositary in such registered ADR holder's name and to deliver such share certificate to the registered ADR holder at the address set forth on the ADR register maintained by the depositary. After providing such instruction to the custodian and delivering a copy of the ADR register to us, the depositary and its agents will perform no further acts under the deposit agreement or the ADRs and shall cease to have any obligations under the deposit agreement and/or the ADRs. After we receive the copy of the ADR register and the shares and/or deposited securities from the depositary, we shall be discharged from all obligations under the deposit agreement

except (i) to distribute the shares to the registered ADR holders entitled thereto and (ii) for its obligations to the depositary and its agents.

If the shares are listed or quoted for trading on a stock exchange or in a securities market as of the date so fixed for termination, then instead of the provisions in the prior paragraph, after the date so fixed for termination, the depositary and its agents will perform no further acts under the deposit agreement or the ADRs, except to receive and hold (or sell) distributions on shares and/or deposited securities and deliver shares and/or deposited securities being withdrawn. As soon as practicable after the date so fixed for termination, the depositary has agreed to use its reasonable efforts to sell the shares and/or deposited securities and shall thereafter (as long as it may lawfully do so) hold in an account (which may be a segregated or unsegregated account) the net proceeds of such sales, together with any other cash then held by it under the deposit agreement, without liability for interest, in trust for the pro rata benefit of the registered ADR holders not theretofore surrendered. After making such sale, the depositary shall be discharged from all obligations in respect of the deposit agreement and the ADRs, except to account for such net proceeds and other cash. After the date so fixed for termination, we shall be discharged from all obligations under the deposit agreement except for our obligations to the depositary and its agents.

Notwithstanding anything to the contrary, in connection with any such termination, the depositary may, in its sole discretion and without notice to us, establish an unsponsored American depositary share program (on such terms as the depositary may determine) for our shares and make available to ADR holders a means to withdraw the shares represented by the ADSs issued under the deposit agreement and to direct the deposit of such shares into such unsponsored American depositary share program, subject, in each case, to receipt by the depositary, at its discretion, of the fees, charges and expenses provided for under the deposit agreement and the fees, charges and expenses applicable to the unsponsored American depositary share program.

Limitations on Obligations and Liability to ADR holders

Limits on our obligations and the obligations of the depositary; limits on liability to ADR holders and holders of ADSs

Prior to the issue, registration, registration of transfer, split-up, combination, or cancellation of any ADRs, or the delivery of any distribution in respect thereof, and from time to time in the case of the production of proofs as described below, we or the depositary or its custodian may require:

- payment with respect thereto of (i) any stock transfer or other tax or other governmental charge, (ii) any stock transfer or registration fees in effect for the registration of transfers of shares or other deposited securities upon any applicable register and (iii) any applicable fees and expenses described in the deposit agreement;
- the production of proof satisfactory to it of (i) the identity of any signatory and genuineness of any signature and (ii) such other information, including without limitation, information as to citizenship, residence, exchange control approval, beneficial or other ownership of, or interest in, any securities, compliance with applicable law, regulations, provisions of or governing deposited securities and terms of the deposit agreement and the ADRs, as it may deem necessary or proper; and
- compliance with such regulations as the depositary may establish consistent with the deposit agreement.

The issuance of ADRs, the acceptance of deposits of shares, the registration, registration of transfer, split-up or combination of ADRs or the withdrawal of shares, may be suspended, generally or in particular instances, when the ADR register or any register for deposited securities is closed or when any such action is deemed advisable by the depositary; provided that the ability to withdraw shares may

only be limited under the following circumstances: (i) temporary delays caused by closing transfer books of the depositary or our transfer books or the deposit of shares in connection with voting at a shareholders' meeting, or the payment of dividends, (ii) the payment of fees, taxes, and similar charges, and (iii) compliance with any laws or governmental regulations relating to ADRs or to the withdrawal of deposited securities.

The deposit agreement expressly limits the obligations and liability of the depositary, ourselves and our respective agents, provided, however, that no disclaimer of liability under the Securities Act of 1933 is intended by any of the limitations of liabilities provisions of the deposit agreement. The deposit agreement provides that each of us, the depositary and our respective agents will:

- incur or assume no liability (including, without limitation, to holders or beneficial owners) if any present or future law, rule, regulation, fiat, order or decree of the Cayman Islands, Hong Kong, the People's Republic of China, the United States or any other country or jurisdiction, or of any governmental or regulatory authority or securities exchange or market or automated quotation system, the provisions of or governing any deposited securities, any present or future provision of our charter, any act of God, war, terrorism, epidemic, pandemic, nationalization, expropriation, currency restrictions, work stoppage, strike, civil unrest, revolutions, rebellions, explosions, computer failure or circumstance beyond our, the depositary's or our respective agents' direct and immediate control shall prevent or delay, or shall cause any of them to be subject to any civil or criminal penalty in connection with, any act which the deposit agreement or the ADRs provide shall be done or performed by us, the depositary or our respective agents (including, without limitation, voting);
- incur or assume no liability (including, without limitation, to holders or beneficial owners) by reason of any non-performance or delay, caused as aforesaid, in the performance of any act or things which by the terms of the deposit agreement it is provided shall or may be done or performed or any exercise or failure to exercise discretion under the deposit agreement or the ADRs including, without limitation, any failure to determine that any distribution or action may be lawful or reasonably practicable;
- incur or assume no liability (including, without limitation, to holders or beneficial owners) if it performs its obligations under the deposit agreement and ADRs without gross negligence or willful misconduct;
- in the case of the depositary and its agents, be under no obligation to appear in, prosecute or defend any action, suit or other proceeding in respect of any deposited securities the ADSs or the ADRs;
- in the case of us and our agents, be under no obligation to appear in, prosecute or defend any action, suit or other proceeding in respect of any deposited securities the ADSs or the ADRs, which in our or our agents' opinion, as the case may be, may involve it in expense or liability, unless indemnity satisfactory to us or our agent, as the case may be against all expense (including fees and disbursements of counsel) and liability be furnished as often as may be requested;
- not be liable (including, without limitation, to holders or beneficial owners) for any action or inaction by it in reliance upon the advice of or information from any legal counsel, any accountant, any person presenting shares for deposit, any registered holder of ADRs, or any other person believed by it to be competent to give such advice or information and/or, in the case of the depositary, us; or
- may rely and shall be protected in acting upon any written notice, request, direction, instruction or document believed by it to be genuine and to have been signed, presented or given by the proper party or parties.

Neither the depositary nor its agents have any obligation to appear in, prosecute or defend any action, suit or other proceeding in respect of any deposited securities, the ADSs or the ADRs. We and our agents shall only be obligated to appear in, prosecute or defend any action, suit or other proceeding in respect of any deposited securities, the ADSs or the ADRs, which in our opinion may involve us in expense or liability, if indemnity satisfactory to us against all expense (including fees and disbursements of counsel) and liability is furnished as often as may be required. The depositary and its agents may fully respond to any and all demands or requests for information maintained by or on its behalf in connection with the deposit agreement, any registered holder or holders of ADRs, any ADRs or otherwise related to the deposit agreement or ADRs to the extent such information is requested or required by or pursuant to any lawful authority, including without limitation laws, rules, regulations, administrative or judicial process, banking, securities or other regulators. The depositary shall not be liable for the acts or omissions made by, or the insolvency of, any securities depository, clearing agency or settlement system. Furthermore, the depositary shall not be responsible for, and shall incur no liability in connection with or arising from, the insolvency of any custodian that is not a branch or affiliate of JPMorgan. Notwithstanding anything to the contrary contained in the deposit agreement or any ADRs, the depositary shall not be responsible for, and shall incur no liability in connection with or arising from, any act or omission to act on the part of the custodian except to the extent that any registered ADR holder has incurred liability directly as a result of the custodian having (i) committed fraud or willful misconduct in the provision of custodial services to the depositary or (ii) failed to use reasonable care in the provision of custodial services to the depositary as determined in accordance with the standards prevailing in the jurisdiction in which the custodian is located. The depositary and the custodian(s) may use third party delivery services and providers of information regarding matters such as, but not limited to, pricing, proxy voting, corporate actions, class action litigation and other services in connection with the ADRs and the deposit agreement, and use local agents to provide services such as, but not limited to, attendance at any meetings of security holders of issuers. Although the depositary and the custodian will use reasonable care (and cause their agents to use reasonable care) in the selection and retention of such third party providers and local agents, they will not be responsible for any errors or omissions made by them in providing the relevant information or services. The depositary shall not have any liability for the price received in connection with any sale of securities, the timing thereof or any delay in action or omission to act nor shall it be responsible for any error or delay in action, omission to act, default or negligence on the part of the party so retained in connection with any such sale or proposed sale.

The depositary has no obligation to inform ADR holders or beneficial owners about the requirements of the laws, rules or regulations or any changes therein or thereto of the Cayman Islands, Hong Kong, the People's Republic of China, the United States or any other country or jurisdiction or of any governmental or regulatory authority or any securities exchange or market or automated quotation system.

Additionally, none of the depositary, the custodian or us, or any of their or our respective directors, officers, employees, agents or affiliates shall be liable for the failure by any registered holder of ADRs or beneficial owner therein to obtain the benefits of credits or refunds of non-U.S. tax paid against such ADR holder's or beneficial owner's income tax liability. The depositary is under no obligation to provide the ADR holders and beneficial owners, or any of them, with any information about our tax status. None of the depositary, custodian, us or any of their or our respective directors, officers, employees, agents or affiliates shall incur any liability for any tax or tax consequences that may be incurred by registered ADR holders or beneficial owners on account of their ownership or disposition of ADRs or ADSs.

Neither the depositary nor its agents will be responsible for any failure to carry out any instructions to vote any of the deposited securities, for the manner in which any voting instructions are given or deemed to be given pursuant to the terms of the deposit agreement, including instructions to

give a discretionary proxy to a person designated by us, for the manner in which any vote is cast, including, without limitation, any vote cast by a person to whom the depositary is instructed (or deemed to have been instructed to grant a discretionary proxy in accordance with the deposit agreement), or for the effect of any such vote. The depositary may rely upon instructions from us or our counsel in respect of any approval or license required for any currency conversion, transfer or distribution. The depositary shall not incur any liability for the content of any information submitted to it by us or on our behalf for distribution to ADR holders or for any inaccuracy of any translation thereof, for any investment risk associated with acquiring an interest in the deposited securities, for the validity or worth of the deposited securities, for the credit-worthiness of any third party, for allowing any rights to lapse upon the terms of the deposit agreement or for the failure or timeliness of any notice from us. The depositary shall not be liable for any acts or omissions made by a successor depositary whether in connection with a previous act or omission of the depositary or in connection with any matter arising wholly after the removal or resignation of the depositary. Subject to certain customary exceptions, neither the depositary nor us, nor any of its or our agents shall be liable to the other for any indirect, special, punitive or consequential damages (excluding reasonable fees and expenses of counsel) or lost profits, in each case of any form incurred by any of them, or liable to any other person or entity (including, without limitation holders or beneficial owners of ADRs and ADSs) for any indirect, special, punitive or consequential damages (including fees and expenses of counsel) or lost profits, whether or not foreseeable and regardless of the type of action in which such a claim may be brought.

In the deposit agreement each party thereto (including, for avoidance of doubt, each ADR holder and beneficial owner) irrevocably waives, to the fullest extent permitted by applicable law, any right it may have to a trial by jury in any suit, action or proceeding against the depositary and/or us directly or indirectly arising out of or relating to the shares or other deposited securities, the ADSs or the ADRs, the deposit agreement or any transaction contemplated therein, or the breach thereof (whether based on contract, tort, common law or any other theory). No provision of the deposit agreement or the ADRs is intended to constitute a waiver or limitation of any rights which an ADR holder or any beneficial owner may have under the Securities Act of 1933 or the Securities Exchange Act of 1934, to the extent applicable.

The depositary and its agents may own and deal in any class of securities of our company and our affiliates and in ADRs.

Disclosure of Interest in ADSs

To the extent that the provisions of or governing any deposited securities may require disclosure of or impose limits on beneficial or other ownership of, or interest in, deposited securities, other shares and other securities and may provide for blocking transfer, voting or other rights to enforce such disclosure or limits, you as ADR holders or beneficial owners agree to comply with all such disclosure requirements and ownership limitations and to comply with any reasonable instructions we may provide in respect thereof. We reserve the right to instruct you as ADS holders to deliver your ADSs for cancellation and withdrawal of the deposited securities so as to permit us to deal directly with the holder thereof as a holder of shares and, by holding an ADS or an interest therein, you and beneficial owners will be agreeing to comply with such instructions.

Books of Depositary

The depositary or its agent will maintain a register for the registration, registration of transfer, combination and split-up of ADRs, which register shall include the depositary's direct registration system. Registered holders of ADRs may inspect such records at the depositary's office at all reasonable times, but solely for the purpose of communicating with other ADR holders in the interest of the business of our company or a matter relating to the deposit agreement. Such register may be

closed at any time or from time to time, when deemed expedient by the depositary or, in the case of the issuance book portion of the ADR Register, when reasonably requested by the Company solely in order to enable the Company to comply with applicable law.

The depositary will maintain facilities for the delivery and receipt of ADRs.

Appointment

In the deposit agreement, each registered holder of ADRs and each beneficial owner, upon acceptance of any ADSs or ADRs (or any interest in any of them) issued in accordance with the terms and conditions of the deposit agreement will be deemed for all purposes to:

- be a party to and bound by the terms of the deposit agreement and the applicable ADR or ADRs,
- appoint the depositary its attorney-in-fact, with full power to delegate, to act on its behalf and to take any and all actions contemplated in the deposit agreement and the applicable ADR or ADRs, to adopt any and all procedures necessary to comply with applicable laws and to take such action as the depositary in its sole discretion may deem necessary or appropriate to carry out the purposes of the deposit agreement and the applicable ADR and ADRs, the taking of such actions to be the conclusive determinant of the necessity and appropriateness thereof; and
- acknowledge and agree that (i) nothing in the deposit agreement or any ADR shall give rise to a partnership or joint venture among the parties thereto, nor establish a fiduciary or similar relationship among such parties, (ii) the depositary, its divisions, branches and affiliates, and their respective agents, may from time to time be in the possession of non-public information about us, ADR holders, beneficial owners and/or their respective affiliates, (iii) the depositary and its divisions, branches and affiliates may at any time have multiple banking relationships with us, ADR holders, beneficial owners and/or the affiliates of any of them, (iv) the depositary and its divisions, branches and affiliates may, from time to time, be engaged in transactions in which parties adverse to us, ADR holders, beneficial owners and/or their respective affiliates may have interests, (v) nothing contained in the deposit agreement or any ADR(s) shall (A) preclude the depositary or any of its divisions, branches or affiliates from engaging in any such transactions or establishing or maintaining any such relationships, or (B) obligate the depositary or any of its divisions, branches or affiliates to disclose any such transactions or relationships or to account for any profit made or payment received in any such transactions or relationships, (vi) the depositary shall not be deemed to have knowledge of any information held by any branch, division or affiliate of the depositary and (vii) notice to an ADR holder shall be deemed, for all purposes of the deposit agreement and the ADRs, to constitute notice to any and all beneficial owners of the ADSs evidenced by such ADR holder's ADRs. For all purposes under the deposit agreement and the ADRs, the ADR holders thereof shall be deemed to have all requisite authority to act on behalf of any and all beneficial owners of the ADSs evidenced by such ADRs.

Governing Law

The deposit agreement, the ADSs and the ADRs are governed by and construed in accordance with the internal laws of the State of New York. In the deposit agreement, we have submitted to the non-exclusive jurisdiction of the courts of the State of New York and appointed an agent for service of process on our behalf. Any action based on the deposit agreement, the ADSs, the ADRs or the transactions contemplated therein or thereby may also be instituted by the depositary against us in any competent court in the Cayman Islands, Hong Kong, the People's Republic of China and the United States.

Under the deposit agreement, by holding or owning an ADR or ADS or an interest therein, ADR holders and beneficial owners each irrevocably agree that any legal suit, action or proceeding against or involving ADR holders or beneficial owners brought by us or the depository, arising out of or based upon the deposit agreement, the ADSs, the ADRs or the transactions contemplated thereby, may be instituted in a state or federal court in New York, New York, irrevocably waive any objection which you may have to the laying of venue of any such proceeding, and irrevocably submit to the non-exclusive jurisdiction of such courts in any such suit, action or proceeding. By holding or owning an ADR or ADS or an interest therein, ADR holders and beneficial owners each also irrevocably agree that any legal suit, action or proceeding against or involving the depository brought by ADR holders or beneficial owners, arising out of or based upon the deposit agreement, the ADSs, the ADRs or the transactions contemplated thereby, may only be instituted in a state or federal court in New York, New York.

Notwithstanding the foregoing, (i) the depository may, in its sole discretion, elect to institute any dispute, suit, action, controversy, claim or proceeding directly or indirectly based on, arising out of or relating to the deposit agreement, the ADSs, the ADRs or the transactions contemplated therein or thereby, including without limitation any question regarding its or their existence, validity, interpretation, performance or termination, against any other party or parties to the deposit agreement (including, without limitation, against ADR holders and beneficial owners of interests in ADSs), by having the matter referred to and finally resolved by an arbitration conducted under the terms described below, and (ii) the depository may in its sole discretion require, by written notice to the relevant party or parties, that any dispute, suit, action, controversy, claim or proceeding against the depository by any party or parties to the deposit agreement (including, without limitation, by ADR holders and beneficial owners of interests in ADSs) shall be referred to and finally settled by an arbitration conducted under the terms described below. Any such arbitration shall be conducted in the English language either in New York, New York in accordance with the Commercial Arbitration Rules of the American Arbitration Association or in Hong Kong following the arbitration rules of the United Nations Commission on International Trade Law (UNCITRAL).

Jury Trial Waiver

In the deposit agreement, each party thereto (including, for the avoidance of doubt, each holder and beneficial owner of, and/or holder of interests in, ADSs or ADRs) irrevocably waives, to the fullest extent permitted by applicable law, any right it may have to a trial by jury in any suit, action or proceeding against the depository and/or us directly or indirectly arising out of or relating to the shares or other deposited securities, the ADSs or the ADRs, the deposit agreement or any transaction contemplated therein, or the breach thereof (whether based on contract, tort, common law or any other theory), including any claim under the U.S. federal securities laws.

If we or the depository were to oppose a jury trial demand based on such waiver, the court would determine whether the waiver was enforceable in the facts and circumstances of that case in accordance with applicable state and federal law, including whether a party knowingly, intelligently and voluntarily waived the right to a jury trial. The waiver to right to a jury trial in the deposit agreement is not intended to be deemed a waiver by any holder or beneficial owner of ADSs of our or the depository's compliance with the U.S. federal securities laws and the rules and regulations promulgated thereunder.

SHARES ELIGIBLE FOR FUTURE SALE

Upon completion of this offering, we will have ADSs outstanding, representing ordinary shares, or approximately % of our outstanding ordinary shares, assuming the underwriters do not exercise their option to purchase additional ADSs. All of the ADSs sold in this offering will be freely transferable by persons other than our "affiliates" without restriction or further registration under the Securities Act. Sales of substantial amounts of our ADSs in the public market could adversely affect prevailing market prices of our ADSs. Prior to this offering, there has been no public market for our ordinary shares or the ADSs, and while our ADSs have been approved for listing on the Nasdaq, we cannot assure you that a regular trading market will develop in the ADSs. Our ordinary shares will not be listed on any exchange or quoted for trading on any over-the-counter trading system. We do not expect that a trading market will develop for our ordinary shares not represented by the ADSs.

Lock-up Agreements

We, [our directors, executive officers and existing shareholders] have agreed with the underwriters, subject to some exceptions, not to sell, transfer or dispose of, directly or indirectly, any of our ordinary shares, in the form of ADSs or otherwise, or any securities convertible into or exchangeable or exercisable for our ordinary shares, in the form of ADSs or otherwise, for a period of [180] days after the date of this prospectus. After the expiration of the [180]-day period, the ordinary shares or ADSs held by our directors, executive officers and our existing shareholders may be sold subject to the restrictions under Rule 144 under the Securities Act or by means of registered public offerings.

Rule 144

All of our ordinary shares outstanding prior to this offering are "restricted shares" as that term is defined in Rule 144 under the Securities Act and may be sold publicly in the United States only if they are subject to an effective registration statement under the Securities Act or pursuant to an exemption from the registration requirements. Under Rule 144 as currently in effect, a person who is not deemed to have been our affiliate at any time during the three months preceding a sale and who has beneficially owned our restricted shares for at least six months is generally entitled to sell the restricted securities without registration under the Securities Act beginning 90 days after the date of this prospectus, subject to certain additional restrictions.

Our affiliates who have beneficially owned "restricted securities" for at least six months may sell within any three-month period a number of restricted shares that does not exceed the greater of the following:

- 1% of the then outstanding ordinary shares of the same class, in the form of ADSs or otherwise, which will equal approximately ordinary shares immediately after this offering, assuming the underwriters do not exercise their option to purchase additional ADSs; or
- the average weekly trading volume of our ordinary shares in the form of ADSs or otherwise on the Nasdaq during the four calendar weeks preceding the date on which notice of the sale on Form 144 is filed with the SEC.

Affiliates who sell restricted securities under Rule 144 may not solicit orders or arrange for the solicitation of orders, and they are also subject to certain manner of sale provisions and notice requirements and the availability of current public information about us. In addition, in each case, shares held by our affiliates would remain subject to lock-up arrangements and would only become eligible for sale when the lock-up period expires.

Persons who are not our affiliates are only subject to one of these additional restrictions, the requirement of the availability of current public information about us, and this additional restriction does not apply if they have beneficially owned our restricted shares for more than one year.

Rule 701

In general, under Rule 701 of the Securities Act as currently in effect, each of our employees, consultants or advisors who purchases our ordinary shares from us in connection with a compensatory stock or option plan or other written agreement relating to compensation is eligible to resell such ordinary shares 90 days after we became a reporting company under the Exchange Act in reliance on Rule 144, but without compliance with some of the restrictions, including the holding period, contained in Rule 144. However, the Rule 701 shares would remain subject to lock-up arrangements and would only become eligible for sale when the lock-up period expires.

Registration Rights

Upon the completion of this offering, holders of our registrable securities will be entitled to request that we register their shares under the Securities Act, following the expiration of the lock-up agreements described above. See "Description of Share Capital—Shareholders' Agreements—Registration Rights."

Form S-8

We intend to file a registration statement on Form S-8 under the Securities Act covering all ordinary shares which are either subject to outstanding options or may be issued upon exercise or vesting of any options or other equity awards which may be granted or issued in the future pursuant to our share incentive plan. We expect to file this registration statement as soon as practicable after the date of this prospectus. Shares registered under any registration statements will be available for sale in the open market, except to the extent that the shares are subject to vesting restrictions with us or the contractual restrictions and the lock-up described below.

TAXATION

The following discussion of Cayman Islands, PRC and U.S. federal income tax consequences of an investment in the ADSs or ordinary shares is based upon laws and relevant interpretations thereof in effect as of the date of this prospectus, all of which are subject to change. This discussion does not deal with all possible tax consequences relating to an investment in the ADSs or ordinary shares, such as the tax consequences under state, local and other tax laws. To the extent that the discussion relates to matters of Cayman Islands tax law, it represents the opinion of Walkers (Hong Kong), our Cayman Islands counsel. To the extent that the discussion relates to matters of PRC tax law, it represents the opinion of Tian Yuan Law Firm, our PRC legal counsel.

Cayman Islands Taxation

The Cayman Islands currently levies no taxes on individuals or corporations based upon profits, income, gains or appreciation, and there is no taxation in the nature of inheritance tax or estate duty. There are no other taxes likely to be material to us or holders of our ADSs or ordinary shares levied by the government of the Cayman Islands, except for stamp duties which may be applicable on instruments executed in, or after execution brought within the jurisdiction of the Cayman Islands. The Cayman Islands is not party to any double tax treaties that are applicable to any payments made to or by our company. There are no exchange control regulations or currency restrictions in the Cayman Islands.

Payments of dividends and capital in respect of the ADSs or ordinary shares will not be subject to taxation in the Cayman Islands and no withholding will be required on the payment of a dividend or capital to any holder of the ADSs or ordinary shares, nor will gains derived from the disposal of the ADSs or ordinary shares be subject to Cayman Islands income or corporation tax.

Material PRC Income Tax Considerations

Under the PRC EIT Law, which became effective on January 1, 2008 and amended on December 29, 2019, an enterprise established outside the PRC with "de facto management bodies" within the PRC is considered a "resident enterprise" for PRC enterprise income tax purposes and is generally subject to a uniform 25% enterprise income tax rate on its worldwide income. Under the implementation rules to the PRC EIT Law, a "de facto management body" is defined as a body that has material and overall management and control over the manufacturing and business operations, personnel and human resources, finances and properties of an enterprise.

In addition, the SAT Circular 82 issued by the SAT in April 2009 specifies that certain offshore incorporated enterprises controlled by PRC enterprises or PRC enterprise groups will be classified as PRC resident enterprises if the following are located or resident in the PRC: (a) senior management personnel and departments that are responsible for daily production, operation and management; (b) financial and personnel decision-making bodies; (c) key properties, accounting books, company seal, minutes of board meetings and shareholders' meetings; and (d) half or more of the senior management or directors having voting rights. Further to SAT Circular 82, the SAT issued the SAT Bulletin 45, which took effect in September 2011, to provide more guidance on the implementation of SAT Circular 82. SAT Bulletin 45 provides for procedures and administration details of determination on resident status and administration on post-determination matters. Our company is a company incorporated outside the PRC. As a holding company, its key assets are its ownership interests in its subsidiaries, and its key assets are located, and its records (including the resolutions of its board of directors and the resolutions of its shareholders) are maintained, outside the PRC. As such, we do not believe that our company meets all of the conditions above or is a PRC resident enterprise for PRC tax purposes. For the same reasons, we believe our other entities outside China are not PRC resident enterprises either. However, the tax resident status of an enterprise is subject to determination by the PRC tax authorities and uncertainties remain with respect to the interpretation of the term "de facto management body." There can be no assurance that the PRC government will ultimately take a view that is consistent with

us. If the PRC tax authorities determine that our Cayman Islands holding company is a PRC resident enterprise for PRC enterprise income tax purposes, a number of unfavorable PRC tax consequences could follow. For example, a 10% withholding tax would be imposed on dividends we pay to our non-PRC enterprise shareholders (including our ADS holders). In addition, nonresident enterprise shareholders (including our ADS holders) may be subject to PRC tax on gains realized on the sale or other disposition of ADSs or ordinary shares, if such income is treated as sourced from within the PRC. Furthermore, if we are deemed a PRC resident enterprise, dividends paid to our non-PRC individual shareholders (including our ADS holders) and any gain realized on the transfer of ADSs or ordinary shares by such shareholders may be subject to PRC tax at a rate of 20% (which, in the case of dividends, may be withheld at source by us). These rates may be reduced by an applicable tax treaty, but it is unclear whether in practice non-PRC shareholders of our company would be able to obtain the benefits of any tax treaties between their country of tax residence and the PRC in the event that we are treated as a PRC resident enterprise. See "Risk Factors—Risks Related to Doing Business in China—If we are classified as a PRC resident enterprise for PRC income tax purposes, such classification could result in unfavorable tax consequences to us and our non-PRC shareholders or ADS holders."

Material U.S. Federal Income Tax Considerations

In the opinion of Davis Polk & Wardwell LLP, the following are material U.S. federal income tax consequences to the U.S. Holders described below of owning and disposing of our ADSs or ordinary shares. This discussion is not a comprehensive description of all of the tax considerations that may be relevant to your decision to acquire the ADSs or ordinary shares. This discussion applies to you only if you are a U.S. Holder that acquires ADSs in this offering and holds the ADSs or underlying ordinary shares as capital assets for U.S. federal income tax purposes. In addition, it does not describe all of the tax consequences that may be relevant in light of your particular circumstances, including alternative minimum and Medicare contribution tax considerations, and tax consequences applicable to you if you are subject to special rules, such as:

- one of certain financial institutions;
- a dealer or trader in securities that uses a mark-to-market method of tax accounting;
- a person holding ADSs or ordinary shares as part of a straddle, conversion transaction, integrated transaction or similar transaction;
- a person whose functional currency for U.S. federal income tax purposes is not the U.S. dollar;
- an entity classified as partnerships for U.S. federal income tax purposes (and investors therein);
- a tax-exempt entity, "individual retirement account" or "Roth IRA";
- a person that owns or is deemed to own 10% or more of our stock by vote or value; or
- a person holding ADSs or ordinary shares in connection with a trade or business conducted outside the United States.

If you are a partnership for U.S. federal income tax purposes, the U.S. federal income tax consequences to your partners will generally depend on their status and your activities. Partnerships holding ADSs or ordinary shares and their partners should consult their tax advisers as to the particular U.S. federal income tax consequences of acquiring, owning or disposing of the ADSs or ordinary shares.

This discussion is based on the Internal Revenue Code of 1986, as amended, or the Code, administrative pronouncements, judicial decisions and final, temporary and proposed Treasury regulations, and the income tax treaty between the United States and the PRC, or the Treaty, all as of the date hereof, any of which is subject to change, possibly with retroactive effect.

For purposes of this discussion, you are a "U.S. Holder" if for U.S. federal income tax purposes you are a beneficial owner of ADSs or ordinary shares and:

- a citizen or individual resident of the United States;
- a corporation or other entity taxable as a corporation for U.S. federal income tax purposes, created or organized in or under the laws of the United States, any state therein or the District of Columbia; or
- an estate or trust the income of which is subject to U.S. federal income taxation regardless of its source.

In general, if you own ADSs you will be treated as the owner of the underlying ordinary shares represented by those ADSs for U.S. federal income tax purposes. Accordingly, no gain or loss will be recognized if you exchange ADSs for the underlying ordinary shares represented by those ADSs.

Taxation of Distributions

Except as described under "*Passive Foreign Investment Company Rules*" below, distributions paid on ADSs or ordinary shares, other than certain pro rata distributions of ordinary shares, will generally be treated as dividends to the extent paid out of our current or accumulated earnings and profits (as determined under U.S. federal income tax principles). Because we do not maintain calculations of our earnings and profits under U.S. federal income tax principles, we expect that any distributions will be reported to you as dividends. Dividends will not be eligible for the dividends-received deduction generally available to United States corporations under the Code. Subject to applicable limitations, dividends paid on our ADSs to certain non-corporate U.S. Holders may be taxable at a favorable rate. Dividends will be included in your income on the date of your, or in the case of ADSs, the depository's, receipt of the dividend. The amount of any dividend income paid in non-U.S. currency will be its U.S. dollar value calculated by reference to the spot rate in effect on the date of receipt, regardless of whether the payment is in fact converted into U.S. dollars. If the dividend is converted into U.S. dollars on the date of receipt, you generally should not be required to recognize foreign currency gain or loss in respect of the dividend income. You may have foreign currency gain or loss, which will be United States source, if the dividend is converted into U.S. dollars after the date of receipt.

Dividends will be treated as foreign-source income for foreign tax credit purposes. As described in "*PRC Taxation*", dividends paid by us may be subject to PRC withholding tax. For U.S. federal income tax purposes, the amount of the dividend income will include any amounts withheld in respect of PRC withholding tax. Subject to applicable limitations, which vary depending upon your circumstances, PRC taxes withheld from dividend payments (at a rate not exceeding the applicable rate provided in the Treaty) generally will be creditable against your U.S. federal income tax liability. The rules governing foreign tax credits are complex and you should consult your tax advisers regarding the creditability of foreign tax credits in your particular circumstances. In lieu of claiming a credit, you may elect to deduct PRC taxes in computing its taxable income, subject to applicable limitations. An election to deduct foreign taxes instead of claiming foreign tax credits must apply to all foreign taxes paid or accrued in the taxable year.

Sale or Other Disposition of ADSs or Ordinary Shares

Except as described under "*Passive Foreign Investment Company Rules*" below, gain or loss realized on the sale or other taxable disposition of ADSs or ordinary shares will be capital gain or loss, and will be long-term capital gain or loss if you held the ADSs or ordinary shares for more than one year. The amount of the gain or loss will equal the difference between your tax basis in the ADSs or ordinary shares disposed of and the amount realized on the disposition, in each case as determined in

U.S. dollars. This gain or loss will generally be U.S.-source gain or loss for foreign tax credit purposes. The deductibility of capital losses is subject to limitations.

As described in "—PRC Taxation" gains on the sale of ADSs or ordinary shares may be subject to PRC taxes. You are entitled to use foreign tax credits to offset only the portion of your U.S. federal income tax liability that is attributable to foreign-source income. Because under the Code capital gains of U.S. persons are generally treated as U.S.-source income, this limitation may preclude you from claiming a credit for all or a portion of any PRC taxes imposed on any such gains. However, if you are eligible for the benefits of the Treaty, you may elect to treat the gain as PRC-source and therefore claim foreign tax credits in respect of PRC taxes on such disposition gains. You should consult your tax advisor regarding their eligibility for the benefits of the Treaty and the creditability of any PRC tax on disposition gains in your particular circumstances.

Passive Foreign Investment Company Rules

In general, a non-U.S. corporation will be a passive foreign investment company, or a PFIC, for any taxable year in which (i) 75% or more of its gross income consists of passive income, or the income test, or (ii) 50% or more of the average value of its assets (generally determined on a quarterly basis) consists of assets that produce, or are held for the production of, passive income, or the asset test. For purposes of the above calculations, a non-U.S. corporation that directly or indirectly owns at least 25% by value of the ordinary shares of another corporation is treated as if it held its proportionate share of the assets of the other corporation and received directly its proportionate share of the income of the other corporation. Passive income generally includes interest, dividends, gains from certain property transactions, rents and royalties (other than certain rents or royalties derived in the active conduct of a trade or business). Cash is a passive asset for PFIC purposes. Goodwill is an active asset under the PFIC rules to the extent attributable to activities that produce active income.

The assets shown on our balance sheet are expected to consist primarily of cash and cash equivalents for the foreseeable future. Therefore, whether we will satisfy the asset test for the current or any future taxable year will depend largely on the value of our goodwill and on how quickly we utilize the cash in our business. We cannot give any assurance as to whether we will be a PFIC for the current or any future taxable year because (i) the value of our goodwill may be determined by reference to the market price of our ADSs, which may be volatile given the nature and early stage of our business, (ii) we expect to hold a significant amount of cash, and (iii) a company's PFIC status is an annual determination that can be made only after the end of each taxable year. In addition, prior to commercialization of our product candidates, we could be a PFIC for any taxable year in which our interest and other investment income constitutes 75% or more of our total gross income.

If we were a PFIC for any taxable year and any of our subsidiaries were also a PFIC (any such entity, a "Lower-tier PFIC"), you would be deemed to own a proportionate amount (by value) of the ordinary shares of each Lower-tier PFIC and would be subject to U.S. federal income tax according to the rules described in the subsequent paragraph on (i) certain distributions by a Lower-tier PFIC and (ii) dispositions of shares of Lower-tier PFICs, in each case as if you held your proportionate share of these shares directly, even though you will not receive the proceeds of those distributions or dispositions.

Generally, if we are a PFIC for any taxable year during which you own ADSs or ordinary shares, gain recognized upon a disposition (including, under certain circumstances, a pledge) of ADSs or ordinary shares will be allocated ratably over your holding period for the ADSs or ordinary shares. The amounts allocated to the taxable year of disposition and to years before we became a PFIC will be taxed as ordinary income. The amount allocated to each other taxable year will be subject to tax at the highest rate in effect for that taxable year for individuals or corporations, as appropriate, and an interest charge will be imposed on the resulting tax liability for each relevant taxable year. Further, to the extent that any distribution received by you on your ADSs or ordinary shares exceeds 125% of the

average of the annual distributions on the ADSs or ordinary shares received during the preceding three years or your holding period, whichever is shorter, that distribution will be subject to taxation in the same manner.

If we are a PFIC for any taxable year during which you own ADSs or ordinary shares, we will generally continue to be treated as a PFIC with respect to you for all succeeding years during which you own ADSs or ordinary shares, even if we cease to meet the threshold requirements for PFIC status. If we are a PFIC for any taxable year but cease to be PFIC for subsequent years, you should consult your tax advisor regarding the availability of a "deemed sale" election that would allow you to eliminate the continuing PFIC status under certain circumstances.

Alternatively, if we are a PFIC and if our ADSs or ordinary shares are "regularly traded" on a "qualified exchange," you may be able to make a mark-to-market election that would result in tax treatment different from the general tax treatment described in the preceding paragraphs. Our ADSs will be treated as "regularly traded" in any calendar year in which more than a *de minimis* quantity of the ADSs, as the case may be, are traded on a qualified exchange on at least 15 days during each calendar quarter. The Nasdaq Global Market, on which the ADSs expected to be listed, is a qualified exchange for this purpose. If you make the mark-to-market election, you generally will recognize as ordinary income any excess of the fair market value of the ADSs at the end of each taxable year over their adjusted tax basis, and will recognize an ordinary loss in respect of any excess of the adjusted tax basis of the ADSs over their fair market value at the end of the taxable year (but only to the extent of the net amount of income previously included as a result of the mark-to-market election). If you make the election, your tax basis in the ADSs will be adjusted to reflect these income or loss amounts. Any gain recognized on the sale or other disposition of ADSs in a year in which we are a PFIC will be treated as ordinary income and any loss will be treated as an ordinary loss (but only to the extent of the net amount of income previously included as a result of the mark-to-market election) and any remaining loss will be capital loss. There is no provision under U.S. federal income tax law that provides for a right to make a mark-to-market election with respect to any lower-tier PFICs that are not publicly traded.

We do not intend to provide the information that would otherwise enable you to make a "qualified electing fund election," which would result in alternate treatment if we were a PFIC for any taxable year.

If you own ADSs or ordinary shares during any year in which we are a PFIC, you generally will be required to file annual reports on Internal Revenue Service Form 8621 (or any successor form) with respect to us. Additionally, if we are a PFIC for the taxable year in which we paid a dividend or the prior taxable year, the favorable rate discussed above with respect to dividends paid to certain non-corporate U.S. Holders would not apply. You should consult your tax adviser regarding our PFIC status for any taxable year and the potential application of the PFIC rules.

Information Reporting and Backup Withholding

Payments of dividends and sales proceeds from the sale or exchange of our ADSs or ordinary shares that are made within the United States or through certain U.S.-related financial intermediaries generally are subject to information reporting, and may be subject to backup withholding, unless (i) you are a corporation or other exempt recipient or (ii) in the case of backup withholding, you provide a correct taxpayer identification number and certify that you are not subject to backup withholding. Backup withholding is not an additional tax. The amount of any backup withholding from a payment to you will be allowed as a credit against your U.S. federal income tax liability and may entitle you to a refund, provided that the required information is timely furnished to the Internal Revenue Service.

Certain U.S. Holders who are individuals (and certain specified entities) may be required to report information relating to their ownership of ordinary shares, or non-U.S. accounts through which ADSs or ordinary shares are held. You should consult your tax adviser regarding your reporting obligations with respect to the ADSs or ordinary shares.

UNDERWRITING

We and the underwriters named below have entered into an underwriting agreement with respect to the ADSs being offered. Subject to certain conditions, each underwriter has severally agreed to purchase the number of ADSs indicated in the following table. Goldman Sachs (Asia) L.L.C., Morgan Stanley & Co. LLC and Jefferies LLC are the representatives of the underwriters.

<u>Underwriters</u>	<u>Number of ADSs</u>
Goldman Sachs (Asia) L.L.C.	
Morgan Stanley & Co. LLC	
Jefferies LLC	
China Renaissance Securities (Hong Kong) Limited	
Total:	

The underwriters are committed to take and pay for all of the ADSs being offered, if any are taken, other than the ADSs covered by the option described below unless and until this option is exercised.

Certain of the underwriters are expected to make offers and sales both inside and outside the United States through their respective selling agents. Any offers or sales in the United States will be conducted by broker-dealers registered with the SEC. Goldman Sachs (Asia) L.L.C. will offer ADSs in the United States through its SEC-registered broker-dealer affiliate in the United States, Goldman Sachs & Co. LLC. China Renaissance Securities (Hong Kong) Limited will offer ADSs in the United States through its SEC-registered broker-dealer affiliate in the United States, China Renaissance Securities (US) Inc.

The underwriters have an option to buy up to an additional ADSs from us to cover sales by the underwriters of a greater number of ADSs than the total number set forth in the table above. They may exercise that option for 30 days. If any ADSs are purchased pursuant to this option, the underwriters will severally purchase ADSs in approximately the same proportion as set forth in the table above.

The following table shows the per ADS and total underwriting discounts and commissions to be paid to the underwriters by us. Such amounts are shown assuming both no exercise and full exercise of the underwriters' option to purchase additional ADSs.

	<u>Paid by Us</u>	
	<u>No Exercise</u>	<u>Full Exercise</u>
Per ADS	US\$	US\$
Total	US\$	US\$

ADSs sold by the underwriters to the public will initially be offered at the initial public offering price set forth on the cover of this prospectus. Any ADSs sold by the underwriters to securities dealers may be sold at a discount of up to US\$ per ADS from the initial public offering price. After the initial offering of the ADSs, the representatives may change the offering price and the other selling terms. The offering of the ADSs by the underwriters is subject to receipt and acceptance and subject to the underwriters' right to reject any order in whole or in part.

[We, our officers, directors and existing shareholders have agreed with the underwriters, subject to certain exceptions, not to dispose of or hedge any of their ordinary shares or ADSs or securities convertible into or exchangeable for ordinary shares or ADSs during the period from the date of this prospectus continuing through the date 180 days after the date of this prospectus, except with the prior

written consent of the representatives. This agreement does not apply to any existing employee benefit plans. See "Shares Eligible for Future Sale" for a discussion of certain transfer restrictions.]

Prior to the offering, there has been no public market for the ADSs. The initial public offering price has been negotiated among the representatives and us. Among the factors to be considered in determining the initial public offering price of the ADSs, in addition to prevailing market conditions, will be our historical performance, estimates of our business potential and earnings prospects, an assessment of our management and the consideration of the above factors in relation to market valuation of companies in related businesses.

An application has been made to quote the ADSs on the Nasdaq Global Market under the symbol "ADAG".

In connection with the offering, the underwriters may purchase and sell ADSs in the open market. These transactions may include short sales, stabilizing transactions and purchases to cover positions created by short sales. Short sales involve the sale by the underwriters of a greater number of ADSs than they are required to purchase in the offering, and a short position represents the amount of such sales that have not been covered by subsequent purchases. A "covered short position" is a short position that is not greater than the amount of additional ADSs for which the underwriters' option described above may be exercised. The underwriters may cover any covered short position by either exercising their option to purchase additional ADSs or purchasing ADSs in the open market. In determining the source of ADSs to cover the covered short position, the underwriters will consider, among other things, the price of ADSs available for purchase in the open market as compared to the price at which they may purchase additional ADSs pursuant to the option described above. "Naked" short sales are any short sales that create a short position greater than the amount of additional ADSs for which the option described above may be exercised. The underwriters must cover any such naked short position by purchasing ADSs in the open market. A naked short position is more likely to be created if the underwriters are concerned that there may be downward pressure on the price of the ADSs in the open market after pricing that could adversely affect investors who purchase in the offering. Stabilizing transactions consist of various bids for or purchases of ADSs made by the underwriters in the open market prior to the completion of the offering.

The underwriters may also impose a penalty bid. This occurs when a particular underwriter repays to the underwriters a portion of the underwriting discount received by it because the representatives have repurchased ADSs sold by or for the account of such underwriter in stabilizing or short covering transactions.

Purchases to cover a short position and stabilizing transactions, as well as other purchases by the underwriters for their own accounts, may have the effect of preventing or retarding a decline in the market price of our ADSs, and together with the imposition of the penalty bid, may stabilize, maintain or otherwise affect the market price of the ADSs. As a result, the price of the ADSs may be higher than the price that otherwise might exist in the open market. The underwriters are not required to engage in these activities and may end any of these activities at any time. These transactions may be effected on the Nasdaq Global Market, in the over-the-counter market or otherwise.

We estimate that our share of the total expenses of the offering, excluding underwriting discounts and commissions, will be approximately US\$.

We have agreed to indemnify the several underwriters against certain liabilities, including liabilities under the Securities Act of 1933.

A prospectus in electronic format may be made available on the websites maintained by one or more underwriters, or selling group members, if any, participating in the offering. The underwriters may agree to allocate a number of ADSs to underwriters and selling group members for sale to their online brokerage account holders. Internet distributions will be allocated by the representatives to

underwriters and selling group members that may make internet distributions on the same basis as other allocations.

The underwriters and their respective affiliates are full service financial institutions engaged in various activities, which may include sales and trading, commercial and investment banking, advisory, investment management, investment research, principal investment, hedging, market making, brokerage and other financial and non-financial activities and services. Certain of the underwriters and their respective affiliates have provided, and may in the future provide, a variety of these services to the issuer and to persons and entities with relationships with the issuer, for which they received or will receive customary fees and expenses.

In the ordinary course of their various business activities, the underwriters and their respective affiliates, officers, directors and employees may purchase, sell or hold a broad array of investments and actively trade securities, derivatives, loans, commodities, currencies, credit default swaps and other financial instruments for their own account and for the accounts of their customers, and such investment and trading activities may involve or relate to assets, securities and/or instruments of the issuer (directly, as collateral securing other obligations or otherwise) and/or persons and entities with relationships with the issuer. The underwriters and their respective affiliates may also communicate independent investment recommendations, market color or trading ideas and/or publish or express independent research views in respect of such assets, securities or instruments and may at any time hold, or recommend to clients that they should acquire, long and/or short positions in such assets, securities and instruments.

The underwriters do not intend sales to discretionary accounts to exceed % of the total number of ADSs offered by them.

The address of Goldman Sachs (Asia) L.L.C. is 68th Floor, Cheung Kong Center, 2 Queen's Road Central, Hong Kong. The address of Morgan Stanley & Co. LLC is 1585 Broadway Avenue, New York, New York 10036, United States. The address of Jefferies LLC is 520 Madison Avenue, New York, New York 10022, United States. The address of China Renaissance Securities (Hong Kong) Limited is Units 8107-08, Level 81, International Commerce Centre, 1 Austin Road West, Kowloon, Hong Kong.

Selling Restrictions

No action may be taken in any jurisdiction other than the United States that would permit a public offering of the ADSs or the possession, circulation or distribution of this prospectus in any jurisdiction where action for that purpose is required. Accordingly, the ADSs may not be offered or sold, directly or indirectly, and neither the prospectus nor any other offering material or advertisements in connection with the ADSs may be distributed or published in or from any country or jurisdiction except under circumstances that will result in compliance with any applicable laws, rules and regulations of any such country or jurisdiction. Persons into whose possession this prospectus comes are advised to inform themselves about and to observe any restrictions relating to the offering and the distribution of this prospectus. This prospectus does not constitute an offer to sell or a solicitation of an offer to buy any securities offered by this prospectus in any jurisdiction in which such an offer or a solicitation is unlawful.

Australia

This document has not been lodged with the Australian Securities & Investments Commission and is only directed to certain categories of exempt persons. Accordingly, if you receive this document in Australia:

- (a) you confirm and warrant that you are either:
 - (i) "sophisticated investor" under section 708(8)(a) or (b) of the Corporations Act 2001 (Cth) of Australia, or the Corporations Act;
 - (ii) "sophisticated investor" under section 708(8)(c) or (d) of the Corporations Act and that you have provided an accountant's certificate to the company which complies with the requirements of section 708(8)(c)(i) or (ii) of the Corporations Act and related regulations before the offer has been made;
 - (iii) person associated with the company under section 708(12) of the Corporations Act; or
 - (iv) "professional investor" within the meaning of section 708(11)(a) or (b) of the Corporations Act;

and to the extent that you are unable to confirm or warrant that you are an exempt sophisticated investor, associated person or professional investor under the Corporations Act, any offer made to you under this document is void and incapable of acceptance;

- (b) you warrant and agree that you will not offer any of the ADSs issued to you pursuant to this document for resale in Australia within 12 months of those ADSs being issued unless any such resale offer is exempt from the requirement to issue a disclosure document under section 708 of the Corporations Act

Bermuda

The ADSs may be offered or sold in Bermuda only in compliance with the provisions of the Investment Business Act of 2003 of Bermuda which regulates the sale of securities in Bermuda.

Additionally, non-Bermudian persons (including companies) may not carry on or engage in any trade or business in Bermuda unless such persons are permitted to do so under applicable Bermuda legislation.

British Virgin Islands

The ADSs are not being, and may not be offered to the public or to any person in the British Virgin Islands for purchase or subscription by us or on our behalf. The ADSs may be offered to companies incorporated under the BVI Business Companies Act, 2004 (British Virgin Islands) (each a BVI Company), but only where the offer will be made to, and received by, the relevant BVI Company entirely outside of the British Virgin Islands.

This prospectus has not been, and will not be, registered with the Financial Services Commission of the British Virgin Islands. No registered prospectus has been or will be prepared in respect of the ADSs for the purposes of the Securities and Investment Business Act 2010, or SIBA, or the Public Issuers Code of the British Virgin Islands.

The ADSs may be offered to persons located in the British Virgin Islands who are "qualified investors" for the purposes of SIBA. Qualified investors include (i) certain entities which are regulated by the Financial Services Commission in the British Virgin Islands, including banks, insurance companies, licensees under SIBA and public, professional and private mutual funds; (ii) a company, any securities of which are listed on a recognized exchange; and (iii) persons defined as "professional investors" under SIBA, which is any person (a) whose ordinary business involves, whether for that

person's own account or the account of others, the acquisition or disposal of property of the same kind as the property, or a substantial part of our property; or (b) who has signed a declaration that he, whether individually or jointly with his spouse, has a net worth in excess of US\$1,000,000 and that he consents to being treated as a professional investor.

Canada

The ADSs may be sold only to purchasers purchasing, or deemed to be purchasing, as principal that are accredited investors, as defined in National Instrument 45-106 *Prospectus Exemptions* or subsection 73.3(1) of the *Securities Act* (Ontario), and are permitted clients, as defined in National Instrument 31-103 Registration Requirements, Exemptions and Ongoing Registrant Obligations. Any resale of the ADSs must be made in accordance with an exemption from, or in a transaction not subject to, the prospectus requirements of applicable securities laws.

Securities legislation in certain provinces or territories of Canada may provide a purchaser with remedies for rescission or damages if this prospectus (including any amendment thereto) contains a misrepresentation, provided that the remedies for rescission or damages are exercised by the purchaser within the time limit prescribed by the securities legislation of the purchaser's province or territory. The purchaser should refer to any applicable provisions of the securities legislation of the purchaser's province or territory for particulars of these rights or consult with a legal advisor.

Pursuant to section 3A.3 (or, in the case of securities issued or guaranteed by the government of a non-Canadian jurisdiction, section 3A.4) of National Instrument 33-105 Underwriting Conflicts, or NI 33-105, the underwriters are not required to comply with the disclosure requirements of NI 33-105 regarding underwriter conflicts of interest in connection with this offering.

Cayman Islands

This prospectus does not constitute an invitation or offer to the public in the Cayman Islands of the ADSs, whether by way of sale or subscription. ADSs or ordinary shares have not been offered or sold, and will not be offered or sold, directly or indirectly, in the Cayman Islands.

Dubai International Finance Center

This document relates to an Exempt Offer, as defined in the Offered Securities Rules module of the DFSA Rulebook, or the OSR, in accordance with the Offered Securities Rules of the Dubai Financial Services Authority. This document is intended for distribution only to Persons, as defined in the OSR, of a type specified in those rules. It must not be delivered to, or relied on by, any other Person. The Dubai Financial Services Authority has no responsibility for reviewing or verifying any documents in connection with Exempt Offers. The Dubai Financial Services Authority has not approved this document nor taken steps to verify the information set out in it, and has no responsibility for it. The ADSs to which this document relates may be illiquid and/or subject to restrictions on their resale. Prospective purchasers of the ADSs offered should conduct their own due diligence on the ADSs. If you do not understand the contents of this document you should consult an authorized financial adviser.

European Economic Area

In relation to each Member State of the European Economic Area (each an "EEA State"), no ADSs have been offered or will be offered pursuant to the offering contemplated by this prospectus to the public in that EEA State prior to the publication of a prospectus in relation to the ADSs which has been approved by the competent authority in that EEA State or, where appropriate, approved in another EEA State and notified to the competent authority in that EEA State, all in accordance with

the EU Prospectus Regulation, except that it may make an offer to the public in that EEA State of any ADSs at any time under the following exemptions under the EU Prospectus Regulation:

- (a) to any legal entity which is a qualified investor as defined under the EU Prospectus Regulation;
- (b) to fewer than 150 natural or legal persons (other than qualified investors as defined under the EU Prospectus Regulation), subject to obtaining the prior consent of underwriters for any such offer; or
- (c) in any other circumstances falling within Article 1(4) of the EU Prospectus Regulation, provided that no such offer of the ADSs shall require the Company or any underwriter to publish a prospectus pursuant to Article 3 of the EU Prospectus Regulation or supplement a prospectus pursuant to Article 23 of the EU Prospectus Regulation.

For the purposes of this provision, the expression an "offer to the public" in relation to the ADSs in any EEA State means the communication in any form and by any means of sufficient information on the terms of the offer and any ADSs to be offered so as to enable an investor to decide to purchase or subscribe for any ADSs, and the expression "**EU Prospectus Regulation**" means Regulation (EU) 2017/1129.

Hong Kong

The ADSs may not be offered or sold in Hong Kong by means of any document other than (i) in circumstances which do not constitute an offer to the public within the meaning of the Companies (Winding Up and Miscellaneous Provisions) Ordinance (Cap.32, Laws of Hong Kong), or (ii) to "professional investors" within the meaning of the Securities and Futures Ordinance (Cap.571, Laws of Hong Kong) and any rules made thereunder, or (iii) in other circumstances which do not result in the document being a "prospectus" within the meaning of the Companies (Winding Up and Miscellaneous Provisions) Ordinance (Cap.32, Laws of Hong Kong), and no advertisement, invitation or document relating to the ADSs may be issued or may be in the possession of any person for the purpose of issue (in each case whether in Hong Kong or elsewhere), which is directed at, or the contents of which are likely to be accessed or read by, the public in Hong Kong (except if permitted to do so under the laws of Hong Kong) other than with respect to ADSs which are or are intended to be disposed of only to persons outside Hong Kong or only to "professional investors" within the meaning of the Securities and Futures Ordinance (Cap.571, Laws of Hong Kong) and any rules made thereunder.

Israel

This document does not constitute a prospectus under the Israeli Securities Law, 5728-1968, and has not been filed with or approved by the Israel Securities Authority. In Israel, this prospectus may be distributed only to, and is directed only at, investors listed in the first addendum, or the Addendum, to the Israeli Securities Law, consisting primarily of joint investment in trust funds; provident funds; insurance companies; banks, portfolio managers, investment advisors, members of the Tel Aviv Stock Exchange Ltd., underwriters, each purchasing for their own account; venture capital funds; entities with equity in excess of NIS 50 million and "qualified individuals," each as defined in the Addendum (as it may be amended from time to time), collectively referred to as qualified investors. Qualified investors shall be required to submit written confirmation that they fall within the scope of the Addendum.

Japan

The ADSs have not been and will not be registered under the Financial Instruments and Exchange Law of Japan, and ADSs will not be offered or sold, directly or indirectly, in Japan or to, or for the benefit of, any resident of Japan (which term as used herein means any person resident in Japan,

including any corporation or other entity organized under the laws of Japan), or to others for re-offering or resale, directly or indirectly, in Japan or to a resident of Japan, except pursuant to any exemption from the registration requirements of, and otherwise in compliance with, the Financial Instruments and Exchange Law and any other applicable laws, regulations and ministerial guidelines of Japan.

Korea

The ADSs may not be offered, sold and delivered directly or indirectly, or offered or sold to any person for reoffering or resale, directly or indirectly, in Korea or to any resident of Korea except pursuant to the applicable laws and regulations of Korea, including the Korea Securities and Exchange Act and the Foreign Exchange Transaction Law and the decrees and regulations thereunder. The ADSs have not been registered with the Financial Services Commission of Korea for public offering in Korea. Furthermore, the ADSs may not be resold to Korean residents unless the purchaser of the ADSs complies with all applicable regulatory requirements (including but not limited to government approval requirements under the Foreign Exchange Transaction Law and its subordinate decrees and regulations) in connection with the purchase of the ADSs.

Kuwait

Unless all necessary approvals from the Kuwait Ministry of Commerce and Industry required by Law No. 31/1990 "Regulating the Negotiation of Securities and Establishment of Investment Funds," its Executive Regulations and the various Ministerial Orders issued pursuant thereto or in connection therewith, have been given in relation to the marketing and sale of the ADSs, these may not be marketed, offered for sale, nor sold in the State of Kuwait. Neither this prospectus (including any related document), nor any of the information contained therein is intended to lead to the conclusion of any contract of whatsoever nature within Kuwait.

Malaysia

No prospectus or other offering material or document in connection with the offer and sale of the securities has been or will be registered with the Securities Commission of Malaysia, or Commission, for the Commission's approval pursuant to the Capital Markets and Services Act 2007. Accordingly, this prospectus and any other document or material in connection with the offer or sale, or invitation for subscription or purchase, of the securities may not be circulated or distributed, nor may the securities be offered or sold, or be made the subject of an invitation for subscription or purchase, whether directly or indirectly, to persons in Malaysia other than (i) a closed end fund approved by the Commission; (ii) a holder of a Capital Markets Services License; (iii) a person who acquires the securities as principal, if the offer is on terms that the securities may only be acquired at a consideration of not less than RM250,000 (or its equivalent in foreign currencies) for each transaction; (iv) an individual whose total net personal assets or total net joint assets with his or her spouse exceeds RM3 million (or its equivalent in foreign currencies), excluding the value of the primary residence of the individual; (v) an individual who has a gross annual income exceeding RM300,000 (or its equivalent in foreign currencies) per annum in the preceding twelve months; (vi) an individual who, jointly with his or her spouse, has a gross annual income of RM400,000 (or its equivalent in foreign currencies), per annum in the preceding twelve months; (vii) a corporation with total net assets exceeding RM10 million (or its equivalent in a foreign currencies) based on the last audited accounts; (viii) a partnership with total net assets exceeding RM10 million (or its equivalent in foreign currencies); (ix) a bank licensee or insurance licensee as defined in the Labuan Financial Services and Securities Act 2010; (x) an Islamic bank licensee or takaful licensee as defined in the Labuan Financial Services and Securities Act 2010; and (xi) any other person as may be specified by the Commission; provided that, in the each of the preceding categories (i) to (xi), the distribution of the securities is made by a holder of

a Capital Markets Services License who carries on the business of dealing in securities. The distribution in Malaysia of this prospectus is subject to Malaysian laws. This prospectus does not constitute and may not be used for the purpose of public offering or an issue, offer for subscription or purchase, invitation to subscribe for or purchase any securities requiring the registration of a prospectus with the Commission under the Capital Markets and Services Act 2007.

People's Republic of China

This prospectus has not been and will not be circulated or distributed in China, and ADSs may not be offered or sold, and will not be offered or sold to any person for re-offering or resale, directly or indirectly, to any PRC resident except pursuant to applicable PRC laws and regulations.

Qatar

In the State of Qatar, the offer contained herein is made on an exclusive basis to the specifically intended recipient thereof, upon that person's request and initiative, for personal use only and shall in no way be construed as a general offer for the sale of securities to the public or an attempt to do business as a bank, an investment company or otherwise in the State of Qatar. This prospectus and the underlying securities have not been approved or licensed by the Qatar Central Bank or the Qatar Financial Centre Regulatory Authority or any other regulator in the State of Qatar. The information contained in this prospectus shall only be shared with any third parties in Qatar on a need to know basis for the purpose of evaluating the contained offer. Any distribution of this prospectus by the recipient to third parties in Qatar beyond the terms hereof is not permitted and shall be at the liability of such recipient.

Saudi Arabia

This prospectus may not be distributed in the Kingdom of Saudi Arabia except to such persons as are permitted under the Offers of Securities Regulations issued by the Capital Market Authority. The Capital Market Authority does not make any representation as to the accuracy or completeness of this prospectus, and expressly disclaims any liability whatsoever for any loss arising from, or incurred in reliance upon, any part of this prospectus. Prospective purchasers of the securities offered hereby should conduct their own due diligence on the accuracy of the information relating to the securities. If you do not understand the contents of this prospectus you should consult an authorized financial adviser.

Singapore

This prospectus has not been registered as a prospectus with the Monetary Authority of Singapore. Accordingly, this prospectus and any other document or material in connection with the offer or sale, or invitation for subscription or purchase, of our ADSs may not be circulated or distributed, nor may our ADSs be offered or sold, or be made the subject of an invitation for subscription or purchase, whether directly or indirectly, to persons in Singapore other than (i) to an institutional investor under Section 274 of the Securities and Futures Act, Chapter 289 of Singapore, or the SFA, (ii) to a relevant person or any person pursuant to Section 275(1A), and in accordance with the conditions specified in Section 275 of the SFA, and in accordance with the conditions specified in Section 275 of the SFA or (iii) otherwise pursuant to, and in accordance with the conditions of, any other applicable provision of the SFA, in each case subject to compliance with conditions set forth in the SFA.

Where our ADSs are subscribed or purchased under Section 275 by a relevant person which is: (a) a corporation (which is not an accredited investor as defined in Section 4A of the SFA) the sole business of which is to hold investments and the entire share capital of which is owned by one or more individuals, each of whom is an accredited investor; or (b) a trust (where the trustee is not an

accredited investor) whose sole purpose is to hold investments and each beneficiary of the trust is an individual who is an accredited investor; shares, debentures and units of shares and debentures of that corporation or the beneficiaries' rights and interest (howsoever described) in that trust shall not be transferred within six months after that corporation or that trust has acquired the ADSs under Section 275 of the SFA, except: (1) to an institutional investor (for corporations under Section 274 of the SFA) or to a relevant person defined in Section 275(2) of the SFA, or to any person pursuant to an offer that is made on terms that such shares, debentures and units of shares and debentures of that corporation or such rights and interest in that trust are acquired at a consideration of not less than US\$200,000 (or its equivalent in a foreign currency) for each transaction, whether such amount is to be paid for in cash or by exchange of securities or other assets, and further for corporations, in accordance with the conditions, specified in Section 275 of the SFA; (2) where no consideration is or will be given for the transfer; or (3) where the transfer is by operation of law.

South Africa

Due to restrictions under the securities laws of South Africa, the ADSs are not offered, and the offer shall not be transferred, sold, renounced or delivered, in South Africa or to a person with an address in South Africa, unless one or other of the following exemptions applies:

- (a) the offer, transfer, sale, renunciation or delivery is to:
 - (i) persons whose ordinary business is to deal in securities, as principal or agent;
 - (ii) the South African Public Investment Corporation;
 - (iii) persons or entities regulated by the Reserve Bank of South Africa;
 - (iv) authorized financial service providers under South African law;
 - (v) financial institutions recognized as such under South African law;
 - (vi) a wholly-owned subsidiary of any person or entity contemplated in (c), (d) or (e), acting as agent in the capacity of an authorized portfolio manager for a pension fund or collective investment scheme (in each case duly registered as such under South African law); or
 - (vii) any combination of the person in (i) to (vi); or

- (b) the total contemplated acquisition cost of the securities, for any single addressee acting as principal is equal to or greater than ZAR1,000,000.

No "offer to the public" (as such term is defined in the South African Companies Act, No. 71 of 2008 (as amended or re-enacted) (the "South African Companies Act")) in South Africa is being made in connection with the issue of the ADSs. Accordingly, this document does not, nor is it intended to, constitute a "registered prospectus" (as that term is defined in the South African Companies Act) prepared and registered under the South African Companies Act and has not been approved by, and/or filed with, the South African Companies and Intellectual Property Commission or any other regulatory authority in South Africa. Any issue or offering of the ADSs in South Africa constitutes an offer of the ADSs in South Africa for subscription or sale in South Africa only to persons who fall within the exemption from "offers to the public" set out in section 96(1)(a) of the South African Companies Act. Accordingly, this document must not be acted on or relied on by persons in South Africa who do not fall within section 96(1)(a) of the South African Companies Act (such persons being referred to as "SA Relevant Persons"). Any investment or investment activity to which this document relates is available in South Africa only to SA Relevant Persons and will be engaged in South Africa only with SA relevant persons.

Switzerland

The ADSs will not be publicly offered in Switzerland and will not be listed on the SIX Swiss Exchange, or SIX, or on any other stock exchange or regulated trading facility in Switzerland. This prospectus has been prepared without regard to the disclosure standards for issuance prospectuses under art. 652a or art. 1156 of the Swiss Code of Obligations or the disclosure standards for listing prospectuses under art. 27 ff. of the SIX Listing Rules or the listing rules of any other stock exchange or regulated trading facility in Switzerland.

Neither this prospectus nor any other offering or marketing material relating to our company or the ADSs have been or will be filed with or approved by any Swiss regulatory authority. In particular, this prospectus will not be filed with, and the offer of the ADSs will not be supervised by, the Swiss Financial Market Supervisory Authority, and the offer of the ADSs has not been and will not be authorized under the Swiss Federal Act on Collective Investment Schemes (the "CISA"). The investor protection afforded to acquirers of interests in collective investment schemes under the CISA does not extend to acquirers of the ADSs.

Taiwan

The ADSs have not been and will not be registered or filed with, or approved by, the Financial Supervisory Commission of Taiwan pursuant to relevant securities laws and regulations and may not be offered or sold in Taiwan through a public offering or in circumstances which constitute an offer within the meaning of the Securities and Exchange Act of Taiwan or relevant laws and regulations that require a registration, filing or approval of the Financial Supervisory Commission of Taiwan. No person or entity in Taiwan has been authorized to offer or sell the ADSs in Taiwan.

United Arab Emirates

This prospectus is not intended to constitute an offer, sale or delivery of shares or other securities under the laws of the United Arab Emirates, or the UAE. The ADSs have not been and will not be registered under Federal Law No. 4 of 2000 Concerning the Emirates Securities and Commodities Authority and the Emirates Security and Commodity Exchange, or with the UAE Central Bank, the Dubai Financial Market, the Abu Dhabi Securities Market or with any other UAE exchange.

The offering, the ADSs and interests therein have not been approved or licensed by the UAE Central Bank or any other relevant licensing authorities in the UAE, and do not constitute a public offer of securities in the UAE in accordance with the Commercial Companies Law, Federal Law No. 8 of 1984 (as amended) or otherwise.

In relation to its use in the UAE, this prospectus is strictly private and confidential and is being distributed to a limited number of investors and must not be provided to any person other than the original recipient, and may not be reproduced or used for any other purpose. The interests in the ADSs may not be offered or sold directly or indirectly to the public in the UAE.

United Kingdom

In relation to the United Kingdom, no ADSs have been offered or will be offered pursuant to the offering contemplated by this prospectus to the public in the United Kingdom prior to the publication of a prospectus in relation to the ADSs which has been approved by the Financial Conduct Authority in accordance with the UK Prospectus Regulation, except that it may make an offer to the public in the United Kingdom of any ADSs at any time under the following exemptions under the UK Prospectus Regulation:

- (a) to any legal entity which is a qualified investor as defined under the UK Prospectus Regulation;

- (b) to fewer than 150 natural or legal persons (other than qualified investors as defined under the UK Prospectus Regulation), subject to obtaining the prior consent of underwriters for any such offer; or
- (c) in any other circumstances falling within Article 1(4) of the UK Prospectus Regulation,

provided that no such offer of the ADSs shall require the Company or any underwriter to publish a prospectus pursuant to Article 3 of the UK Prospectus Regulation or supplement a prospectus pursuant to Article 23 of the UK Prospectus Regulation.

In the United Kingdom, the offer is only addressed to, and is directed only at, "qualified investors" within the meaning of Article 2(e) of the UK Prospectus Regulation, who are also (i) persons having professional experience in matters relating to investments who fall within the definition of "investment professionals" in Article 19(5) of the Financial Services and Markets Act 2000 (Financial Promotion) Order 2005 (the "**Order**"); (ii) high net worth bodies corporate, unincorporated associations and partnerships and trustees of high value trusts as described in Article 49(2) of the Order; or (iii) persons to whom it may otherwise lawfully be communicated (all such persons being referred to as "**relevant persons**"). This document must not be acted on or relied on by persons who are not relevant persons. Any investment or investment activity to which this document relates is available only to relevant persons and will be engaged in only with relevant persons.

For the purposes of this provision, the expression an "offer to the public" in relation to the ADSs in the United Kingdom means the communication in any form and by any means of sufficient information on the terms of the offer and any ADSs to be offered so as to enable an investor to decide to purchase or subscribe for any ADSs, and the expression "**UK Prospectus Regulation**" means the UK version of Regulation (EU) No 2017/1129 as amended by The Prospectus (Amendment etc.) (EU Exit) Regulations 2019, which is part of UK law by virtue of the European Union (Withdrawal) Act 2018.

EXPENSES RELATING TO THIS OFFERING

Set forth below is an itemization of the total expenses, excluding underwriting discounts and commissions, that we expect to incur in connection with this offering. With the exception of the SEC registration fee, the FINRA filing fee and the Nasdaq listing fee, all amounts are estimates.

SEC Registration Fee	US\$
Nasdaq Listing Fee	US\$
FINRA Filing Fee	US\$
Printing and Engraving Expenses	US\$
Legal Fees and Expenses	US\$
Accounting Fees and Expenses	US\$
Miscellaneous	US\$
Total	<u>US\$</u>

LEGAL MATTERS

We are being represented by Davis Polk & Wardwell LLP with respect to certain legal matters of U.S. federal securities and New York state law. Certain legal matters with respect to U.S. federal and New York State law in connection with this offering will be passed upon for the underwriters by Latham & Watkins LLP. The validity of the ordinary shares represented by the ADSs offered in this offering and other certain legal matters as to Cayman Islands law will be passed upon for us by Walkers (Hong Kong). Legal matters as to PRC law will be passed upon for us by Tian Yuan Law Firm and for the underwriters by Commerce & Finance Law Offices. Davis Polk & Wardwell LLP may rely upon Walkers (Hong Kong) with respect to matters governed by Cayman Islands law and Tian Yuan Law Firm with respect to matters governed by PRC law. Latham & Watkins LLP may rely upon Commerce & Finance Law Offices with respect to matters governed by PRC law.

EXPERTS

The financial statements as of December 31, 2018 and 2019 and for the years then ended included in this Prospectus have been so included in reliance on the report of PricewaterhouseCoopers Zhong Tian LLP, an independent registered public accounting firm, given on the authority of said firm as experts in auditing and accounting.

The offices of PricewaterhouseCoopers Zhong Tian LLP are located at 11th Floor, PricewaterhouseCoopers Center, Link Square 2, 202 Hu Bin Road, Huangpu District, Shanghai 200021, the People's Republic of China.

WHERE YOU CAN FIND ADDITIONAL INFORMATION

We have filed with the SEC a registration statement (including amendments and exhibits to the registration statement) on Form F-1 under the Securities Act. This prospectus, which is part of the registration statement, does not contain all of the information set forth in the registration statement and the exhibits and schedules to the registration statement. For further information, we refer you to the registration statement and the exhibits and schedules filed as part of the registration statement. If a document has been filed as an exhibit to the registration statement, we refer you to the copy of the document that has been filed. Each statement in this prospectus relating to a document filed as an exhibit is qualified in all respects by the filed exhibit.

Upon completion of this offering, we will become subject to the informational requirements of the Exchange Act. Accordingly, we will be required to file reports and other information with the SEC, including annual reports on Form 20-F and reports on Form 6-K. The SEC maintains an Internet site at www.sec.gov that contains reports, proxy and information statements and other information we have filed electronically with the SEC.

As a foreign private issuer, we are exempt under the Exchange Act from, among other things, the rules prescribing the furnishing and content of proxy statements, and our executive officers, directors and principal shareholders are exempt from the reporting and short-swing profit recovery provisions contained in Section 16 of the Exchange Act. In addition, we will not be required under the Exchange Act to file periodic reports and financial statements with the SEC as frequently or as promptly as U.S. companies whose securities are registered under the Exchange Act.

ADAGENE INC.

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Report of Independent Registered Public Accounting Firm

To the Board of Directors and Shareholders of Adagene Inc.

Opinion on the Financial Statements

We have audited the accompanying consolidated balance sheets of Adagene Inc. and its subsidiaries (the "Company") as of December 31, 2019 and 2018, and the related consolidated statements of comprehensive loss, of changes in shareholders' deficit and of cash flows for the years then ended, including the related notes (collectively referred to as the "consolidated financial statements"). In our opinion, the consolidated financial statements present fairly, in all material respects, the financial position of the Company as of December 31, 2019 and 2018, and the results of its operations and its cash flows for the years then ended in conformity with accounting principles generally accepted in the United States of America.

Basis for Opinion

These consolidated financial statements are the responsibility of the Company's management. Our responsibility is to express an opinion on the Company's consolidated financial statements based on our audits. We are a public accounting firm registered with the Public Company Accounting Oversight Board (United States) (PCAOB) and are required to be independent with respect to the Company in accordance with the U.S. federal securities laws and the applicable rules and regulations of the Securities and Exchange Commission and the PCAOB.

We conducted our audits of these consolidated financial statements in accordance with the standards of the PCAOB. Those standards require that we plan and perform the audit to obtain reasonable assurance about whether the consolidated financial statements are free of material misstatement, whether due to error or fraud.

Our audits included performing procedures to assess the risks of material misstatement of the consolidated financial statements, whether due to error or fraud, and performing procedures that respond to those risks. Such procedures included examining, on a test basis, evidence regarding the amounts and disclosures in the consolidated financial statements. Our audits also included evaluating the accounting principles used and significant estimates made by management, as well as evaluating the overall presentation of the consolidated financial statements. We believe that our audits provide a reasonable basis for our opinion.

/s/ PricewaterhouseCoopers Zhong Tian LLP

Shanghai, the People's Republic of China
September 22, 2020

We have served as the Company's auditor since 2020.

ADAGENE INC.

CONSOLIDATED BALANCE SHEETS

AS OF DECEMBER 31, 2018 AND 2019

	Notes	As of December 31,	
		2018	2019
		US\$	US\$
ASSETS			
Current assets:			
Cash and cash equivalents		16,058,455	92,532,788
Short-term investments		33,000,000	8,000,000
Accounts receivable, net	3	—	480,000
Amounts due from related parties	14	1,170,029	1,433,186
Prepayments and other current assets	4	1,588,489	1,476,973
Total current assets		51,816,973	103,922,947
Property, equipment and software, net	5	2,550,756	1,879,325
Other non-current assets		49,187	87,227
TOTAL ASSETS		54,416,916	105,889,499
LIABILITIES, MEZZANINE EQUITY AND SHAREHOLDERS' DEFICIT			
Current liabilities:			
Accounts payable		558,977	712,714
Contract liabilities		—	993,378
Amounts due to related parties	14	3,674,248	1,895,779
Accruals and other current liabilities	6	2,574,441	2,540,164
Warrant liabilities	8	1,207,415	—
Short-term borrowings	7	2,331,274	716,723
Current portion of long-term borrowings	7	—	322,525
Total current liabilities		10,346,355	7,181,283
Long-term borrowings	7	—	1,515,868
Other non-current liabilities		142,114	—
TOTAL LIABILITIES		10,488,469	8,697,151
Commitments and contingencies	15		
LIABILITIES, MEZZANINE EQUITY AND SHAREHOLDERS' DEFICIT (CONTINUED)			
Mezzanine equity:			
Series A-1 convertible redeemable preferred shares (par value of US\$0.0001 per share; 5,473,957 and 5,473,957 shares authorized, issued and outstanding as of December 31, 2018 and 2019, respectively)		5,473,957	5,473,957
Series A-2 convertible redeemable preferred shares (par value of US\$0.0001 per share; 2,370,414 and 2,370,414 shares authorized, issued and outstanding as of December 31, 2018 and 2019, respectively)		3,000,000	3,000,000
Series B convertible redeemable preferred shares (par value of US\$0.0001 per share; 7,494,537 and 7,494,537 shares authorized, issued and outstanding as of December 31, 2018 and 2019, respectively)		27,999,995	27,999,995
Series C-1 convertible redeemable preferred shares (par value of US\$0.0001 per share; 5,597,354 and 5,597,354 shares authorized, issued and outstanding as of December 31, 2018 and 2019, respectively)		48,481,159	48,727,343
Series C-2 convertible redeemable preferred shares (par value of US\$0.0001 per share; nil and 1,861,121 shares authorized, issued and outstanding as of December 31, 2018 and 2019, respectively)		—	18,999,999
Series C-3 convertible redeemable preferred shares (par value of US\$0.0001 per share; nil and 4,452,441 shares authorized, issued and outstanding as of December 31, 2018 and 2019, respectively)		—	50,000,000
Total mezzanine equity		84,955,111	154,201,294
Shareholders' deficit:			
Ordinary shares (par value of US\$0.0001 per share; 500,000,000 and 500,000,000 shares authorized; 15,159,136 and 15,193,136 shares issued and outstanding as of December 31, 2018 and 2019, respectively)		1,516	1,519
Subscriptions receivable from shareholders		(197,068)	(197,068)
Additional paid-in capital		6,405,318	6,789,542
Accumulated other comprehensive loss		(410,693)	(344,894)
Accumulated deficit		(46,825,737)	(63,258,045)
Total shareholders' deficit		(41,026,664)	(57,008,946)
TOTAL LIABILITIES, MEZZANINE EQUITY AND SHAREHOLDERS' DEFICIT		54,416,916	105,889,499

The accompanying notes are an integral part of these consolidated financial statements.

ADAGENE INC.
CONSOLIDATED STATEMENTS OF COMPREHENSIVE LOSS
FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

	Notes	For the years ended December 31,	
		2018 US\$	2019 US\$
Revenues			
Licensing revenue	10	1,511,168	480,000
Expenses			
Research and development expenses		(16,080,560)	(16,211,750)
Third parties		(8,575,969)	(10,507,444)
Related parties		(7,504,591)	(5,704,306)
Administrative expenses		(2,765,134)	(3,437,900)
Loss from operations		(17,334,526)	(19,169,650)
Interest income		710,711	922,680
Interest expense		(91,085)	(138,096)
Other income		901,713	723,476
Foreign exchange gain, net		12,698	21,867
Change in fair value of warrant liabilities		534,305	1,207,415
Loss before income tax		(15,266,184)	(16,432,308)
Income tax expense	11	—	—
Net loss attributable to Adagene Inc.'s shareholders		(15,266,184)	(16,432,308)
Other comprehensive income (loss)			
Foreign currency translation adjustments, net of nil tax		(11,288)	65,799
Total comprehensive loss attributable to Adagene Inc.'s shareholders		(15,277,472)	(16,366,509)
Net loss attributable to Adagene Inc.'s shareholders		(15,266,184)	(16,432,308)
Deemed contribution from convertible redeemable preferred shareholders	8	1,186,187	—
Accretion of convertible redeemable preferred shares to redemption value	8	(222,846)	(246,184)
Net loss attributable to ordinary shareholders		(14,302,843)	(16,678,492)
Weighted average number of ordinary shares used in per share calculation:			
—Basic	12	15,159,136	15,178,232
—Diluted	12	15,159,136	15,178,232
Net loss per ordinary share			
—Basic	12	(0.94)	(1.10)
—Diluted	12	(0.94)	(1.10)

The accompanying notes are an integral part of these consolidated financial statements.

ADAGENE INC.

CONSOLIDATED STATEMENTS OF CHANGES IN SHAREHOLDERS' DEFICIT

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

	Ordinary shares		Subscriptions receivable from shareholders	Additional paid-in capital	Accumulated other comprehensive loss	Accumulated deficit	Total shareholders' deficit
	Number of shares	Amount					
		US\$	US\$	US\$	US\$	US\$	US\$
Balance as of January 1, 2018	15,159,136	1,516	(197,068)	7,687,811	(399,405)	(32,745,740)	(25,652,886)
Net loss	—	—	—	—	—	(15,266,184)	(15,266,184)
Other comprehensive loss	—	—	—	—	(11,288)	—	(11,288)
Share-based compensation	—	—	—	126,540	—	—	126,540
Modification of convertible redeemable preferred shares	—	—	—	(1,186,187)	—	1,186,187	—
Accretion of convertible redeemable preferred shares to redemption value	—	—	—	(222,846)	—	—	(222,846)
Balance as of December 31, 2018	15,159,136	1,516	(197,068)	6,405,318	(410,693)	(46,825,737)	(41,026,664)
Net loss	—	—	—	—	—	(16,432,308)	(16,432,308)
Other comprehensive income	—	—	—	—	65,799	—	65,799
Exercise of share options (Note 9)	34,000	3	—	18,697	—	—	18,700
Share-based compensation	—	—	—	611,711	—	—	611,711
Accretion of convertible redeemable preferred shares to redemption value	—	—	—	(246,184)	—	—	(246,184)
Balance as of December 31, 2019	15,193,136	1,519	(197,068)	6,789,542	(344,894)	(63,258,045)	(57,008,946)

The accompanying notes are an integral part of these consolidated financial statements.

ADAGENE INC.

CONSOLIDATED STATEMENTS OF CASH FLOWS

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

	For the years ended	
	December 31,	
	2018	2019
	US\$	US\$
Cash flows from operating activities:		
Net loss	(15,266,184)	(16,432,308)
Adjustments to reconcile net loss to net cash used in operating activities:		
Depreciation and amortization	909,002	816,686
Gain on disposal of property, equipment and software	(846)	(1,841)
Share-based compensation	126,540	611,711
Change in fair value of warrant liabilities	(534,305)	(1,207,415)
Foreign exchange gain, net	(12,698)	(21,867)
Changes in operating assets and liabilities:		
Accounts receivable, net	—	(480,000)
Prepayments and other current assets	(1,337,063)	111,516
Amount due from related parties	268,671	(704,253)
Other non-current assets	(28,808)	(38,040)
Accounts payable	389,124	153,737
Contract liabilities	—	993,378
Amount due to related parties	2,135,397	(1,778,469)
Accruals and other current liabilities	(913,495)	(34,277)
Other non-current liabilities	—	(142,114)
Net cash used in operating activities	(14,264,665)	(18,153,556)
Cash flows from investing activities:		
Placement of short-term investments	(58,000,000)	(19,000,000)
Withdrawal of short-term investments	29,000,000	44,000,000
Proceeds from disposal of property, equipment and software	5,166	7,697
Purchase of property, equipment and software	(514,703)	(151,829)
Net cash (used in) generated from investing activities	(29,509,537)	24,855,868
Cash flows from financing activities:		
Proceeds from borrowings	2,417,868	2,651,874
Proceeds from issuance of convertible redeemable preferred shares and warrants	50,000,033	68,999,999
Proceeds from exercise of share options	—	459,796
Repayment of borrowings	(1,360,051)	(2,417,192)
Net cash generated from financing activities	51,057,850	69,694,477
Effect of exchange rate on cash and cash equivalents	38,703	77,544
Net increase in cash and cash equivalents	7,322,351	76,474,333
Cash and cash equivalents at the beginning of year	8,736,104	16,058,455
Cash and cash equivalents at the end of year	16,058,455	92,532,788
Supplemental cashflow disclosures:		
Interest paid	91,085	142,058
Non-cash activities:		
Accretion of convertible redeemable preferred shares to redemption value	222,846	246,184
Deemed contribution from convertible redeemable preferred shareholders	1,186,187	—

The accompanying notes are an integral part of these consolidated financial statements.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

1. ORGANIZATION AND BASIS OF PRESENTATION

Adagene Inc. (the "Company") is a limited liability company incorporated in the Cayman Islands on February 25, 2011. The Company, together with its subsidiaries (collectively, the "Group") are principally engaged in research, development and production of monoclonal antibody drugs for cancers.

As of December 31, 2019, the Company's principal subsidiaries are as follows:

Entity	Date of incorporation	Place of incorporation	Percentage of legal ownership by the Company	Principal activities
Adagene (Hong Kong) Limited	December 12, 2011	Hong Kong	100%	Investment holding
Adagene Incorporated	September 20, 2017	The United States of America	100%	Research and development of innovative medicines
Adagene (Suzhou) Limited	February 28, 2012	The People's Republic of China ("PRC" or "China")	100%	Research and development of innovative medicines
Adagene Australia PTY Ltd.	May 30, 2018	Australia	100%	Research and development of innovative medicines

2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES***Basis of presentation***

The accompany consolidated financial statements have been prepared in accordance with U.S. generally accepted accounting principles ("U.S. GAAP").

Principles of Consolidation

The consolidated financial statements of the Group include the financial statements of the Company and its subsidiaries. All significant intercompany balances and transactions have been eliminated upon consolidation.

Use of estimates

The preparation of consolidated financial statements in conformity with U.S. GAAP requires management to make estimates and assumptions that affect the reported amounts of assets and liabilities, disclosures of contingent assets and liabilities at the balance sheet dates and the reported amounts of expenses during the reporting periods. Significant estimates and assumptions reflected in the Group's consolidated financial statements include, but are not limited to, the useful lives and impairment of long-lived assets, tax valuation allowance, share-based compensation expenses and the fair value of warrant liabilities. Management bases the estimates on historical experience and various other assumptions that are believed to be reasonable, the results of which form the basis for making judgments about the carrying values of assets and liabilities. Actual results could materially differ from those estimates.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)

Foreign currency translation

The functional currency of the Company, Adagene (Hong Kong) Limited and Adagene Incorporated is the United States dollar ("US\$"). The functional currency of the Company's PRC subsidiary is Renminbi ("RMB"). The functional currency of the Company's Australia subsidiary is Australian dollar ("AU\$"). The determination of the respective functional currency is based on the criteria stated in Accounting Standard Codification ("ASC") 830, *Foreign Currency Matters*. The Company uses US\$ as its reporting currency. The financial statements of the Company's PRC and Australia subsidiaries are translated from the functional currency to the reporting currency.

Transactions denominated in foreign currencies are remeasured into the functional currency at the exchange rates quoted by the People's Bank of China (the "PBOC") prevailing on the transaction dates. Monetary assets and liabilities denominated in foreign currencies are re-measured at the exchange rates prevailing at the balance sheet date. Non-monetary items that are measured in terms of historical costs in foreign currency are re-measured using the exchange rates at the dates of the initial transactions. Exchange gains and losses are included in the consolidated statements of comprehensive loss.

Assets and liabilities are translated at the exchange rates at the balance sheet date, equity accounts are translated at historical exchange rates and revenues, expenses, gains and losses are translated using the average rate for the year. Translation adjustments are reported as accumulated comprehensive loss and are shown as a separate component of other comprehensive loss in the consolidated statements of comprehensive loss.

Cash and cash equivalents

Cash and cash equivalents primarily consist of cash and demand deposits which are highly liquid. The Group considers highly liquid investments that are readily convertible to known amounts of cash and with original maturities from the date of purchase of three months or less to be cash equivalents. All cash and cash equivalents are unrestricted as to withdrawal and use.

Short-term investments

Short-term investments are deposits at bank with maturities of greater than three months, but less than twelve months. Short-term investments are stated at cost, which approximates fair value. Interest earned is included in interest income.

Accounts receivable and allowance for doubtful accounts

Account receivable is recorded when the Group has an unconditional right to consideration. A right to consideration is unconditional if only the passage of time is required before payment of that consideration is due. Accounts receivable are carried at net realizable value. An allowance for doubtful accounts is recorded in the period when collection of the amount is no longer probable. In evaluating the collectability of receivable balances, the Group considers specific evidence including the aging of the receivable, the customer's payment history, its current credit-worthiness and other factors. Accounts receivable are written off when management determines a balance is uncollectable after all collection efforts have ceased.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)

Fair value measurements

The Group applies ASC 820, *Fair Value Measurements and Disclosures*. ASC 820 defines fair value, establishes a framework for measuring fair value and expands disclosures about fair value measurements. ASC 820 requires disclosures to be provided for fair value measurements. ASC 820 establishes a three-tier fair value hierarchy, which prioritizes the inputs used in measuring fair value as follows:

Level 1—Observable inputs that reflect quoted prices (unadjusted) for identical assets or liabilities in active markets.

Level 2—Includes other inputs that are directly or indirectly observable in the marketplace.

Level 3—Unobservable inputs which are supported by little or no market activity.

ASC 820 describes three main approaches to measuring the fair value of assets and liabilities: (1) market approach; (2) income approach; and (3) cost approach. The market approach uses prices and other relevant information generated from market transactions involving identical or comparable assets or liabilities. The income approach uses valuation techniques to convert future amounts to a single present value amount. The measurement is based on the value indicated by current market expectations about those future amounts. The cost approach is based on the amount that would currently be required to replace an asset.

The carrying amounts of cash and cash equivalent, short-term investments, accounts receivable, amounts due to related parties and other current assets, accounts payable, amounts due to related parties, accrued liabilities and other current liabilities and short-term borrowings approximate their fair values because of their generally short maturities. The carrying amount of long-term borrowings approximate their fair values since they bear interest rates which approximate market interest rates.

As more fully described in Note 8, the Group has issued warrants to purchase its preferred shares. The Group measured its warrant liabilities at fair value on a recurring basis. As the Group's warrants are not traded in an active market with readily observable prices, the Group uses significant unobservable inputs to measure the fair value of warrant liabilities. These instruments are categorized in the Level 3 valuation hierarchy based on the significance of unobservable factors in the overall fair value measurement.

The following table presents a reconciliation of all financial instruments measured at fair value on a recurring basis using Level 3 unobservable inputs:

	<u>Warrant liabilities</u>
	US\$
Initial recognition during the year ended December 31, 2018	1,741,720
Fair value change	(534,305)
Balance as of December 31, 2018	1,207,415
Fair value change	(1,207,415)
Balance as of December 31, 2019	—

ADAGENE INC.**NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)****FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019****2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)**

The Group did not transfer any assets or liabilities in or out of Level 3 during the years ended December 31, 2018 and 2019.

The Group had no financial assets and liabilities measured and recorded at fair value on a nonrecurring basis as of December 31, 2018 and 2019.

Property, equipment and software

Property and equipment and software are stated at cost less accumulated depreciation and amortization. Depreciation and amortization is computed using the straight-line method over the estimated useful lives of the assets as follows:

Category	Estimated Useful Life
Machinery and laboratory equipment	5 years
Vehicles	4 years
Furniture and tools	3 - 5 years
Electronic equipment	3 years
Computer software	3 - 5 years
Leasehold improvements	Lesser of lease terms or estimated useful lives of the assets

Repair and maintenance costs are charged to expense as incurred, whereas the cost of renewals and betterments that extend the useful lives of property, equipment and software are capitalized as additions to the related assets. Retirements, sales and disposals of assets are recorded by removing the cost and accumulated depreciation and amortization from the asset and accumulated depreciation and amortization accounts with any resulting gain or loss reflected in the consolidated statements of comprehensive loss.

Impairment of long-lived assets

The Group evaluates the recoverability of its long-lived assets, including fixed assets and intangible assets with finite lives, for impairment whenever events or changes in circumstances indicate that the carrying amount of an asset may not be fully recoverable. When these events occur, the Group measures impairment by comparing the carrying amount of the assets to the estimated undiscounted future cash flows expected to result from the use of the assets and their eventual disposition. If the sum of the expected undiscounted cash flows is less than the carrying amount of the assets, the Group recognizes an impairment loss based on the excess of the carrying amount of the assets over their fair value. Fair value is generally determined by discounting the cash flows expected to be generated by the assets, when the market prices are not readily available. The adjusted carrying amount of the assets is the new cost basis and is depreciated over the assets' remaining useful lives. Long-lived assets are grouped with other assets and liabilities at the lowest level for which identifiable cash flows are largely independent of the cash flows of other assets and liabilities.

No impairment loss was recorded for the years ended December 31, 2018 and 2019.

ADAGENE INC.**NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)****FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019****2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)*****Segment reporting***

In accordance with ASC 280, *Segment Reporting*, the Group's chief operating decision maker ("CODM") has been identified as the Chief Executive Officer. The Group's CODM reviews the consolidated results of operations when making decisions about allocating resources and assessing performance of the Group. The Group operates and manages its business as a single segment. No geographical segments are presented as substantially all of the Group's long-lived assets are located in the PRC.

Revenue recognition

At contract inception of collaboration and out-licensing arrangements, the Group analyzes its arrangements to assess whether they are within the scope of ASC 808, *Collaborative Arrangements* ("ASC 808") to determine whether such arrangements involve joint operating activities performed by parties that are both active participants in the activities and exposed to significant risks and rewards dependent on the commercial success of such activities. For collaboration arrangements within the scope of ASC 808 that contain multiple elements, the Group first determines which elements of the collaboration are deemed to be within the scope of ASC 808 and those that are reflective of a vendor-customer relationship and therefore within the scope of ASC 606. For elements of collaboration arrangements that are accounted for pursuant to ASC 808, an appropriate recognition method is determined and applied consistently. Under the criteria of Accounting Standard Codification ("ASC") 606, *Revenue from Contracts with Customers* (Topic 606) ("ASC 606"), the Group recognizes revenue to depict the transfer of control of promised goods or services to customers in an amount that reflects the consideration to which the entity expects to receive in exchange for those goods or services.

The Group adopted ASC 606 for all periods presented. To determine revenue recognition for arrangements that an entity determines are within the scope of ASC 606, the entity performs the following five steps: (i) identify the contract(s) with a customer; (ii) identify the performance obligations in the contract; (iii) determine the transaction price, including variable consideration, if any; (iv) allocate the transaction price to the performance obligations in the contract; and (v) recognize revenue when (or as) the entity satisfies a performance obligation. The Group only applies the five-step model to contracts when it is probable that the entity will collect substantially all the consideration to which it is entitled in exchange for the goods or services it transfers to the customer. The Group reviews the contract to determine which performance obligations are distinct and represent a promise to provide distinct goods or services or a series of distinct goods or services as defined by the standard. The Group recognizes as revenue the amount of the transaction price that is allocated to each performance obligation as and when that performance obligation is satisfied.

Licenses of Intellectual Property: Upfront non-refundable payments for licensing the Group's intellectual property are evaluated to determine if the license is distinct from the other performance obligations identified in the arrangement. For licenses determined to be distinct, the Group recognizes revenues from non-refundable, up-front fees allocated to the license at a point in time, when the transfer of control of the license to the licensee occurs and the licensee is able to use and benefit from the license.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)

Milestone Payments: At the inception of each arrangement that includes development, commercialization, and regulatory milestone payments, the Group evaluates whether the milestones are considered probable of being reached and to the extent that a significant reversal of cumulative revenue would not occur in future periods, estimates the amount to be included in the transaction price using the most likely amount method. The transaction price is then allocated to each performance obligation on a relative stand-alone selling price basis, for which the Group recognizes revenue as or when the performance obligations under the contract are satisfied. At the end of each subsequent reporting period, the Group re-evaluates the probability of achieving such development milestones and any related constraint, and if necessary, adjust the estimate of the overall transaction price. Any such adjustments are recorded on a cumulative catch-up basis, which would affect revenues and earnings in the period of adjustment.

Royalties: For arrangements that include sales-based royalties, including milestone payments based on the level of sales, and the license is deemed to be the predominant item to which the royalties relate, the Group recognizes revenue at the later of (i) when the related sales occur, or (ii) when the performance obligation to which some or all of the royalty has been allocated has been satisfied (or partially satisfied).

To date, no milestone payments or royalty payments were received. Substantially all of the Group's revenue has been derived from its out-licensing agreements with respect to licensed products such as DNA sequences, cell lines, etc., and such revenues are recognized when the customer obtains control of the licensed product, which occurs at a point in time, upon delivery to the customer.

Contract assets and contract liabilities

When a customer pays consideration before the Group transfers products or services, the Group records its obligation as a contract liability; When the Group satisfies its performance obligations by providing products or services to a customer before the customer pays consideration and before payment is due, the Group recognizes its rights to consideration as a contract asset.

Research and development expenses

Elements of research and development expenses primarily include (1) payroll and other related costs of personnel engaged in research and development activities, (2) costs related to pre-clinical testing of the Group's technologies under development and clinical trials such as payments to contract research organizations ("CRO") and contract manufacturing organizations ("CMO"), investigators and clinical trial sites that conduct the clinical studies; (3) costs to develop the product candidates, including raw materials and supplies, product testing, depreciation and amortization, and facility related expenses, (4) other research and development expenses. Research and development expenses are charged to expense as incurred when these expenditures relate to the Group's research and development services and have no alternative future uses. As of December 31, 2019, the Group has several ongoing clinical studies in various clinical trial stages. The contracts with CRO and CMO are generally cancellable, with notice, at the Group's option. The Group did not record any accrued expenses related to cancellation of CRO or CMO contracts as of December 31, 2019 as the Group did not have any plan to cancel the existing CRO or CMO contracts.

ADAGENE INC.**NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)****FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019****2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)*****Government subsidies***

Government subsidies primarily consist of financial subsidies received from provincial and local governments for operating a business in their jurisdictions and compliance with specific policies promoted by the governments. The Group's PRC based subsidiary received government subsidies from certain local government. The Group's government subsidies consist of specific subsidies and other subsidies. Specific subsidies are subsidies that the local government has set certain conditions for the subsidies. Other subsidies are the subsidies that the local government has not set any conditions and are not tied to future trends or performance of the Group, receipt of such subsidy income is not contingent upon any further actions or performance of the Group and the amounts do not have to be refunded under any circumstances. The Group recorded specific subsidies as other non-current liabilities when received and recognized as other income when the conditions are met. Other subsidies are recognized as other income upon receipt as further performance by the Group is not required.

Leases

Leases are classified at the inception date as either a capital lease or an operating lease. The Group assesses a lease to be a capital lease if any of the following conditions exists: a) ownership is transferred to the lessee by the end of the lease term, b) there is a bargain purchase option, c) the lease term is at least 75% of the property's estimated remaining economic life or d) the present value of the minimum lease payments at the beginning of the lease term is 90% or more of the fair value of the leased property to the lessor at the inception date. A capital lease is accounted for as if there was an acquisition of an asset and an incurrence of an obligation at the inception of the lease.

All other leases are accounted for as operating leases wherein rental payments are expensed on a straight-line basis over their respective lease terms. The Group leases certain office space under non-cancelable operating lease agreements. Certain lease agreements contain rent holidays. Rent holidays are considered in determining the straight-line rent expense to be recorded over the lease term. The lease term begins on the date of initial possession of the leased property for purpose of recognizing lease expense on straight-line basis over the term of the lease.

Comprehensive loss

Comprehensive loss is defined as the changes in equity of the Group during a period from transactions and other events and circumstances excluding transactions resulting from investments by shareholders and distributions to shareholders. Accumulated other comprehensive loss of the Group includes foreign currency translation adjustments related to the Group and its subsidiaries whose functional currency is not US\$.

Income taxes

The Group follows the liability method of accounting for income taxes in accordance with ASC 740, *Income Taxes* ("ASC 740"). Under this method, deferred tax assets and liabilities are determined based on the difference between the financial reporting and tax bases of assets and liabilities using enacted tax rates that will be in effect in the period in which the differences are expected to reverse. The Group records a valuation allowance to offset deferred tax assets if based on

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)

the weight of available evidence, it is more likely than not that some portion, or all, of the deferred tax assets will not be realized. The effect on deferred taxes of a change in tax rate is recognized in tax expense in the period that includes the enactment date of the change in tax rate.

The Group evaluates its uncertain tax positions using the provisions of ASC 740, which prescribes a recognition threshold that a tax position is required to meet before being recognized in the consolidated financial statements.

The Group recognizes in the consolidated financial statements the benefit of a tax position which is "more likely than not" to be sustained under examination based solely on the technical merits of the position assuming a review by tax authorities having all relevant information. Tax positions that meet the recognition threshold are measured using a cumulative probability approach, at the largest amount of tax benefit that has a greater than fifty percent likelihood of being realized upon settlement. It is the Group's policy to recognize interest and penalties related to unrecognized tax benefits, if any, as a component of income tax expense.

Borrowings

Borrowings are recognized initially at fair value, net of transaction costs incurred. Borrowings are subsequently stated at amortized cost; any difference between the proceeds (net of transaction costs) and the redemption value is recognized in the consolidated statements of comprehensive loss over the period of the borrowings using the effective interest method.

Share-based compensation

The Group applies ASC 718, *Compensation—Stock Compensation* ("ASC 718"), to account for its employee share-based payments awards granted to certain directors, executives and employees. Share options granted are classified as equity awards and are measured based on the grant date fair value of the equity instrument issued, and recognized as compensation costs using the straight-line method over the requisite service period, which is generally the vesting period of the share options, with a corresponding impact reflected in additional paid-in capital.

A change in any of the terms or conditions of share-based awards is accounted for as a modification of the awards. The Group calculates incremental compensation expense of a modification as the excess of the fair value of the modified awards over the fair value of the original awards immediately before its terms are modified at the modification date. For vested awards, the Group recognizes incremental compensation cost in the period when the modification occurs. For awards not being fully vested, the Group recognizes the sum of the incremental compensation expense and the remaining unrecognized compensation expense for the original awards over the remaining requisite service period after modification.

Net loss per share

In accordance with ASC 260, *Earnings Per Share*, basic net loss per share is computed by dividing net loss attributable to ordinary shareholders by the weighted average number of unrestricted ordinary shares outstanding during the year using the two-class method. Under the two-class method, net loss is allocated between ordinary shares and other participating securities based on dividends declared (or

ADAGENE INC.**NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)****FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019****2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)**

accumulated) and participating rights in undistributed earnings as if all the earnings for the reporting period had been distributed. The Company's convertible redeemable preferred shares are participating securities because they are entitled to receive dividends or distributions on an as converted basis. Diluted net loss per share is calculated by dividing net loss attributable to ordinary shareholders, as adjusted for the effect of dilutive ordinary equivalent shares, if any, by the weighted average number of ordinary and dilutive ordinary equivalent shares outstanding during the period. Ordinary equivalent shares include ordinary shares issuable upon the conversion of the convertible redeemable preferred shares using the if-converted method, and ordinary shares issuable upon the exercise of share options, using the treasury stock method. Ordinary share equivalents are excluded from the computation of diluted earnings per share if their effects are anti-dilutive. For the periods presented herein, the computation of basic net loss per share using the two-class method is not applicable as the Group is in a net loss position and the participating securities do not have contractual rights and obligations to share in the losses of the Group.

Employee defined contribution plan

As stipulated by the regulations of the PRC, full-time employees of the Group are entitled to staff welfare benefits including medical care, welfare subsidies, unemployment insurance and pension benefits through a PRC government-mandated multi-employer defined contribution plan. The Group is required to accrue for these benefits based on certain percentages of the qualified employees' salaries. The Group is required to make contributions to the plans out of the amounts accrued. The PRC government is responsible for the medical benefits and the pension liability to be paid to these employees and the Group's obligations are limited to the amounts contributed. The Group has no further payment obligations once the contributions have been paid. The Group recorded employee benefit expenses of US\$947,244 and US\$1,145,165 for the years ended December 31, 2018 and 2019, respectively.

Concentration of risks***Concentration of credit risk***

As of December 31, 2018 and 2019, the aggregate amount of cash and cash equivalents and short-term investments of US\$422,210 and US\$801,923 respectively, were held at major financial institutions located in the PRC, and US\$48,636,245 and US\$99,730,865, respectively, were deposited with major financial institutions located outside the PRC. Management believes that these financial institutions are of high credit quality and continually monitors the credit worthiness of these financial institutions.

Accounts receivable are typically unsecured and denominated in US\$ and are derived from revenues earned from customers. As of December 31, 2019, the accounts receivable balance is from one customer. The Group manages credit risk of accounts receivable through ongoing monitoring of the outstanding balances.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)

Concentration of suppliers

A significant portion of the Group's research and development services were purchased from its one supplier, who collectively accounted for 40.67% and 23.13% of the Group's total research and development services purchases for the years ended December 31, 2018 and 2019, respectively.

Business and economic risk

The Group believes that changes in any of the following areas could have a material adverse effect on the Group's future consolidated financial position, results of operations or cash flows: changes in the overall demand for services; competitive pressures due to new entrants; advances and new trends in new technologies and industry standards; changes in certain strategic relationships; regulatory considerations and risks associated with the Group's ability to attract employees necessary to support its growth. The Group's operations could also be adversely affected by significant political, regulatory, economic and social uncertainties in the PRC.

Foreign currency exchange rate risk

A significant portion of the Group's businesses are transacted in RMB, which is not a freely convertible currency. On January 1, 1994, the PRC government abolished the dual rate system and introduced a single rate of exchange as quoted daily by the People's Bank of China (the "PBOC"). However, the unification of the exchange rates does not imply that the RMB may be readily convertible into US\$ or other foreign currencies. All foreign exchange transactions continue to take place either through the PBOC or other banks authorized to buy and sell foreign currencies at the exchange rates quoted by the PBOC. Approval of foreign currency payments by the PBOC or other institutions requires submitting a payment application form together with suppliers' invoices, shipping documents and signed contracts.

From July 21, 2005, the RMB is permitted to fluctuate within a narrow and managed band against a basket of certain foreign currencies. For U.S. dollar against RMB, there was appreciation of approximately 5.7% and 1.3% in the years ended December 31, 2018 and 2019, respectively. It is difficult to predict how market forces or PRC or U.S. government policy may impact the exchange rate between the RMB and the U.S. dollar in the future.

The functional currency and the reporting currency of the Company are the US\$. However, most of the expenses of the Group are denominated in RMB. Any significant fluctuation of the valuation of RMB may materially affect the Group's cash flows, expenses, losses and financial position, and the value of any dividends payable on the American Depositary Shares in US\$.

Recently issued accounting pronouncements

The Group is an emerging growth company ("EGC") as defined by the Jumpstart Our Business Startups Act ("JOBS Act"). The JOBS Act provides that an EGC can take advantage of extended transition periods for complying with new or revised accounting standards. This allows an EGC to delay adoption of certain accounting standards until those standards would otherwise apply to private companies. The Group elected to take advantage of the extended transition periods. However, this election will not apply should the Group cease to be classified as an EGC.

ADAGENE INC.**NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)****FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019****2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)**

In February 2016, the FASB issued ASU No. 2016-02 ("ASU 2016-02"), Leases (Topic 842), which modifies lease accounting for lessees to increase transparency and comparability by recording lease assets and liabilities for operating leases and disclosing key information about leasing arrangements. In July 2018, the FASB issued ASU No. 2018-10 ("ASU 2018-10"), Codification Improvements to Topic 842, Leases, which clarifies certain aspects of the guidance issued in ASU 2016-02; and ASU No. 2018-11 ("ASU 2018-11"), Leases (Topic 842): Targeted Improvements, which provides entities with an additional (and optional) transition method to adopt the new leases standard. Under this new transition method, an entity initially applies the new leases standard at the adoption date and recognizes a cumulative-effect adjustment to the opening balance of retained earnings in the period of adoption. Consequently, an entity's reporting for the comparative periods presented in the financial statements in which it adopts the new leases standard will continue to be in accordance with current GAAP (Topic 840, Leases). In November 2019, the FASB issued ASU No. 2019-10, Financial Instruments—Credit Losses (Topic 326), Derivatives and Hedging (Topic 815), and Leases (Topic 842), Effective Dates ("ASU 2019-10"), which extends the adoption date for certain registrants. The updated guidance is effective for the Group for annual reporting periods beginning January 1, 2021 and interim periods within annual periods beginning January 1, 2022. Early adoption is permitted. The Group does not plan to early adopt the new lease standards and the Group expects that applying the ASU 2016-02 would materially increase the assets and liabilities due to the recognition of right-of-use assets and lease liabilities on its consolidated balance sheets, with an immaterial impact on its consolidated statements of comprehensive loss and consolidated statements of cash flows.

In June 2016, the FASB issued ASU No. 2016-13, Financial Instruments—Credit Losses (Topic 326): Measurement of Credit Losses on Financial Instruments ("ASU 2016-13"). ASU 2016-13 is intended to improve financial reporting by requiring timelier recording of credit losses on loans and other financial instruments held by financial institutions and other organizations. This ASU requires the measurement of all expected credit losses for financial assets held at the reporting date based on historical experience, current conditions, and reasonable and supportable forecasts. This ASU requires enhanced disclosures to help investors and other financial statement users better understand significant estimates and judgments used in estimating credit losses, as well as the credit quality and underwriting standards of the Group's portfolio. These disclosures include qualitative and quantitative requirements that provide additional information about the amounts recorded in the financial statements. In November 2019, the FASB issued ASU 2019-10, which extends the adoption date for certain registrants. The amendments in ASU 2016-13 are effective for fiscal years beginning after December 15, 2022, including interim periods within fiscal years beginning after December 15, 2023. The Group does not plan to early adopt ASU 2016-13 and is currently in the process of evaluating the impact of adoption of this guidance on its consolidated financial statements.

In June 2018, the FASB issued ASU No. 2018-07, Compensation—Stock Compensation (Topic 718): Improvements to nonemployee share-based payment accounting ("ASU 2018-07"). The amendments in this update expand the scope of Topic 718 to include share-based payment transactions for acquiring goods and services from nonemployees. The amendments are effective for fiscal years beginning after December 15, 2019, and interim periods within fiscal years beginning after December 15, 2020. Early adoption is permitted, but no earlier than an entity's adoption date of Topic 606. The Group adopted on January 1, 2018 this guidance which do not have a significant impact on the consolidated financial statements.

ADAGENE INC.**NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)****FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019****2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)**

In August 2018, the FASB issued ASU No. 2018-13, Fair Value Measurement (Topic 820): Disclosure Framework—Changes to the Disclosure Requirements for Fair Value Measurement ("ASU 2018-13"). ASU 2018-13 modifies the disclosure requirements for fair value measurements by removing, modifying, or adding certain disclosures. The amendments in ASU 2018-13 are effective for all entities for fiscal years, and interim periods within those fiscal years, beginning after December 15, 2019. The amendments on changes in unrealized gains and losses, the range and weighted average of significant unobservable inputs used to develop Level 3 fair value measurements, and the narrative description of measurement uncertainty should be applied prospectively for only the most recent interim or annual period presented in the initial fiscal year of adoption. All other amendments should be applied retrospectively to all periods presented upon their effective date. Early adoption is permitted upon issuance of ASU 2018-13. An entity is permitted to early adopt any removed or modified disclosures upon issuance of ASU 2018-13 and delay adoption of the additional disclosures until their effective date. The Group elected to early adopt this ASU and applied this guidance retrospectively to all periods presented. The impact of this ASU to the consolidated financial statements is immaterial.

In November 2018, the FASB issued ASU 2018-18, Collaborative Arrangements (Topic 808): Clarifying the Interaction between Topic 808 and Topic 606. This update clarifies that certain transactions between participants in a collaborative arrangement should be accounted for under ASC 606 when the counterparty is a customer and precludes an entity from presenting consideration from a transaction in a collaborative arrangement as revenue from contracts with customers if the counterparty is not a customer for that transaction. The update is effective in fiscal years beginning after December 15, 2021, and interim periods therein, and early adoption is permitted for entities that have adopted ASC 606. This guidance should be applied retrospectively to the date of initial application of Topic 606. The Group elected to early adopt this ASU and applied this guidance retrospectively to all periods presented. The impact of this ASU to the consolidated financial statements is immaterial.

In December 2019, the FASB issued ASU 2019-12, Income Taxes (Topic 740): Simplifying the Accounting for Income Taxes. This update simplifies the accounting for income taxes as part of the FASB's overall initiative to reduce complexity in accounting standards. The amendments include removal of certain exceptions to the general principles of ASC 740, *Income taxes*, and simplification in several other areas such as accounting for a franchise tax (or similar tax) that is partially based on income. The update is effective in fiscal years beginning after December 15, 2022, and interim periods therein, and early adoption is permitted. Certain amendments in this update should be applied retrospectively or modified retrospectively, all other amendments should be applied prospectively. The Group does not plan to early adopt ASU 2019-12 and is currently evaluating the impact on its financial statements of adopting this guidance.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

3. ACCOUNTS RECEIVABLE, NET

	As of December 31,	
	2018 US\$	2019 US\$
Accounts receivable	—	480,000
Allowance for doubtful accounts	—	—
	<u>—</u>	<u>480,000</u>

4. PREPAYMENTS AND OTHER CURRENT ASSETS

Prepayments and other current assets consist of the following:

	As of December 31,	
	2018 US\$	2019 US\$
Deposits(a)	970,407	970,394
Interest receivables	260,173	227,278
Prepayments	295,476	211,435
Others	62,433	67,866
	<u>1,588,489</u>	<u>1,476,973</u>

Note (a): The deposits represented the amounts that the Group paid to its CRO vendors for various outsourced research and development programs according to the terms of respective CRO agreement. The Group expects to recover the deposits when the programs end.

5. PROPERTY, EQUIPMENT AND SOFTWARE

Property, equipment and software consist of the following:

	As of December 31,	
	2018 US\$	2019 US\$
Machinery and laboratory equipment	3,408,277	3,462,343
Leasehold improvements	779,837	767,205
Electronic equipment	576,421	605,882
Furniture and tools	100,016	98,396
Vehicles	81,452	80,133
Software	64,761	71,056
Total property, equipment and software	5,010,764	5,085,015
Less: accumulated depreciation and amortization	(2,460,008)	(3,205,690)
Net book value	<u>2,550,756</u>	<u>1,879,325</u>

Depreciation and amortization expenses recognized for the years ended December 31, 2018 and 2019 were US\$909,002 and US\$816,686, respectively.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

6. ACCRUALS AND OTHER CURRENT LIABILITIES

Accrued liabilities and other current liabilities consist of the following:

	As of December 31,	
	2018	2019
	US\$	US\$
Payroll and related liabilities	2,230,898	2,370,523
Professional service fees	93,801	145,157
Utility and maintenance	102,120	4,595
Other taxes and surcharge	32,982	—
Others	114,640	19,889
	<u>2,574,441</u>	<u>2,540,164</u>

7. BORROWINGS

	As of December 31,	
	2018	2019
	US\$	US\$
Current		
Short-term borrowings:		
Bank loans	2,331,274	716,723
Current portion of long-term borrowings	—	322,525
Total current borrowings	<u>2,331,274</u>	<u>1,039,248</u>
Non-Current		
Long-term borrowings:		
Bank loans	—	1,515,868
Total non-current borrowings	<u>—</u>	<u>1,515,868</u>
Total borrowings	<u>2,331,274</u>	<u>2,555,116</u>

Short-term borrowings

In April 2017, the Group borrowed a loan with amount of RMB4,000,000 (equivalent to approximately US\$612,164) from Agricultural Bank of China Limited for a term of one year and at the interest rate of 4.79% per annum. The borrowing was guaranteed by Peter Luo, who is the Chairman, Chief Executive Officer and a principal shareholder of the Company. The borrowing was repaid in March 2018.

In May 2017, the Group borrowed a loan with the amount of RMB3,000,000 (equivalent to approximately US\$459,123) from Bank of Ningbo Co., Ltd. for a term of one year and at the interest rate of 4.79% per annum and the borrowing was repaid in May 2018. In June 2017, the Group borrowed a loan with the amount of RMB2,000,000 (equivalent to approximately US\$306,082) from Bank of Ningbo Co., Ltd. for a term of one year and at the interest rate of 4.79% per annum. The borrowing was repaid in June 2018.

ADAGENE INC.**NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)****FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019****7. BORROWINGS (Continued)**

In March 2018, the Group borrowed a loan with the amount of RMB3,000,000 (equivalent to approximately US\$437,114) from Bank of Jiangsu Co., Ltd. for a term of one year and at the interest rate of 5.22% per annum. The borrowing was repaid in March 2019. In June 2018, the Group borrowed a loan with the amount of RMB2,000,000 (equivalent to approximately US\$291,409) from Bank of Jiangsu Co., Ltd. for a term of one year and at the interest rate of 5.22% per annum. The borrowing was repaid in June 2019. These borrowings with Bank of Jiangsu Co., Ltd were guaranteed by Peter Luo and Kristine She. Peter Luo is the Chairman, Chief Executive Officer and a principal shareholder of the Company. Kristine She is one of the senior management personnel of the Company.

In May 2018, the Group borrowed a loan with the amount of RMB3,000,000 (equivalent to approximately US\$437,114) from Bank of Ningbo Co., Ltd. for a term of one year and at the interest rate of 5.00% per annum. The borrowing was repaid in May 2019. In June 2018, the Group borrowed a loan with the amount of RMB2,000,000 (equivalent to approximately US\$291,409) from Bank of Ningbo Co., Ltd. for a term of one year and at the interest rate of 5.00% per annum. The borrowing was repaid in June 2019.

In July 2018, the Group borrowed a loan with amount of RMB6,000,000 (equivalent to approximately US\$874,228) from Agricultural Bank of China Limited for a term of one year and at the interest rate of 5.22% per annum. The borrowing was guaranteed by Peter Luo, who is the Chairman, Chief Executive Officer and a principal shareholder of the Company. The borrowing was repaid in July 2019.

In September 2019, the Group borrowed a loan with amount of RMB5,000,000 (equivalent to approximately US\$716,723) from Bank of Ningbo Co., Ltd. for a term of one year and at the interest rate of 4.35% per annum.

Long-term borrowings

In February 2019, the Group borrowed a loan with amount of RMB7,500,000 (equivalent to approximately US\$1,075,084) from Shanghai Pudong Development Bank Co., Ltd. for a term of three years and at the interest rate of 5.46% per annum. The Group repaid RMB375,000 (equivalent to approximately US\$53,754) in August 2019. As of December 31, 2019, RMB1,250,000 (equivalent to approximately US\$179,181) repayable within twelve months for this agreement was classified as "Current portion of long-term borrowing".

In June 2019, the Group borrowed a loan with amount of RMB6,000,000 (equivalent to approximately US\$860,067) from Shanghai Pudong Development Bank Co., Ltd. for a term of three years and at the interest rate of 5.23% per annum. The Group repaid RMB300,000 (equivalent to approximately US\$43,004) in December 2019. As of December 31, 2019, RMB1,000,000 (equivalent to approximately US\$143,344) repayable within twelve months for this agreement was classified as "Current portion of long-term borrowing".

ADAGENE INC.**NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)****FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019****7. BORROWINGS (Continued)***Future maturities of short-term borrowings and long-term borrowings*

Future principal maturities of short-term borrowings and long-term borrowings as of December 31, 2019 are as followings:

	US\$
2020	1,039,248
2021	870,818
2022	645,050
	<u>2,555,116</u>

8. CONVERTIBLE REDEEMABLE PREFERRED SHARES AND WARRANTS

In November 2011, the Company issued convertible notes ("Series Pre-A Convertible Notes") to certain investors in the amount of 4,590,908. The notes carried a simple interest (non-compounding) of 6% per annum as set out in the note purchase agreement. All outstanding principal balance and accrued but unpaid interest of the notes should be automatically converted into the convertible redeemable preferred shares of the Company at a price no more than US\$1 per share.

In November 2014, the Company issued 5,473,957 Series A-1 convertible redeemable preferred shares ("Series A-1 Preferred Shares") to certain investors upon conversion of the Company's Series Pre-A convertible notes at a conversion price of US\$1 per share. Concurrently, the Company issued 2,370,414 Series A-2 convertible redeemable preferred shares ("Series A-2 Preferred Shares") to certain investors at US\$1.27 per share for a total consideration of US\$3,000,000. Series A-1 Preferred Shares and Series A-2 Preferred Shares are collectively referred to as the Series A Preferred Shares.

From January through June 2016, the Company issued 7,494,537 Series B convertible redeemable preferred shares ("Series B Preferred Shares") to certain investors at US\$3.74 per share for a total consideration of US\$27,999,995.

From February through May 2018, the Company issued 5,597,354 Series C-1 convertible redeemable preferred shares ("Series C-1 Preferred Shares") to certain investors at US\$8.93 per share for a total consideration of US\$50,000,033. Concurrently, in February 2018, the Company also issued warrants to two Series C-1 investors at nil consideration ("Series C-1 Warrants"). The Series C-1 Warrants allowed the holders to purchase Series C-2 Preferred Shares (defined below) at the exercise price of US\$10.21 per share for a total consideration of up to US\$7,500,000. Series C-1 Warrants were exercisable, in whole or in part, at any time from the warrant issuance date to the earlier of i) April 1, 2019, ii) a deemed liquidation event or iii) the closing of the Qualified IPO. Series C-1 Warrants expired on April 1, 2019.

From June through November 2019, the Company issued 1,861,121 Series C-2 convertible redeemable preferred shares ("Series C-2 Preferred Shares") to certain investors at US\$10.21 per share for a total consideration of US\$18,999,999.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

8. CONVERTIBLE REDEEMABLE PREFERRED SHARES AND WARRANTS (Continued)

In December 2019, the Company issued 4,452,441 Series C-3 convertible redeemable preferred shares ("Series C-3 Preferred Shares") to a certain investor at US\$11.23 per share for a total consideration of US\$50,000,000.

Series C-1 Preferred Shares, Series C-2 Preferred Shares and Series C-3 Preferred Shares are collectively referred to as the Series C Preferred Shares.

The key features of the Series A Preferred Shares, Series B Preferred Shares and Series C Preferred Shares (collectively the "Preferred Shares") are as follows:

Dividends

Each holder of the Preferred Shares will be entitled to receive non-cumulative dividends when declared by the Board of Directors prior and in preference to ordinary shareholders. The dividend should be paid at the rate of 6% of the original issue price per share per annum on each Preferred Shares in the sequence of Series C Preferred Shares and Preferred Shares other than the Series C Preferred Shares. After the preferential dividends relating to the Preferred Shares have been paid in full or declared and set apart in any fiscal year of the Company, any additional dividends out of funds or assets legally available therefore may be declared in that fiscal year for the Shares and, if such additional dividends are declared, the preferred shareholders shall be entitled to participate on an as converted-basis pro-rata in any dividends or distributions paid to the ordinary shareholders.

Voting

Each Preferred Share has voting rights equivalent to the number of ordinary shares to which it is convertible at the record date. The Preferred Shares shall vote separately as a class with respect to certain specified matters. Otherwise, the preferred shareholders and ordinary shareholders shall vote together as a single class.

Liquidation preference

In the event of any liquidation, dissolution or winding up of the Company, or the cessation of the business of the Company or of a substantial portion of the business of the Company, whether voluntary or involuntary, or any deemed liquidation event (unless waived by the preferred shareholders), the preferred shareholders shall be entitled to receive a per share amount equal to 100% of the original issue price of the respective series of the Preferred Shares, in the sequence of Series C Preferred Shares, Series B Preferred Shares and Series A Preferred Shares. After such liquidation amounts have been paid in full, the preferred shareholders are entitled to receive a simple interest accruing on each Preferred Share at 6% of its original issue price per annum from the date of issuance of such Preferred Share to the date of distribution of such amount, in the sequence of Series C Preferred Shares, Series B Preferred Shares and Series A Preferred Shares. After such interest amounts have been paid in full, any remaining funds or assets of the Company legally available for distribution to shareholders shall be distributed on a pro rata basis among the preferred shareholders, on an as-converted basis, together with the ordinary shareholders.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

8. CONVERTIBLE REDEEMABLE PREFERRED SHARES AND WARRANTS (Continued)

Conversion

Each Preferred Share may be converted at any time into ordinary shares at the option of the preferred shareholders based on the then-effective conversion price. The initial conversion ratio is 1:1, subject to adjustment in the event of share splits, share combinations, ordinary share dividends or distributions, other dividends, reorganizations, mergers, consolidations, reclassifications, exchanges, substitutions, or dilutive issuance.

All Preferred Shares are converted automatically into ordinary shares at the then effective applicable conversion price upon the earlier of a Qualified Public Offering (public offering of the Company's shares with an offering price (exclusive of underwriting commissions and expenses) that reflects a market capitalization (immediately prior to the public offering) of not less than US\$650,000,000 and with an aggregate proceeds of no less than US\$75 million) or a date specified by written consent or agreement of the holders of at least 80% of the voting power of the then outstanding Preferred Shares.

Redemption

The Preferred Shares are redeemable upon request by the holders of the majority outstanding Preferred Shares if the Company fails to consummate a Qualified Public Offering or complete a deemed liquidation event on or before March 31, 2025 at the redemption price equal to the original issue price plus any declared but unpaid dividends.

Accounting for Preferred Shares

The Preferred Shares are classified as mezzanine equity in the consolidated balance sheets because they are contingently redeemable upon the occurrence of an event outside of the Company's control (e.g. the Company not achieving a Qualified Public Offering or a deemed liquidation event before March 31, 2025 ("Target QIPO Date")). The Preferred Shares were determined to be mezzanine equity with no embedded feature to be bifurcated and no beneficial conversion features to be recognized. The Preferred Shares are initially recorded at their respective issuance date fair value, net of issuance cost and fair value allocated to the detachable warrants. The Company did not incur material issuance cost for any Preferred Shares issued.

The Company concluded that the Preferred Shares are not currently redeemable, but are probable to become redeemable. The Company accreted changes in the redemption value over the period from the date of issuance to the earliest redemption date using the interest method. No accretion charge was recorded as the redemption value is fixed to original issue price for the years presented, except for Series C-1 Preferred Shares issued with detachable warrants.

Modification of Preferred Shares

The Company made several amendments to the Preferred Shares, mainly including: 1) added redemption rights for Series A Preferred Shares upon the issuance of the Series B Preferred Shares; 2) extended the Target QIPO Date upon the issuance of the Series C-1 Preferred Shares and the Series C-3 Preferred Shares. These amendments are accounted for as modifications rather than extinguishments as the fair values of these Preferred Shares immediately after the amendments were

ADAGENE INC.
NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)
FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019
8. CONVERTIBLE REDEEMABLE PREFERRED SHARES AND WARRANTS (Continued)

not significantly different from their respective fair values immediately before the amendment. When Preferred Shares are modified and such modification results in value transfer between preferred shareholders and ordinary shareholders, the value transferred is treated as a deemed dividend to or deemed contribution from the preferred shareholders.

On February 2, 2018, the Target QIPO Date was extended from January 19, 2023 to February 2, 2025 (7th anniversary of Series C-1 closing) upon issuance of Series C-1 Preferred Shares. On the date of the modifications, the Company assessed the total fair value of preferred shares immediately before and after the change of the terms with the assistance from an independent third-party appraiser. The combined change in fair value of Preferred Shares immediately before and after the modification was US\$1,186,187. The increase in fair value of the ordinary shares of is US\$1,186,187, in substance, a transfer of wealth mostly from the preferred shareholders to the ordinary shareholder, and therefore are recorded as deemed contribution from the Preferred Shareholders.

The Company's Preferred Shares activities for the periods presented are summarized below:

Mezzanine equity	Series A-1	Series A-2	Series B	Series C-1	Series C-2	Series C-3	Total
	US\$	US\$	US\$	US\$	US\$	US\$	US\$
Balance as of							
December 31, 2017	5,473,957	3,000,000	27,999,995	—	—	—	36,473,952
Issuance of Series C-1 Preferred Shares	—	—	—	48,258,313	—	—	48,258,313
Accretion of Series C-1 Preferred Shares to redemption value	—	—	—	222,846	—	—	222,846
Balance as of							
December 31, 2018	5,473,957	3,000,000	27,999,995	48,481,159	—	—	84,955,111
Issuance of Series C-2 Preferred Shares	—	—	—	—	18,999,999	—	18,999,999
Issuance of Series C-3 Preferred Shares	—	—	—	—	—	50,000,000	50,000,000
Accretion of Series C-1 Preferred Shares to redemption value	—	—	—	246,184	—	—	246,184
Balance as of							
December 31, 2019	<u>5,473,957</u>	<u>3,000,000</u>	<u>27,999,995</u>	<u>48,727,343</u>	<u>18,999,999</u>	<u>50,000,000</u>	<u>154,201,294</u>

The warrants are freestanding instruments and classified as liabilities in accordance with ASC 480. The warrants are initially recognized at fair value, with subsequent changes in fair value recorded currently in earnings. The Company recognized gains from the decrease in fair value of the warrants of US\$534,305 and US\$1,207,415 for the years ended December 31, 2018 and 2019, respectively.

The Company has measured the warrant liabilities at fair values on a recurring basis using significant unobservable inputs (Level 3) for the years ended December 31, 2018. The Group used the Black-Scholes option pricing model to estimate the fair value of warrant liabilities using the following assumptions:

	For the year ended December 31, 2018
Risk-free interest rate	1.92%, 2.37%
Exercise price	US\$10.2089
Maturity date	01/04/2019
Estimated volatility rate	63.78%, 71.00%

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

8. CONVERTIBLE REDEEMABLE PREFERRED SHARES AND WARRANTS (Continued)

The model requires the input of highly subjective assumptions including the risk-free rate interest rate, maturity date, estimated volatility rate and fair value of underlying preferred shares. The risk-free rate for periods within the contractual life is based on the US treasury strip bond with maturity similar to the maturity of the warrants as of valuation dates. For expected volatilities, the Company has made reference to the historical daily stock prices volatilities of ordinary shares of several comparable companies in the same industry as the Company. The estimated fair value of the preferred shares was determined with assistance from an independent third-party valuation firm.

The significant unobservable inputs used in the fair value measurement of the warrant liabilities include risk-free interest rate, interval between valuation date and maturity date, estimated volatility rate and fair value of underlying preferred shares. Significant decreases in interval between valuation date and maturity date, estimated volatility rate and fair value of underlying preferred shares would result in a significantly lower fair value measurement. Significant increases in risk-free interest rate would result in a significantly lower fair value measurement.

9. SHARE-BASED COMPENSATION

On November 7, 2015, the Company adopted a share incentive plan ("2015 Plan"). Under the 2015 Plan, the Company's Board of Directors has approved that a maximum aggregate number of shares that may be issued pursuant to all awards granted shall be 4,336,126. In September 2017, the Company replaced 2015 Plan with the amended and restated share incentive plan ("2017 Plan") and increased the maximum number of shares issuable to 6,336,126. In December 2019, the Company replaced 2017 Plan with the second amended and restated share incentive plan ("2019 Plan") and further increased the maximum number of shares issuable to 11,391,131. The terms of the 2015 Plan, 2017 Plan and 2019 Plan are substantially the same other than the maximum aggregate number of shares the Company may issue under the respective plan.

Share options granted to each grantee under the Share Incentive Plan will generally be exercisable upon the grantee renders service to the Company in accordance with a stipulated vesting schedule. Grantees are generally subject to a vesting schedule of no longer than five years, under which the grantee earns an entitlement to vest a certain percentage of his option grants at the end of each month

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

9. SHARE-BASED COMPENSATION (Continued)

or year of completed service. The share option awards shall expire no more than 10 years from their grant dates.

	Number of Options	Weighted- Average Exercise Price US\$ per option	Weighted- Average Grant Date Fair Value US\$ per option	Weighted Average Remaining Contractual Term Years	Aggregate Intrinsic Value US\$
Outstanding at January 1, 2018	848,828	0.38	3.75	8.56	4,907,569
Forfeited	(274,500)	0.55	5.71	—	—
Outstanding at January 1, 2019	574,328	0.30	2.80	7.14	3,734,644
Granted	372,500	1.26	5.46	—	—
Exercised	(34,000)	0.55	5.71	—	—
Forfeited	(46,800)	1.22	5.75	—	—
Outstanding at December 31, 2019	866,028	0.65	3.67	7.27	6,328,171
Vested and expected to vest at December 31, 2019	866,028	0.65	3.67	7.27	6,328,171
Exercisable at December 31, 2019	527,278	0.34	2.60	6.26	4,018,019

The aggregate intrinsic value in the table above represents the difference between the exercise price of the awards and the fair value of the underlying ordinary shares at each reporting date, for those awards that had exercise price below the estimated fair value of the relevant ordinary shares.

The aggregate fair value of the equity awards vested during the years ended December 31, 2018 and 2019 was US\$165,609 and US\$366,113, respectively. As of December 31, 2019, there was US\$1,554,053 of total unrecognized employee share-based compensation expense, may be adjusted for actual forfeitures occurring in the future. Total unrecognized compensation cost will be recognized over a weighted-average period of 3.32 years.

Fair value of share options

The fair value of share options was determined using the binomial option valuation model, with the assistance from an independent third-party appraiser. The binomial model requires the input of highly subjective assumptions, including the expected volatility, the exercise multiple, the risk-free rate and the dividend yield. For expected volatility, the Group has made reference to historical volatility of several comparable companies in the same industry. The exercise multiple was estimated as the average ratio of the stock price to the exercise price of when employees would decide to voluntarily exercise their vested share options. The risk-free rate for periods within the contractual life of the share options is based on the market yield of U.S. Treasury Bonds in effect at the time of grant. The dividend yield is based on the expected dividend policy over the contractual life of the share options. The estimated fair value of the ordinary shares, at the share option grant dates, was determined with the assistance from an independent third-party appraiser.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

9. SHARE-BASED COMPENSATION (Continued)

The assumptions used to estimate the fair value of the share options granted are as follows:

	For the year ended December 31, 2019
Risk-free interest rate	1.78% - 2.73%
Dividend yield	0%
Expected volatility range	67.5% - 71.0%
Exercise multiple	2.2 - 2.8
Contractual life	10 years

Total share-based compensation expenses recognized for the years ended December 31, 2018 and 2019 were as follows:

	For the years ended December 31,	
	2018	2019
	US\$	US\$
Research and development expenses	126,540	404,620
Administrative expenses	—	207,091
Total share-based compensation expenses	<u>126,540</u>	<u>611,711</u>

Up to the date of the issuance of these consolidated financial statements, some proceeds of the subscription capital arising from the exercise of vested share options by certain employees remained outstanding and such amount was presented as subscriptions receivable, a contra-equity balance on the consolidated balance sheets as of December 31, 2018 and 2019, respectively.

10. COLLABORATION ARRANGEMENTS

Guilin Sanjin Pharmaceutical Co., Ltd. License Agreement

In December 2018, the Group entered into (i) a collaboration agreement (the "Sanjin Greater China Agreement") that covers Greater China with Guilin Sanjin Pharmaceutical Co., Ltd. ("Sanjin") and certain of its subsidiaries (collectively, "Sanjin Parties") and (ii) a collaboration agreement (the "Sanjin ROW Agreement", together with the Sanjin Greater China Agreement, the "2018 Sanjin Agreements") that covers the regions other than Greater China with Sanjin. Pursuant to the Sanjin Greater China Agreement, the Group licensed the Chinese intellectual property directly related to a monospecific antibody molecule that binds to the PD-L1 target (the "PD-L1 Project"), including patent rights, patent application rights and technologies based on the core sequence of the molecule, to Sanjin Parties. Sanjin Parties will own all the Chinese intellectual property developed in the exercise of Sanjin Parties' rights under the agreement, including but not limited to improvements (including combination products), clinical trials, regulatory filings, and commercialization rights relating thereto. The Group also granted Sanjin Parties a royalty-free license to use our other existing intellectual property and improvements thereto which are related to the PD-L1 Project for the purposes of exploiting its rights and performing its obligations under the agreement. Sanjin Parties will enjoy all the economic benefits

ADAGENE INC.**NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)****FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019****10. COLLABORATION ARRANGEMENTS (Continued)**

deriving from the PD-L1 Project in Greater China, including but not limited to patent transfer fee, licensing fee, sales revenue and sales commission, etc. Sanjin Parties will pay the Group (i) single-digit percentage of net sales of the products that use the licensed antibody after such products enter the market and (ii) a low to mid-low double-digit percentage of the profits resulting from any transfer of the license to any third parties depending on the timing of the transfer relative to the development stage of the product. The Group also received RMB10,000,000 (equivalent to approximately US\$1,511,168) upfront fee upon the effectiveness of the agreement from Sanjin Parties.

Pursuant to the Sanjin ROW Agreement, the Group granted Sanjin a royalty-free license to use all intellectual property relating to (i) the collaboration under the agreement that the Group controlled before the Group entered into the agreement or acquired independently of the agreement and (ii) improvements thereto for the purposes of exploiting its rights and performing its obligations under the agreement. Any intellectual property generated independently by a party under the agreement will be solely owned by that party who generated such intellectual property, and any intellectual property generated from cooperation between the Group and Sanjin's affiliates in connection with the collaboration will be jointly owned. The Group retain the ownership of patent rights of key intellectual property pertaining to PD-L1 outside of the Greater China. In addition, all the results obtained by Sanjin relating to the research and development of any new antibody developed under the agreement will be owned by Sanjin. The Group retain a majority of the economic benefits derived from the Sanjin ROW Agreement, including but not limited to any patent transfer fee, licensing fee and gains realized under such transfer. In case the Group intend to transfer to a third party our share of economic interests in any country outside of Greater China, the Group must notify Sanjin and Sanjin will receive a right of first refusal if it pays the Group a deposit equal to a low double-digit percentage of the consideration that the Group expect to receive from such third party. If Sanjin waives the right of first refusal, the Group can proceed with the transfer, provided that the final transaction price with the third party is not lower than the amount of the offering price that was included in our notice to Sanjin.

The Group agreed not to (i) independently develop any monospecific antibodies that bind to the PD-L1 target or (ii) grant any rights associated with such antibodies to any third parties during the three-year period from the effective date of the agreement. The exclusivity obligation does not prevent the Group from (i) developing or granting any licenses to third parties for intellectual property that covers bispecific antibodies, ADCs, diagnostic antibodies, nano-particles and masked antibody against PD-L1 target and (ii) continuing to provide antibody screening service that were commenced before the execution of the Sanjin Greater China Agreement and either party has the independent right to conduct combination therapy studies outside of the Greater China. Either non-breaching party may terminate the 2018 Sanjin Agreements if the other party's ability to comply with its respective obligations under the agreements is negatively affected by contingencies such as failure to maintain operation or changes in core project management and the other party fails to take effective remedial measures. Each agreement automatically terminates upon the termination of the other agreement. Upon the rescission or termination, Sanjin Parties will return to the Group all the intellectual property, documents and data provided by the Group under the 2018 Sanjin Agreements.

In the event that the failure of the development of the product candidate solely arises from the Group's research and development basis specified under this agreement, Sanjin has the right to claim

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

10. COLLABORATION ARRANGEMENTS (Continued)

back all the payment made to the Group. The Group considers the possibility of occurrence of such event is very remote.

For the year ended December 31, 2018, the Group recognized revenue of RMB10,000,000 (equivalent to approximately US\$1,511,168) for this agreement.

Dragon Boat Biopharmaceutical (Shanghai) Limited License Agreement

In May 2019, the Group entered into (i) a collaboration agreement that covers Greater China (the "Dragon Boat Greater China Agreement") and (ii) a collaboration agreement that covers the regions other than Greater China (the "Dragon Boat ROW Agreement," together with the Dragon Boat Greater China Agreement, the "2019 Dragon Boat Agreements"), with Dragon Boat Biopharmaceutical (Shanghai) Limited ("Dragon Boat"), a subsidiary of Sanjin. Pursuant to the Dragon Boat Greater China Agreement, the Group will license the Chinese intellectual property directly related to a certain monospecific antibody molecule that binds to a specified target (the "Specified Project"), including the patent rights, patent application rights and technologies based on the core sequence of the molecule, to Dragon Boat. Dragon Boat will own all the Chinese intellectual property developed in the exercise of Dragon Boat's rights under the agreement, including but not limited to improvements (including combination products), clinical trials, regulatory filings, and commercialization rights relating thereto. The Group also granted Dragon Boat a royalty-free license to use our other existing intellectual property and improvements thereto which are related to the Specified Project for the purposes of exploiting its rights and performing its obligations under the agreement. Dragon Boat will enjoy all the economic benefits deriving from the Specified Project in Greater China, including but not limited to patent transfer fee, licensing fee, sales revenue and sales commission, etc. and will pay the Group (i) certain high-six figure dollar milestone payments upon the achievement of certain milestones (including milestones of launch of pre-clinical safety evaluation animal test, obtaining Investigational New Drug ("IND") approval in PRC and completion of clinical phase I test in PRC) and (ii) a single-digit percentage of net sales of the products that use the licensed antibody after such products enter the market.

Pursuant to the Dragon Boat ROW Agreement, the Group granted Dragon Boat a royalty-free license to use all intellectual property relating to (i) the collaboration under the agreement that the Group controlled before the Group entered into the agreement or acquired independently of the agreement and (ii) improvements thereto for the purposes of exploiting its rights and performing its obligations under the agreement. Any intellectual property generated independently by a party under the agreement will be solely owned by that party who generated such intellectual property, and any intellectual property generated from cooperation between the Group and Dragon Boat in connection with the collaboration will be jointly owned. The Group retain the ownership of patent rights of key intellectual property pertaining to the specified target outside of the Greater China. In addition, all the results obtained by Dragon Boat relating to the research and development of any new antibody developed under the agreement will be owned by Dragon Boat. The Group retains a majority of the economic benefits derived from the Dragon Boat ROW Agreement, including but not limited to any patent transfer fee, licensing fee and gains realized under such transfer. In case the Group intend to transfer to a third party our share of economic interests in any country outside of Greater China, the Group must notify Dragon Boat and Dragon Boat will receive a right of first refusal if it pays the

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

10. COLLABORATION ARRANGEMENTS (Continued)

Group a deposit equal to a low double-digit percentage of the consideration that the Group expects to receive from such third party. If Dragon Boat waives the right of first refusal, the Group can proceed with the transfer, provided that the final transaction price with the third party is not lower than the amount of the offering price that was included in our notice to Dragon Boat.

Under the 2019 Dragon Boat Agreements, the Group agreed not to (i) independently develop any monospecific antibodies that bind to the specified target or (ii) grant any rights associated with such antibodies to any third parties during the three-year period from the effective date of the agreements. The exclusivity obligation does not prevent the Group from (i) developing or granting any licenses to third parties for intellectual property that covers bispecific antibodies, ADCs, diagnostic antibodies, nano-particles and masked antibody against the specific target and (ii) continuing to provide antibody screening service that were commenced before the execution of the Dragon Boat Greater China Agreement and either party has the independent right to conduct combination therapy studies outside of the Greater China. Either nonbreaching party may terminate the 2019 Dragon Boat Agreements if the other party's ability to comply with its obligations under the agreements is negatively affected by contingencies such as failure to maintain operation or changes in core project management and the other party fails to take effective remedial measures. Each agreement automatically terminates upon the termination of the other agreement. Upon the rescission or termination, Dragon Boat will return to the Group all the intellectual property, documents and data provided by the Group under the 2019 Dragon Boat Agreements.

In the event that the failure of the development of the product candidate solely arises from the Group's research and development basis specified under this agreement, Dragon Boat has the right to claim back all the payment made to the Group.

For the year ended December 31, 2019, no revenue was recognized for this agreement since the licensed product has not been transferred to Dragon Boat.

As of December 31, 2019, upfront fee of RMB4,000,000 (equivalent to approximately US\$573,378) that was received by the Group was recorded as contract liabilities in the consolidated balance sheets, as the performance obligation had not been satisfied by the Group.

ADC Therapeutics SA License and Collaboration Agreements

In April 2019, the Group entered into a material transfer and collaboration agreement (the "ADCT Collaboration Agreement") and a license agreement (the "ADCT License Agreement") with ADC Therapeutics SA ("ADC Therapeutics"). These two agreements are combined as a single contract as the agreements were negotiated as a package with a single commercial objective.

ADCT Collaboration Agreement

Pursuant to the ADCT Collaboration Agreement, the Group agreed to generate masked antibodies with respect to up to two exclusive targets selected by ADC Therapeutics. Upon our delivery of certain initial results, ADC Therapeutics has the option to license the Group's technology with respect to one or both targets as further detailed below. ADC Therapeutics has not yet exercised such options as of December 31, 2019.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

10. COLLABORATION ARRANGEMENTS (Continued)

Under the ADCT Collaboration Agreement, the Group is eligible to receive up to a low-seven-figure dollar amount in consideration for the Group's exclusivity obligations, upon achievement of certain development milestones (including milestones of delivery of certain amino acid sequences and successful outcome of the first in-vivo study) and upon ADC Therapeutics' election to proceed with development for the two elected targets. Apart from performance obligation to deliver the amino acid sequences of the corresponding masking peptides, the Group is not required to perform any additional research and development services. ADC Therapeutics has the right to terminate the ADCT Collaboration Agreement at any time and for any reason in its entirety or on a target-by-target basis upon thirty days' prior written notice to the Group. Either party may terminate the ADCT Collaboration Agreement, in its entirety or on a target-by-target basis, upon the other party's uncured material breach of the agreement or the other party's insolvency-related events.

The Group also granted ADC Therapeutics an exclusive target reservation right for one year from the commencement of the agreement and an option to renewal for another year with a consideration of low-six-figure dollar amount.

ADCT License Agreement

Subject to the exercise of the options contained in the ADCT Collaboration Agreement, the Group has granted ADC Therapeutics, with respect to each elected target, an exclusive, worldwide, perpetual and irrevocable (subject only to the termination provisions) license (with the right to grant sublicenses) to develop, make, use, commercialize and import the antibody drug conjugates that comprise masked antibodies generated by the Group under these programs.

Under the ADCT License Agreement, if ADC Therapeutics exercises both of its options granted thereunder, the Group could be eligible to receive up to a low-nine-figure dollar amount in development and regulatory milestone payments upon the achievement of certain milestones (including milestones of successful completion of Good Laboratory Practice Toxicology studies, launch of clinical trials and start of commercial sales in difference countries and etc.) and up to a mid-eight-figure dollar amount in sales milestone payments, in addition to mid-single-digit percentage net sales-based tiered royalties on products licensed under the ADCT License Agreement, subject to certain reductions. Royalties, if any, will be payable on a country-by-country and product-by-product basis, until the earlier of (i) the tenth anniversary of the first commercial sale of such product or (ii) the expiration of the last-to-expire patent licensed under the agreement in such country, unless earlier terminated by the parties, following which any licenses granted to ADC Therapeutics under the ADCT License Agreement shall become fully paid up, perpetual and irrevocable.

ADC Therapeutics has the right to terminate the ADCT License Agreement before the expiration of the royalty term on a product-by-product basis or in its entirety (i) for any reason or no reason upon thirty days' written notice to the Group, or (ii) if ADC Therapeutics chooses to discontinue the development or sale of the applicable licensed product worldwide. Each party has certain rights to terminate the ADCT License Agreement with prior written notice upon the other party's uncured material breach or insolvency.

For the years ended December 31, 2018 and 2019, no revenue was recognized for this agreement since the licensed product has not been transferred to ADC Therapeutics.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

10. COLLABORATION ARRANGEMENTS (Continued)

As of December 31, 2019, upfront fee of US\$225,000 and exclusive target reservation right of US\$100,000 that were received by the Group were recorded as contract liabilities in the consolidated balance sheets, as the performance obligation had not been satisfied by the Group.

Signal Pharmaceuticals LLC

In January 2019, the Group entered into an agreement with Signal Pharmaceuticals LLC, a subsidiary of Celgene Corporation, for a purchase order of delivery of certain sequences with total consideration of US\$480,000. For the year ended December 31, 2019, the Group recognized revenue of US\$480,000 upon delivery of such sequences.

11. INCOME TAX EXPENSE

PRC

Effective from January 1, 2008, the PRC's statutory, Enterprise Income Tax ("EIT") rate is 25%. In accordance with the implementation rules of EIT Law, a qualified "Technology Advanced Service Enterprises" ("TASE") is eligible for a preferential tax rate of 15%. The TASE certificate is effective for three years. An entity must file required supporting documents with the tax authority and ensure fulfillment of the relevant TASE criteria before using the preferential rate. An entity could apply for the TASE certificate every year.

Adagene (Suzhou) Limited was first recognized as a qualified TASE in March 2015 and renewed in December 2018. Adagene (Suzhou) Limited was authorized to enjoy the preferential tax rate of 15% from 2015 to 2021.

Cayman Islands

Adagene Inc. is incorporated in the Cayman Islands. Under the current laws of the Cayman Islands Adagene Inc. is not subject to tax on income or capital gain. Additionally, the Cayman Islands does not impose a withholding tax on payments of dividends to shareholders.

Hong Kong

Adagene (Hong Kong) Limited is incorporated in Hong Kong. Companies registered in Hong Kong are subject to Hong Kong profits tax on the taxable income as reported in their respective statutory financial statements adjusted in accordance with the relevant Hong Kong tax laws. The applicable tax rate in Hong Kong is 16.5%. For the years ended December 31, 2018 and 2019, Adagene (Hong Kong) Limited did not make any provisions for Hong Kong profit tax as there were no assessable profits derived from or earnings in Hong Kong for any of the periods presented. Under the Hong Kong tax law, Adagene (Hong Kong) Limited is exempted from income tax on its foreign-derived income and there are no withholding taxes in Hong Kong on remittance of dividends.

ADAGENE INC.**NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)****FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019****11. INCOME TAX EXPENSE (Continued)***Australia*

Adagene Australia Pty Ltd. is incorporated in Australia. Companies registered in Australia are subject to Australia profits tax on the taxable income as reported in their respective statutory financial statements adjusted in accordance with the relevant Australia tax laws. The applicable tax rate in Australia is 30%. Adagene Australia Pty Ltd. has no taxable income for all periods presented, therefore, no provision for income taxes is required.

United States

Adagene Incorporated is incorporated in U.S. and is subject to U.S. federal corporate income tax at a rate of 21%. Adagene Incorporated is also subject to state income tax in California of 8.84%. Adagene Incorporated has no taxable income for all periods presented, therefore, no provision for income taxes is required. Reconciliation between the income tax expense computed by applying the statutory tax rate to loss before income tax and the actual provision for income tax is as follows:

	For the years ended	
	December 31,	
	2018	2019
	US\$	US\$
Loss before income tax	(15,266,184)	(16,432,308)
Income tax computed at respective applicable tax rate	(30,560)	27,875
Research and development super-deduction ^(a)	(579,090)	(230,126)
Non-deductible expenses	4,010	3,306
Changes in valuation allowance	605,640	198,945
Income tax expense	—	—

Note (a): Due to the impacts of research and development super-deduction, the Group's subsidiary, Adagene (Suzhou) Limited did not have any taxable profit for the years ended December 31, 2018 and 2019.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

11. INCOME TAX EXPENSE (Continued)

Deferred tax assets and liabilities

Deferred taxes were measured using the enacted tax rates for the periods in which the temporary differences are expected to be reversed. The tax effects of temporary differences that give rise to the deferred tax balances as of December 31, 2018 and 2019 are as follows:

	For the years ended	
	December 31,	
	2018	2019
	US\$	US\$
Deferred tax assets:		
Net operating loss carry forward	735,717	928,989
Depreciation and amortization of property, equipment and software	4,213	9,886
Gross deferred tax assets	<u>739,930</u>	<u>938,875</u>
Less: valuation allowance	<u>(739,930)</u>	<u>(938,875)</u>
Total deferred tax assets, net	<u>—</u>	<u>—</u>

Movement of the valuation allowance is as follows:

	For the years ended	
	December 31,	
	2018	2019
	US\$	US\$
Balance as of January 1	<u>134,290</u>	<u>739,930</u>
Addition	605,640	198,945
Balance as of December 31	<u>739,930</u>	<u>938,875</u>

A valuation allowance is provided to reduce the amount of deferred tax assets if it is considered more likely than not that some portion or all of the deferred tax assets will not be realized in the foreseeable future. In making such determination, the Group evaluates a variety of positive and negative factors including the Group's operating history, accumulated deficit, the existence of taxable temporary differences and reversal periods.

The Group has incurred net accumulated operating losses for income tax purposes since its inception. The Group believes that it is more likely than not that these net accumulated operating losses will not be utilized in the future. Therefore, the Group has provided full valuation allowances for the deferred tax assets as of December 31, 2018 and 2019.

The Group evaluates each uncertain tax position (including the potential application of interest and penalties) based on the technical merits, and measure the unrecognized benefits associated with the tax positions. As of December 31, 2018 and 2019, the Group did not have any significant unrecognized uncertain tax positions.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

12. NET LOSS PER SHARE

Basic and diluted net loss per share for the years ended December 31, 2018 and 2019 are calculated as follows:

	For the years ended	
	December 31,	
	2018	2019
	US\$	US\$
Numerator:		
Net loss attributable to Adagene Inc.'s shareholders	(15,266,184)	(16,432,308)
Deemed contribution from convertible redeemable preferred shareholders	1,186,187	—
Accretion of convertible redeemable preferred shares to redemption value	(222,846)	(246,184)
Net loss attributable to ordinary shareholders	<u>(14,302,843)</u>	<u>(16,678,492)</u>
Denominator:		
Weighted-average number of ordinary shares outstanding—basic and diluted	15,159,136	15,178,232
Net Loss per share—basic and diluted	(0.94)	(1.10)

The effects of all outstanding convertible redeemable preferred shares and share options have been excluded from the computation of diluted loss per share for the years ended December 31, 2018 and 2019 as their effects would be anti-dilutive.

The potentially dilutive securities that have not been included in the calculation of diluted net loss per share as their inclusion would be anti-dilutive are as follows:

	For the years ended	
	December 31,	
	2018	2019
Convertible redeemable preferred shares	20,430,200	21,977,914
Share options	293,133	582,526

13. UNAUDITED PRO FORMA NET LOSS PER SHARE

The unaudited pro forma net loss per ordinary share is computed using the weighted-average number of ordinary shares outstanding and the automatic conversion of all of the Group's outstanding mezzanine equity into ordinary shares upon the closing of the Group's Qualified Public Offering, as if it had occurred on January 1, 2019. The Group believes the unaudited pro forma net loss per share provides material information to investors, as the automatic conversion of the Group's outstanding mezzanine equity. The disclosure of pro forma net loss per ordinary share provides an indication of net loss per ordinary share that is comparable to what will be reported by the Group as a public company following the closing of the Qualified Public Offering.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

13. UNAUDITED PRO FORMA NET LOSS PER SHARE (Continued)

The unaudited basic and diluted pro forma net loss per share is calculated as follows:

	For the year ended December 31, 2019 US\$ (Unaudited)
Numerator:	
Net loss attributable to ordinary shareholders in computing pro forma net loss per share—basic and diluted	(16,678,492)
Add back accretion of convertible redeemable preferred shares to redemption value	246,184
Numerator for pro-forma basic and diluted net loss per share	(16,432,308)
Denominator:	
Weighted-average number of ordinary shares outstanding—basic and diluted	15,178,232
Add: adjustment to reflect assumed effect of automatic conversion of convertible redeemable preferred shares	21,977,914
Pro forma weighted average number of shares outstanding—basic and diluted	37,156,146
Pro forma net loss per share—basic and diluted	(0.44)

14. RELATED PARTY TRANSACTIONS

a) *Related Parties*

Name of related parties	Relationship
Peter Luo	Chairman, Chief Executive Officer and a principal shareholder of the Company
Four senior management personnel	Management and ordinary shareholders of the Company
WuXi AppTec Co., Ltd. ("WuXi AppTec Group")	A principal shareholder of the Group
WuXi Biologics (Shanghai) Co., Ltd.	Controlled by the ultimate controlling party of a principal shareholder of the Group

b) *The Group had the following related party balances at the end of the year:*

	As of December 31,	
	2018 US\$	2019 US\$
WuXi AppTec Group	39,250	739,051
Four senior management personnel(i)	622,420	350,865
Peter Luo(i)	508,359	338,818
WuXi Biologics (Shanghai) Co., Ltd.	—	4,452
Total amounts due from related parties	1,170,029	1,433,186

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

14. RELATED PARTY TRANSACTIONS (Continued)

	As of December 31,	
	2018	2019
	US\$	US\$
WuXi Biologics (Shanghai) Co., Ltd.	3,368,735	1,379,741
WuXi AppTec Group	220,888	432,784
Peter Luo(ii)	84,625	83,254
Total amounts due to related parties	3,674,248	1,895,779

c) The Group had the following related party transactions:

	For the years ended	
	December 31,	
	2018	2019
	US\$	US\$
Receipt of CRO and CMO services:		
WuXi Biologics (Shanghai) Co., Ltd.	6,535,512	3,567,962
WuXi AppTec Group	969,079	2,136,344
	7,504,591	5,704,306

- (i) In October and November 2017, Peter Luo and other four senior management personnel elected to exercise the vested share options that granted under 2015 Plan. As of December 31, 2018, the balance of amounts due from Peter Luo and other four senior management personnel represented the receivables arising from the exercise of share options and related withholding individual income tax amounts. The receivables arising from the exercise of share options were subsequently received in the year ended December 31, 2019. As of December 31, 2019, the balance of amounts due from Peter Luo and other four senior management personnel represented withholding individual income tax amounts.
- (ii) As of December 31, 2018 and 2019, the balance of amounts due to Peter Luo represented the Group's receipt of personal subsidy on behalf of Peter Luo, which was subsequently remitted to Peter Luo in May 2020.

15. COMMITMENTS AND CONTINGENCIES

Operating lease commitments

Future minimum payments under non-cancelable operating leases with initial terms in excess of one year consist of the following as of December 31, 2019:

	US\$
For the years ending:	
2020	173,854
2021	83,305
Total	257,159

Payments under operating leases are expensed on a straight-line basis over the periods of their respective leases. The Group's lease arrangements have no renewal options, rent escalation clauses, restrictions or contingent rents and are all executed with third parties. For the years ended

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

15. COMMITMENTS AND CONTINGENCIES (Continued)

December 31, 2018 and 2019, total rental related expenses for all operating leases amounted to US\$176,268 and US\$175,812, respectively.

Contingencies

The Group is currently not involved in any legal or administrative proceedings that may have a material adverse impact on the Group's business, financial position or results of operations.

16. RESTRICTED NET ASSETS

The Group's ability to pay dividends may depend on the Group receiving distributions of funds from its PRC subsidiary. Relevant PRC statutory laws and regulations permit payments of dividends by the Group's PRC subsidiary only out of its retained earnings, if any, as determined in accordance with PRC accounting standards and regulations. The results of operations reflected in the consolidated financial statements prepared in accordance with U.S. GAAP differ from those reflected in the statutory financial statements of the Group's PRC subsidiary.

In accordance with the Company law of the PRC, a domestic enterprise is required to provide statutory reserves of at least 10% of its annual after-tax profit until such reserve has reached 50% of its respective registered capital based on the enterprise's PRC statutory accounts. A domestic enterprise is also required to provide discretionary surplus reserve, at the discretion of the Board of Directors, from the profits determined in accordance with the enterprise's PRC statutory accounts. The aforementioned reserves can only be used for specific purposes and are not distributable as cash dividends. The Group's PRC subsidiary was established as domestic invested enterprise and therefore is subject to the above mentioned restrictions on distributable profits.

As of December 31, 2018 and 2019, the total restricted net assets of the Company's subsidiary incorporated in PRC and subjected to restriction amounted to approximately US\$1,267,900 and US\$1,267,900, respectively. Other subsidiaries are not subjected to such restriction.

As a result of these PRC laws and regulations subject to the limit discussed above that require annual appropriations of 10% of after-tax income to be set aside, prior to payment of dividends, as general reserve fund, the Group's PRC subsidiary is restricted in their ability to transfer a portion of their net assets to the Group.

Foreign exchange and other regulations in the PRC further restrict the Company's PRC subsidiaries from transferring funds to the Company in the form of dividends, loans and advances.

Since the Group has a consolidated shareholders' deficit, its net asset base for purposes of calculating the proportionate share of restricted net assets of consolidated subsidiaries should be zero. Therefore, the restrictions placed on the net assets of the Company's PRC subsidiaries with positive equity would result in the 25 percent threshold being exceeded and a corresponding requirement to provide parent company financial information (Note 18).

17. SUBSEQUENT EVENTS

The Group evaluated subsequent events through September 22, 2020, the date these consolidated financial statements were issued.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

17. SUBSEQUENT EVENTS (Continued)

In March and August 2020, pursuant to the 2019 Plan, the Board of Directors of the Company passed resolutions and granted 1,944,565 and 4,028,808 share options to certain employees, respectively.

In June 2020, the Group borrowed a loan with amount of RMB10,000,000 (equivalent to approximately US\$1,412,529) from Agricultural Bank of China Limited for a term of one year and at the interest rate of 4.20% per annum.

Beginning in January 2020, the emergence and wide spread of the novel Coronavirus ("COVID-19") has resulted in quarantines, travel restrictions, and the temporary closure of stores and facilities in China, US and elsewhere. Substantially all of the Group's operating and workforce are concentrated in China and US. Consequently, the COVID-19 outbreak could potentially delay patient's access to hospital and the progress of clinical trials of the Group, which may adversely affect the Group's business operations, financial condition and operating results for 2020. The extent to which COVID-19 impacts the business and financial results of the Group in the longer term will depend on future developments, which are uncertain and cannot be predicted, including new information which may emerge concerning the severity of the coronavirus and the actions to contain the coronavirus or treat its impact, among others. The Group will continue to evaluate the impact on the results of operation, financial position and cash flows of the Group and react actively as the situation evolves.

Events subsequent to original issuance of consolidated financial statements (Unaudited)

In October 2020, pursuant to the 2019 Plan, the Board of Directors of the Company passed resolutions and granted 340,000 share options to certain employees with an exercise price of US\$2.26 per share. The fair values of these share options ranged from US\$9.02 to US\$9.23 per option and were determined by using the binomial option valuation model. The share-based compensation expense related to these share options is expected to be recognized over a weighted-average period of 4.06 years.

On November 9, 2020, the Company passed a board resolution to waive the vesting schedules and conditions of 2,375,000 share options granted to certain management members. Pursuant to this board resolution, these management members exercised all these share options at the original exercise price per share ranging from US\$1.83 to US\$2.26 and as a result, 2,375,000 ordinary shares were issued to them. These management members paid the exercise price by issuing recourse promissory notes in the total amount of US\$5,197,650. These ordinary shares are subject to the Company's repurchase rights before the share awards meet certain conditions. The management assessed and considered that such change in terms met the exception for modification accounting (as defined in ASU 2017-09, Scope of Modification Accounting), hence no additional amount of compensation expense was recognized.

In November 2020, the Group borrowed two loans with the amount of RMB5,000,000 (equivalent to approximately US\$760,086) and RMB5,000,000 (equivalent to US\$760,086) from China Merchants Bank Co., Ltd., respectively. Both of these two loans have a term of one year and bear an interest rate of 4.10% per annum.

On January 16, 2021, the Company passed a board resolution whereby certain management members surrendered a total of 491,119 ordinary shares as repayment for their respective promissory notes issued in connection with the exercising of options granted to them. In the meantime, the Company further granted these management members the right to repurchase the surrendered shares

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

17. SUBSEQUENT EVENTS (Continued)

by way of new option grants with an exercise price of US\$13.85 per share. The fair values of these share options were US\$7.85 per option, determined by using the binomial option valuation model. The share-based compensation expense related to these share options is expected to be recognized immediately upon grants.

Pursuant to the same board resolution, the 2021 Performance Incentive Plan was also adopted, under which an aggregate of 2,994,000 ordinary shares shall be reserved for issuance, and the share limit will automatically increase on the first trading day in January of each year (commencing with 2022) by an amount equal to (1) 5% of the total number of the Company's outstanding ordinary shares on December 31 of the prior year, or (2) such lesser number as determined by the board of directors.

On February 1, 2021, the Company entered into a collaboration and license agreement (the "Exelixis Agreement") with Exelixis, Inc., pursuant to which the Company agreed to generate masked antibodies with the Company's SAFEbody technology against an initial target and a second target to be selected by Exelixis. Under the Exelixis Agreement, the Company are entitled to receive an US\$11,000,000 upfront payment. In addition, the Company could be eligible to receive, on a target-by-target basis, up to US\$127,500,000 in development and regulatory milestone payments upon the achievement of specified development and regulatory milestones and up to US\$262,500,000 in sales milestone payments upon the achievement of specified commercial milestones. In the aggregate, the Company could be eligible to receive up to US\$255,000,000 in development and regulatory milestone payments and up to US\$525,000,000 in sales milestone payments for both targets under the Exelixis Agreement. In addition, the Company are entitled to receive mid-single-digit percentage net sales-based royalties on products developed under the Exelixis Agreement, subject to certain reductions.

18. CONDENSED FINANCIAL INFORMATION OF THE PARENT COMPANY

The Company performed a test on the restricted net assets of consolidated subsidiaries in accordance with Securities and Exchange Commission Regulation S-X Rule 4-08 I(3), "General Notes to Financial Statements" and concluded that it was applicable for the Company to disclose the financial statements for the parent company.

The subsidiaries did not pay any dividends to the Company for the years presented. For the purpose of presenting parent company only financial information, the Company records its investments in its subsidiaries under the equity method of accounting. Such investments are presented on the separate condensed balance sheets of the Company as "Investments (deficit) in subsidiaries" and the loss of the subsidiaries is presented as "share of losses of subsidiaries". Certain information and footnote disclosures generally included in financial statements prepared in accordance with U.S. GAAP have been condensed and omitted. The footnote disclosures contain supplemental information relating to the operations of the Company, as such, these statements should be read in conjunction with the notes to the consolidated financial statements of the Company.

The Company did not have significant capital and other commitments, long-term obligations, other long-term debt, or guarantees as of December 31, 2018 and 2019.

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

18. CONDENSED FINANCIAL INFORMATION OF THE PARENT COMPANY (Continued)

Balance sheets

	Notes	As of December 31,	
		2018	2019
		US\$	US\$
ASSETS			
Current assets:			
Cash and cash equivalents		10,748,281	91,168,159
Short-term investments		33,000,000	8,000,000
Amounts due from related parties ^(a)		7,822,425	6,362,724
Prepayments and other current assets		1,237,469	1,211,891
Total current assets		52,808,175	106,742,774
Investments in subsidiaries		2,172,601	2,507,052
Other non-current assets		9,916	4,299
TOTAL ASSETS		54,990,692	109,254,125
LIABILITIES, MEZZANINE EQUITY AND SHAREHOLDERS' DEFICIT			
Current liabilities:			
Accounts payable		391,385	326,762
Contract liabilities		—	325,000
Amounts due to related parties ^(a)		9,383,956	11,247,674
Accruals and other current liabilities		79,489	162,341
Warrant liabilities		1,207,415	—
Total current liabilities		11,062,245	12,061,777
TOTAL LIABILITIES		11,062,245	12,061,777

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

18. CONDENSED FINANCIAL INFORMATION OF THE PARENT COMPANY (Continued)

Balance sheets (Continued)

	As of December 31,	
	2018	2019
	US\$	US\$
LIABILITIES, MEZZANINE EQUITY AND SHAREHOLDERS' DEFICIT		
(CONTINUED)		
Mezzanine equity:		
Series A-1 convertible redeemable preferred shares (par value of US\$0.0001 per share; 5,473,957 and 5,473,957 shares authorized, issued and outstanding as of December 31, 2018 and 2019, respectively)	5,473,957	5,473,957
Series A-2 convertible redeemable preferred shares (par value of US\$0.0001 per share; 2,370,414 and 2,370,414 shares authorized, issued and outstanding as of December 31, 2018 and 2019, respectively)	3,000,000	3,000,000
Series B convertible redeemable preferred shares (par value of US\$0.0001 per share; 7,494,537 and 7,494,537 shares authorized, issued and outstanding as of December 31, 2018 and 2019, respectively)	27,999,995	27,999,995
Series C-1 convertible redeemable preferred shares (par value of US\$0.0001 per share; 5,597,354 and 5,597,354 shares authorized, issued and outstanding as of December 31, 2018 and 2019, respectively)	48,481,159	48,727,343
Series C-2 convertible redeemable preferred shares (par value of US\$0.0001 per share; nil and 1,861,121 shares authorized, issued and outstanding as of December 31, 2018 and 2019, respectively)	—	18,999,999
Series C-3 convertible redeemable preferred shares (par value of US\$0.0001 per share; nil and 4,452,441 shares authorized, issued and outstanding as of December 31, 2018 and 2019, respectively)	—	50,000,000
Total mezzanine equity	84,955,111	154,201,294
Shareholders' deficit:		
Ordinary shares (par value of US\$0.0001 per share; 500,000,000 and 500,000,000 shares authorized; 15,159,136 and 15,193,136 shares issued and outstanding as of December 31, 2018 and 2019, respectively)	1,516	1,519
Subscriptions receivable from shareholders	(197,068)	(197,068)
Additional paid-in capital	6,405,318	6,789,542
Accumulated other comprehensive loss	(410,693)	(344,894)
Accumulated deficit	(46,825,737)	(63,258,045)
Total shareholders' deficit	(41,026,664)	(57,008,946)
TOTAL LIABILITIES, MEZZANINE EQUITY AND SHAREHOLDERS' DEFICIT	54,990,692	109,254,125

ADAGENE INC.

NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019

18. CONDENSED FINANCIAL INFORMATION OF THE PARENT COMPANY (Continued)

Statements of comprehensive loss

	For the years ended	
	December 31,	
	2018	2019
	US\$	US\$
Revenues		
Service revenue from a related party ^(b)	—	288,983
Expenses		
Research and development expenses	(15,803,341)	(18,318,724)
Third parties	(1,810,899)	(2,844,722)
Related parties	(13,992,442)	(15,474,002)
Administrative expenses	(556,382)	(787,568)
Loss from operations	(16,359,723)	(18,817,309)
Interest income	691,448	908,981
Foreign exchange loss, net	—	(47)
Change in fair value of warrant liabilities	534,305	1,207,415
Equity in income (share of losses) of subsidiaries	(132,214)	268,652
Loss before income tax	(15,266,184)	(16,432,308)
Income tax expense	—	—
Net loss attributable to Adagene Inc.'s shareholders	(15,266,184)	(16,432,308)
Other comprehensive income (loss)		
Foreign currency translation adjustments, net of nil tax	(11,288)	65,799
Total comprehensive loss attributable to Adagene Inc.'s shareholders	(15,277,472)	(16,366,509)
Net loss attributable to Adagene Inc.'s shareholders	(15,266,184)	(16,432,308)
Deemed contribution from convertible redeemable preferred shareholders	1,186,187	—
Accretion of convertible redeemable preferred shares to redemption value	(222,846)	(246,184)
Net loss attributable to ordinary shareholders	(14,302,843)	(16,678,492)

ADAGENE INC.
NOTES TO THE CONSOLIDATED FINANCIAL STATEMENTS (Continued)
FOR THE YEARS ENDED DECEMBER 31, 2018 AND 2019
18. CONDENSED FINANCIAL INFORMATION OF THE PARENT COMPANY (Continued)
Statements of cash flows

	For the years ended	
	December 31,	
	2018	2019
	US\$	US\$
Net cash used in operating activities	(16,346,700)	(14,521,997)
Third parties	(2,919,363)	(2,660,397)
Related parties	(13,427,337)	(11,861,600)
Net cash generated from (used in) investing activities	(28,535,658)	25,941,876
Net cash generated from financing activities	50,018,733	68,999,999
Net increase in cash and cash equivalents	5,136,375	80,419,878
Cash and cash equivalents at the beginning of year	5,611,906	10,748,281
Cash and cash equivalents at the end of year	10,748,281	91,168,159

(a) The Company had the following related party balances at the end of the year:

	As of December 31,	
	2018	2019
	US\$	US\$
Adagene (Hong Kong) Limited ⁽ⁱ⁾	2,522,425	3,032,766
Adagene Incorporated ⁽ⁱ⁾	5,300,000	2,790,233
WuXi AppTec Group	—	517,168
Adagene Australia PTY Ltd. ⁽ⁱ⁾	—	22,557
Total amounts due from related parties	7,822,425	6,362,724

	As of December 31,	
	2018	2019
	US\$	US\$
Adagene (Suzhou) Limited ⁽ⁱⁱ⁾	8,812,400	9,126,474
Adagene Incorporated ⁽ⁱⁱ⁾	571,543	1,743,435
WuXi AppTec Group	—	377,752
Adagene (Hong Kong) Limited	13	13
Total amounts due to related parties	9,383,956	11,247,674

(i) As of December 31, 2018 and 2019, the amounts due from Adagene (Hong Kong) Limited, Adagene Incorporated and Adagene Australia PTY Ltd. represented the receivables arising from the expenses paid by the Company on behalf of these subsidiaries.

(ii) As of December 31, 2018 and 2019, the amounts due to Adagene (Suzhou) Limited and Adagene Incorporated mainly represented the payables arising from the research and development services provided by these two subsidiaries.

(b) For the years ended December 31, 2019, the service revenue from a related party was for Adagene Incorporated, with zero gross margin rate charged.

ADAGENE INC.

**CONSOLIDATED BALANCE SHEET AS OF DECEMBER 31, 2019 AND
UNAUDITED INTERIM CONDENSED CONSOLIDATED BALANCE SHEET
AS OF SEPTEMBER 30, 2020**

	Notes	As of	As of September 30,	
		December 31, 2019	2020	2020 US\$ (Pro forma) (Note 13)
		US\$	US\$	
ASSETS				
Current assets:				
Cash and cash equivalents		92,532,788	82,894,840	82,894,840
Short-term investments		8,000,000	—	—
Accounts receivable, net	3	480,000	—	—
Amounts due from related parties	14	1,433,186	407,381	407,381
Prepayments and other current assets	4	1,476,973	3,303,782	3,303,782
Total current assets		103,922,947	86,606,003	86,606,003
Property, equipment and software, net	5	1,879,325	1,844,382	1,844,382
Other non-current assets		87,227	626,394	626,394
TOTAL ASSETS		105,889,499	89,076,779	89,076,779
LIABILITIES, MEZZANINE EQUITY AND SHAREHOLDERS' DEFICIT				
Current liabilities:				
Accounts payable		712,714	1,207,305	1,207,305
Contract liabilities		993,378	887,363	887,363
Amounts due to related parties	14	1,895,779	2,386,690	2,386,690
Accruals and other current liabilities	6	2,540,164	2,789,161	2,789,161
Short-term borrowings	7	716,723	2,202,611	2,202,611
Current portion of long-term borrowings	7	322,525	943,452	943,452
Total current liabilities		7,181,283	10,416,582	10,416,582
Long-term borrowings	7	1,515,868	3,135,049	3,135,049
Other non-current liabilities		—	58,736	58,736
TOTAL LIABILITIES		8,697,151	13,610,367	13,610,367
Commitments and contingencies	15			
LIABILITIES, MEZZANINE EQUITY AND SHAREHOLDERS' DEFICIT (CONTINUED)				
Mezzanine equity:				
Series A-1 convertible redeemable preferred shares (par value of US\$0.0001 per share; 5,473,957 and 5,473,957 shares authorized, issued and outstanding as of December 31, 2019 and September 30, 2020 respectively; and none outstanding on a pro-forma basis as of September 30, 2020)		5,473,957	5,473,957	—
Series A-2 convertible redeemable preferred shares (par value of US\$0.0001 per share; 2,370,414 and 2,370,414 shares authorized, issued and outstanding as of December 31, 2019 and September 30, 2020 respectively; and none outstanding on a pro-forma basis as of September 30, 2020)		3,000,000	3,000,000	—
Series B convertible redeemable preferred shares (par value of US\$0.0001 per share; 7,494,537 and 7,494,537 shares authorized, issued and outstanding as of December 31, 2019 and September 30, 2020 respectively; and none outstanding on a pro-forma basis as of September 30, 2020)		27,999,995	27,999,995	—
Series C-1 convertible redeemable preferred shares (par value of US\$0.0001 per share; 5,597,354 and 5,597,354 shares authorized, issued and outstanding as of December 31, 2019 and September 30, 2020 respectively; and none outstanding on a pro-forma basis as of September 30, 2020)		48,727,343	48,912,970	—
Series C-2 convertible redeemable preferred shares (par value of US\$0.0001 per share; 1,861,121 and 1,861,121 shares authorized, issued and outstanding as of December 31, 2019 and September 30, 2020 respectively; and none outstanding on a pro-forma basis as of September 30, 2020)		18,999,999	18,999,999	—
Series C-3 convertible redeemable preferred shares (par value of US\$0.0001 per share; 4,452,441 and 4,452,441 shares authorized, issued and outstanding as of December 31, 2019 and September 30, 2020 respectively; and none outstanding on a pro-forma basis as of September 30, 2020)		50,000,000	50,000,000	—
Total mezzanine equity		154,201,294	154,386,921	—
Shareholders' deficit:				
Ordinary shares (par value of US\$0.0001 per share; 500,000,000 and 500,000,000 shares authorized; 15,193,136 and 16,466,897 shares issued and outstanding as of December 31, 2019 and September 30, 2020)		1,519	1,646	4,372
Subscriptions receivable from shareholders		(197,068)	(1,974,542)	(1,974,542)
Additional paid-in capital		6,789,542	16,499,520	170,883,715
Accumulated other comprehensive loss		(344,894)	(559,674)	(559,674)
Accumulated deficit		(63,258,045)	(92,887,459)	(92,887,459)
Total shareholders' equity (deficit)		(57,008,946)	(78,920,509)	75,466,412
TOTAL LIABILITIES, MEZZANINE EQUITY AND SHAREHOLDERS' DEFICIT		105,889,499	89,076,779	89,076,779

The accompanying notes are an integral part of these unaudited interim condensed consolidated financial statements.

ADAGENE INC.

UNAUDITED INTERIM CONDENSED CONSOLIDATED STATEMENTS
OF COMPREHENSIVE LOSS

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

	Notes	For the nine months ended September 30,	
		2019 US\$	2020 US\$
Revenues			
Licensing revenue	10	—	309,500
Expenses			
Research and development expenses		(11,517,923)	(23,567,779)
Third parties		(7,388,214)	(16,133,614)
Related parties		(4,129,709)	(7,434,165)
Administrative expenses		(2,268,038)	(7,448,069)
Loss from operations		(13,785,961)	(30,706,348)
Interest income		711,964	612,898
Interest expense		(101,506)	(113,733)
Other income		217,404	654,755
Foreign exchange loss, net		(67,673)	(76,986)
Change in fair value of warrant liabilities		1,207,415	—
Loss before income tax		(11,818,357)	(29,629,414)
Income tax expense	11	—	—
Net loss attributable to Adagene Inc.'s shareholders		(11,818,357)	(29,629,414)
Other comprehensive income			
Foreign currency translation adjustments, net of nil tax		281,021	(214,780)
Total comprehensive loss attributable to Adagene Inc.'s shareholders		(11,537,336)	(29,844,194)
Net loss attributable to Adagene Inc.'s shareholders		(11,818,357)	(29,629,414)
Accretion of convertible redeemable preferred shares to redemption value	8	(184,014)	(185,627)
Net loss attributable to ordinary shareholders		(12,002,371)	(29,815,041)
Weighted average number of ordinary shares used in per share calculation:			
—Basic	12	15,172,711	15,757,463
—Diluted	12	15,172,711	15,757,463
Net loss per ordinary share			
—Basic	12	(0.79)	(1.89)
—Diluted	12	(0.79)	(1.89)

The accompanying notes are an integral part of these unaudited interim condensed consolidated financial statements.

ADAGENE INC.

UNAUDITED INTERIM CONDENSED CONSOLIDATED STATEMENTS OF CHANGES IN SHAREHOLDERS' DEFICIT
FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

	Ordinary shares		Subscriptions receivable from shareholders US\$	Additional paid-in capital US\$	Accumulated other comprehensive loss US\$	Accumulated deficit US\$	Total shareholders' deficit US\$
	Number of shares	Amount US\$					
Balance as of							
January 1, 2019	15,159,136	1,516	(197,068)	6,405,318	(410,693)	(46,825,737)	(41,026,664)
Net loss	—	—	—	—	—	(11,818,357)	(11,818,357)
Other comprehensive income	—	—	—	—	281,021	—	281,021
Exercise of share options (Note 9)	34,000	3	—	18,697	—	—	18,700
Share-based compensation	—	—	—	438,888	—	—	438,888
Accretion of convertible redeemable preferred shares to redemption value	—	—	—	(184,014)	—	—	(184,014)
Balance as of							
September 30, 2019	15,193,136	1,519	(197,068)	6,678,889	(129,672)	(58,644,094)	(52,290,426)
Balance as of							
January 1, 2020	15,193,136	1,519	(197,068)	6,789,542	(344,894)	(63,258,045)	(57,008,946)
Net loss	—	—	—	—	—	(29,629,414)	(29,629,414)
Other comprehensive income	—	—	—	—	(214,780)	—	(214,780)
Exercise of share options (Note 9)	1,273,761	127	(1,777,474)	1,849,894	—	—	72,547
Share-based compensation	—	—	—	8,045,711	—	—	8,045,711
Accretion of convertible redeemable preferred shares to redemption value	—	—	—	(185,627)	—	—	(185,627)
Balance as of							
September 30, 2020	16,466,897	1,646	(1,974,542)	16,499,520	(559,674)	(92,887,459)	(78,920,509)

The accompanying notes are an integral part of these unaudited interim condensed consolidated financial statements.

ADAGENE INC.

UNAUDITED INTERIM CONDENSED CONSOLIDATED STATEMENTS OF CASH FLOWS

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

	For the nine months ended	
	September 30,	
	2019	2020
	US\$	US\$
Cash flows from operating activities:		
Net loss	(11,818,357)	(29,629,414)
Adjustments to reconcile net loss to net cash used in operating activities:		
Depreciation and amortization	602,687	647,220
Loss on disposal of property, equipment and software	—	323
Share-based compensation	438,888	8,045,711
Change in fair value of warrant liabilities	(1,207,415)	—
Foreign exchange loss, net	67,673	76,986
Changes in operating assets and liabilities:		
Accounts receivable, net	—	480,000
Prepayments and other current assets	(420,268)	(1,826,809)
Amounts due from related parties	(64,915)	1,025,805
Other non-current assets	6,334	4,759
Accounts payable	(70,722)	494,591
Contract liabilities	985,539	(106,015)
Amounts due to related parties	(540,456)	490,911
Accruals and other current liabilities	230,081	248,997
Other non-current liabilities	—	58,736
Net cash used in operating activities	(11,790,931)	(19,988,199)
Cash flows from investing activities:		
Placement of short-term investments	(19,000,000)	—
Withdrawal of short-term investments	38,000,000	8,000,000
Proceeds from disposal of property, equipment and software	—	8,040
Purchase of property, equipment and software	(57,545)	(574,447)
Net cash generated from investing activities	18,942,455	7,433,593
Cash flows from financing activities:		
Proceeds from employee's exercise of share-options	—	72,547
Proceeds from borrowings	2,699,114	4,505,342
Proceeds from issuance of convertible redeemable preferred shares and warrants	16,000,001	—
Payment of deferred initial public offering cost	—	(543,926)
Repayment of borrowings	(2,389,081)	(936,825)
Net cash generated from financing activities	16,310,034	3,097,138
Effect of exchange rate on cash and cash equivalents	152,933	(180,480)
Net increase in cash and cash equivalents	23,614,491	(9,637,948)
Cash and cash equivalents at the beginning of period	16,058,455	92,532,788
Cash and cash equivalents at the end of period	39,672,946	82,894,840
Supplemental cashflow disclosures:		
Interest paid	101,506	113,733
Non-cash activities:		
Accretion of convertible redeemable preferred shares to redemption value	184,014	185,627
Payables for deferred initial public offering cost	—	21,450

The accompanying notes are an integral part of these unaudited interim condensed consolidated financial statements.

ADAGENE INC.

NOTES TO THE UNAUDITED INTERIM CONDENSED

CONSOLIDATED FINANCIAL STATEMENTS

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

1. ORGANIZATION

Adagene Inc. (the "Company") is a limited liability company incorporated in the Cayman Islands on February 25, 2011. The Company, together with its subsidiaries (collectively, the "Group") are principally engaged in research, development and production of monoclonal antibody drugs for cancers.

As of September 30, 2020, the Company's principal subsidiaries are as follows:

<u>Entity</u>	<u>Date of incorporation</u>	<u>Place of incorporation</u>	<u>Percentage of legal ownership by the Company</u>	<u>Principal activities</u>
Adagene (Hong Kong) Limited	December 12, 2011	Hong Kong	100%	Investment holding
Adagene Incorporated	September 20, 2017	The United States of America	100%	Research and development of innovative medicines
Adagene (Suzhou) Limited	February 28, 2012	The People's Republic of China ("PRC" or "China")	100%	Research and development of innovative medicines
Adagene Australia PTY Ltd.	May 30, 2018	Australia	100%	Research and development of innovative medicines
Adagene PTE. Ltd.	March 27, 2020	Singapore	100%	Research and development of innovative medicines
Adagene AG	August 31, 2020	Switzerland	100%	Research and development of innovative medicines

2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES***Basis of presentation***

The accompanying unaudited interim condensed consolidated financial statements have been prepared in accordance with U.S. generally accepted accounting principles ("U.S. GAAP") and applicable rules and regulations of the Securities and Exchange Commission regarding financial reporting that are consistent with those used in the preparation of the Company's audited consolidated financial statements for the year ended December 31, 2019. Accordingly, these unaudited interim condensed consolidated financial statements do not include all of the information and footnotes required by U.S. GAAP for annual financial statements.

In the opinion of management, the Group's unaudited interim condensed consolidated financial statements and accompanying notes include all adjustments (consisting of normal recurring adjustments) considered necessary for the fair statement of the Group's financial position as of September 30, 2020, and results of operations and cash flows for the nine months ended September 30, 2019 and 2020. Interim results of operations are not necessarily indicative of the results for the full year or for any future period. These unaudited interim condensed consolidated financial statements should be read in conjunction with the audited consolidated financial statements as of and for the year ended December 31, 2019, and related notes included in the Group's audited consolidated financial statements. The financial information as of December 31, 2019 presented in the unaudited interim

ADAGENE INC.

NOTES TO THE UNAUDITED INTERIM CONDENSED

CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)

condensed consolidated financial statements is derived from the audited consolidated financial statements as of December 31, 2019. Significant accounting policies followed by the Group in the preparation of the accompanying unaudited interim condensed consolidated financial statements are summarized below.

Revenue recognition

At contract inception of collaboration and out-licensing arrangements, the Group analyzes its arrangements to assess whether they are within the scope of ASC 808, Collaborative Arrangements ("ASC 808") to determine whether such arrangements involve joint operating activities performed by parties that are both active participants in the activities and exposed to significant risks and rewards dependent on the commercial success of such activities. For collaboration arrangements within the scope of ASC 808 that contain multiple elements, the Group first determines which elements of the collaboration are deemed to be within the scope of ASC 808 and those that are reflective of a vendor-customer relationship and therefore within the scope of ASC 606. For elements of collaboration arrangements that are accounted for pursuant to ASC 808, an appropriate recognition method is determined and applied consistently. Under the criteria of Accounting Standard Codification ("ASC") 606, Revenue from Contracts with Customers (Topic 606) ("ASC 606"), the Group recognizes revenue to depict the transfer of control of promised goods or services to customers in an amount that reflects the consideration to which the entity expects to receive in exchange for those goods or services.

The Group adopted ASC 606 for all periods presented. To determine revenue recognition for arrangements that an entity determines are within the scope of ASC 606, the entity performs the following five steps: (i) identify the contract(s) with a customer; (ii) identify the performance obligations in the contract; (iii) determine the transaction price, including variable consideration, if any; (iv) allocate the transaction price to the performance obligations in the contract; and (v) recognize revenue when (or as) the entity satisfies a performance obligation. The Group only applies the five-step model to contracts when it is probable that the entity will collect substantially all the consideration to which it is entitled in exchange for the goods or services it transfers to the customer. The Group reviews the contract to determine which performance obligations are distinct and represent a promise to provide distinct goods or services or a series of distinct goods or services as defined by the standard. The Group recognizes as revenue the amount of the transaction price that is allocated to each performance obligation as and when that performance obligation is satisfied.

Licenses of Intellectual Property: Upfront non-refundable payments for licensing the Group's intellectual property are evaluated to determine if the license is distinct from the other performance obligations identified in the arrangement. For licenses determined to be distinct, the Group recognizes revenues from non-refundable, up-front fees allocated to the license at a point in time, when the transfer of control of the license to the licensee occurs and the licensee is able to use and benefit from the license.

Milestone Payments: At the inception of each arrangement that includes development, commercialization, and regulatory milestone payments, the Group evaluates whether the milestones are considered probable of being reached and to the extent that a significant reversal of cumulative

ADAGENE INC.

NOTES TO THE UNAUDITED INTERIM CONDENSED

CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)

revenue would not occur in future periods, estimates the amount to be included in the transaction price using the most likely amount method. The transaction price is then allocated to each performance obligation on a relative stand-alone selling price basis, for which the Group recognizes revenue as or when the performance obligations under the contract are satisfied. At the end of each subsequent reporting period, the Group re-evaluates the probability of achieving such development milestones and any related constraint, and if necessary, adjust the estimate of the overall transaction price. Any such adjustments are recorded on a cumulative catch-up basis, which would affect revenues and earnings in the period of adjustment.

Royalties: For arrangements that include sales-based royalties, including milestone payments based on the level of sales, and the license is deemed to be the predominant item to which the royalties relate, the Group recognizes revenue at the later of (i) when the related sales occur, or (ii) when the performance obligation to which some or all of the royalty has been allocated has been satisfied (or partially satisfied).

To date, no milestone payments or royalty payments were received. Substantially all of the Group's revenue has been derived from its out-licensing agreements with respect to licensed products such as DNA sequences, cell lines, etc., and such revenues are recognized when the customer obtains control of the licensed product, which occurs at a point in time, upon delivery to the customer.

Contract assets and contract liabilities

When a customer pays consideration before the Group transfers products or services, the Group records its obligation as a contract liability; When the Group satisfies its performance obligations by providing products or services to a customer before the customer pays consideration and before payment is due, the Group recognizes its rights to consideration as a contract asset.

Research and development expenses

Elements of research and development expenses primarily include (1) payroll and other related costs of personnel engaged in research and development activities, (2) costs related to pre-clinical testing of the Group's technologies under development and clinical trials such as payments to contract research organizations ("CRO") and contract manufacturing organization ("CMO"), investigators and clinical trial sites that conduct the clinical studies; (3) costs to develop the product candidates, including raw materials and supplies, product testing, depreciation and amortization, and facility related expenses, (4) other research and development expenses. Research and development expenses are charged to expense as incurred when these expenditures relate to the Group's research and development services and have no alternative future uses.

Share-based compensation

The Company grants restricted shares and stock options to eligible employees and nonemployees and accounts for share-based compensation in accordance with ASC 718, Compensation—Stock Compensation.

ADAGENE INC.

NOTES TO THE UNAUDITED INTERIM CONDENSED

CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)

Share-based compensation awards are measured at the grant date fair value of the awards and recognized as expenses a) immediately at the grant date if no vesting conditions are required; or b) for share-based awards granted with only service conditions, using the straight-line method over the vesting period; or c) for share-based awards granted with service conditions and performance conditions, using the graded vesting method over the vesting period if and when the company concludes that it is probable that the performance conditions will be achieved.

A change in any of the terms or conditions of share-based awards is accounted for as a modification of the awards. The Group calculates incremental compensation expense of a modification as the excess of the fair value of the modified awards over the fair value of the original awards immediately before its terms are modified at the modification date. For vested awards, the Group recognizes incremental compensation cost in the period when the modification occurs. For awards not being fully vested, the Group recognizes the sum of the incremental compensation expense and the remaining unrecognized compensation expense for the original awards over the remaining requisite service period after modification.

Employee defined contribution plan

As stipulated by the regulations of the PRC, full-time employees of the Group are entitled to staff welfare benefits including medical care, welfare subsidies, unemployment insurance and pension benefits through a PRC government-mandated multi-employer defined contribution plan. The Group is required to accrue for these benefits based on certain percentages of the qualified employees' salaries. The Group is required to make contributions to the plans out of the amounts accrued. The PRC government is responsible for the medical benefits and the pension liability to be paid to these employees and the Group's obligations are limited to the amounts contributed. The Group has no further payment obligations once the contributions have been paid. The Group recorded employee benefit expenses of US\$885,255 and US\$663,034 for the nine months ended September 30, 2019 and 2020, respectively.

Borrowings

Borrowings are recognized initially at fair value, net of transaction costs incurred. Borrowings are subsequently stated at amortized cost; any difference between the proceeds (net of transaction costs) and the redemption value is recognized in the consolidated statements of comprehensive loss over the period of the borrowings using the effective interest method.

Recently issued accounting pronouncements

The Group is an emerging growth company ("EGC") as defined by the Jumpstart Our Business Startups Act ("JOBS Act"). The JOBS Act provides that an EGC can take advantage of extended transition periods for complying with new or revised accounting standards. This allows an EGC to delay adoption of certain accounting standards until those standards would otherwise apply to private companies. The Group elected to take advantage of the extended transition periods. However, this election will not apply should the Group cease to be classified as an EGC.

ADAGENE INC.

NOTES TO THE UNAUDITED INTERIM CONDENSED

CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)

In February 2016, the FASB issued ASU No. 2016-02 ("ASU 2016-02"), Leases (Topic 842), which modifies lease accounting for lessees to increase transparency and comparability by recording lease assets and liabilities for operating leases and disclosing key information about leasing arrangements. In July 2018, the FASB issued ASU No. 2018-10 ("ASU 2018-10"), Codification Improvements to Topic 842, Leases, which clarifies certain aspects of the guidance issued in ASU 2016-02; and ASU No. 2018-11 ("ASU 2018-11"), Leases (Topic 842): Targeted Improvements, which provides entities with an additional (and optional) transition method to adopt the new leases standard. Under this new transition method, an entity initially applies the new leases standard at the adoption date and recognizes a cumulative-effect adjustment to the opening balance of retained earnings in the period of adoption. Consequently, an entity's reporting for the comparative periods presented in the financial statements in which it adopts the new leases standard will continue to be in accordance with current GAAP (Topic 840, Leases). In November 2019, the FASB issued ASU No. 2019-10, Financial Instruments—Credit Losses (Topic 326), Derivatives and Hedging (Topic 815), and Leases (Topic 842), Effective Dates ("ASU 2019-10"), which extends the adoption date for certain registrants. The updated guidance is effective for the Group for annual reporting periods beginning January 1, 2021 and interim periods within annual periods beginning January 1, 2022. Early adoption is permitted. The Group does not plan to early adopt the new lease standards and the Group expects that applying the ASU 2016-02 would materially increase the assets and liabilities due to the recognition of right-of-use assets and lease liabilities on its interim condensed consolidated balance sheets, with an immaterial impact on its interim condensed consolidated statements of comprehensive loss and interim condensed consolidated statements of cash flows.

In June 2016, the FASB issued ASU No. 2016-13, Financial Instruments—Credit Losses (Topic 326): Measurement of Credit Losses on Financial Instruments ("ASU 2016-13"). ASU 2016-13 is intended to improve financial reporting by requiring timelier recording of credit losses on loans and other financial instruments held by financial institutions and other organizations. This ASU requires the measurement of all expected credit losses for financial assets held at the reporting date based on historical experience, current conditions, and reasonable and supportable forecasts. This ASU requires enhanced disclosures to help investors and other financial statement users better understand significant estimates and judgments used in estimating credit losses, as well as the credit quality and underwriting standards of the Group's portfolio. These disclosures include qualitative and quantitative requirements that provide additional information about the amounts recorded in the financial statements. In November 2019, the FASB issued ASU 2019-10, which extends the adoption date for certain registrants. The amendments in ASU 2016-13 are effective for fiscal years beginning after December 15, 2022, including interim periods within fiscal years beginning after December 15, 2023. The Group does not plan to early adopt ASU 2016-13 and is currently in the process of evaluating the impact of adoption of this guidance on its interim condensed consolidated financial statements.

In June 2018, the FASB issued ASU No. 2018-07, Compensation—Stock Compensation (Topic 718): Improvements to nonemployee share-based payment accounting ("ASU 2018-07"). The amendments in this update expand the scope of Topic 718 to include share-based payment transactions for acquiring goods and services from nonemployees. The amendments are effective for fiscal years beginning after December 15, 2019, and interim periods within fiscal years beginning after

ADAGENE INC.

NOTES TO THE UNAUDITED INTERIM CONDENSED

CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)

December 15, 2020. Early adoption is permitted, but no earlier than an entity's adoption date of Topic 606. The Group adopted on January 1, 2018 this guidance which do not have a significant impact on the interim condensed consolidated financial statements.

In November 2018, the FASB issued ASU 2018-18, Collaborative Arrangements (Topic 808): Clarifying the Interaction between Topic 808 and Topic 606. This update clarifies that certain transactions between participants in a collaborative arrangement should be accounted for under ASC 606 when the counterparty is a customer and precludes an entity from presenting consideration from a transaction in a collaborative arrangement as revenue from contracts with customers if the counterparty is not a customer for that transaction. The update is effective in fiscal years beginning after December 15, 2021, and interim periods therein, and early adoption is permitted for entities that have adopted ASC 606. This guidance should be applied retrospectively to the date of initial application of Topic 606. The Group does not plan to early adopt ASU 2018-18 and is currently evaluating the impact on its financial statements of adopting this guidance.

In December 2019, the FASB issued ASU 2019-12, Income Taxes (Topic 740): Simplifying the Accounting for Income Taxes. This update simplifies the accounting for income taxes as part of the FASB's overall initiative to reduce complexity in accounting standards. The amendments include removal of certain exceptions to the general principles of ASC 740, *Income taxes*, and simplification in several other areas such as accounting for a franchise tax (or similar tax) that is partially based on income. The update is effective in fiscal years beginning after December 15, 2022, and interim periods therein, and early adoption is permitted. Certain amendments in this update should be applied retrospectively or modified retrospectively, all other amendments should be applied prospectively. The Group does not plan to early adopt ASU 2019-12 and is currently evaluating the impact on its financial statements of adopting this guidance.

3. ACCOUNTS RECEIVABLE, NET

	<u>As of</u> <u>December 31,</u> <u>2019</u>	<u>As of</u> <u>September 30,</u> <u>2020</u>
	US\$	US\$
Accounts receivable	480,000	—
Allowance for doubtful accounts	—	—
	<u>480,000</u>	<u>—</u>

ADAGENE INC.**NOTES TO THE UNAUDITED INTERIM CONDENSED****CONSOLIDATED FINANCIAL STATEMENTS (Continued)****FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020****4. PREPAYMENTS AND OTHER CURRENT ASSETS**

Prepayments and other current assets consist of the following:

	<u>As of</u> <u>December 31,</u> <u>2019</u>	<u>As of</u> <u>September 30,</u> <u>2020</u>
	US\$	US\$
Deposits	970,394	1,744,176
Prepayments(a)	211,435	1,417,962
Interest receivables	227,278	5,620
Others	67,866	136,024
	<u>1,476,973</u>	<u>3,303,782</u>

Note (a): The prepayments mainly represented the Group's prepaid service fees to its CRO and CMO vendors.

5. PROPERTY, EQUIPMENT AND SOFTWARE

Property, equipment and software consist of the following:

	<u>As of</u> <u>December 31,</u> <u>2019</u>	<u>As of</u> <u>September 30,</u> <u>2020</u>
	US\$	US\$
Machinery and laboratory equipment	3,462,343	3,791,884
Electronic equipment	605,882	851,996
Leasehold improvements	767,205	785,917
Furniture and tools	98,396	112,602
Computer software	71,056	102,986
Vehicles	80,133	82,087
	<u>5,085,015</u>	<u>5,727,472</u>
Accumulated depreciation and amortization	<u>(3,205,690)</u>	<u>(3,883,090)</u>
	<u>1,879,325</u>	<u>1,844,382</u>

Depreciation and amortization expenses recognized for the nine months ended September 30, 2019 and 2020 were US\$602,687 and US\$647,220, respectively.

ADAGENE INC.

NOTES TO THE UNAUDITED INTERIM CONDENSED

CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

6. ACCRUALS AND OTHER CURRENT LIABILITIES

Accrued liabilities and other current liabilities consist of the following:

	As of December 31, 2019	As of September 30, 2020
	US\$	US\$
Payroll and related liabilities	2,370,523	2,407,232
Professional service fees	145,157	331,493
Utility and maintenance	4,595	19,176
Others	19,889	31,260
	<u>2,540,164</u>	<u>2,789,161</u>

7. BORROWINGS

	As of December 31, 2019	As of September 30, 2020
	US\$	US\$
Current		
Short-term borrowings:		
Bank loans	716,723	2,202,611
Current portion of long-term borrowings	322,525	943,452
Total current borrowings	<u>1,039,248</u>	<u>3,146,063</u>
Non-Current		
Long-term borrowings:		
Bank loans	1,515,868	3,135,049
Total non-current borrowings	<u>1,515,868</u>	<u>3,135,049</u>
Total borrowings	<u><u>2,555,116</u></u>	<u><u>6,281,112</u></u>

Short-term borrowing

In March 2018, the Group borrowed a loan with the amount of RMB3,000,000 (equivalent to approximately US\$437,114) from Bank of Jiangsu Co., Ltd. for a term of one year and at the interest rate of 5.22% per annum. The borrowing was repaid in March 2019. In June 2018, the Group borrowed a loan with the amount of RMB2,000,000 (equivalent to approximately US\$291,409) from Bank of Jiangsu Co., Ltd. for a term of one year and at the interest rate of 5.22% per annum. The borrowing was repaid in June 2019. These borrowings with Bank of Jiangsu Co., Ltd were guaranteed by Peter Luo and Kristine She. Peter Luo is the Chairman, Chief Executive Officer and a principal shareholder of the Company. Kristine She is one of the senior management personnel of the Company.

In May 2018, the Group borrowed a loan with the amount of RMB3,000,000 (equivalent to approximately US\$437,114) from Bank of Ningbo Co., Ltd. for a term of one year and at the interest

ADAGENE INC.

NOTES TO THE UNAUDITED INTERIM CONDENSED

CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

7. BORROWINGS (Continued)

rate of 5.00% per annum. The borrowing was repaid in May 2019. In June 2018, the Group borrowed a loan with the amount of RMB2,000,000 (equivalent to approximately US\$291,409) from Bank of Ningbo Co., Ltd. for a term of one year and at the interest rate of 5.00% per annum. The borrowing was repaid in June 2019.

In July 2018, the Group borrowed a loan with amount of RMB6,000,000 (equivalent to approximately US\$874,228) from Agricultural Bank of China Limited for a term of one year and at the interest rate of 5.22% per annum. The borrowing was guaranteed by Peter Luo, who is the Chairman, Chief Executive Officer and a principal shareholder of the Company. The borrowing was repaid in July 2019.

In September 2019, the Group borrowed a loan with amount of RMB5,000,000 (equivalent to approximately US\$716,723) from Bank of Ningbo Co., Ltd. for a term of one year and at the interest rate of 4.35% per annum. The borrowing was repaid in September 2020.

In June 2020, the Group borrowed a loan with amount of RMB10,000,000 (equivalent to approximately US\$1,412,529) from Agricultural Bank of China Limited for a term of one year and at the interest rate of 4.20% per annum.

In September 2020, the Group borrowed a loan with amount of RMB5,000,000 (equivalent to approximately US\$734,204) from Bank of Ningbo Co., Ltd. for a term of one year and at the interest rate of 4.20% per annum.

Long-term borrowings

In February 2019, the Group borrowed a loan with amount of RMB7,500,000 (equivalent to approximately US\$1,075,084) from Shanghai Pudong Development Bank Co., Ltd. for a term of three years and at the interest rate of 5.46% per annum. The Group repaid RMB375,000 (equivalent to approximately US\$53,754), RMB375,000 (equivalent to approximately US\$53,365) and RMB 875,000 (equivalent to approximately US\$125,148), in August 2019, February 2020 and August 2020 respectively. As of December 31, 2019 and September 30, 2020, RMB1,250,000 (equivalent to approximately US\$179,181) and RMB3,375,000 (equivalent to approximately US\$495,587) repayable within twelve months for this agreement was classified as "Current portion of long-term borrowing", respectively.

In June 2019, the Group borrowed a loan with amount of RMB6,000,000 (equivalent to approximately US\$860,067) from Shanghai Pudong Development Bank Co., Ltd. for a term of three years and at the interest rate of 5.23% per annum. The Group repaid RMB300,000 (equivalent to approximately US\$43,003) and RMB300,000 (equivalent to approximately US\$42,908) in December 2019 and June 2020, respectively. As of December 31, 2019 and September 30, 2020, RMB1,000,000 (equivalent to approximately US\$143,344) and RMB1,400,000 (equivalent to approximately US\$205,577) repayable within twelve months for this agreement was classified as "Current portion of long-term borrowing", respectively.

In September 2020, the Group borrowed a loan with amount of RMB16,500,000 (equivalent to approximately US\$2,422,872) from Shanghai Pudong Development Bank Co., Ltd. for a term of three years and at the interest rate of 4.27% per annum. As of September 30, 2020, RMB1,650,000

ADAGENE INC.**NOTES TO THE UNAUDITED INTERIM CONDENSED****CONSOLIDATED FINANCIAL STATEMENTS (Continued)****FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020****7. BORROWINGS (Continued)**

(equivalent to approximately US\$242,287) repayable within twelve months for this agreement was classified as "Current portion of long-term borrowings".

Future maturities of short-term borrowings and long-term borrowings

Future principal maturities of short-term borrowings and long-term borrowings as of September 30, 2020 are as follows:

	US\$
Remaining three months of 2020	102,789
2021	3,336,955
2022	1,226,120
2023	1,615,248
Total	<u>6,281,112</u>

8. CONVERTIBLE REDEEMABLE PREFERRED SHARES AND WARRANTS

In November 2011, the Company issued convertible notes ("Series Pre-A Convertible Notes") to certain investors in the amount of 4,590,908. The notes carried a simple interest (non-compounding) of 6% per annum as set out in the note purchase agreement. All outstanding principal balance and accrued but unpaid interest of the notes should be automatically converted into the convertible redeemable preferred shares of the Company at a price no more than US\$1 per share.

In November 2014, the Company issued 5,473,957 Series A-1 convertible redeemable preferred shares ("Series A-1 Preferred Shares") to certain investors upon conversion of the Company's Series Pre-A convertible notes at a conversion price of US\$1 per share. Concurrently, the Company issued 2,370,414 Series A-2 convertible redeemable preferred shares ("Series A-2 Preferred Shares") to certain investors at US\$1.27 per share for a total consideration of US\$3,000,000. Series A-1 Preferred Shares and Series A-2 Preferred Shares are collectively referred to as the Series A Preferred Shares.

From January through June 2016, the Company issued 7,494,537 Series B convertible redeemable preferred shares ("Series B Preferred Shares") to certain investors at US\$3.74 per share for a total consideration of US\$27,999,995.

From February through May 2018, the Company issued 5,597,354 Series C-1 convertible redeemable preferred shares ("Series C-1 Preferred Shares") to certain investors at US\$8.93 per share for a total consideration of US\$50,000,033. Concurrently, in February 2018, the Company also issued warrants to two Series C-1 investors at nil consideration ("Series C-1 Warrants"). The Series C-1 Warrants allowed the holders to purchase Series C-2 Preferred Shares (defined below) at the exercise price of US\$10.21 per share for a total consideration of up to US\$7,500,000. Series C-1 Warrants were exercisable, in whole or in part, at any time from the warrant issuance date to the earlier of i) April 1, 2019, ii) a deemed liquidation event or iii) the closing of the Qualified IPO. Series C-1 Warrants expired on April 1, 2019.

ADAGENE INC.

NOTES TO THE UNAUDITED INTERIM CONDENSED

CONSOLIDATED FINANCIAL STATEMENTS (Continued)

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8. CONVERTIBLE REDEEMABLE PREFERRED SHARES AND WARRANTS (Continued)

From June through November 2019, the Company issued 1,861,121 Series C-2 convertible redeemable preferred shares ("Series C-2 Preferred Shares") to certain investors at US\$10.21 per share for a total consideration of US\$18,999,999.

In December 2019, the Company issued 4,452,441 Series C-3 convertible redeemable preferred shares ("Series C-3 Preferred Shares") to a certain investor at US\$11.23 per share for a total consideration of US\$50,000,000.

Series C-1 Preferred Shares, Series C-2 Preferred Shares and Series C-3 Preferred Shares are collectively referred to as the Series C Preferred Shares.

The key features of the Series A Preferred Shares, Series B Preferred Shares and Series C Preferred Shares (collectively the "Preferred Shares") are as follows:

Dividends

Each holder of the Preferred Shares will be entitled to receive non-cumulative dividends when declared by the Board of Directors prior and in preference to ordinary shareholders. The dividend should be paid at the rate of 6% of the original issue price per share per annum on each Preferred Shares in the sequence of Series C Preferred Shares and Preferred Shares other than the Series C Preferred Shares. After the preferential dividends relating to the Preferred Shares have been paid in full or declared and set apart in any fiscal year of the Company, any additional dividends out of funds or assets legally available therefore may be declared in that fiscal year for the Shares and, if such additional dividends are declared, the preferred shareholders shall be entitled to participate on an as converted-basis pro-rata in any dividends or distributions paid to the ordinary shareholders.

Voting

Each Preferred Share has voting rights equivalent to the number of ordinary shares to which it is convertible at the record date. The Preferred Shares shall vote separately as a class with respect to certain specified matters. Otherwise, the preferred shareholders and ordinary shareholders shall vote together as a single class.

Liquidation preference

In the event of any liquidation, dissolution or winding up of the Company, or the cessation of the business of the Company or of a substantial portion of the business of the Company, whether voluntary or involuntary, or any deemed liquidation event (unless waived by the preferred shareholders), the preferred shareholders shall be entitled to receive a per share amount equal to 100% of the original issue price of the respective series of the Preferred Shares, in the sequence of Series C Preferred Shares, Series B Preferred Shares and Series A Preferred Shares. After such liquidation amounts have been paid in full, the preferred shareholders are entitled to receive a simple interest accruing on each Preferred Share at 6% of its original issue price per annum from the date of issuance of such Preferred Share to the date of distribution of such amount, in the sequence of Series C Preferred Shares, Series B Preferred Shares and Series A Preferred Shares. After such interest amounts have been paid in full, any remaining funds or assets of the Company legally available for distribution to shareholders shall be distributed on a pro rata basis among the preferred shareholders, on an as-converted basis, together with the ordinary shareholders.

ADAGENE INC.

NOTES TO THE UNAUDITED INTERIM CONDENSED

CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

8. CONVERTIBLE REDEEMABLE PREFERRED SHARES AND WARRANTS (Continued)

Conversion

Each Preferred Share may be converted at any time into ordinary shares at the option of the preferred shareholders based on the then-effective conversion price. The initial conversion ratio is 1:1, subject to adjustment in the event of share splits, share combinations, ordinary share dividends or distributions, other dividends, reorganizations, mergers, consolidations, reclassifications, exchanges, substitutions, or dilutive issuance.

All Preferred Shares are converted automatically into ordinary shares at the then effective applicable conversion price upon the earlier of a Qualified Public Offering (public offering of the Company's shares with an offering price (exclusive of underwriting commissions and expenses) that reflects a market capitalization (immediately prior to the public offering) of not less than US\$650,000,000 and with an aggregate proceeds of no less than US\$75 million) or a date specified by written consent or agreement of the holders of at least 80% of the voting power of the then outstanding Preferred Shares.

Redemption

The Preferred Shares are redeemable upon request by the holders of the majority outstanding Preferred Shares if the Company fails to consummate a Qualified Public Offering or complete a deemed liquidation event on or before March 31, 2025 at the redemption price equal to the original issue price plus any declared but unpaid dividends.

Accounting for Preferred Shares

The Preferred Shares are classified as mezzanine equity in the consolidated balance sheets because they are contingently redeemable upon the occurrence of an event outside of the Company's control (e.g. the Company not achieving a Qualified Public Offering or a deemed liquidation event before March 31, 2025 ("Target QIPO Date")). The Preferred Shares were determined to be mezzanine equity with no embedded feature to be bifurcated and no beneficial conversion features to be recognized. The Preferred Shares are initially recorded at their respective issuance date fair value, net of issuance cost and fair value allocated to the detachable warrants. The Company did not incur material issuance cost for any Preferred Shares issued.

The Company concluded that the Preferred Shares are not currently redeemable, but are probable to become redeemable. The Company accreted changes in the redemption value over the period from the date of issuance to the earliest redemption date using the interest method. No accretion charge was recorded as the redemption value is fixed to original issue price for the years presented, except for Series C-1 Preferred Shares issued with detachable warrants.

Modification of Preferred Shares

The Company made several amendments to the Preferred Shares, mainly including: 1) added redemption rights for Series A Preferred Shares upon the issuance of the Series B Preferred Shares; 2) extended the Target QIPO Date upon the issuance of the Series C-1 Preferred Shares and the

ADAGENE INC.

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CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

8. CONVERTIBLE REDEEMABLE PREFERRED SHARES AND WARRANTS (Continued)

Series C-3 Preferred Shares. These amendments are accounted for as modifications rather than extinguishments as the fair values of these Preferred Shares immediately after the amendments were not significantly different from their respective fair values immediately before the amendment. When Preferred Shares are modified and such modification results in value transfer between preferred shareholders and ordinary shareholders, the value transferred is treated as a deemed dividend to or deemed contribution from the preferred shareholders.

The Company's Preferred Shares activities for the periods presented are summarized below:

Mezzanine equity	Series A-1 US\$	Series A-2 US\$	Series B US\$	Series C-1 US\$	Series C-2 US\$	Series C-3 US\$	Total US\$
Balance as of							
December 31, 2018	5,473,957	3,000,000	27,999,995	48,481,159	—	—	84,955,111
Issuance of Series C-2 Preferred Shares	—	—	—	—	16,000,001	—	16,000,001
Accretion of Series C-1 Preferred Shares to redemption value	—	—	—	184,014	—	—	184,014
Balance as of							
September 30, 2019	5,473,957	3,000,000	27,999,995	48,665,173	16,000,001	—	101,139,126
Balance as of							
December 31, 2019	5,473,957	3,000,000	27,999,995	48,727,343	18,999,999	50,000,000	154,201,294
Accretion of Series C-1 Preferred Shares to redemption value	—	—	—	185,627	—	—	185,627
Balance as of							
September 30, 2020	5,473,957	3,000,000	27,999,995	48,912,970	18,999,999	50,000,000	154,386,921

9. SHARE-BASED COMPENSATION

On November 7, 2015, the Company adopted a share incentive plan ("2015 Plan"). Under the 2015 Plan, the Company's Board of Directors has approved that a maximum aggregate number of shares that may be issued pursuant to all awards granted shall be 4,336,126. In September 2017, the Company replaced 2015 Plan with the amended and restated share incentive plan ("2017 Plan") and increased the maximum number of shares issuable to 6,336,126. In December 2019, the Company replaced 2017 Plan with the second amended and restated share incentive plan ("2019 Plan") and further increased the maximum number of shares issuable to 11,391,131. The terms of the 2015 Plan, 2017 Plan and 2019 Plan are substantially the same other than the maximum aggregate number of shares the Company may issue under the respective plan.

Share options containing only service conditions granted to each grantee under the share incentive plan will generally be exercisable upon the grantee renders service to the Company in accordance with a stipulated vesting schedule. Grantees are generally subject to a vesting schedule of no longer than five years, under which the grantee earns an entitlement to vest a certain percentage of his option grants at the end of each month or year of completed service. The share option awards shall expire no more than 10 years from their grant dates.

Share options containing both service conditions and performance conditions granted to each grantee under the share incentive plan shall become eligible for vesting upon the occurrence of their applicable performance conditions (including but not limited to the occurrence of an initial public offering ("IPO"), the completion of business and operational goals, etc.). Upon being eligible for

ADAGENE INC.

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9. SHARE-BASED COMPENSATION (Continued)

vesting, each such performance-based option shall vest in a series of four equal and continuous annual installments for each full year of services completed following the award date of such option.

As of September 30, 2020, the Company has granted 2,178,808 share options containing both service conditions and performance conditions. All other share options granted contain only service conditions.

On August 15, 2020, pursuant to board resolution, 184,692 share options that granted to a senior management was modified and the vesting period of the 184,692 share options has been shortened from 5 years to 4 years. Since the fair value of the modified award is lower than the fair value of the original award immediately before the modification, there is no incremental compensation cost to be recognized.

	Number of Options	Weighted- Average Exercise Price US\$ per option	Weighted- Average Grant Date Fair Value US\$ per option	Weighted Average Remaining Contractual Term Years	Aggregate Intrinsic Value US\$
Outstanding, January 1, 2019	574,328	0.30	2.80	7.14	3,734,644
Granted	372,500	1.26	5.46	—	—
Exercised	(34,000)	0.55	5.71	—	—
Forfeited	(46,800)	1.22	5.75	—	—
Outstanding, December 31, 2019	866,028	0.65	3.67	7.27	6,328,171
Granted	5,973,373	2.04	7.09	—	—
Exercised	(1,273,761)	1.45	5.55	—	—
Forfeited	(14,000)	1.39	6.45	—	—
Outstanding, September 30, 2020	5,551,640	1.96	6.91	9.55	37,495,526
Vested and expected to vest at September 30, 2020	5,551,640	1.96	6.91	9.55	37,495,526
Exercisable at September 30, 2020	412,387	0.91	4.86	7.32	3,217,567

The aggregate intrinsic value in the table above represents the difference between the exercise price of the awards and the fair value of the underlying ordinary shares at each reporting date, for those awards that had exercise price below the estimated fair value of the relevant ordinary shares.

The aggregate fair value of the equity awards vested during the nine months ended September 30, 2019 and 2020 was US\$ 243,927 and US\$7,699,049, respectively. As of September 30, 2020, there was US\$ 20,941,290 of total unrecognized employee share-based compensation expense, may be adjusted for actual forfeitures occurring in the future. Total unrecognized compensation cost will be recognized over a weighted-average period of 3.42 years.

ADAGENE INC.

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CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

9. SHARE-BASED COMPENSATION (Continued)

Fair value of share options

The fair value of share options was determined using the binomial option valuation model, with the assistance from an independent third-party appraiser. The binomial model requires the input of highly subjective assumptions, including the expected volatility, the exercise multiple, the risk-free rate and the dividend yield. For expected volatility, the Group has made reference to historical volatility of several comparable companies in the same industry. The exercise multiple was estimated as the average ratio of the stock price to the exercise price of when employees would decide to voluntarily exercise their vested share options. The risk-free rate for periods within the contractual life of the share options is based on the market yield of U.S. Treasury Bonds in effect at the time of grant. The dividend yield is based on the expected dividend policy over the contractual life of the share options. The estimated fair value of the ordinary shares, at the share option grant dates, was determined with the assistance from an independent third-party appraiser.

The assumptions used to estimate the fair value of the share options granted are as follows:

	For the nine months ended September 30,	
	2019	2020
Risk-free interest rate	2.73%	0.71% - 0.83%
Dividend yield	—	—
Expected volatility range	71.0%	72.3% - 73.1%
Exercise multiple	2.2	2.2 - 2.8
Contractual life	10 years	10 years

Total share-based compensation expenses recognized for the nine months ended September 30, 2019 and 2020 were as follows:

	For the nine months ended September 30,	
	2019	2020
	US\$	US\$
Research and development expenses	244,846	4,971,881
Administrative expenses	194,042	3,073,830
Total share-based compensation expenses	<u>438,888</u>	<u>8,045,711</u>

10. COLLABORATION ARRANGEMENTS

Dragon Boat Biopharmaceutical (Shanghai) Limited License Agreement

In May 2019, the Group entered into (i) a collaboration agreement that covers Greater China (the "Dragon Boat Greater China Agreement") and (ii) a collaboration agreement that covers the regions other than Greater China (the "Dragon Boat ROW Agreement," together with the Dragon Boat Greater China Agreement, the "2019 Dragon Boat Agreements"), with Dragon Boat Biopharmaceutical (Shanghai) Limited ("Dragon Boat"), a subsidiary of Sanjin. Pursuant to the Dragon Boat Greater

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CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

10. COLLABORATION ARRANGEMENTS (Continued)

China Agreement, the Group will license the Chinese intellectual property directly related to a certain monospecific antibody molecule that binds to a specified target (the "Specified Project"), including the patent rights, patent application rights and technologies based on the core sequence of the molecule, to Dragon Boat. Dragon Boat will own all the Chinese intellectual property developed in the exercise of Dragon Boat's rights under the agreement, including but not limited to improvements (including combination products), clinical trials, regulatory filings, and commercialization rights relating thereto. The Group also granted Dragon Boat a royalty-free license to use our other existing intellectual property and improvements thereto which are related to the Specified Project for the purposes of exploiting its rights and performing its obligations under the agreement. Dragon Boat will enjoy all the economic benefits deriving from the Specified Project in Greater China, including but not limited to patent transfer fee, licensing fee, sales revenue and sales commission, etc. and will pay the Group (i) certain high-six figure dollar milestone payments upon the achievement of certain milestones (including milestones of launch of pre-clinical safety evaluation animal test, obtaining Investigational New Drug ("IND") approval in PRC and completion of clinical phase I test in PRC) and (ii) a single-digit percentage of net sales of the products that use the licensed antibody after such products enter the market.

Pursuant to the Dragon Boat ROW Agreement, the Group granted Dragon Boat a royalty-free license to use all intellectual property relating to (i) the collaboration under the agreement that the Group controlled before the Group entered into the agreement or acquired independently of the agreement and (ii) improvements thereto for the purposes of exploiting its rights and performing its obligations under the agreement. Any intellectual property generated independently by a party under the agreement will be solely owned by that party who generated such intellectual property, and any intellectual property generated from cooperation between the Group and Dragon Boat in connection with the collaboration will be jointly owned. The Group retain the ownership of patent rights of key intellectual property pertaining to the specified target outside of the Greater China. In addition, all the results obtained by Dragon Boat relating to the research and development of any new antibody developed under the agreement will be owned by Dragon Boat. The Group retains a majority of the economic benefits derived from the Dragon Boat ROW Agreement, including but not limited to any patent transfer fee, licensing fee and gains realized under such transfer. In case the Group intend to transfer to a third party our share of economic interests in any country outside of Greater China, the Group must notify Dragon Boat and Dragon Boat will receive a right of first refusal if it pays the Group a deposit equal to a low double-digit percentage of the consideration that the Group expects to receive from such third party. If Dragon Boat waives the right of first refusal, the Group can proceed with the transfer, provided that the final transaction price with the third party is not lower than the amount of the offering price that was included in our notice to Dragon Boat.

Under the 2019 Dragon Boat Agreements, the Group agreed not to (i) independently develop any monospecific antibodies that bind to the specified target or (ii) grant any rights associated with such antibodies to any third parties during the three-year period from the effective date of the agreements. The exclusivity obligation does not prevent the Group from (i) developing or granting any licenses to third parties for intellectual property that covers bispecific antibodies, ADCs, diagnostic antibodies, nano-particles and masked antibody against the specific target and (ii) continuing to provide antibody

ADAGENE INC.

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CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

10. COLLABORATION ARRANGEMENTS (Continued)

screening service that were commenced before the execution of the Dragon Boat Greater China Agreement and either party has the independent right to conduct combination therapy studies outside of the Greater China. Either nonbreaching party may terminate the 2019 Dragon Boat Agreements if the other party's ability to comply with its obligations under the agreements is negatively affected by contingencies such as failure to maintain operation or changes in core project management and the other party fails to take effective remedial measures. Each agreement automatically terminates upon the termination of the other agreement. Upon the rescission or termination, Dragon Boat will return to the Group all the intellectual property, documents and data provided by the Group under the 2019 Dragon Boat Agreements.

For the nine months ended September 30, 2019 and 2020, no revenue was recognized for this agreement respectively since the licensed product has not been transferred to Dragon Boat.

As of September 30, 2020, upfront fee of RMB4,000,000 (equivalent to approximately US\$579,836) that received by the Group was recorded as contract liabilities in the consolidated balance sheets, as the performance obligation had not been satisfied by the Group.

ADC Therapeutics SA License and Collaboration Agreements

In April 2019, the Group entered into a material transfer and collaboration agreement (the "ADCT Collaboration Agreement") and a license agreement (the "ADCT License Agreement") with ADC Therapeutics SA ("ADC Therapeutics"). These two agreements are combined as a single contract as the agreements were negotiated as a package with a single commercial objective.

ADCT Collaboration Agreement

Pursuant to the ADCT Collaboration Agreement, the Group agreed to generate masked antibodies with respect to up to two exclusive targets selected by ADC Therapeutics. Upon our delivery of certain initial results, ADC Therapeutics has the option to license the Group's technology with respect to one or both targets as further detailed below. ADC Therapeutics has not yet exercised such options as of September 30, 2020.

Under the ADCT Collaboration Agreement, the Group is eligible to receive up to a low-seven-figure dollar amount in consideration for the Group's exclusivity obligations, upon achievement of certain development milestones (including milestones of delivery of certain amino acid sequences and successful outcome of the first in-vivo study) and upon ADC Therapeutics' election to proceed with development for the two elected targets. Apart from performance obligation to deliver the amino acid sequences of the corresponding masking peptides, the Group is not required to perform any additional research and development services.

ADC Therapeutics has the right to terminate the ADCT Collaboration Agreement at any time and for any reason in its entirety or on a target-by-target basis upon thirty days' prior written notice to the Group. Either party may terminate the ADCT Collaboration Agreement, in its entirety or on a target-by-target basis, upon the other party's uncured material breach of the agreement or the other party's insolvency-related events.

ADAGENE INC.

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CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

10. COLLABORATION ARRANGEMENTS (Continued)

The Group also granted ADC Therapeutics an exclusive target reservation right for one year from the commencement of the agreement and an option to renewal for another year with a consideration of low-six-figure dollar amount.

ADCT License Agreement

Subject to the exercise of the options contained in the ADCT Collaboration Agreement, the Group has granted ADC Therapeutics, with respect to each elected target, an exclusive, worldwide, perpetual and irrevocable (subject only to the termination provisions) license (with the right to grant sublicenses) to develop, make, use, commercialize and import the antibody drug conjugates that comprise masked antibodies generated by the Group under these programs.

Under the ADCT License Agreement, if ADC Therapeutics exercises both of its options granted thereunder, the Group could be eligible to receive up to a low-nine-figure dollar amount in development and regulatory milestone payments upon the achievement of certain milestones (including milestones of successful completion of Good Laboratory Practice Toxicology studies, launch of clinical trials and start of commercial sales in difference countries and etc.) and up to a mid-eight-figure dollar amount in sales milestone payments, in addition to mid-single-digit percentage net sales-based tiered royalties on products licensed under the ADCT License Agreement, subject to certain reductions. Royalties, if any, will be payable on a country-by-country and product-by-product basis, until the earlier of (i) the tenth anniversary of the first commercial sale of such product or (ii) the expiration of the last-to-expire patent licensed under the agreement in such country, unless earlier terminated by the parties, following which any licenses granted to ADC Therapeutics under the ADCT License Agreement shall become fully paid up, perpetual and irrevocable.

ADC Therapeutics has the right to terminate the ADCT License Agreement before the expiration of the royalty term on a product-by-product basis or in its entirety (i) for any reason or no reason upon thirty days' written notice to the Group, or (ii) if ADC Therapeutics chooses to discontinue the development or sale of the applicable licensed product worldwide. Each party has certain rights to terminate the ADCT License Agreement with prior written notice upon the other party's uncured material breach or insolvency.

For the nine months ended September 30, 2019 and 2020, no revenue was recognized for this agreement respectively since the licensed product has not been transferred to ADC Therapeutics. For the nine months ended September 30, 2020, the Group recognized US\$100,000 as other income due to the expiration of exclusive target reservation right, which is not related to the Group's major operation activity.

As of September 30, 2020, upfront fee of US\$225,000 that received by the Group was recorded as contract liabilities in the consolidated balance sheets, as the performance obligation had not been satisfied by the Group.

ADAGENE INC.

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CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

11. INCOME TAX EXPENSE

The Group has incurred net accumulated operating losses for income tax purposes since its inception. The Group believes that it is more likely than not that these net accumulated operating losses will not be utilized in the future. Therefore, the Group has provided full valuation allowances for the deferred tax assets as of December 31, 2019 and September 30, 2020.

12. NET LOSS PER SHARE

Basic and diluted net loss per share for the nine months ended September 30, 2019 and 2020 are calculated as follows:

	For the nine months ended September 30,	
	2019 US\$	2020 US\$
Numerator:		
Net loss attributable to Adagene Inc.'s shareholders	(11,818,357)	(29,629,414)
Accretion of convertible redeemable preferred shares to redemption value	(184,014)	(185,627)
Net loss attributable to ordinary shareholders	(12,002,371)	(29,815,041)
Denominator:		
Weighted-average number of ordinary shares outstanding—basic and diluted	15,172,711	15,757,463
Net loss per share—basic and diluted	(0.79)	(1.89)

The effects of all outstanding convertible redeemable preferred shares and share options were excluded from the computation of diluted net loss per share for the nine months ended September 30, 2019 and 2020 as their effects would be anti-dilutive.

The potentially dilutive securities that have not been included in the calculation of diluted net loss per share as their inclusion would be anti-dilutive are as follows:

	For the nine months ended September 30,	
	2019	2020
Convertible redeemable preferred shares	21,562,018	27,249,824
Share options	526,910	1,238,252

13. UNAUDITED PRO FORMA BALANCE SHEET AND NET LOSS PER SHARE

The unaudited pro forma balance sheet information as of September 30, 2020 assumes the automatic conversion of all of the outstanding convertible redeemable preferred shares into ordinary shares at a conversion ratio of 1:1, as if the conversion and expiry had occurred as of September 30, 2020.

The unaudited pro forma net loss per ordinary share is computed using the weighted-average number of ordinary shares outstanding and the automatic conversion of all of the Group's outstanding mezzanine equity into ordinary shares upon the closing of the Group's Qualified Public Offering, as if

ADAGENE INC.

NOTES TO THE UNAUDITED INTERIM CONDENSED

CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

13. UNAUDITED PRO FORMA BALANCE SHEET AND NET LOSS PER SHARE (Continued)

it had occurred on January 1, 2020. The Group believes the unaudited pro forma net loss per share provides material information to investors, as the automatic conversion of the Group's outstanding mezzanine equity. The disclosure of pro forma net loss per ordinary share provides an indication of net loss per ordinary share that is comparable to what will be reported by the Group as a public company following the closing of the Qualified Public Offering.

The unaudited basic and diluted pro forma net loss per share is calculated as follows:

	<u>For the six months ended September 30, 2020</u> US\$
Numerator:	
Net loss attributable to ordinary shareholders in computing pro forma net loss per share—basic and diluted	(29,815,041)
Add back accretion of convertible redeemable preferred shares to redemption value	185,627
Numerator for pro-forma basic and diluted net loss per share	<u>(29,629,414)</u>
Denominator:	
Weighted-average number of ordinary shares outstanding—basic and diluted	15,757,463
Add: adjustment to reflect assumed effect of automatic conversion of convertible redeemable preferred shares	27,249,824
Pro forma weighted average number of shares outstanding—basic and diluted	<u>43,007,287</u>
Pro forma net loss per share—basic and diluted	<u>(0.69)</u>

14. RELATED PARTY TRANSACTIONS

a) *Related Parties*

<u>Name of related parties</u>	<u>Relationship</u>
Peter Luo	Chairman, Chief Executive Officer and a principal shareholder of the Company
Four senior management personnel	Management and ordinary shareholders of the Company
WuXi AppTec Co., Ltd. ("WuXi AppTec Group")	A principal shareholder of the Group
WuXi Biologics (Shanghai) Co., Ltd.	Controlled by the ultimate controlling party of a principal shareholder of the Group

ADAGENE INC.

NOTES TO THE UNAUDITED INTERIM CONDENSED

CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

14. RELATED PARTY TRANSACTIONS (Continued)

b) The Group had the following related party balances at the end of the year/period:

	As of December 31, 2019	As of September 30, 2020
	US\$	US\$
WuXi Biologics (Shanghai) Co., Ltd.	4,452	367,900
WuXi AppTec Group	739,051	39,481
Four senior management personnel(i)	350,865	—
Peter Luo(i)	338,818	—
Total amounts due from related parties	1,433,186	407,381

	As of December 31, 2019	As of September 30, 2020
	US\$	US\$
WuXi Biologics (Shanghai) Co., Ltd.	1,379,741	2,032,352
WuXi AppTec Group	432,784	354,338
Peter Luo(ii)	83,254	—
Total amounts due to related parties	1,895,779	2,386,690

c) The Group had the following related party transactions during the periods:

	For the nine months ended September 30,	
	2019	2020
	US\$	US\$
Receipt of CRO and CMO services:		
WuXi Biologics (Shanghai) Co., Ltd.	2,815,862	5,107,176
WuXi AppTec Group	1,313,847	2,326,989
	4,129,709	7,434,165

(i) In October and November 2017, Peter Luo and other four senior management personnel elected to exercise the vested share options that granted under 2015 Plan. As of December 31, 2019, the balance of amounts due from Peter Luo and other four senior management personnel represented the receivables arising from withholding individual income tax amounts. The balance of amount due from four senior management personnel and Peter Luo was settled in September 2020.

(ii) In May 2014 the Group received a subsidy from the local government for attracting high skilled personnel on behalf of Peter Luo, which was settled in May 2020.

ADAGENE INC.**NOTES TO THE UNAUDITED INTERIM CONDENSED****CONSOLIDATED FINANCIAL STATEMENTS (Continued)****FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020****15. COMMITMENTS AND CONTINGENCIES*****Operating lease commitments***

Future minimum payments under non-cancelable operating leases with initial terms in excess of one year consist of the following as of September 30, 2020:

	<u>US\$</u>
Remaining three months of 2020	80,459
2021	230,084
2022	144,949
2023	104,289
Total	<u>559,781</u>

Payments under operating leases are expensed on a straight-line basis over the periods of their respective leases. The Group's lease arrangements have no renewal options, rent escalation clauses, restrictions or contingent rents and are all executed with third parties. For the nine months ended September 30, 2019 and 2020, total rental related expenses for all operating leases amounted to US\$135,750 and US\$116,061, respectively.

Contingencies

The Group is currently not involved in any legal or administrative proceedings that may have a material adverse impact on the Group's business, financial position or results of operations

16. SUBSEQUENT EVENTS

The Group evaluated subsequent events through February 1, 2021, the date these interim condensed consolidated financial statements were issued.

Beginning in January 2020, the emergence and wide spread of the novel Coronavirus ("COVID-19") has resulted in quarantines, travel restrictions, and the temporary closure of stores and facilities in China, US and elsewhere. Substantially all of the Group's operating and workforce are concentrated in China and US. Consequently, the COVID-19 outbreak could potentially delay patient's access to hospital and the progress of clinical trials of the Group, which may adversely affect the Group's business operations, financial condition and operating results for 2020. The extent to which COVID-19 impacts the business and financial results of the Group in the longer term will depend on future developments, which are uncertain and cannot be predicted, including new information which may emerge concerning the severity of the coronavirus and the actions to contain the coronavirus or treat its impact, among others. The Group will continue to evaluate the impact on the results of operation, financial position and cash flows of the Group and react actively as the situation evolves.

In October 2020, pursuant to the 2019 Plan, the Board of Directors of the Company passed resolutions and granted 340,000 share options to certain employees with an exercise price of US\$2.26 per share. The fair values of these share options ranged from US\$9.02 to US\$9.23 per option and were determined by using the binomial option valuation model. The share-based compensation expense

ADAGENE INC.

NOTES TO THE UNAUDITED INTERIM CONDENSED

CONSOLIDATED FINANCIAL STATEMENTS (Continued)

FOR THE NINE MONTHS ENDED SEPTEMBER 30, 2019 AND 2020

16. SUBSEQUENT EVENTS (Continued)

related to these share options is expected to be recognized over a weighted-average period of 4.06 years.

On November 9, 2020, the Company passed a board resolution to waive the vesting schedules and conditions of 2,375,000 share options granted to certain management members. Pursuant to this board resolution, these management members exercised all these share options at the original exercise price per share ranging from US\$1.83 to US\$2.26 and as a result, 2,375,000 ordinary shares were issued to them. These management members paid the exercise price by issuing recourse promissory notes in the total amount of US\$5,197,650. These ordinary shares are subject to the Company's repurchase rights before the share awards meet certain conditions. The management assessed and considered that such change in terms met the exception for modification accounting (as defined in ASU 2017-09, Scope of Modification Accounting), hence no additional amount of compensation expense was recognized.

In November 2020, the Group borrowed two loans with the amount of RMB5,000,000 (equivalent to approximately US\$760,086) and RMB5,000,000 (equivalent to US\$760,086) from China Merchants Bank Co., Ltd., respectively. Both of these two loans have a term of one year and bear an interest rate of 4.10% per annum.

On January 16, 2021, the Company passed a board resolution whereby certain management members surrendered a total of 491,119 ordinary shares as repayment for their respective promissory notes issued in connection with the exercising of options granted to them. In the meantime, the Company further granted these management members the right to repurchase the surrendered shares by way of new option grants with an exercise price of US\$13.85 per share. The fair values of these share options were US\$7.85 per option, determined by using the binomial option valuation model. The share-based compensation expense related to these share options is expected to be recognized immediately upon grants.

Pursuant to the same board resolution, the 2021 Performance Incentive Plan was also adopted, under which an aggregate of 2,994,000 ordinary shares shall be reserved for issuance, and the share limit will automatically increase on the first trading day in January of each year (commencing with 2022) by an amount equal to (1) 5% of the total number of the Company's outstanding ordinary shares on December 31 of the prior year, or (2) such lesser number as determined by the board of directors.

On February 1, 2021, the Company entered into a collaboration and license agreement (the "Exelixis Agreement") with Exelixis, Inc., pursuant to which the Company agreed to generate masked antibodies with the Company's SAFEbody technology against an initial target and a second target to be selected by Exelixis. Under the Exelixis Agreement, the Company are entitled to receive an US\$11,000,000 upfront payment. In addition, the Company could be eligible to receive, on a target-by-target basis, up to US\$127,500,000 in development and regulatory milestone payments upon the achievement of specified development and regulatory milestones and up to US\$262,500,000 in sales milestone payments upon the achievement of specified commercial milestones. In the aggregate, the Company could be eligible to receive up to US\$255,000,000 in development and regulatory milestone payments and up to US\$525,000,000 in sales milestone payments for both targets under the Exelixis Agreement. In addition, the Company are entitled to receive mid-single-digit percentage net sales-based royalties on products developed under the Exelixis Agreement, subject to certain reductions.

PART II**INFORMATION NOT REQUIRED IN THE PROSPECTUS****Item 6. Indemnification of Directors and Officers**

Cayman Islands law does not limit the extent to which a company's articles of association may provide for indemnification of officers and directors, except to the extent any such provision may be held by the Cayman Islands courts to be contrary to public policy, such as to provide indemnification against civil fraud or the consequences or committing a crime. Under our post-offering memorandum and articles of association, which will become effective immediately prior to the completion of this offering, to the fullest extent permissible under Cayman Islands law every director and officer of our company shall be indemnified against all actions, proceedings, costs, charges, expenses, losses, damages or liabilities incurred or sustained by him in connection with the execution or discharge of his duties, powers, authorities or discretions as a director or officer of our company, including without prejudice to the generality of the foregoing, any costs, expenses, losses or liabilities incurred by him in defending (whether successfully or otherwise) any civil proceedings concerning our company or its affairs in any court whether in the Cayman Islands or elsewhere.

Pursuant to the form of indemnification agreements to be filed as Exhibit 10.2 to this Registration Statement, we will agree to indemnify our directors and executive officers against certain liabilities and expenses that they incur in connection with claims made by reason of their being a director or officer of our company.

The Underwriting Agreement, the form of which will be filed as Exhibit 1.1 to this Registration Statement, will also provide for indemnification of us and our officers and directors.

Insofar as indemnification for liabilities arising under the Securities Act of 1933, as amended, may be permitted to directors, officers or persons controlling us pursuant to the foregoing provisions, we have been informed that in the opinion of the SEC such indemnification is against public policy as expressed in the Securities Act and is therefore unenforceable.

Item 7. Recent Sales of Unregistered Securities

During the past three years, we have issued the following securities (including options to acquire our ordinary shares) without registering the securities under the Securities Act. We believe that each of the following issuances was exempt from registration under the Securities Act in reliance on Regulation S under the Securities Act regarding sales by an issuer in offshore transactions or pursuant to Section 4(a)(2) of the Securities Act regarding transactions not involving a public offering. None of the transactions involved an underwriter.

<u>Purchaser</u>	<u>Date of Issuance</u>	<u>Title and Number of Securities</u>	<u>Consideration</u>
SCC Venture VI Holdco, Ltd.	February 2, 2018	1,679,206 series C-1 preferred shares	US\$15,000,011.36
Gopher Harvest Co-Investment Fund LP	February 2, 2018	559,735 series C-1 preferred shares	US\$5,000,000.80
AVICT Global Holdings Limited 航信环球控股有限公司	February 2, 2018	1,119,471 series C-1 preferred shares	US\$10,000,010.55
King Star Med LP	March 19, 2018	1,119,471 series C-1 preferred shares	US\$10,000,000.00
WEALTHY TECHNOLOGIES LIMITED	March 19, 2018	559,735 series C-1 preferred shares	US\$5,000,000.00

<u>Purchaser</u>	<u>Date of Issuance</u>	<u>Title and Number of Securities</u>	<u>Consideration</u>
Chief Strategic International Limited	May 16, 2018	559,736 series C-1 preferred shares	US\$5,000,010.00
Mega Prime Development Limited	June 13, 2019	685,676 series C-2 preferred shares	US\$6,999,998.00
Poly Platinum Enterprises Limited	June 13, 2019	489,769 series C-2 preferred shares	US\$5,000,003.00
Chief Strategic International Limited	June 13, 2019	391,815 series C-2 preferred shares	US\$4,000,000.00
MODEST CHAMPION LIMITED 冠謙有限公司	November 21, 2019	293,861 series C-2 preferred shares	US\$2,999,998.00
General Atlantic Singapore AI Pte. Ltd.	December 19, 2019	4,452,441 series C-3 preferred shares	US\$50,000,000.00
Certain directors, officers and employees	From August 17, 2017 to January 19, 2021	6,761,534 ordinary shares [#]	US\$7,912,715.40

Options and Warrants

SCC Venture VI Holdco, Ltd.	February 2, 2018	Warrant to purchase up to US\$5,625,000 worth of series C-2 preferred shares*	N/A
Gopher Harvest Co-Investment Fund LP	February 2, 2018	Warrant to purchase up to US\$1,875,000 worth of series C-2 preferred shares*	N/A
Certain directors, officers and employees	From August 17, 2017 to January 19, 2021	Option to purchase 7,154,992 ordinary shares	Past and future services provided by these individuals to us

Notes:

The number of ordinary shares includes the 2,262,500 ordinary shares, issued but deemed to be not outstanding as of the date of this prospectus, held by Great Han Fortune LP.

* These warrants expired on April 1, 2019.

Item 8. Exhibits and Financial Statement Schedules

(a) Exhibits:

See Exhibit Index for a complete list of all exhibits filed as part of this registration, which Exhibit Index is incorporated herein by reference.

(b) Financial Statement Schedules

Schedules have been omitted because the information required to be set forth therein is not applicable or is shown in the consolidated financial statements and the notes thereto.

Item 9. Undertakings

The undersigned hereby undertakes:

(a) The undersigned registrant hereby undertakes to provide to the underwriters at the closing specified in the underwriting agreements, certificates in such denominations and registered in such names as required by the underwriters to permit prompt delivery to each purchaser.

(b) Insofar as indemnification for liabilities arising under the Securities Act of 1933 may be permitted to directors, officers and controlling persons of the registrant pursuant to the foregoing provisions, or otherwise, the registrant has been advised that in the opinion of the U.S. Securities and Exchange Commission such indemnification is against public policy as expressed in the Securities Act of 1933 and is, therefore, unenforceable. In the event that a claim for indemnification against such liabilities (other than the payment by the registrant of expenses incurred or paid by a director, officer, or controlling person of the registrant in the successful defense of any action, suit or proceeding) is asserted by such director, officer or controlling person in connection with the securities being registered, the registrant will, unless in the opinion of its counsel the matter has been settled by controlling precedent, submit to a court of appropriate jurisdiction the question of whether such indemnification by it is against public policy as expressed in the Securities Act of 1933 and will be governed by the final adjudication of such issue.

(c) The undersigned registrant hereby undertakes that:

(1) For purposes of determining any liability under the Securities Act of 1933, the information omitted from the form of prospectus filed as part of this registration statement in reliance upon Rule 430A and contained in a form of prospectus filed by the registrant pursuant to Rule 424(b)(1) or (4) or 497(h) under the Securities Act of 1933 shall be deemed to be part of this registration statement as of the time it was declared effective.

(2) For the purpose of determining any liability under the Securities Act of 1933, each post-effective amendment that contains a form of prospectus shall be deemed to be a new registration statement relating to the securities offered therein, and the offering of such securities at that time shall be deemed to be the initial bona fide offering thereof.

ADAGENE INC.

EXHIBIT INDEX

<u>Exhibit Number</u>	<u>Description of Document</u>
1.1*	Form of Underwriting Agreement
3.1†	Sixth Amended and Restated Memorandum and Articles of Association of the Registrant, as currently in effect
3.2	Form of Seventh Amended and Restated Memorandum and Articles of Association of the Registrant, as effective immediately prior to the completion of this offering
4.1†	Form of Registrant's Specimen American Depositary Receipt (included in Exhibit 4.3)
4.2†	Registrant's Specimen Certificate for Ordinary Shares
4.3†	Form of Deposit Agreement between the Registrant, the depository and holders of the American Depositary Shares
4.4†	Fifth Amended and Restated Shareholders Agreement by and among Adagene Inc. and shareholders of Adagene Inc. named therein dated December 19, 2019
4.5†	Fourth Amended and Restated Right of First Refusal and Co-Sale Agreement by and between Adagene Inc., non-investor shareholders and investors named therein dated December 19, 2019
5.1†	Opinion of Walkers (Hong Kong) regarding the validity of the ordinary shares being registered
8.1†	Opinion of Walkers (Hong Kong) regarding certain Cayman Island tax matters (included in Exhibit 5.1)
8.2†	Opinion of Tian Yuan Law Firm regarding certain PRC tax matters (included in Exhibit 99.2)
8.3†	Opinion of Davis Polk & Wardwell LLP regarding material U.S. federal income tax consequences
10.1†	Adagene Inc. Second Amended and Restated Share Incentive Plan
10.2†	Form of Indemnification Agreement between the Registrant and each of its directors and executive officers
10.3†	Form of Employment Agreement between the Registrant and an executive officer of the Registrant
10.4†	Shares Purchase Agreement by and among Adagene Inc., its subsidiaries and General Atlantic Singapore AI Pte. Ltd. dated October 15, 2019
10.5†	Shares Purchase Agreement by and among Adagene Inc., its subsidiaries and certain investors of Adagene Inc. named therein dated June 9, 2019
10.6†	Shares Purchase Agreement by and among Adagene Inc., its subsidiaries and certain investors of Adagene Inc. named therein dated February 2, 2018
10.7#†	English translation of Cooperation Agreement on the PD-L1 Project by and among Guilin Sanjin Pharmaceutical Co., Ltd. and its affiliates and Adagene (Suzhou) Limited dated December 2018

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<u>Exhibit Number</u>	<u>Description of Document</u>
10.8#†	English translation of Cooperation Agreement on International Interests of PD-L1 Project between Guilin Sanjin Pharmaceutical Co., Ltd. and Adagene Inc. dated December 2018
10.9#†	English translation of Cooperation Agreement on the Undisclosed Project between Dragon Boat Biopharmaceutical (Shanghai) Limited and Adagene (Suzhou) Limited dated May 2019
10.10#†	English translation of Cooperation Agreement on International Interests of Undisclosed Project between Dragon Boat Biopharmaceutical (Shanghai) Limited and Adagene Inc. dated May 2019
10.11#†	Material Transfer and Collaboration Agreement between ADC Therapeutics SA and Adagene Inc. dated April 11, 2019
10.12#†	License Agreement Among ADC Therapeutics SA and Adagene Inc. dated April 11, 2019
10.13†	Adagene Inc. 2021 Performance Incentive Plan
10.14#	Collaboration and License Agreement between Exelixis, Inc. and Adagene Incorporated dated February 1, 2021
21.1†	List of subsidiaries of the Registrant
23.1	Consent of PricewaterhouseCoopers Zhong Tian LLP, Independent Registered Public Accounting Firm
23.2†	Consent of Walkers (Hong Kong),(included in Exhibit 5.1)
23.3†	Consent of Tian Yuan Law Firm (included in Exhibit 99.2)
23.4†	Consent of Davis Polk & Wardwell LLP (included in Exhibit 8.3)
24.1†	Powers of Attorney (included on signature page)
99.1†	Code of Business Conduct and Ethics of the Registrant
99.2†	Opinion of Tian Yuan Law Firm regarding certain PRC law matters
99.3†	Consent of Frost & Sullivan
99.4†	Consent of Andy (Yiu Leung) Cheung
99.5†	Consent of Min Li
99.6†	Consent of Man Kin (Raymond) Tam
99.7†	Representation under Item 8.A.4 of Form 20-F

* To be filed by amendment.

Portions of this exhibit have been omitted because they are both (i) not material and (ii) would likely cause competitive harm to the Company if publicly disclosed.

† Previously filed.

SIGNATURES

Pursuant to the requirements of the Securities Act of 1933, the registrant certifies that it has reasonable grounds to believe that it meets all of the requirements for filing on Form F-1 and has duly caused this registration statement to be signed on its behalf by the undersigned, thereunto duly authorized, in the City of San Francisco, State of California, on February 1, 2021.

Adagene Inc.

By: /s/ PETER (PEIZHI) LUO

Name: Peter (Peizhi) Luo
Title: Chief Executive Officer

Pursuant to the requirements of the Securities Act of 1933, as amended, this registration statement has been signed by the following persons on February 1, 2021 in the capacities indicated:

<u>Signature</u>	<u>Title</u>
<u>/s/ PETER (PEIZHI) LUO</u> Peter (Peizhi) Luo	Chief Executive Officer, Director (principal executive officer)
<u>*</u>	
<u>Yunxia Yang</u>	Director
<u>*</u>	
<u>Yu Miao</u>	Director
<u>*</u>	
<u>Lefei Sun</u>	Director
<u>/s/ MAN KIN (RAYMOND) TAM</u> Man Kin (Raymond) Tam	Chief Financial Officer (principal financial officer and principal accounting officer)

*By: /s/ PETER (PEIZHI) LUO

Name: Peter (Peizhi) Luo
Attorney-in-fact

SIGNATURE OF AUTHORIZED REPRESENTATIVE IN THE UNITED STATES

Pursuant to the Securities Act of 1933, the undersigned, the duly authorized representative in the United States of Adagene Inc., has signed this registration statement or amendment thereto in New York on February 1, 2021.

Authorized U.S. Representative

By: /s/ COLLEEN A. DE VRIES

Name: Colleen A. De Vries

Title: *Senior Vice President*

THE COMPANIES ACT (AS AMENDED)
OF THE CAYMAN ISLANDS
COMPANY LIMITED BY SHARES
SEVENTH AMENDED AND RESTATED
MEMORANDUM AND ARTICLES OF ASSOCIATION
OF
ADAGENE INC.
(ADOPTED BY SPECIAL RESOLUTION PASSED ON JANUARY 19, 2021 AND EFFECTIVE CONDITIONAL AND IMMEDIATELY
PRIOR TO THE COMPLETION OF THE COMPANY'S INITIAL PUBLIC OFFERING OF AMERICAN DEPOSITARY SHARES
REPRESENTING ITS ORDINARY SHARES)

THE COMPANIES ACT (AS AMENDED)
OF THE CAYMAN ISLANDS
COMPANY LIMITED BY SHARES

SEVENTH AMENDED AND RESTATED
MEMORANDUM OF ASSOCIATION

OF

ADAGENE INC.

(ADOPTED BY SPECIAL RESOLUTION PASSED ON JANUARY 19, 2021 AND EFFECTIVE CONDITIONAL AND IMMEDIATELY
PRIOR TO THE COMPLETION OF THE COMPANY'S INITIAL PUBLIC OFFERING OF AMERICAN DEPOSITARY SHARES
REPRESENTING ITS ORDINARY SHARES)

1. The name of the Company is Adagene Inc. (the "**Company**").
 2. The registered office of the Company will be situated at the offices of Vistra (Cayman) Limited, P. O. Box 31119 Grand Pavilion, Hibiscus Way, 802 West Bay Road, Grand Cayman, KY1 - 1205 Cayman Islands or at such other location as the Directors may from time to time determine.
 3. The objects for which the Company is established are unrestricted and the Company shall have full power and authority to carry out any object not prohibited by any law as provided by Section 7(4) of the Companies Act (as amended) of the Cayman Islands (the "**Companies Act**").
 4. The Company shall have and be capable of exercising all the functions of a natural person of full capacity irrespective of any question of corporate benefit as provided by Section 27(2) of the Companies Act.
 5. The Company will not trade in the Cayman Islands with any person, firm or corporation except in furtherance of the business of the Company carried on outside the Cayman Islands; provided that nothing in this section shall be construed as to prevent the Company effecting and concluding contracts in the Cayman Islands, and exercising in the Cayman Islands all of its powers necessary for the carrying on of its business outside the Cayman Islands.
 6. The liability of the shareholders of the Company is limited to the amount from time to time, if any, unpaid on the shares respectively held by them.
 7. The authorised share capital of the Company is US\$80,000 divided into 800,000,000 shares comprising (i) 640,000,000 Ordinary Shares of a nominal or par value of US\$0.0001 each, and (ii) 160,000,000 shares of a nominal or par value of US\$0.0001 each of such class or classes (however designated) as the board of directors may determine in accordance with Article 9 of the Articles. Subject to the Companies Act and the Articles of Association, the Company shall have power to redeem or purchase any of its shares, and to sub-divide or consolidate the said shares or any of them and to issue all or any part of its capital whether original, redeemed, increased or reduced with or without any preference, priority, special privilege or other rights or subject to any postponement of rights or to any conditions or restrictions whatsoever and so that unless the conditions of issue shall otherwise expressly provide every issue of shares whether stated to be ordinary, preference or otherwise shall be subject to the powers on the part of the Company hereinbefore provided.
 8. The Company may exercise the power contained in Section 206 of the Companies Act to deregister in the Cayman Islands and be registered by way of continuation in some other jurisdiction.
-

9. Capitalized terms that are not defined in this Memorandum of Association bear the same meaning as those given in the Articles of Association of the Company.

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THE COMPANIES ACT (AS AMENDED)
OF THE CAYMAN ISLANDS
COMPANY LIMITED BY SHARES

SEVENTH AMENDED AND RESTATED
ARTICLES OF ASSOCIATION

OF

ADAGENE INC.

(ADOPTED BY SPECIAL RESOLUTION PASSED ON JANUARY 19, 2021 AND EFFECTIVE CONDITIONAL AND IMMEDIATELY PRIOR TO THE COMPLETION OF THE COMPANY'S INITIAL PUBLIC OFFERING OF AMERICAN DEPOSITARY SHARES REPRESENTING ITS ORDINARY SHARES)

TABLE A

The Regulations contained or incorporated in Table 'A' in the First Schedule of the Companies Act shall not apply to the Company and the following Articles shall comprise the Articles of Association of the Company.

INTERPRETATION

1. In these Articles the following defined terms will have the meanings ascribed to them, if not inconsistent with the subject or context:

“**ADS**” means an American depositary share, each representing such number of Ordinary Shares as set out in the registration statements of the Company;

“**Affiliate**” means in respect of a Person, any other Person that, directly or indirectly, through (1) one or more intermediaries, Controls, is Controlled by, or is under common Control with, such Person, and (i) in the case of a natural person, shall include, without limitation, such person's spouse, parents, children, siblings, mother-in-law, father-in-law, brothers-in-law and sisters-in-law, a trust for the benefit of any of the foregoing, and a corporation, partnership or any other entity wholly or jointly owned by any of the foregoing, and (ii) in the case of an entity, shall include a partnership, a corporation or any other entity or any natural person which directly, or indirectly through one or more intermediaries, Controls, is Controlled by, or is under common Control with, such entity;

“**Appointment Period**” means the period commencing upon the effectiveness of the Company's Registration Statement and ending on the earliest of (i) the date upon which Peter Peizhi Luo beneficially owns (as defined under Rule 13d-3 of the Exchange Act) less than five per cent (5%) of the Shares outstanding on a fully diluted basis, (ii) the death or legal determination of Peter Peizhi Luo's incapacity and (iii) the termination of Peter Peizhi Luo as an executive officer or principal scientific advisor to the Company or any Group Company for cause (as determined under his related employment or consulting arrangements and subject to all related cure provisions);

“**Articles**” means these articles of association of the Company, as amended or substituted from time to time;

“**Board**” and “**Board of Directors**” and “**Directors**” means the directors of the Company for the time being, or as the case may be, the directors assembled as a board or as a committee thereof;

“**Chairman**” means the chairman of the Board of Directors;

“**Class**” or “**Classes**” means any class or classes of Shares as may from time to time be issued by the Company;

“**Commission**” means Securities and Exchange Commission of the United States of America or any other federal agency for the time being administering the Securities Act;

“**Companies Act**” means the Companies Act (as amended) of the Cayman Islands;

“**Company**” means Adagene Inc., a Cayman Islands exempted company;

“**Company’s Website**” means the website of the Company, the address or domain name of which has been notified to Shareholders;

“**Control**” means the ownership, directly or indirectly, of shares possessing more than fifty per cent (50%) of the voting power of the corporation, partnership or other entity (other than, in the case of a corporation, securities having such power only by reason of the happening of a contingency), or having the power to control the management or elect a majority of members to the board of directors or equivalent decision-making body of such corporation, partnership or other entity;

“**day**” means calendar day;

“**Designated Stock Exchange**” means the national stock exchange in the United States on which any Shares and ADSs are listed for trading;

“**Designated Stock Exchange Rules**” means the relevant code, rules and regulations, as amended, from time to time, applicable as a result of the original and continued listing of any Shares or ADSs on the Designated Stock Exchange;

“**electronic**” means the meaning given to it in the Electronic Transactions Law (as amended) of the Cayman Islands and any amendment thereto or re-enactments thereof for the time being in force and includes every other law incorporated therewith or substituted therefor;

“**electronic communication**” means electronic posting to the Company’s Website, transmission to any number, address or internet website or other electronic delivery methods as otherwise decided and approved by not less than a majority of the vote of the Board;

“**Exchange Act**” means the Securities Exchange Act of 1934, as amended;

“**Group Company**” means each of the Company and all of its direct or indirect Subsidiaries, and “**Group**” refers to all of the Group Companies collectively.

“**Independent Director**” means a Director who is an independent director as defined in the Designated Stock Exchange Rules;

“**Memorandum of Association**” means the memorandum of association of the Company, as amended or substituted from time to time;

“**month**” means calendar month;

“**Office**” means the registered office of the Company as required by the Companies Act;

“**Officer**” means the offices for the time being and from time to time of the Company;

“**Ordinary Resolution**” means a resolution:

- (a) passed by a simple majority of the votes cast by such Shareholders as, being entitled to do so, vote in person or, where proxies are allowed, by proxy at a general meeting of the Company and where a poll is taken regard shall be had in computing a majority to the number of votes to which each Shareholder is entitled by these Articles; or
- (b) approved in writing by all of the Shareholders entitled to vote at a general meeting of the Company in one or more instruments each signed by one or more of the Shareholders and the effective date of the resolution so adopted shall be the date on which the instrument, or the last of such instruments, if more than one, is executed;

“**Ordinary Shares**” means an ordinary share of par value of US\$0.0001 each in the capital of the Company having the rights and subject to the restrictions set out in these Articles, including a fraction of a share;

“**paid up**” means paid up as to the par value in respect of the issue of any Shares and includes credited as paid up;

“**Person**” means any natural person, firm, company, joint venture, partnership, corporation, association or other entity (whether or not having a separate legal personality) or any of them as the context so requires, other than in respect of a Director or Officer in which circumstances Person shall mean any person or entity permitted to act as such in accordance with the laws of the Cayman Islands;

“**Register**” means the register of Shareholders of the Company required to be kept pursuant to the Companies Act;

“**Registration Statement**” means the Company’s registration statement on Form F-1 (as it may be amended or supplemented) filed with the SEC under the Securities Act in connection with the initial public offering of American depository shares representing its Ordinary Shares;

“**Seal**” means the common seal of the Company (if adopted) including any facsimile thereof;

“**SEC**” means the United States Securities and Exchange Commission;

“**Secretary**” means any Person appointed by the Directors to perform any of the duties of the secretary of the Company;

“**Securities Act**” means the United States Securities Act of 1933, as amended, or any similar federal statute and the rules and regulations of the Commission thereunder, all as the same shall be in effect at the time;

“**Share**” means a share in the capital of the Company. All references to “Shares” herein shall be deemed to be Shares of any or all Classes as the context may require. For the avoidance of doubt in these Articles the expression “Share” shall include a fraction of a Share;

“**Shareholder**” means a Person who is registered as the holder of Shares in the Register and includes each subscriber to the Memorandum of Association pending entry in the Register of such subscriber;

“**Share Premium Account**” means the share premium account established in accordance with these Articles and the Companies Act;

“**signed**” means bearing a signature or representation of a signature affixed by mechanical means or an electronic symbol or process attached to or logically associated with an electronic communication and executed or adopted by a person with the intent to sign the electronic communication;

“**Special Resolution**” means a special resolution of the Company passed in accordance with the Companies Act being a resolution:

- (a) passed by not less than two-thirds of the votes cast by such Shareholders as, being entitled to do so, vote in person or, where proxies are allowed, by proxy at a general meeting of the Company of which notice specifying the intention to propose the resolution as a special resolution has been duly given and where a poll is taken regard shall be had in computing a majority to the number of votes to which each Shareholder is entitled; or
- (b) approved in writing by all of the Shareholders entitled to vote at a general meeting of the Company in one or more instruments each signed by one or more of the Shareholders and the effective date of the special resolution so adopted shall be the date on which the instrument or the last of such instruments, if more than one, is executed;

“**Subsidiary**” means, with respect to any given Person, any other Person that is Controlled directly or indirectly by such given Person;

“**Treasury Shares**” means Shares that were previously issued but were purchased, redeemed, surrendered or otherwise acquired by the Company and not cancelled;

“**United States**” means the United States of America, its territories, its possessions and all areas subject to its jurisdiction; and

“**year**” means calendar year.

2. In these Articles, save where the context requires otherwise:

- (a) words importing the singular number shall include the plural number and vice versa;
- (b) words importing the masculine gender only shall include the feminine gender and any Person as the context may require;
- (c) the word “may” shall be construed as permissive and the word “shall” shall be construed as imperative;
- (d) reference to a dollar or dollars (or US\$) and to a cent or cents is reference to dollars and cents of the United States of America;
- (e) reference to a statutory enactment shall include reference to any amendment or re-enactment thereof for the time being in force;
- (f) reference to any determination by the Directors shall be construed as a determination by the Directors in their sole and absolute discretion and shall be applicable either generally or in any particular case;
- (g) reference to “in writing” shall be construed as written or represented by any means reproducible in writing, including any form of print, lithograph, email, facsimile, photograph or telex or represented by any other substitute or format for storage or transmission for writing including in the form of an electronic record or partly one and partly another;
- (h) any requirements as to delivery under the Articles include delivery in the form of an electronic record or an electronic communication; and
- (i) any requirements as to execution or signature under the Articles, including the execution of the Articles themselves, can be satisfied in the form of an electronic signature as defined in the Electronic Transaction Law (as amended). Sections 8 and 19 of the Electronic Transactions Law (as amended) shall not apply.

3. Subject to the last two preceding Articles, any words defined in the Companies Act shall, if not inconsistent with the subject or context, bear the same meaning in these Articles.

PRELIMINARY

4. The business of the Company may be conducted as the Directors see fit.
5. The Office shall be at such address in the Cayman Islands as the Directors may from time to time determine. The Company may in addition establish and maintain such other offices and places of business and agencies in such places as the Directors may from time to time determine.
6. The expenses incurred in the formation of the Company and in connection with the offer for subscription and issue of Shares shall be paid by the Company. Such expenses may be amortised over such period as the Directors may determine and the amount so paid shall be charged against income and/or capital in the accounts of the Company as the Directors shall determine.
7. The Directors shall keep, or cause to be kept, the Register at such place or (subject to compliance with the Companies Act and these Articles) places as the Directors may from time to time determine. In the absence of any such determination, the Register shall be kept at the Office.

SHARES

8. Subject to these Articles, all Shares for the time being unissued shall be under the control of the Directors who may, in their absolute discretion and without the approval of the Shareholders, cause the Company to:
- (a) issue, allot or dispose of the same (whether in certificated form or non-certificated form, including fractions of a Share) to such Persons, in such manner, on such terms and having such rights and being subject to such restrictions as they may from time to time determine;
 - (b) grant rights over Shares or other securities to be issued in one or more classes or series as they deem necessary or appropriate and determine the designations, powers, preferences, privileges and other rights attaching to such Shares or securities, including dividend rights, voting rights, conversion rights, terms of redemption and liquidation preferences, any or all of which may be greater than the powers, preferences, privileges and rights associated with the then issued and outstanding Shares, at such times and on such other terms as they think proper; and
 - (c) grant options with respect to such Shares and issue warrants, convertible securities or similar instruments with respect thereto;
- and, for such purposes, the Directors may reserve an appropriate number of Shares for the time being unissued.
9. The Directors, or the Shareholders by Ordinary Resolution, may authorise the division of Shares into any number of Classes and sub-classes and the different Classes and sub-classes shall be authorised, established and designated (or re-designated as the case may be) and the variations in the relative rights (including, without limitation, voting, dividend and redemption rights), restrictions, preferences, privileges and payment obligations as between the different Classes (if any) may be fixed and determined by the Directors or by the Shareholders by Ordinary Resolution. The Directors may issue Shares with such preferred or other rights, all or any of which may be greater than the rights of Ordinary Shares, at such time and on such terms as they may think appropriate. Notwithstanding Article 12, the Directors may issue from time to time, out of the authorised share capital of the Company (other than the authorised but unissued Ordinary Shares), series of preferred shares in their absolute discretion and without approval of the Shareholders; provided, however, before any preferred shares of any such series are issued, the Directors shall by resolution of Directors determine, with respect to any series of preferred shares, the terms and rights of that series, including:

- (a) the designation of such series, the number of preferred shares to constitute such series and the subscription price thereof if different from the par value thereof;
- (b) whether the preferred shares of such series shall have voting rights, in addition to any voting rights provided by law, and, if so, the terms of such voting rights, which may be general or limited;
- (c) the dividends, if any, payable on such series, whether any such dividends shall be cumulative, and, if so, from what dates, the conditions and dates upon which such dividends shall be payable, and the preference or relation which such dividends shall bear to the dividends payable on any shares of any other class or any other series of shares;
- (d) whether the preferred shares of such series shall be subject to redemption by the Company, and, if so, the times, prices and other conditions of such redemption;
- (e) whether the preferred shares of such series shall have any rights to receive any part of the assets available for distribution amongst the Shareholders upon the liquidation of the Company, and, if so, the terms of such liquidation preference, and the relation which such liquidation preference shall bear to the entitlements of the holders of shares of any other class or any other series of shares;
- (f) whether the preferred shares of such series shall be subject to the operation of a retirement or sinking fund and, if so, the extent to and manner in which any such retirement or sinking fund shall be applied to the purchase or redemption of the preferred shares of such series for retirement or other corporate purposes and the terms and provisions relative to the operation thereof;
- (g) whether the preferred shares of such series shall be convertible into, or exchangeable for, shares of any other class or any other series of preferred shares or any other securities and, if so, the price or prices or the rate or rates of conversion or exchange and the method, if any, of adjusting the same, and any other terms and conditions of conversion or exchange;
- (h) the limitations and restrictions, if any, to be effective while any preferred shares of such series are outstanding upon the payment of dividends or the making of other distributions on, and upon the purchase, redemption or other acquisition by the Company of, the existing shares or shares of any other class of shares or any other series of preferred shares;
- (i) the conditions or restrictions, if any, upon the creation of indebtedness of the Company or upon the issue of any additional shares, including additional shares of such series or of any other class of shares or any other series of preferred shares; and
- (j) any other powers, preferences and relative, participating, optional and other special rights, and any qualifications, limitations and restrictions thereof;

and, for such purposes, the Directors may reserve an appropriate number of Shares for the time being unissued. The Company is not obliged to issue, allot or dispose of shares if it is, in the opinion of the Directors, unlawful or impracticable. The Company shall not issue Shares to bearer.

10. The Company may insofar as may be permitted by law, pay a commission to any Person in consideration of his subscribing or agreeing to subscribe whether absolutely or conditionally for any Shares. Such commissions may be satisfied by the payment of cash or the lodgement of fully or partly paid-up Shares or partly in one way and partly in the other. The Company may also pay such brokerage as may be lawful on any issue of Shares.

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11. The Directors may refuse to accept any application for Shares, and may accept any application in whole or in part, for any reason or for no reason.

MODIFICATION OF RIGHTS

12. Whenever the capital of the Company is divided into different Classes (and as otherwise determined by the Directors), the rights attached to any such Class may, subject to any rights or restrictions for the time being attached to any Class, only be materially adversely varied or abrogated with the consent in writing of the holders of not less than a majority of the issued Shares of the relevant Class, or with the sanction of a special resolution passed at a separate meeting of the holders of the Shares of such Class by a majority of two-thirds of the votes cast at such a meeting. To every such separate meeting all the provisions of these Articles relating to general meetings of the Company or to the proceedings thereat shall, *mutatis mutandis*, apply, except that the necessary quorum shall be one or more Persons at least holding or representing by proxy one-third in nominal or par value amount of the issued Shares of the relevant Class (but so that if at any adjourned meeting of such holders a quorum as above defined is not present, those Shareholders who are present shall form a quorum) and that, subject to any rights or restrictions for the time being attached to the Shares of that Class, every Shareholder of the Class shall on a poll have one vote for each Share of the Class held by him. For the purposes of this Article the Directors may treat all the Classes or any two or more Classes as forming one Class if they consider that all such Classes would be affected in the same way by the proposals under consideration, but in any other case shall treat them as separate Classes.
13. The rights conferred upon the holders of the Shares of any Class issued with preferred or other rights shall not, subject to any rights or restrictions for the time being attached to the Shares of that Class, be deemed to be materially adversely varied or abrogated by, *inter alia*, the creation, allotment or issue of further Shares ranking *pari passu* with or subsequent to them or the redemption or purchase of any Shares of any Class by the Company. The rights of the holders of Shares shall not be deemed to be materially adversely varied by the creation or issue of Shares with preferred or other rights including, without limitation, the creation of Shares with enhanced or weighted voting rights.

CERTIFICATES

14. Every Person whose name is entered as a Shareholder in the Register shall, without payment, be entitled to a certificate within one month after allotment or lodgement of transfer (or within such other period as the conditions of issue shall provide) in the form determined by the Directors. All certificates shall specify the Share or Shares held by that person and the amount paid up thereon, provided that in respect of a Share or Shares held jointly by several persons the Company shall not be bound to issue more than one certificate, and delivery of a certificate for a Share to one of several joint holders shall be sufficient delivery to all. All certificates for Shares shall be delivered personally or sent through the post addressed

to the Shareholder entitled thereto at such Shareholder's address notified by such Shareholder to the Company prior to such delivery, or if absent of such notification, at the registered address of such Shareholder as appearing in the Register.

15. Every share certificate of the Company shall bear legends required under the applicable laws, including the Securities Act.
16. Any two or more certificates representing Shares of any one Class held by any Shareholder may at the Shareholder's request be cancelled and a single new certificate for such Shares issued in lieu on payment (if the Directors shall so require) of US\$1.00 or such smaller sum as the Directors shall determine.

17. If a share certificate shall be damaged or defaced or alleged to have been lost, stolen or destroyed, a new certificate representing the same Shares may be issued to the relevant Shareholder upon request subject to delivery up of the old certificate or (if alleged to have been lost, stolen or destroyed) compliance with such conditions as to evidence and indemnity and the payment of out-of-pocket expenses of the Company in connection with the request as the Directors may think fit.
18. In the event that Shares are held jointly by several persons, any request may be made by any one of the joint holders and if so made shall be binding on all of the joint holders.

FRACTIONAL SHARES

19. The Directors may issue fractions of a Share and, if so issued, a fraction of a Share shall be subject to and carry the corresponding fraction of liabilities (whether with respect to nominal or par value, premium, contributions, calls or otherwise), limitations, preferences, privileges, qualifications, restrictions, rights (including, without prejudice to the generality of the foregoing, voting and participation rights) and other attributes of a whole Share. If more than one fraction of a Share of the same Class is issued to or acquired by the same Shareholder such fractions shall be accumulated.

LIEN

20. The Company has a first and paramount lien on every Share (whether or not fully paid) for all amounts (whether presently payable or not) payable at a fixed time or called in respect of that Share. The Company also has a first and paramount lien on every Share registered in the name of a Person indebted or under liability to the Company (whether he is the sole registered holder of a Share or one of two or more joint holders) for all amounts owing by him or his estate to the Company (whether or not presently payable). The Directors may at any time declare a Share to be wholly or in part exempt from the provisions of this Article. The Company's lien on a Share extends to any amount payable in respect of it.
21. The Company may sell, in such manner as the Directors may determine, any Share on which the Company has a lien, but no sale shall be made unless an amount in respect of which the lien exists is presently payable nor until the expiration of fourteen days after a notice in writing, demanding payment of such part of the amount in respect of which the lien exists as is presently payable, has been given to the registered holder for the time being of the Share, or the Persons entitled thereto by reason of his death or bankruptcy.
22. For giving effect to any such sale the Directors may authorise some Person to transfer the Shares sold to the purchaser thereof. The purchaser shall be registered as the holder of the Shares comprised in any such transfer and he shall not be bound to see to the application of the purchase money, nor shall his title to the Shares be affected by any irregularity or invalidity in the proceedings in reference to the sale.
23. The proceeds of the sale after deduction of expenses, fees and commission incurred by the Company shall be received by the Company and applied in payment of such part of the amount in respect of which the lien exists as is presently payable, and the residue shall (subject to a like lien for sums not presently payable as existed upon the Shares prior to the sale) be paid to the Person entitled to the Shares immediately prior to the sale.

CALLS ON SHARES

24. The Directors may from time to time make calls upon the Shareholders in respect of any moneys unpaid on their Shares, and each Shareholder shall (subject to receiving at least fourteen days' notice specifying the time or times of payment) pay to the Company at the time or times so specified the amount called on such Shares.
25. The joint holders of a Share shall be jointly and severally liable to pay calls in respect thereof.

26. If a sum called in respect of a Share is not paid before or on the day appointed for payment thereof, the Person from whom the sum is due shall pay interest upon the sum at the rate of eight percent per annum from the day appointed for the payment thereof to the time of the actual payment, but the Directors shall be at liberty to waive payment of that interest wholly or in part.
27. The provisions of these Articles as to the liability of joint holders and as to payment of interest shall apply in the case of non-payment of any sum which, by the terms of issue of a Share, becomes payable at a fixed time, whether on account of the amount of the Share, or by way of premium, as if the same had become payable by virtue of a call duly made and notified.
28. The Directors may make arrangements on the issue of partly paid Shares for a difference between the Shareholders, or the particular Shares, in the amount of calls to be paid and in the times of payment.
29. The Directors may, if they think fit, receive from any Shareholder willing to advance the same all or any part of the moneys uncalled and unpaid upon any partly paid Shares held by him, and upon all or any of the moneys so advanced may (until the same would, but for such advance, become presently payable) pay interest at such rate (not exceeding without the sanction by Ordinary Resolution, eight percent per annum) as may be agreed upon between the Shareholder paying the sum in advance and the Directors. No such sum paid in advance of calls shall entitle the Shareholder paying such sum to any portion of a dividend declared in respect of any period prior to the date upon which such sum would, but for such payment, become presently payable.

FORFEITURE OF SHARES

30. If a Shareholder fails to pay any call or instalment of a call in respect of partly paid Shares on the day appointed for payment, the Directors may, at any time thereafter during such time as any part of such call or instalment remains unpaid, serve a notice on him requiring payment of so much of the call or instalment as is unpaid, together with any interest which may have accrued.
31. The notice shall name a further day (not earlier than the expiration of fourteen days from the date of the notice) on or before which the payment required by the notice is to be made, and shall state that in the event of non-payment at or before the time appointed the Shares in respect of which the call was made will be liable to be forfeited.
32. If the requirements of any such notice as aforesaid are not complied with, any Share in respect of which the notice has been given may at any time thereafter, before the payment required by notice has been made, be forfeited by a resolution of the Directors to that effect.
33. A forfeited Share may be sold or otherwise disposed of on such terms and in such manner as the Directors think fit, and at any time before a sale or disposition the forfeiture may be cancelled on such terms as the Directors think fit.
34. A Person whose Shares have been forfeited shall cease to be a Shareholder in respect of the forfeited Shares, but shall, notwithstanding, remain liable to pay to the Company all moneys which at the date of forfeiture were payable by him to the Company in respect of the Shares forfeited, but his liability shall cease if and when the Company receives payment in full of the amount unpaid on the Shares forfeited. A certificate in writing under the hand of a Director that a Share has been duly forfeited on a date stated in the certificate, shall be conclusive evidence of the facts in the certificate as against all Persons claiming to be entitled to the Share.
35. The Company may receive the consideration, if any, given for a Share on any sale or disposition thereof pursuant to the provisions of these Articles as to forfeiture and may execute a transfer of the Share in favour of the Person to whom the Share is sold or disposed of and that Person shall be registered as the holder of the Share, and shall not be bound to see to the application of the purchase money, if any, nor shall his title to the Shares be affected by any irregularity or invalidity in the proceedings in reference to the disposition or sale.

36. The provisions of these Articles as to forfeiture shall apply in the case of non-payment of any sum which by the terms of issue of a Share becomes due and payable, whether on account of the amount of the Share, or by way of premium, as if the same had been payable by virtue of a call duly made and notified.

TRANSFER OF SHARES

37. The instrument of transfer of any Share shall be in any usual or common form or such other form as the Directors may determine and be executed by or on behalf of the transferor and if in respect of a nil or partly paid up Share, or if so required by the Directors, shall also be executed on behalf of the transferee and shall be accompanied by the certificate (if any) of the Shares to which it relates and such other evidence as the Directors may reasonably require to show the right of the transferor to make the transfer. The transferor shall be deemed to remain a Shareholder until the name of the transferee is entered in the Register in respect of the relevant Shares.
38. (a) Subject to the terms of issue thereof, the Directors may determine to decline to register any transfer of Shares without assigning any reason therefor.
- (b) The Directors may also decline to register any transfer of any Share unless:
- (i) the instrument of transfer is lodged with the Company, accompanied by the certificate for the Shares to which it relates and such other evidence as the Board may reasonably require to show the right of the transferor to make the transfer;
 - (ii) the instrument of transfer is in respect of only one Class of Shares;
 - (iii) the instrument of transfer is properly stamped, if required;
 - (iv) in the case of a transfer to joint holders, the number of joint holders to whom the Share is to be transferred does not exceed four;
 - (v) the Shares are free from any lien in favour of the Company; or
 - (vi) a fee of such maximum sum as the Designated Stock Exchange may determine to be payable or such lesser sum as the Board of Directors may from time to time require, is paid to the Company in respect thereof.
39. The registration of transfers may, on ten days' notice being given by advertisement in such one or more newspapers or by electronic means or by any other means in accordance with the Designated Stock Exchange Rules, be suspended and the Register closed at such times and for such periods as the Directors may, in their absolute discretion, from time to time determine, provided always that such registration of transfer shall not be suspended nor the Register closed for more than 30 days in any year.
40. All instruments of transfer that are registered shall be retained by the Company. If the Directors refuse to register a transfer of any Shares, they shall within three months after the date on which the transfer was lodged with the Company send to each of the transferor and the transferee notice of the refusal.

TRANSMISSION OF SHARES

41. The legal personal representative of a deceased sole holder of a Share shall be the only Person recognised by the Company as having any title to the Share. In the case of a Share registered in the name of two or more holders, the survivors or survivor, or the legal personal representatives of the deceased holder of the Share, shall be the only Person recognised by the Company as having any title to the Share.
42. Any Person becoming entitled to a Share in consequence of the death or bankruptcy of a Shareholder shall upon such evidence being produced as may from time to time be required by the Directors, have the right either to be registered as a Shareholder in respect of the Share or, instead of being registered himself, to make such transfer of the Share as the deceased or bankrupt Person could have made; but the Directors shall, in either case, have the same right to decline or suspend registration as they would have had in the case of a transfer of the Share by the deceased or bankrupt Person before the death or bankruptcy.

43. A Person becoming entitled to a Share by reason of the death or bankruptcy of a Shareholder shall be entitled to the same dividends and other advantages to which he would be entitled if he were the registered Shareholder, except that he shall not, before being registered as a Shareholder in respect of the Share, be entitled in respect of it to exercise any right conferred by membership in relation to meetings of the Company, provided however, that the Directors may at any time give notice requiring any such person to elect either to be registered himself or to transfer the Share, and if the notice is not complied with within ninety days, the Directors may thereafter withhold payment of all dividends, bonuses or other monies payable in respect of the Share until the requirements of the notice have been complied with.

REGISTRATION OF EMPOWERING INSTRUMENTS

44. The Company shall be entitled to charge a fee not exceeding one dollar (US\$1.00) on the registration of every probate, letters of administration, certificate of death or marriage, power of attorney, notice in lieu of distringas, or other instrument.

ALTERATION OF SHARE CAPITAL

45. The Company may from time to time by Ordinary Resolution increase the share capital by such sum, to be divided into Shares of such Classes and amount, as the resolution shall prescribe and with such rights, priorities and privileges annexed thereto, as the Company in general meeting may determine.
46. The Company may by Ordinary Resolution:
- (a) consolidate and divide all or any of its share capital into Shares of a larger amount than its existing Shares;
 - (b) subdivide its existing Shares, or any of them into Shares of a smaller amount provided that in the subdivision the proportion between the amount paid and the amount, if any, unpaid on each reduced Share shall be the same as it was in case of the Share from which the reduced Share is derived;
 - (c) convert all or any of its paid up Shares into stock and reconvert that stock into paid up Shares of any denomination; and
 - (d) cancel any Shares that, at the date of the passing of the resolution, have not been taken or agreed to be taken by any Person and diminish the amount of its share capital by the amount of the Shares so cancelled.
47. The Company may by Special Resolution reduce its share capital and any capital redemption reserve in any manner authorised by law.

REDEMPTION, PURCHASE AND SURRENDER OF SHARES

48. Subject to the Companies Act and these Articles, the Company may:
- (a) issue Shares on terms that they are to be redeemed or are liable to be redeemed at the option of the Company or the Shareholder on such terms and in such manner as may be determined, before the issue of such Shares, by either the Board or by the Shareholders by Ordinary Resolution;

- (b) redeem or purchase its own Shares (including any redeemable Shares) on such terms and in such manner as have been approved by the Board or the Shareholders by Ordinary Resolution, or are otherwise authorised by these Articles;
 - (c) make a payment in respect of the redemption or purchase of its own Shares in any manner permitted by the Companies Act, including out of its capital; and
 - (d) accept the surrender for no consideration of any paid up Share (including any redeemable Share) on such terms and in such manner as the Directors may determine.
49. The holder of the Shares being purchased shall be bound to deliver up to the Company the certificate(s) (if any) thereof. Any Share in respect of which notice of redemption has been given shall not be entitled to participate in the profits of the Company in respect of the period after the date specified as the date of redemption in the notice of redemption.
50. The redemption, purchase or surrender of any Share shall not be deemed to give rise to the redemption, purchase or surrender of any other Share.
51. The Directors may when making payments in respect of redemption or purchase of Shares, if authorised by the terms of issue of the Shares being redeemed or purchased or with the agreement of the holder of such Shares, make such payment either in cash or in specie including without limitation, interests in a special purpose vehicle holding assets of the Company or holding entitlement to the proceeds of assets held by the Company or in a liquidation structure.

TREASURY SHARES

52. Shares that the Company purchases, redeems or acquires (by way of surrender or otherwise) may, at the option of the Company, be cancelled immediately or held as Treasury Shares in accordance with the Companies Act. In the event that the Directors do not specify that the relevant Shares are to be held as Treasury Shares, such Shares shall be cancelled.
53. No dividend may be declared or paid, and no other distribution (whether in cash or otherwise) of the Company's assets (including any distribution of assets to Shareholders on a winding up) may be declared or paid in respect of a Treasury Share.
54. The Company shall be entered in the Register as the holder of the Treasury Shares provided that:
- (a) the Company shall not be treated as a Shareholder for any purpose and shall not exercise any right in respect of the Treasury Shares, and any purported exercise of such a right shall be void;
 - (b) a Treasury Share shall not be voted, directly or indirectly, at any meeting of the Company and shall not be counted in determining the total number of issued shares at any given time, whether for the purposes of these Articles or the Companies Act, save that an allotment of Shares as fully paid bonus shares in respect of a Treasury Share is permitted and Shares allotted as fully paid bonus shares in respect of a treasury share shall be treated as Treasury Shares.
55. Treasury Shares may be disposed of by the Company on such terms and conditions as determined by the Directors.

GENERAL MEETINGS

56. All general meetings other than annual general meetings shall be called extraordinary general meetings.
57. (a) The Company may, but shall not (unless required by the Companies Act) be obliged to hold a general meeting in each year as its annual general meeting and shall specify the meeting as such in the notices calling it. The annual general meeting shall be held at such time and place as may be determined by the Directors.

- (b) At these meetings the report of the Directors (if any) shall be presented.
58. (a) The Chairman or a majority of the Directors may call general meetings, and they shall on a Shareholders' requisition forthwith proceed to convene an extraordinary general meeting of the Company.
- (b) A Shareholders' requisition is a requisition of Shareholders of the Company holding at the date of deposit of the requisition Shares which carry in aggregate not less than twenty percent (20%) of all votes attaching to all issued and outstanding Shares of the Company that as at the date of the deposit carry the right of voting at general meetings of the Company.
- (c) The requisition must state the objects of the meeting and must be signed by the requisitionists and deposited at the registered office of the Company, and may consist of several documents in like form each signed by one or more requisitionists.
- (d) If the Directors do not within twenty-one (21) days from the date of the deposit of the requisition duly proceeded to convene a general meeting to be held within a further twenty-one (21) days, the requisitionists, or any of them representing more than one-half of the total voting rights of all of them, may themselves convene a general meeting, but any meeting so convened shall not be held after the expiration of three months after the expiration of the said twenty-one (21) days.
- (e) A general meeting convened as aforesaid by requisitionists shall be convened in the same manner as nearly as possible as that in which general meetings are to be convened by Directors.

NOTICE OF GENERAL MEETINGS

59. At least seven (7) days' notice shall be given for any general meeting. Every notice shall be exclusive of the day on which it is given or deemed to be given and of the day for which it is given and shall specify the place, the day and the hour of the meeting and the general nature of the business and shall be given in the manner hereinafter mentioned or in such other manner if any as may be prescribed by the Company, provided that a general meeting of the Company shall, whether or not the notice specified in this Article has been given and whether or not the provisions of these Articles regarding general meetings have been complied with, be deemed to have been duly convened if it is so agreed:
- (a) in the case of an annual general meeting by all the Shareholders (or their proxies) entitled to attend and vote thereat; and
- (b) in the case of an extraordinary general meeting by a majority in number of the Shareholders (or their proxies) having a right to attend and vote at the meeting, being a majority together holding not less than seventy five per cent (75%) in par value of the Shares giving that right.
60. The accidental omission to give notice of a meeting to or the non-receipt of a notice of a meeting by any Shareholder shall not invalidate the proceedings at any meeting.

PROCEEDINGS AT GENERAL MEETINGS

61. No business shall be transacted at any general meeting without the consent of all Shareholders entitled to receive notice of that meeting unless notice of such business has been given in the notice convening that meeting.
62. No business shall be transacted at any general meeting unless a quorum of Shareholders is present at the time when the meeting proceeds to business. The holders of Shares being not less than an aggregate of one-half of all Shares in issue present in person or by proxy and entitled to vote shall be a quorum for all purposes.

63. If within half an hour from the time appointed for the meeting a quorum is not present, the meeting, if convened upon the requisition of Shareholders, shall be dissolved. In any other case it shall stand adjourned to the same day in the next week, at the same time and place, and if at the adjourned meeting a quorum is not present within half an hour from the time appointed for the meeting, the Shareholder or Shareholders present and entitled to vote shall form a quorum.
64. If the Directors wish to make this facility available for a specific general meeting or all general meetings of the Company, participation in any general meeting of the Company may be by means of a telephone or similar communication equipment by way of which all Persons participating in such meeting can communicate with each other and such participation shall be deemed to constitute presence in person at the meeting.
65. The chairman, if any, of the Directors shall preside as chairman at every general meeting of the Company.
66. If there is no such chairman, or if at any general meeting he is not present within fifteen minutes after the time appointed for holding the meeting or is unwilling to act as chairman, any Director or Person nominated by the Directors shall preside as chairman, failing which the Shareholders present in person or by proxy shall choose any Person present to be chairman of that meeting.
- The chairman of the general meeting may with the consent of any general meeting at which a quorum is present (and shall if so directed by the meeting) adjourn a meeting from time to time and from place to place, but no business shall be transacted at any adjourned meeting other than the business left unfinished at the meeting from which the adjournment took place. When a meeting, or adjourned meeting, is adjourned for fourteen days or more, notice of the adjourned meeting shall be given in the manner provided for the original meeting. Save as aforesaid, it shall not be necessary to give any notice of an adjournment or of the business to be transacted at an adjourned meeting.
67. The Directors may cancel or postpone any duly convened general meeting at any time prior to such meeting, except for general meetings requisitioned by the Shareholders in accordance with these Articles, for any reason or for no reason, upon notice in writing to Shareholders. A postponement may be for a stated period of any length or indefinitely as the Directors may determine.
68. At any general meeting a resolution put to the vote of the meeting shall be decided by way of a poll save that the chairman of the meeting may, in good faith, allow a resolution which relates purely to a procedural or administrative matter to be voted on by a show of hands, and where a show of hands is allowed pursuant to these Articles, before or on the declaration of the result of the show of hands, a poll may be demanded by the chairman of the meeting or any Shareholder present in person or by proxy. Where a resolution is voted on by a show of hands, a declaration by the chairman of the meeting that a resolution has, on a show of hands, been carried, or carried unanimously, or by a particular majority, or lost, and an entry to that effect in the book of the proceedings of the Company, shall be conclusive evidence of the fact, without proof of the number or proportion of the votes recorded in favour of, or against, that resolution. For the purposes of these Articles, procedural and administrative matters are (i) the proposal of new resolutions which are not already set out in the notice convening the general meeting or any proposed amendments to the resolutions; and (ii) those that relate to the chairman's duties to maintain the orderly conduct of the meeting and/or allow the business of the meeting to be properly and effectively dealt with, whilst allowing all Shareholders a reasonable opportunity to express their views.
69. If a poll is required or is duly demanded it shall be taken in such manner as the chairman of the meeting directs, and the result of the poll shall be deemed to be the resolution of the meeting at which the poll was required or demanded.

70. In the case of an equality of votes, whether on a show of hands or on a poll, the chairman of the meeting at which the show of hands takes place or at which the poll is demanded, shall be entitled to a second or casting vote.
71. All questions submitted to a meeting shall be decided by an Ordinary Resolution except where a greater majority is required by these Articles or by the Law. A poll demanded on the election of a chairman of the meeting or on a question of adjournment shall be taken forthwith. A poll demanded on any other question shall be taken at such time as the chairman of the meeting directs.

VOTES OF SHAREHOLDERS

72. Subject to any rights and restrictions for the time being attached to any Share, on a show of hands every Shareholder present in person and every Person representing a Shareholder by proxy (or, if a corporation or other non-natural person, by its duly authorized representative or proxy) shall, at a general meeting or extraordinary general meeting of the Company, each have one (1) vote and on a poll every Shareholder and every Person representing a Shareholder by proxy (or, if a corporation or other non-natural person, by its duly authorized representative or proxy) shall have one (1) vote for each Share of which he or the Person represented by proxy is the holder.
73. In the case of joint holders the vote of the senior who tenders a vote, whether in person or by proxy (or, if a corporation or other non-natural person, by its duly authorized representative or proxy), shall be accepted to the exclusion of the votes of the other joint holders and for this purpose seniority shall be determined by the order in which the names stand in the Register.
74. Shares carrying the right to vote which are held by a Shareholder of unsound mind, or in respect of whom an order has been made by any court having jurisdiction in lunacy, may be voted in respect of Shares carrying the right to vote held by him, whether on a show of hands or on a poll, by his committee, or other Person in the nature of a committee appointed by that court, and any such committee or other Person, may vote in respect of such Shares by proxy.
75. No Shareholder shall be entitled to vote at any general meeting of the Company unless he is registered as a Shareholder on the record date for such meeting or unless all calls, if any, or other sums presently payable by him in respect of Shares carrying the right to vote held by him have been paid.
76. On a poll votes may be given either personally or by proxy.
77. Each Shareholder, other than a recognised clearing house (or its nominee(s)) or depositary (or its nominee(s)), may only appoint one proxy on a show of hand and on a poll, each such proxy is under no obligation to cast all his votes in the same way. The instrument appointing a proxy shall be in writing under the hand of the appointor or of his attorney duly authorised in writing or, if the appointor is a corporation, either under Seal or under the hand of an officer or attorney duly authorised. A proxy need not be a Shareholder. On a poll a Shareholder entitled to more than one vote need not use all his votes or cast all his votes in the same way.
78. An instrument appointing a proxy may be in any usual or common form or such other form as the Directors may approve and may be expressed to be for a particular meeting or any adjournment thereof or generally until revoked.
79. The instrument appointing a proxy shall be deposited at the Office or at such other place as is specified for that purpose in the notice convening the meeting, or in any instrument of proxy sent out by the Company:
- (a) not less than forty-eight (48) hours before the time for holding the meeting or adjourned meeting at which the person named in the instrument proposes to vote; or

- (b) in the case of a poll taken more than forty-eight (48) hours after it is demanded, be deposited as aforesaid after the poll has been demanded and not less than twenty-four (24) hours before the time appointed for the taking of the poll; or
- (c) where the poll is not taken forthwith but is taken not more than forty-eight (48) hours after it was demanded be delivered at the meeting at which the poll was demanded to the chairman or to the secretary or to any director;

provided that the Directors may in the notice convening the meeting, or in an instrument of proxy sent out by the Company, direct that the instrument appointing a proxy may be deposited (no later than the time for holding the meeting or adjourned meeting) at the Registered Office or at such other place as is specified for that purpose in the notice convening the meeting, or in any instrument of proxy sent out by the Company. The Chairman may in any event at his discretion direct that an instrument of proxy shall be deemed to have been duly deposited. An instrument of proxy that is not deposited in the manner permitted shall be invalid.

- 80. The instrument appointing a proxy shall be deemed to confer authority to demand or join in demanding a poll.
- 81. A resolution in writing signed by all the Shareholders for the time being entitled to receive notice of and to attend and vote at general meetings of the Company (or being corporations by their duly authorised representatives) shall be as valid and effective as if the same had been passed at a general meeting of the Company duly convened and held.
- 82. Votes given in accordance with the terms of an instrument of proxy, which has been deposited in accordance with Article 79, shall be valid notwithstanding the previous death or insanity of the principal or revocation of the proxy or of the authority under which the proxy was executed, or the transfer of the Share in respect of which the proxy is given unless notice in writing of such death, insanity, revocation or transfer was received by the Company at the Registered Office before the commencement of the general meeting, or adjourned meeting at which it is sought to use the proxy.

CORPORATIONS ACTING BY REPRESENTATIVES AT MEETINGS

- 83. Any corporation which is a Shareholder or a Director may by resolution of its directors or other governing body authorise such Person as it thinks fit to act as its representative at any meeting of the Company or of any meeting of holders of a Class or of the Directors or of a committee of Directors, and the Person so authorised shall be entitled to exercise the same powers on behalf of the corporation which he represents as that corporation could exercise if it were an individual Shareholder or Director.

CLEARING HOUSES

- 84. If a clearing house (or its nominee) is a Shareholder of the Company it may, by resolution of its directors or other governing body or by power of attorney, authorise such person or persons as it thinks fit to act as its representative or representatives at any general meeting of the Company or at any general meeting of any class of Shareholders of the Company provided that, if more than one person is so authorised, the authorisation shall specify the number and class of Shares in respect of which each such person is so authorised. A person so authorised pursuant to this Article shall be entitled to exercise the same powers on behalf of the clearing house (or its nominee) which he represents as that clearing house (or its nominee) could exercise if it were an individual Shareholder holding the number and Class of Shares specified in such authorisation.

DIRECTORS

- 85. (a) Unless otherwise determined by the Company in general meeting, the number of Directors shall not be more than nine (9). The Board of Directors shall have the right to change the size of the Board of Directors and determine the exact number of Directors.

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- (b) Subject to Article 85(f), the Board of Directors shall have a Chairman elected and appointed by a majority of the Directors then in office. The period for which the Chairman will hold office will also be determined by a majority of all of the Directors then in office. The Chairman shall preside as chairman at every meeting of the Board of Directors. To the extent the Chairman is not present at a meeting of the Board of Directors within fifteen minutes after the time appointed for holding the same, or if the Chairman is unable to or unwilling to act as the chairman of a meeting of the Directors, the attending Directors may choose one of their number to be the chairman of the meeting.
 - (c) For so long as Wuxi Pharmatech Healthcare Fund I L.P. holds no less than 5% of all the Shares in issue on a fully-diluted basis, it shall be entitled, by notice in writing to the Company, to nominate one (1) independent director (as contemplated by the terms of applicable Designated Stock Exchange Rules and/or SEC rules with respect to audit committee independence) and any such Person shall be appointed and agreed by the Board.
 - (d) For so long as JSR Limited holds no less than 5% of all the Shares in issue on a fully-diluted basis, it shall be entitled, by notice in writing to the Company, to appoint one (1) Person as a director of the Company and to remove, replace and/or reappoint any such Person so appointed.
 - (e) For so long as General Atlantic Singapore AI Pte. Ltd. and Affiliates thereof collectively hold no less than 5% of all the Shares in issue on a fully-diluted basis, General Atlantic Singapore AI Pte. Ltd. and Affiliates thereof shall be entitled, by notice in writing to the Company, to appoint one (1) Person as a director of the Company and to remove, replace and reappoint any such Person so appointed.
 - (f) For as long as Peter Peizhi Luo holds or beneficially owns (as defined under Rule 13d-3 of the Exchange Act) any Shares of the Company or is employed by the Company or any of its Subsidiaries, Peter Peizhi Luo shall occupy one (1) seat of the Board (the "**Founder Director**") and shall serve as the Chairman of the Board of Directors.

- (g) In addition to the Founder Director, during the Appointment Period, Peter Peizhi Luo shall be entitled, by notice in writing to the Company, to appoint one (1) Person as director of the Company and to remove, replace and/or reappoint such Person so appointed (the “**Additional Founder Director**”, together with the Founder Director, the “**Founder Directors**”).
- (h) In addition to the Founder Directors, one (1) seat on the Board shall be reserved for and occupied by an executive officer of the Company who is responsible for clinical development or research and development.
- (i) Subject to Article 85(d), (e), and (g), the Board of Directors may by the affirmative vote of a simple majority of the remaining Directors present and voting at a Board meeting, appoint any person to be a Director either to fill a vacancy on the Board arising from the office of any Director being vacated in any of the circumstances described in Article 108 or as an addition to the existing Board. In the event of a vacancy arising from the office of an Independent Director being vacated, the Board may only appoint another Independent Director with the affirmative vote of the Founder Director to fill such vacancy.
- (j) An appointment of a Director may be on terms that the Director shall automatically retire from office (unless he or she has sooner vacated office) upon any specified event or after any specified period in a written agreement between the Company and the Director, if any; but no such term shall be implied in the absence of express provision.

- (k) Each Director whose term of office expires shall be eligible for re-election at a meeting of the Shareholders or re-appointment by the Board; provided that no re-appointment of any Independent Director shall be effective without the affirmative vote of the Founder Director.
86. Subject to Article 85, any vacancy, including newly created directorships resulting from any increase in the authorised number of Directors or amendment of these Articles, and vacancies created by removal or resignation of a Director, may be filled by the consent of a majority of the Directors then in office, though less than a quorum, or by a sole remaining director, and the Directors so chosen shall hold office until the next annual election and until their successors are duly elected and shall qualify, unless sooner displaced; provided, however, that where such vacancy occurs among the Directors elected by the holders of a class or series of shares under Article 85, such vacancy shall be filled and appointed in accordance with Article 85.
87. A Director (other than such Director appointed pursuant to Article 85) may be removed from office by Ordinary Resolution of the Company, notwithstanding anything in these Articles or in any agreement between the Company and such Director (but without prejudice to any claim for damages under such agreement) but with respect to a Director appointed pursuant to Article 85 and other shareholder agreement(s), only upon the vote or written consent of the Shareholders entitled to appoint such Director. Any vacancies created by the resignation, removal or death of a Director appointed pursuant to Article 85 shall be filled in such manner as stipulated in Article 85. The notice of any meeting at which a resolution to remove a Director shall be proposed or voted upon must contain a statement of the intention to remove that Director and such notice must be served on that Director not less than ten (10) days before the meeting. Such Director is entitled to attend the meeting and be heard on the motion for his removal.
88. The Board may, from time to time, and except as required by applicable law or Designated Stock Exchange Rules, adopt, institute, amend, modify or revoke the corporate governance policies or initiatives, which shall be intended to set forth the policies of the Company and the Board on various corporate governance related matters as the Board shall determine by resolution from time to time.
89. Subject to applicable law, Designated Stock Exchange Rules and the Articles, the Board may establish any committee of the Board as it deems appropriate from time to time, and committees of the Board shall have the rights, powers and privileges delegated to such committees by the Board from time to time.
90. A Director shall not be required to hold any Shares in the Company by way of qualification. A Director who is not a Shareholder of the Company shall nevertheless be entitled to attend and speak at general meetings.
91. The remuneration of the Directors may be determined by the Directors or by Ordinary Resolution.
92. The Directors shall be entitled to be paid their travelling, hotel and other expenses properly incurred by them in going to, attending and returning from meetings of the Directors, or any committee of the Directors, or general meetings of the Company, or otherwise in connection with the business of the Company, or to receive such fixed allowance in respect thereof as may be determined by the Directors from time to time, or a combination partly of one such method and partly the other.

ALTERNATE DIRECTOR

93. Any Director may in writing appoint another Person to be his alternate and, save to the extent provided otherwise in the form of appointment, such alternate shall have authority to sign written resolutions on behalf of the appointing Director, but shall not be authorised to sign such written resolutions where they have been signed by the appointing Director, and to act in such Director's place at any meeting of the Directors. Every such alternate shall be entitled to attend and vote at meetings of the Directors as the alternate of the Director appointing him and where he is a Director to have a separate vote in addition to his own vote. A Director may at any time in writing revoke the appointment of an alternate appointed by him. Such alternate shall not be an Officer solely as a result of his appointment as an alternate other than in respect of such times as the alternate acts as a Director. The remuneration of such alternate shall be payable out of the remuneration of the Director appointing him and the proportion thereof shall be agreed between them.

POWERS AND DUTIES OF DIRECTORS

94. Subject to the Companies Act, these Articles and to any resolutions passed in a general meeting, the business of the Company shall be managed by the Directors, who may pay all expenses incurred in setting up and registering the Company and may exercise all powers of the Company. No resolution passed by the Company in general meeting shall invalidate any prior act of the Directors that would have been valid if that resolution had not been passed.
95. Subject to these Articles, the Directors may from time to time appoint any Person, whether or not a Director to hold such office in the Company as the Directors may think necessary for the administration of the Company, including but not limited to, chief executive officers, one or more other executive officers, the office of president, one or more vice-presidents, treasurer, assistant treasurer, manager or controller, and for such term and at such remuneration (whether by way of salary or commission or participation in profits or partly in one way and partly in another), and with such powers and duties as the Directors may think fit. Any Person so appointed by the Directors may be removed by the Directors. The Directors may also appoint one or more of their number to the office of managing director upon like terms, but any such appointment shall ipso facto terminate if any managing director ceases from any cause to be a Director, or if the Company by Ordinary Resolution resolves that his tenure of office be terminated.
96. The Directors may appoint any Person to be a Secretary (and if need be an assistant Secretary or assistant Secretaries) who shall hold office for such term, at such remuneration and upon such conditions and with such powers as they think fit. Any Secretary or assistant Secretary so appointed by the Directors may be removed by the Directors or by the Company by Ordinary Resolution.
97. The Directors may delegate any of their powers to committees consisting of such member or members of their body as they think fit; any committee so formed shall in the exercise of the powers so delegated conform to any regulations that may be imposed on it by the Directors.
98. The Directors may from time to time and at any time by power of attorney (whether under Seal or under hand) or otherwise appoint any company, firm or Person or body of Persons, whether nominated directly or indirectly by the Directors, to be the attorney or attorneys or authorised signatory (any such person being an “**Attorney**” or “**Authorised Signatory**”, respectively) of the Company for such purposes and with such powers, authorities and discretion (not exceeding those vested in or exercisable by the Directors under these Articles) and for such period and subject to such conditions as they may think fit, and any such power of attorney or other appointment may contain such provisions for the protection and convenience of Persons dealing with any such Attorney or Authorised Signatory as the Directors may think fit, and may also authorise any such Attorney or Authorised Signatory to delegate all or any of the powers, authorities and discretion vested in him.
99. The Directors may from time to time provide for the management of the affairs of the Company in such manner as they shall think fit and the provisions contained in the three next following Articles shall not limit the general powers conferred by this Article.
100. The Directors from time to time and at any time may establish any committees, local boards or agencies for managing any of the affairs of the Company and may appoint any Person to be a member of such committees or local boards and may appoint any managers or agents of the Company and may fix the remuneration of any such Person.

101. The Directors from time to time and at any time may delegate to any such committee, local board, manager or agent any of the powers, authorities and discretions for the time being vested in the Directors and may authorise the members for the time being of any such local board, or any of them to fill any vacancies therein and to act notwithstanding vacancies and any such appointment or delegation may be made on such terms and subject to such conditions as the Directors may think fit and the Directors may at any time remove any Person so appointed and may annul or vary any such delegation, but no Person dealing in good faith and without notice of any such annulment or variation shall be affected thereby.
102. Any such delegates as aforesaid may be authorised by the Directors to sub-delegate all or any of the powers, authorities, and discretion for the time being vested in them.
103. Subject to the Companies Act and other provisions of these Articles (including Article 48), the Directors may agree with a Shareholder to waive or modify the terms applicable to such Shareholder's subscription for Shares without obtaining the consent of any other Shareholder; provided that such waiver or modification does not amount to a variation or abrogation of the rights attaching to the Shares of any other Shareholders.

BORROWING POWERS OF DIRECTORS

104. The Directors may from time to time at their discretion exercise all the powers of the Company to raise or borrow money and to mortgage or charge its undertaking, property and assets (present and future) and uncalled capital or any part thereof, or to otherwise provide for a security interest to be taken in such undertaking, property or uncalled capital, and to issue debentures, debenture stock, bonds and other securities, whether outright or as collateral, whenever money is borrowed or as security for any debt, liability or obligation of the Company or of any third party.

THE SEAL

105. The Seal shall not be affixed to any instrument except by the authority of a resolution of the Directors provided always that such authority may be given prior to or after the affixing of the Seal and if given after may be in general form confirming a number of affixings of the Seal. The Seal shall be affixed in the presence of a Director or a Secretary (or an assistant Secretary) or in the presence of any one or more Persons as the Directors may appoint for the purpose and every Person as aforesaid shall sign every instrument to which the Seal is so affixed in their presence.
106. The Company may maintain a facsimile of the Seal in such countries or places as the Directors may appoint and such facsimile Seal shall not be affixed to any instrument except by the authority of a resolution of the Directors provided always that such authority may be given prior to or after the affixing of such facsimile Seal and if given after may be in general form confirming a number of affixings of such facsimile Seal. The facsimile Seal shall be affixed in the presence of such Person or Persons as the Directors shall for this purpose appoint and such Person or Persons as aforesaid shall sign every instrument to which the facsimile Seal is so affixed in their presence and such affixing of the facsimile Seal and signing as aforesaid shall have the same meaning and effect as if the Seal had been affixed in the presence of and the instrument signed by a Director or a Secretary (or an assistant Secretary) or in the presence of any one or more Persons as the Directors may appoint for the purpose.
107. Notwithstanding the foregoing, a Secretary or any assistant Secretary shall have the authority to affix the Seal, or the facsimile Seal, to any instrument for the purposes of attesting authenticity of the matter contained therein but which does not create any obligation binding on the Company.

DISQUALIFICATION OF DIRECTORS

108. The office of a Director shall be vacated, if the Director:
- (a) becomes bankrupt or makes any arrangement or composition with his creditors;

- (b) dies or is found to be or becomes of unsound mind;
- (c) resigns his office by notice in writing to the Company;
- (d) is prohibited by any applicable Law or Designated Stock Exchange Rules from being a Director;
- (e) without special leave of absence from the Board, is absent from meetings of the Board for three consecutive meetings and the Board resolves that his office be vacated; or
- (f) is removed from office pursuant to any other provision of these Articles.

PROCEEDINGS OF DIRECTORS

109. The Directors may meet together (either within or outside the Cayman Islands) for the despatch of business, adjourn, and otherwise regulate their meetings and proceedings as they think fit. Questions arising at any meeting shall be decided by a majority of votes of the Directors present at a meeting. At any meeting of the Directors, each Director present in person or represented by his alternate or proxy shall be entitled to one vote. In case of an equality of votes the Chairman shall have a second or casting vote. A Director may, and a Secretary or assistant Secretary on the requisition of a Director shall, at any time summon a meeting of the Directors. Any Director may appoint any Person, whether or not a Director, to be the proxy of that Director to attend and vote on his behalf, in accordance with instructions given by that Director, or in the absence of such instructions at the discretion of the proxy, at a meeting of the Board which that Director is unable to attend personally. The instrument appointing the proxy shall be in writing under the hand of the appointing Director and shall be in any usual or common form or such other form as the Board may approve, and must be lodged with the chairman of the meeting of the Board at which such proxy is to be used, or first used, prior to the commencement of the meeting.
110. A Director may participate in any meeting of the Directors, or of any committee appointed by the Directors of which such Director is a member, by means of telephone or similar communication equipment by way of which all Persons participating in such meeting can communicate with each other and such participation shall be deemed to constitute presence in person at the meeting.
111. The quorum necessary for the transaction of the business of the Directors shall be not less than three (3) Directors provided that any quorum shall require the presence of the Founder Director; *provided*, however, that if such quorum cannot be obtained for a Board meeting after two (2) consecutive notices of such meeting have been sent by the Company with the first notice providing not less than seven (7) days' prior notice and the second notice providing not less than five (5) days' prior notice, then the attendance of any three (3) Directors shall constitute a quorum, *provided* further that matters discussed in such adjourned meeting shall be limited to those stated in the written notices and agendas of the Board meeting, and resolutions passed in such meeting shall be subject to notices and agendas of such meeting as well as copies of all board papers shall be sent to all the Directors and each Shareholder who appointed a director to the Board at least seven (7) days prior to the relevant Board meeting. If at any time there is only a sole Director, the quorum shall be one Director. A Director represented by an alternate Director or by proxy at any meeting shall be deemed to be present for the purposes of determining whether or not a quorum is present.
112. A Director who is in any way, whether directly or indirectly, interested in a contract or transaction or proposed contract or transaction with the Company shall declare the nature of his interest at a meeting of the Directors. A general notice given to the Directors by any Director to the effect that he is a member of any specified company or firm or is to be regarded as interested in any contract or other arrangement which may thereafter be made with that company or firm shall be deemed a sufficient declaration of interest in regard to any contract so made. Subject to the Designated Stock Exchange Rules and disqualification by the chairman of the relevant Board meeting, a Director may vote in respect of any contract or proposed contract or arrangement notwithstanding that he may be interested therein and if he does so his vote shall be counted and he may be counted in the quorum at any meeting of the Directors at which any such contract or proposed contract or arrangement shall come before the meeting for consideration.

113. A Director may hold any other office or place of profit under the Company (other than the office of auditor) in conjunction with his office of Director for such period and on such terms (as to remuneration and otherwise) as the Directors may determine and no Director or intending Director shall be disqualified by his office from contracting with the Company either with regard to his tenure of any such other office or place of profit or as vendor, purchaser or otherwise, nor shall any such contract or arrangement entered into by or on behalf of the Company in which any Director is in any way interested, be liable to be avoided, nor shall any Director so contracting or being so interested be liable to account to the Company for any profit realised by any such contract or arrangement by reason of such Director holding that office or of the fiduciary relation thereby established. A Director, notwithstanding his interest, may be counted in the quorum present at any meeting of the Directors whereat he or any other Director is appointed to hold any such office or place of profit under the Company or whereat the terms of any such appointment are arranged and he may vote on any such appointment or arrangement.
114. Any Director may act by himself or his firm in a professional capacity for the Company, and he or his firm shall be entitled to remuneration for professional services as if he were not a Director; provided that nothing herein contained shall authorise a Director or his firm to act as auditor to the Company.
115. The Directors shall cause minutes to be made in books or loose-leaf folders provided for the purpose of recording:
- (a) all appointments of Officers made by the Directors;
 - (b) the names of the Directors present at each meeting of the Directors and of any committee of the Directors; and
 - (c) all resolutions and proceedings at all meetings of the Company, and of the Directors and of committees of Directors.
116. When the chairman of a meeting of the Directors signs the minutes of such meeting where a quorum was present the same shall be deemed to have been duly held notwithstanding that all the Directors have not actually come together or that there may have been a technical defect in the proceedings.
117. A resolution in writing signed by all the Directors or all the members of a committee of Directors entitled to receive notice of a meeting of Directors or committee of Directors, as the case may be (an alternate Director, subject as provided otherwise in the terms of appointment of the alternate Director, being entitled to sign such a resolution on behalf of his appointer), shall be as valid and effectual as if it had been passed at a duly called and constituted meeting of Directors or committee of Directors, as the case may be. When signed a resolution may consist of several documents each signed by one or more of the Directors or his duly appointed alternate.
118. The continuing Directors may act notwithstanding any vacancy in their body but if and for so long as their number is reduced below the number fixed by or pursuant to these Articles as the necessary quorum of Directors, the continuing Directors may act for the purpose of increasing the number, or of summoning a general meeting of the Company, but for no other purpose.
119. Subject to any regulations imposed on it by the Directors, a committee appointed by the Directors may elect a chairman of its meetings. If no such chairman is elected, or if at any meeting the chairman is not present within fifteen minutes after the time appointed for holding the meeting, or if the Chairman is unable to or unwilling to act as the chairman, the committee members present may choose one of their number to be chairman of the meeting.
120. A committee appointed by the Directors may meet and adjourn as it thinks proper. Subject to any regulations imposed on it by the Directors, questions arising at any meeting shall be determined by a majority of votes of the committee members present and in case of an equality of votes the chairman shall have a second or casting vote.

121. All acts done by any meeting of the Directors or of a committee of Directors, or by any Person acting as a Director, shall notwithstanding that it be afterwards discovered that there was some defect in the appointment of any such Director or Person acting as aforesaid, or that they or any of them were disqualified, be as valid as if every such Person had been duly appointed and was qualified to be a Director.

PRESUMPTION OF ASSENT

122. A Director of the Company who is present at a meeting of the Board of Directors at which action on any Company matter is taken shall be presumed to have assented to the action taken unless his dissent shall be entered in the Minutes of the meeting or unless he shall file his written dissent from such action with the person acting as the chairman or secretary of the meeting before the adjournment thereof or shall forward a written form of such dissent to such person immediately after the adjournment of the meeting. Such right to dissent shall not apply to a Director who voted in favour of such action.

DIVIDENDS

123. Subject to any rights and restrictions for the time being attached to any Shares, or as otherwise provided for in the Companies Act and these Articles, the Directors may from time to time declare dividends (including interim dividends) and other distributions on Shares in issue and authorise payment of the same out of the funds of the Company lawfully available therefor.
124. Subject to any rights and restrictions for the time being attached to any Shares, the Company by Ordinary Resolution may declare dividends, but no dividend shall exceed the amount recommended by the Directors.
125. The Directors may determine, before recommending or declaring any dividend, to set aside out of the funds legally available for distribution such sums as they think proper as a reserve or reserves which shall, in the absolute discretion of the Directors, be applicable for meeting contingencies, or for equalising dividends or for any other purpose to which those funds may be properly applied and pending such application may in the absolute discretion of the Directors, either be employed in the business of the Company or be invested in such investments as the Directors may from time to time think fit.
126. Any dividend may be paid in any manner as the Directors may determine. If paid by cheque it will be sent through the post to the registered address of the Shareholder or Person entitled thereto, or in the case of joint holders, to any one of such joint holders at his registered address or to such Person and such address as the Shareholder or Person entitled, or such joint holders as the case may be, may direct. Every such cheque shall be made payable to the order of the Person to whom it is sent or to the order of such other Person as the Shareholder or Person entitled, or such joint holders as the case may be, may direct.
127. The Directors when paying dividends to the Shareholders in accordance with the foregoing provisions of these Articles may make such payment either in cash or in specie and may determine the extent to which amounts may be withheld therefrom (including, without limitation, any taxes, fees, expenses or other liabilities for which a Shareholder (or the Company, as a result of any action or inaction of the Shareholder) is liable).
128. Subject to any rights and restrictions for the time being attached to any Shares, all dividends shall be declared and paid according to the amounts paid up on the Shares, but if and for so long as nothing is paid up on any of the Shares dividends may be declared and paid according to the par value of the Shares. No amount paid on a Share in advance of calls shall, while carrying interest, be treated for the purposes of this Article as paid on the Share.
129. If several Persons are registered as joint holders of any Share, any of them may give effectual receipts for any dividend or other moneys payable on or in respect of the Share.

130. No dividend shall bear interest against the Company.
131. Any dividend unclaimed after a period of six years from the date of declaration of such dividend may be forfeited by the Board of Directors and, if so forfeited, shall revert to the Company.

ACCOUNTS, AUDIT AND ANNUAL RETURN AND DECLARATION

132. The books of account relating to the Company's affairs shall be kept in such manner as may be determined from time to time by the Directors.
133. The books of account shall be kept at the Office, or at such other place or places as the Directors think fit, and shall always be open to the inspection of the Directors.
134. The Directors may from time to time determine whether and to what extent and at what times and places and under what conditions or regulations the accounts and books of the Company or any of them shall be open to the inspection of Shareholders not being Directors, and no Shareholder (not being a Director) shall have any right of inspecting any account or book or document of the Company except as conferred by law or authorised by the Directors or by Ordinary Resolution.
135. The accounts relating to the Company's affairs shall be audited in such manner and with such financial year end as may be determined from time to time by the Directors or failing any determination as aforesaid shall not be audited.
136. The Directors may appoint an Auditor of the Company who shall hold office until removed from office by a resolution of the Directors and may fix his or their remuneration.
137. Every Auditor of the Company shall have a right of access at all times to the books and accounts and vouchers of the Company and shall be entitled to require from the Directors and Officers of the Company such information and explanation as may be necessary for the performance of the duties of the auditors.
138. Auditors shall, if so required by the Directors, make a report on the accounts of the Company during their tenure of office at the next annual general meeting following their appointment, and at any time during their term of office, upon request of the Directors or any general meeting of the Shareholders.
139. The Directors in each year shall prepare, or cause to be prepared, an annual return and declaration setting forth the particulars required by the Companies Act and deliver a copy thereof to the Registrar of Companies in the Cayman Islands.

CAPITALISATION OF RESERVES

140. Subject to the Companies Act and these Articles, the Directors may:
- (a) resolve to capitalise an amount standing to the credit of reserves (including a Share Premium Account, capital redemption reserve and profit and loss account), whether or not available for distribution;
 - (b) appropriate the sum resolved to be capitalised to the Shareholders in proportion to the nominal amount of Shares (whether or not fully paid) held by them respectively and apply that sum on their behalf in or towards:
 - (i) paying up the amounts (if any) for the time being unpaid on Shares held by them respectively, or
 - (ii) paying up in full unissued Shares or debentures of a nominal amount equal to that sum, and allot the Shares or debentures, credited as fully paid, to the Shareholders (or as they may direct) in those proportions, or partly in one way and partly in the other, but the Share Premium Account, the capital redemption reserve and profits which are not available for distribution may, for the purposes of this Article, only be applied in paying up unissued Shares to be allotted to Shareholders credited as fully paid;

- (c) make any arrangements they think fit to resolve a difficulty arising in the distribution of a capitalised reserve and in particular, without limitation, where Shares or debentures become distributable in fractions the Directors may deal with the fractions as they think fit;
 - (d) authorise a Person to enter (on behalf of all the Shareholders concerned) into an agreement with the Company providing for either:
 - (i) the allotment to the Shareholders respectively, credited as fully paid, of Shares or debentures to which they may be entitled on the capitalisation, or
 - (ii) the payment by the Company on behalf of the Shareholders (by the application of their respective proportions of the reserves resolved to be capitalised) of the amounts or part of the amounts remaining unpaid on their existing Shares,
 and any such agreement made under this authority being effective and binding on all those Shareholders; and
 - (e) generally do all acts and things required to give effect to any of the actions contemplated by this Article.
141. Notwithstanding any provisions in these Articles, the Directors may resolve to capitalise any sum standing to the credit of any of the Company's reserve accounts or funds (including the Share Premium Account and capital redemption reserve fund) or any sum standing to the credit of the profit and loss account or otherwise available for distribution by applying such sum in paying up in full unissued Shares to be allotted and issued to:
- (a) employees (including Directors) or service providers of the Company or its Affiliates upon exercise or vesting of any options or awards granted under any share incentive scheme or employee benefit scheme or other arrangement which relates to such persons that has been adopted or approved by the Directors or the Shareholders;
 - (b) any trustee of any trust or administrator of any share incentive scheme or employee benefit scheme to whom shares are to be allotted and issued by the Company in connection with the operation of any share incentive scheme or employee benefit scheme or other arrangement which relates to such persons that has been adopted or approved by the Directors or Shareholders; or
 - (a) any depository of the Company for the purposes of the issue, allotment and delivery by the depository of ADSs to employees (including Directors) or service providers of the Company or its Affiliates upon exercise or vesting of any options or awards granted under any share incentive scheme or employee benefit scheme or other arrangement which relates to such persons that has been adopted or approved by the Directors or the Shareholders.

SHARE PREMIUM ACCOUNT

142. The Directors shall in accordance with the Companies Act establish a Share Premium Account and shall carry to the credit of such account from time to time a sum equal to the amount or value of the premium paid on the issue of any Share.
143. There shall be debited to any Share Premium Account on the redemption or purchase of a Share the difference between the nominal value of such Share and the redemption or purchase price provided always that at the determination of the Directors such sum may be paid out of the profits of the Company or, if permitted by the Companies Act, out of capital.

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NOTICES

144. Except as otherwise provided in these Articles, any notice or document may be served by the Company or by the Person entitled to give notice to any Shareholder either personally, or by posting it by airmail or air courier service in a prepaid letter addressed to such Shareholder at his address such Shareholder may have specified in writing for the purpose of such service of notices (or if absent of such notification, at his address as appearing in the Register), or by electronic mail to any electronic mail address such Shareholder may have specified in writing for the purpose of such service of notices, or by facsimile or by placing it on the Company's Website should the Directors deem it appropriate provided that the Company shall notify the Shareholders of the placement of such notice by any of the means set out above. In the case of joint holders of a Share, all notices shall be given to that one of the joint holders whose name stands first in the Register in respect of the joint holding, and notice so given shall be sufficient notice to all the joint holders.
145. Notices posted to addresses outside the Cayman Islands shall be forwarded by prepaid airmail.
146. Any Shareholder present, either personally or by proxy, at any meeting of the Company shall for all purposes be deemed to have received due notice of such meeting and, where requisite, of the purposes for which such meeting was convened.
147. Any notice or other document, if served by:
- (a) post, shall be deemed to have been served five days after the time when the letter containing the same is posted;
 - (b) facsimile, shall be deemed to have been served upon production by the transmitting facsimile machine of a report confirming transmission of the facsimile in full to the facsimile number of the recipient;
 - (c) recognised courier service, shall be deemed to have been served 48 hours after the time when the letter containing the same is delivered to the courier service; or
 - (d) electronic mail, shall be deemed to have been served immediately upon the time of the transmission by electronic mail.

In proving service by post or courier service it shall be sufficient to prove that the letter containing the notice or documents was properly addressed and duly posted or delivered to the courier service.

148. Any notice or document delivered or sent by post to or left at the registered address of any Shareholder in accordance with the terms of these Articles shall notwithstanding that such Shareholder be then dead or bankrupt, and whether or not the Company has notice of his death or bankruptcy, be deemed to have been duly served in respect of any Share registered in the name of such Shareholder as sole or joint holder, unless his name shall at the time of the service of the notice or document, have been removed from the Register as the holder of the Share, and such service shall for all purposes be deemed a sufficient service of such notice or document on all Persons interested (whether jointly with or as claiming through or under him) in the Share.
149. Notice of every general meeting of the Company shall be given to:
- (a) all Shareholders holding Shares with the right to receive notice and who have supplied to the Company an address for the giving of notices to them; and

- (b) every Person entitled to a Share in consequence of the death or bankruptcy of a Shareholder, who but for his death or bankruptcy would be entitled to receive notice of the meeting.

No other Person shall be entitled to receive notices of general meetings.

INFORMATION

150. No Shareholder shall be entitled to require discovery of any information in respect of any detail of the Company's trading or any information which is or may be in the nature of a trade secret or secret process which may relate to the conduct of the business of the Company and which in the opinion of the Board would not be in the interests of the Shareholders of the Company to communicate to the public.
151. The Board shall be entitled to release or disclose any information in its possession, custody or control regarding the Company or its affairs to any of its Shareholders including, without limitation, information contained in the Register and transfer books of the Company.

INDEMNITY

152. Every Director (including for the purposes of this Article any alternate Director appointed pursuant to the provisions of these Articles), Secretary, assistant Secretary, or other Officer (but not including the Company's auditors) and the personal representatives of the same (each an "**Indemnified Person**") shall be indemnified and secured harmless, to the fullest extent permitted by applicable law, against all actions, proceedings, costs, charges, expenses, losses, damages or liabilities incurred or sustained by such Indemnified Person, other than by reason of such Indemnified Person's own dishonesty, wilful default or fraud as determined by a court of competent jurisdiction, in or about the conduct of the Company's business or affairs (including as a result of any mistake of judgment) or in the execution or discharge of his duties, powers, authorities or discretions, including without prejudice to the generality of the foregoing, any costs, expenses, losses or liabilities incurred by such Indemnified Person in defending (whether successfully or otherwise) any civil proceedings concerning the Company or its affairs in any court whether in the Cayman Islands or elsewhere.
153. No Indemnified Person shall be liable:
- (a) for the acts, receipts, neglects, defaults or omissions of any other Director or Officer or agent of the Company; or
 - (b) for any loss on account of defect of title to any property of the Company; or
 - (c) on account of the insufficiency of any security in or upon which any money of the Company shall be invested; or
 - (d) for any loss incurred through any bank, broker or other similar Person; or
 - (e) for any loss occasioned by any negligence, default, breach of duty, breach of trust, error of judgement or oversight on such Indemnified Person's part; or
 - (f) for any loss, damage or misfortune whatsoever which may happen in or arise from the execution or discharge of the duties, powers, authorities, or discretions of such Indemnified Person's office or in relation thereto;

unless the same shall happen through such Indemnified Person's own dishonesty, wilful default or fraud as determined by a court of competent jurisdiction.

FINANCIAL YEAR

154. Unless the Directors otherwise prescribe, the financial year of the Company shall end on December 31st in each year and shall begin on January 1st in each year.

NON-RECOGNITION OF TRUSTS

155. No Person shall be recognised by the Company as holding any Share upon any trust and the Company shall not, unless required by law, be bound by or be compelled in any way to recognise (even when having notice thereof) any equitable, contingent, future or partial interest in any Share or (except only as otherwise provided by these Articles or as the Companies Act requires) any other right in respect of any Share except an absolute right to the entirety thereof in each Shareholder registered in the Register.

WINDING UP

156. If the Company shall be wound up the liquidator may, with the sanction of a Special Resolution of the Company and any other sanction required by the Companies Act, divide amongst the Shareholders in kind the whole or any part of the assets of the Company (whether they shall consist of property of the same kind or not) and may for that purpose value any assets and determine how the division shall be carried out as between the Shareholders or different classes of Shareholders. The liquidator may, with the like sanction, vest the whole or any part of such assets in trustees upon such trusts for the benefit of the Shareholders as the liquidator, with the like sanction, shall think fit, but so that no Shareholder shall be compelled to accept any asset upon which there is a liability.
157. If the Company shall be wound up, and the assets available for distribution amongst the Shareholders shall be insufficient to repay the whole of the share capital, such assets shall be distributed so that, as nearly as may be, the losses shall be borne by the Shareholders in proportion to the par value of the Shares held by them. If in a winding up the assets available for distribution amongst the Shareholders shall be more than sufficient to repay the whole of the share capital at the commencement of the winding up, the surplus shall be distributed amongst the Shareholders in proportion to the par value of the Shares held by them at the commencement of the winding up subject to a deduction from those Shares in respect of which there are monies due, of all monies payable to the Company for unpaid calls or otherwise. This Article is without prejudice to the rights of the holders of Shares issued upon special terms and conditions.

AMENDMENT OF ARTICLES OF ASSOCIATION

158. Subject to the Companies Act and the rights attaching to the various Classes, the Company may at any time and from time to time by Special Resolution alter or amend these Articles in whole or in part.

CLOSING OF REGISTER OR FIXING RECORD DATE

159. For the purpose of determining those Shareholders that are entitled to receive notice of, attend or vote at any meeting of Shareholders or any adjournment thereof, or those Shareholders that are entitled to receive payment of any dividend, or in order to make a determination as to who is a Shareholder for any other purpose, the Directors may provide that the Register shall be closed for transfers for a stated period which shall not exceed in any case 30 days. If the Register shall be so closed for the purpose of determining those Shareholders that are entitled to receive notice of, attend or vote at a meeting of Shareholders the Register shall be so closed for at least ten days immediately preceding such meeting and the record date for such determination shall be the date of the closure of the Register.
160. In lieu of or apart from closing the Register, the Directors may fix in advance a date as the record date for any such determination of those Shareholders that are entitled to receive notice of, attend or vote at a meeting of the Shareholders and for the purpose of determining those Shareholders that are entitled to receive payment of any dividend the Directors may, at or within 90 days prior to the date of declaration of such dividend, fix a subsequent date as the record date for such determination.

161. If the Register is not so closed and no record date is fixed for the determination of those Shareholders entitled to receive notice of, attend or vote at a meeting of Shareholders or those Shareholders that are entitled to receive payment of a dividend, the date on which notice of the meeting is posted or the date on which the resolution of the Directors declaring such dividend is adopted, as the case may be, shall be the record date for such determination of Shareholders. When a determination of those Shareholders that are entitled to receive notice of, attend or vote at a meeting of Shareholders has been made as provided in this Article, such determination shall apply to any adjournment thereof.

REGISTRATION BY WAY OF CONTINUATION

162. The Company may by Special Resolution resolve to be registered by way of continuation in a jurisdiction outside the Cayman Islands or such other jurisdiction in which it is for the time being incorporated, registered or existing. In furtherance of a resolution adopted pursuant to this Article, the Directors may cause an application to be made to the Registrar of Companies to deregister the Company in the Cayman Islands or such other jurisdiction in which it is for the time being incorporated, registered or existing and may cause all such further steps as they consider appropriate to be taken to effect the transfer by way of continuation of the Company.

DISCLOSURE

163. The Directors, or any service providers (including the Officers, the Secretary and the registered office agent of the Company) specifically authorised by the Directors, shall be entitled to disclose to any regulatory or judicial authority or to any stock exchange on which securities of the Company may from time to time be listed any information regarding the affairs of the Company including without limitation information contained in the Register and books of the Company.

*** CERTAIN MATERIAL (INDICATED BY THREE ASTERISKS IN BRACKETS) HAS BEEN OMITTED FROM THIS DOCUMENT BECAUSE IT IS BOTH (1) NOT MATERIAL AND (2) WOULD BE COMPETITIVELY HARMFUL IF PUBLICLY DISCLOSED.

COLLABORATION AND LICENSE AGREEMENT

THIS COLLABORATION AND LICENSE AGREEMENT (the “**Agreement**”) is entered into as of February 1, 2021 (the “**Effective Date**”), by and between EXELIXIS, INC., a Delaware corporation having an address at 1851 Harbor Bay Parkway, Alameda, CA 94502, USA (“**Exelixis**”) and ADAGENE INCORPORATED, a Delaware corporation having an address at 315 Montgomery Street, Suite 900, San Francisco, CA 94104 (“**Adagene**”). Exelixis and Adagene may be referred to herein individually as a “**Party**” or collectively as the “**Parties**”.

RECITALS

WHEREAS, Adagene, a biotechnology company, has developed proprietary technology platforms to enable innovative antibody discovery and engineering;

WHEREAS, Exelixis and Adagene desire to form a collaboration to discover masked antibody candidates for further drug and product development and commercialization, under which collaboration Adagene will be responsible for generating such masked antibody candidates using Adagene’s SAFEbody® platform technology, and Exelixis will be responsible for the further development and commercialization of the candidates; and

WHEREAS, the Parties wish to establish such collaboration, all on the terms and conditions set forth below.

AGREEMENT

NOW, THEREFORE, in consideration of the foregoing premises and the mutual covenants contained herein, and for other good and valuable consideration, the receipt and sufficiency of which are hereby acknowledged, Exelixis and Adagene hereby agree as follows:

1. DEFINITIONS

1.1 “**Accounting Standards**” means with respect to Adagene, that Adagene shall maintain records and books of accounts in accordance with United States Generally Accepted Accounting Principles (U.S. GAAP).

1.2 “[***]” means [***].

1.3 “**Adagene Indemnitee**” has the meaning set forth in Section 11.1.

1.4 “**Adagene Know-How**” means all Know-How that Adagene or its Affiliates Control as of the Effective Date or during the Term that is necessary or reasonably useful for the research, Development, manufacture, use, importation, offer for sale, sale, or

other Commercialization of any Lead Compound or Product. Without limiting the generality of the foregoing, the Adagene Know-How existing at the Effective Date is described in Exhibit A.

1.5 “Adagene Patents” means all Patents (other than Adagene Platform Patents) that Adagene or its Affiliates Control as of the Effective Date or during the Term that (a) describe or claim any Adagene Know-How or (b) would be infringed, absent a license or other right to practice granted under such Patents, by the research, Development, manufacture, use, importation, offer for sale, sale, or other Commercialization of any Lead Compound or Product (considering patent applications to be issued with the then-pending claims).

1.6 “Adagene Platform Patents” means all Patents that describe or claim any Adagene Platform Technology and do not specifically describe or claim a Compound or a Product and do not use and are not based on any Exelixis mAB, Exelixis Technology, or any of Exelixis’ Confidential Information.

1.7 “Adagene Platform Technology” means Adagene’s proprietary antibody engineering platform that is known as the SAFEbody[®] technology.

1.8 “Adagene Technology” means the Adagene Know-How, the Adagene Patents, the Adagene Platform Patents, and Adagene’s interest in the Joint Inventions and Joint Patents.

1.9 “Affiliate” means, with respect to any party, any entity that, directly or indirectly through one or more intermediaries, controls, is controlled by, or is under common control with such party, but for only so long as such control exists. As used in this Section 1.9, “control” means (a) to possess, directly or indirectly, the power to direct the management or policies of an entity, whether through ownership of voting securities, by contract relating to voting rights or corporate governance; or (b) direct or indirect beneficial ownership of more than fifty percent (50%) (or such lesser percentage which is the maximum allowed to be owned by a foreign corporation in a particular jurisdiction) of the voting share capital or other equity interest in such entity.

1.10 “Alliance Manager” has the meaning set forth in Section 3.6.

1.11 “Applicable Laws” means the applicable provisions of any and all national, supranational, regional, state, and local laws, treaties, statutes, rules, regulations, administrative codes, guidance, ordinances, judgments, decrees, directives, injunctions, orders, permits (including MAAs) of or from any court, Regulatory Authority, or governmental agency or authority having jurisdiction over or related to the subject item.

1.12 “Auditor” has the meaning set forth in Section 8.6.

1.13 “Available” means, with respect to a Target, that such Target is not Unavailable.

1.14 “Binder” means the antigen binding portion of an antibody, such as Fab and CDR fragments, single chain antibodies (scFv), and single domain antibodies (sdAb) (including any humanized versions thereof).

1.15 “Biosimilar Product” means, with respect to a product in a country, any pharmaceutical product that (a) is biosimilar to, or interchangeable with, such product or may otherwise be legally substituted for such product and for which the BLA references or relies on such product, (b) is approved for the same indication as such product, and (c) is sold in such country by a Third Party that is not a Sublicensee.

1.16 “BLA” means a Biologics License Application as described in 21 C.F.R. §601.2, including any amendments thereto, or any corresponding application in a country or jurisdiction in the Territory outside the U.S.

1.17 “Calendar Quarter” means each respective period of three (3) consecutive months ending on March 31, June 30, September 30, and December 31, or such other quarterly dates as Exelixis may elect by written notice to Adagene to correspond with Exelixis’ fiscal year.

1.18 “Calendar Year” means each respective period of twelve (12) consecutive months ending on December 31.

1.19 “Claim” has the meaning set forth in Section 11.1.

1.20 “Clinical Trial” means Phase 1 Clinical Trial, Phase 2 Clinical Trial, Phase 3 Clinical Trial, or other clinical study, as the context dictates.

1.21 “Collaboration Program” means the performance of both Target Program Plans under this Agreement.

1.22 “Collaboration Target” means each of the (a) Initial Target and (b) first Target designated by Exelixis to be the subject of a Target Program Plan pursuant to Section 4.2.

1.23 “Collaboration Target Designation Notice” has the meaning set forth in Section 4.2(c).

1.24 “Combination Product” means any pharmaceutical product containing (a) a Product and (b) at least one other active pharmaceutical ingredient, either co-formulated or packaged together and sold as a single unit for a single price. For clarity, “Combination Product” does not include any combination therapy that includes (x) the Product and (y) at least one other product which is not a Product, either administered or prescribed jointly, but which are not sold together for a single price.

1.25 “Commercialization” means the conduct of all activities undertaken before and after Regulatory Approval relating to the promotion, sales, marketing, medical support, and distribution (including importing, exporting, transporting, customs clearance, warehousing, invoicing, handling, and delivering products to customers) of products, including sales force efforts, detailing, advertising, market research, market access (including price setting and reimbursement activities), medical education and information services, publication, scientific and medical affairs, advisory and collaborative activities with opinion leaders and professional societies including symposia, marketing, sales force training, and sales (including receiving,

accepting, and filling product orders) and distribution. “**Commercialize**” and “**Commercializing**” have correlative meanings.

1.26 “Commercially Reasonable Efforts” means: (a) with respect to Exelixis’ obligations to Develop, seek MAA Approval, and Commercialize a Product under this Agreement, [***]; and (b) with respect to Adagene’s performance of each Target Program Plan, [***].

1.27 “Committee” means the JSC or any subcommittee established by the JSC, as applicable.

1.28 “Compound” means any Masked Exelixis mAB that binds to a Collaboration Target in the intended tumor microenvironment and is synthesized or created by Adagene using the Adagene Platform Technology, but excluding (a) Third Party-controlled compounds that are tested for comparison purposes and (b) compounds (including Exelixis mABs) provided to Adagene by or on behalf of Exelixis which are not further developed by Adagene under this Agreement.

1.29 “Compound Invention” has the meaning set forth in Section 9.1(b).

1.30 “Compound Research Activities” means, with respect to a Target Program Plan, (a) the activities required to be conducted to generate Compounds directed to the applicable Collaboration Target (such activities referred to as “**Part 1**” of the Target Program Plan) (b) the *in vitro* studies of the Compounds to be conducted by Adagene (such activities referred to as “**Part 2**” of the Target Program Plan), and (c) *in vivo* studies of the Lead Compounds to be conducted by the Parties (such activities referred to as “**Part 3**” of the Target Program Plan).

1.31 “Compulsory Sublicense” means, with respect to a Product in a country, a license or sublicense granted to a Third Party (a “**Compulsory Licensee**”) through the order, decree, or grant of a Governmental Authority in such country, authorizing such Compulsory Licensee to make, use, sell, offer for sale, import, or otherwise Commercialize such Product in such country. A Compulsory Licensee shall not be deemed to be a Sublicensee.

1.32 “Confidentiality Agreement” means that certain Mutual Confidential Disclosure Agreement by and between Exelixis and Adagene dated as of August 26, 2018.

1.33 “Confidential Information” means all Know-How and other proprietary scientific, marketing, financial, or commercial information or data that is generated by or on behalf of a Party or its Affiliates or which one Party or any of its Affiliates has supplied or otherwise made available to the other Party or its Affiliates, whether made available orally, in writing, or in electronic form, including information comprising or relating to concepts, discoveries, inventions, data, designs, or formulae in relation to this Agreement; provided that all Compound Inventions, Data, Exelixis mABs, and Exelixis Technology will be deemed Exelixis’ Confidential Information, all Adagene Technology and Adagene Platform Technology will be deemed Adagene’s Confidential Information, and all Joint Inventions and Joint Patents will be deemed both Parties’ Confidential Information. Confidential Information shall include: (a) the terms and

conditions of this Agreement, and (b) Confidential Information disclosed by either Party pursuant to the Confidentiality Agreement.

1.34 “**Control**” or “**Controlled**” means, with respect to any Know-How, Patents, or other intellectual property rights, the legal authority or right (whether by ownership, license, or otherwise but without taking into account any licenses granted by one Party to the other Party pursuant to this Agreement) of a Party to grant access, a license, or a sublicense of or under such Know-How, Patents, or other intellectual property rights to the other Party, or to otherwise disclose proprietary or trade secret information to such other Party, without breaching the terms of any agreement with a Third Party, or misappropriating the proprietary or trade secret information of a Third Party or triggering any payment obligation to a Third Party. Notwithstanding the foregoing, Know-How, Patents, or other intellectual property rights will be “Controlled” by a Party hereunder if such Know-How, Patents, or other intellectual property rights are owned or in-licensed by a Third Party that becomes an Affiliate of such Party after the Effective Date as a result of such Party (a) acquiring such Third Party or a portion of the business of such Third Party or (b) being acquired by such Third Party (in each case, whether by merger, stock purchase or purchase of assets).

1.35 “**Data**” means any and all scientific, technical, test, or other data pertaining to any Compound or Product that is generated by or on behalf of Adagene under any Target Program Plan, or is generated by on behalf of Exelixis (including by Adagene), or its Affiliates and sublicensees during the Term, including research data, clinical pharmacology data, chemistry-manufacture-controls (CMC) data (including analytical, and stability data), pre-clinical data, clinical data, clinical study reports, and all data used in association with submissions of an IND or MAA with respect to any Lead Compound or Product. For clarity, Data shall not include any data that Adagene generates to the extent that such data is only generally applicable to the Adagene Platform Technology or its use or to Platform Inventions and does not also pertain to any Compound or Collaboration Target.

1.36 “**Data Package Part 1**” has the meaning set forth in Section 4.4(b).

1.37 “**Data Package Part 2**” has the meaning set forth in Section 4.4(c).

1.38 “**Development**” means all development activities for a product (whether alone or for use together, or in combination, with another active agent or pharmaceutical product as a combination product or combination therapy) that are directed to obtaining Regulatory Approval(s) of such product and lifecycle management of such product in any country in the world, including all non-clinical, preclinical, and clinical testing and studies of such product; toxicology, pharmacokinetic, and pharmacological studies; statistical analyses; assay development; protocol design and development; the preparation, filing, and prosecution of any MAA for such product; development activities directed to label expansion and/or obtaining Regulatory Approval for one or more additional indications following initial Regulatory Approval; development activities conducted after receipt of Regulatory Approval; and all regulatory affairs related to any of the foregoing. “**Develop**” and “**Developing**” have correlative meanings.

1.39 “**Development Criteria**” means with respect to a Compound, the criteria for designating such Compound a Lead Compound, as reviewed, discussed, decided, and updated (as necessary) from time to time by the JSC.

1.40 “**Development Costs**” has the meaning set forth in Section 7.2(a).

1.41 “**Disputed Matter**” has the meaning set forth in Section 14.2.

1.42 “**EMA**” means the European Medicines Agency, or any successor agency thereto.

1.43 “**Executive Officers**” the Chief Scientific Officer of Exelixis and the Chief Executive Officer of Adagene.

1.44 “**Exelixis Indemnitee**” has the meaning set forth in Section 11.2.

1.45 “**Exelixis Know-How**” means all Know-How that Exelixis or its Affiliates Control as of the Effective Date or during the Term, including the Compound Inventions, that is necessary or reasonably useful for the research, Development, manufacture, use, importation, offer for sale, sale, or other Commercialization of any Lead Compound or Product.

1.46 “**Exelixis mAB**” means any mAB that is (a) directed to a specific Collaboration Target, and (b) is provided to Adagene by Exelixis for use in the generation of Compounds pursuant to Section 4.4(b) under the Target Program Plan for such Collaboration Target.

1.47 “**Exelixis Patents**” means all Patents in the Territory that Exelixis or its Affiliates Control as of the Effective Date or during the Term, including the Product Patents, that (a) describe or claim any Exelixis Know-How or (b) would be infringed, absent a license or other right to practice granted under such Patents, by the research, Development, manufacture, use, importation, offer for sale, sale, or other Commercialization of any Compound or Product (considering patent applications to be issued with the then-pending claims).

1.48 “**Exelixis Technology**” means the Exelixis Know-How and the Exelixis Patents, including Exelixis’ interest in the Joint Inventions and Joint Patents.

1.49 “**FDA**” means the U.S. Food and Drug Administration, or any successor agency thereto.

1.50 “**Field**” means any and all uses.

1.51 “**First Commercial Sale**” means, on a Product-by-Product and country-by-country basis, the first sale by Exelixis or any of its Affiliates or Sublicensees to a Third Party for end use of such Product in such country after Regulatory Approval has been granted with respect to such Product in such country.

1.52 “**FTE**” means the equivalent of a full-time Adagene (including its Affiliates) employee’s work for a twelve (12) month period (consisting of a total of at least [***] per year of dedicated effort). Any such employee who devotes fewer than [***] per year on the applicable activities shall be treated as an FTE on a pro-rata basis, based upon the actual number of hours worked by such employee on such activities, divided by [***]. For clarity, only hours devoted by qualified scientific, medical, manufacturing, technical, regulatory, and other similar employees of Adagene, as measured in accordance with Adagene’s normal time allocation practices, shall apply toward an FTE, and in no event shall “FTE” include hours devoted by personnel performing administrative, managerial, or corporate functions (e.g., human resources, finance, legal, and investor relations).

1.53 “**FTE Costs**” means the FTE Rate multiplied by the number of FTEs applied by Adagene (including its Affiliates) to the performance of Compound Research Activities in accordance with a Target Program Plan and Target Program Budget.

1.54 “**FTE Rate**” means: (a) for any work carried out in the PRC, an initial rate of [***] per FTE per year, and (b) for any work carried out in the United States, an initial rate of [***] per FTE per year; in each case of (a) and (b) which rate shall apply through [***]. Thereafter, the FTE Rate shall be changed annually on a Calendar Year basis to reflect any year-to-year percentage increase or decrease (as the case may be) in the Consumer Price Index for All Urban Consumers for the U.S., as published by the U.S. Department of Labor, Bureau of Labor Statistics (based on the change from the most recent applicable index available as of the Effective Date to the most recent applicable index available as of the date of the calculation of such revised FTE Rate).

1.55 “**GLP**” means in respect of laboratory activities, the then current standards required by Applicable Law of: (a) the United States, including the FDA’s Good Laboratory Practice regulations at 21 C.F.R. Part 58; (b) the European Union, including the Directive 2004/9/EC, Directive 2004/10/EC, guidance published by the European Commission or EMA in relation to such Directives and any local laws, rules and regulations that implement such Directives and guidance; and (c) such other countries in which a Licensed Product is tested.

1.56 “**GLP Toxicity Study**” means, with respect to a Product, an in vivo toxicology study conducted under conditions of GLP that is required for filing an IND for such Product with the FDA.

1.57 “**Governmental Authority**” means any national, international, federal, state, provincial, or local government, or political subdivision thereof, or any multinational organization or any authority, agency, or commission entitled to exercise any administrative, executive, judicial, legislative, police, regulatory, or taxing authority or power, any court or tribunal (or any department, bureau or division thereof, or any governmental arbitrator or arbitral body)

1.58 “**ICH**” means the International Conference on Harmonization (of Technical Requirements for Registration of Pharmaceuticals for Human Use).

1.59 “**IND**” means an investigational new drug application or equivalent application filed with the applicable Regulatory Authority, which application is required to commence human clinical trials in the applicable country.

- 1.60 “**Indemnified Party**” has the meaning set forth in Section 11.3.
- 1.61 “**Indemnifying Party**” has the meaning set forth in Section 11.3.
- 1.62 “**Initial Target**” means the undisclosed target already agreed between the parties.
- 1.63 “**Initiation**” means, with respect to the GLP Toxicity Study, the first dosing of the first animal subject in such GLP Toxicity Study, and with respect to a Clinical Trial, the first dosing of the first human subject in such Clinical Trial.
- 1.64 “**Injunctive Relief**” has the meaning set forth in Section 14.3(b).
- 1.65 “**Inventions**” means all inventions, whether or not patentable, discovered, made, conceived, or reduced to practice in the course of activities contemplated by this Agreement.
- 1.66 “**Joint Inventions**” has the meaning set forth in Section 9.1(d).
- 1.67 “**Joint Patents**” has the meaning set forth in Section 9.1(d).
- 1.68 “**Joint Steering Committee**” or “**JSC**” has the meaning set forth in Section 3.1.
- 1.69 “**Know-How**” means all technical information, know-how, and data, including inventions, discoveries, trade secrets, specifications, instructions, processes, formulae, compositions of matter, cells, cell lines, assays, animal models and other physical, biological, or chemical materials, expertise and other technology applicable to, development, manufacture, registration, use, or marketing or to methods of assaying or testing them, and including all biological, chemical, pharmacological, biochemical, toxicological, pharmaceutical, physical, and analytical, safety, nonclinical, and clinical data, regulatory documents, data and filings, instructions, processes, formulae, expertise and information, relevant to the research, development, use, importation, offering for sale or sale of, or which may be useful in studying, testing, or developing products. Know-How excludes Patents.
- 1.70 “**Lead Compound**” has the meaning set forth in Section 4.4(b).
- 1.71 “**Losses**” has the meaning set forth in Section 11.1.
- 1.72 “**MAA**” means a marketing authorization application or equivalent application, and all amendments and supplements thereto, filed with the applicable Regulatory Authority in the Territory.

1.73 “**MAA Approval**” means approval of an MAA by the applicable Regulatory Authority for marketing and sale of a product, but excluding any pricing and/or reimbursement approval.

1.74 “**mAB**” means a monoclonal, bispecific, or multispecific antibody.

1.75 “**Mask**” means a masking moiety (peptide) that binds to a mAB via a linker with a protease site so as to inhibit such a mAB from binding to the intended Target except in desired microenvironments.

1.76 “**Masked Exelixis mAB**” means an Exelixis mAB that contains a Mask.

1.77 “**MHLW**” means Japan’s Ministry of Health, Labour and Welfare, or any successor agency thereto.

1.78 “**Net Sales**” means, with respect to any Product, the gross amounts invoiced for sales or other dispositions of such Product by or on behalf of Exelixis or its Affiliates or Sublicensees to Third Parties, less the following deductions to the extent included in the gross invoiced sales price for such Product and determined in each case in accordance with generally accepted accounting principles in the U.S. (“**U.S. GAAP**”) or otherwise directly paid or incurred by Exelixis or its Affiliates or Sublicensees, as applicable, with respect to the sale or other disposition of such Product:

- (a) [***];
- (b) [***];
- (c) [***];
- (d) [***];
- (e) [***];
- (f) [***];
- (g) [***];
- (h) [***]; and
- (i) [***].

In no event will any particular amount identified above be deducted more than once in calculating Net Sales. Sales of a Product between Exelixis and its Affiliates or Sublicensees for resale shall be excluded from the computation of Net Sales, but the subsequent resale of such Product to a Third Party shall be included within the computation of Net Sales.

[***].

In the event a Product is sold as part of a Combination Product in a country, the Net Sales with respect to the Combination Product in such country shall be determined [***].

For clarity, Net Sales will be calculated on an accrual basis, in a manner consistent with U.S. GAAP and this definition of Net Sales. To the extent any accrued amounts used in the calculation of Net Sales are estimates, such estimates shall be true-up in accordance with Exelixis' internal accounting policies, consistently applied, and Net Sales and related payments under this Agreement shall be reconciled as appropriate.

For clarity, no deductions from Net Sales shall be made [***].

1.79 "NMPA" means National Medical Products Administration of China (formerly known as the China Food and Drug Administration), or any successor agency thereto.

1.80 "Part 1" has the meaning set forth in Section 1.30(a).

1.81 "Part 2" has the meaning set forth in Section 1.30(b).

1.82 "Part 3" has the meaning set forth in Section 1.30(c).

1.83 "Patents" means (a) all patents, certificates of invention, applications for certificates of invention, priority patent filings, and patent applications, and (b) any renewals, divisions, continuations (in whole or in part), or requests for continued examination of any of such patents, certificates of invention and patent applications, and any all patents or certificates of invention issuing thereon, and any and all reissues, reexaminations, extensions, supplementary protection certificates, divisions, renewals, substitutions, confirmations, registrations, revalidations, revisions, and additions of or to any of the foregoing.

1.84 "Phase 1 Clinical Trial" means a clinical trial conducted in a small number of patients designed or intended to establish an initial safety profile, pharmacodynamics, or pharmacokinetics of a pharmaceutical product. A Phase 1 Clinical Trial can be further divided into two separate trials:

(a) "Phase 1a Clinical Trial" means a single dose or multiple dose ascending clinical trial intended to evaluate safety and tolerability and determination of dose(s) for further clinical evaluation; and

(b) "Phase 1b Clinical Trial" means a clinical trial of a product, the principal purpose of which is a further determination of safety and pharmacokinetics of such product after an initial Phase 1a Clinical Trial, including as a single agent expansion cohort, prior to commencement of Phase 2 Clinical Trials (or phase 2 of a Phase 1/2 clinical trial) or Phase 3 Clinical Trials.

1.85 "Phase 2 Clinical Trial" means a clinical trial of a pharmaceutical product in human patients in any country to determine initial efficacy and safety and dose range finding. A

Phase 2 Clinical Trial is typically conducted before embarking on a Phase 3 Clinical Trial, but may be registrational.

1.86 "Phase 3 Clinical Trial" means a pivotal clinical trial of a pharmaceutical product in human patients in any country with a defined dose or a set of defined doses of such product designed to ascertain efficacy and safety of such product for the purpose of submitting MAAs for such product to the competent Regulatory Authorities.

1.87 "Platform Inventions" has the meaning set forth in Section 9.1(c).

1.88 "PRC" means the People's Republic of China.

1.89 "Product" means any pharmaceutical product containing a Lead Compound as an active ingredient, whether alone or in combination with other active ingredients, in any form, presentation, dosage, or formulation, and for any mode of administration.

1.90 "Product Infringement" has the meaning set forth in Section 9.3(a).

1.91 "Product Patent" means a Patent that claims a Compound Invention.

1.92 "Public Official or Entity" means (a) any officer, employee (including physicians, hospital administrators, or other healthcare professionals), agent, representative, department, agency, de facto official, representative, corporate entity, instrumentality or subdivision of any government, military, or international organization, including any ministry or department of health or any state-owned or affiliated company or hospital, or (b) any candidate for political office, any political party, or any official of a political party.

1.93 "R&D Cap" has the meaning set forth in Section 7.2(a).

1.94 "Registration Trial" means, with respect to a pharmaceutical product, a clinical study of such product in human patients (a) with a defined dose or a set of defined doses of such product designed to establish statistically significant efficacy and safety of such product for the purpose of enabling the preparation and submission of an MAA for such product to the competent Regulatory Authorities in a country or other jurisdiction, and (b) that would satisfy the requirements of 21 C.F.R. § 312.21(c), or its foreign equivalent.

1.95 "Regulatory Approval" means any and all approvals (including MAA Approval and, where required or commercially beneficial, pricing and reimbursement approval), licenses, registrations, permits, notifications, and authorizations (or waivers) of any Regulatory Authority that are necessary for the manufacture, use, storage, import, transport, promotion, marketing, distribution, offer for sale, sale, or other commercialization of a product in any country or jurisdiction.

1.96 “**Regulatory Authority**” means any Governmental Authority that has responsibility in its applicable jurisdiction over the testing, development, manufacture, use, storage,

import, transport, promotion, marketing, distribution, offer for sale, sale or other commercialization of pharmaceutical products in a given jurisdiction, including any Governmental Authority whose review or approval of pricing or reimbursement of such product is required.

1.97 “Regulatory Exclusivity” means any exclusive marketing rights or data exclusivity rights conferred by any Regulatory Authority with respect to a Product other than patents, including rights conferred in the U.S. under the Hatch-Waxman Act or the FDA Modernization Act of 1997 (including pediatric exclusivity), or rights similar thereto outside the U.S., such as Directive 2001/83/EC (as amended) in the EU.

1.98 “Regulatory Filing” means all applications, filings, submissions, approvals, licenses, registrations, permits, notifications, and authorizations (or waivers) with respect to the testing, Development, manufacture, or Commercialization of any pharmaceutical product made to or received from any Regulatory Authority in a given country, including any INDs and MAAs.

1.99 “Royalty Term” has the meaning set forth in Section 7.5(b).

1.100 “Rules” has the meaning set forth in Section 14.3(a).

1.101 “SEC” means the U.S. Securities and Exchange Commission, or any successor entity or its foreign equivalent in the Territory, as applicable.

1.102 “Sole Inventions” has the meaning set forth in Section 9.1(d).

1.103 “Sublicensee” means a Third Party to whom Exelixis grants a sublicense to offer for sale, sell, have sold, or otherwise Commercialize any Product in the Field in the Territory in accordance with Section 2.2, beyond the mere right to purchase Products from Exelixis and its Affiliates, and excluding wholesalers, full-service distributors that do not promote the sale of the Product, and other similar physical distributors. In no event shall Adagene or any of its Affiliates be deemed a Sublicensee.

1.104 “Target” means any human gene, together with the protein(s) encoded thereby, the expression or modulation of which is intended to, as its primary effect, treat, prevent, or otherwise have a disease-modifying effect.

1.105 “Target Nomination Notice” has the meaning set forth in Section 4.2(a).

1.106 “Target Program Budget” has the meaning set forth in Section 4.3(a).

1.107 “Target Program Plan” has the meaning set forth in Section 4.3(a).

1.108 “Target Selection Term” means the period of time commencing on the Effective Date and ending on the first (1st) anniversary of the Effective Date.

1.109 “**Term**” has the meaning set forth in Section 13.1.

1.110 “**Territory**” means worldwide.

1.111 “**Third Party**” means any entity other than Exelixis or Adagene or an Affiliate of Exelixis or Adagene.

1.112 “**Third Party Infringement Claim**” has the meaning set forth in Section 9.4.

1.113 “**Unavailable**” means, with respect to a Target, that such Target is the subject of (a) [***], or (b) [***]. Notwithstanding the foregoing, [***].

1.114 “**U.S.**” means the United States of America, including its territories and possessions (including Puerto Rico).

1.115 “**Valid Claim**” means (a) a claim of an issued and unexpired patent that has not been revoked or held unenforceable, unpatentable, or invalid by a decision of a court or other governmental agency of competent jurisdiction that is not appealable or has not been appealed within the time allowed for appeal, and that has not been abandoned, disclaimed, denied, or admitted to be invalid or unenforceable through reissue, re-examination, disclaimer, or otherwise, or (b) a claim of a pending patent application that has not been cancelled, withdrawn, abandoned, or finally rejected by an administrative agency action from which no appeal can be taken and that has not been pending for more than [***] from the date of filing of the earliest priority patent application to which such pending patent application is entitled to claim benefit.

1.116 “**VAT**” has the meaning set forth in Section 8.4(c).

2. GRANT OF LICENSES

2.1 **Exelixis Exclusive License.** Subject to the terms and conditions of this Agreement, Adagene hereby grants to Exelixis, during the Term, an exclusive (even as to Adagene), royalty-bearing license, with the right to grant sublicenses as provided in Section 2.2, under the Adagene Technology to research, Develop, use, make, have made, sell, offer for sale, import, and otherwise Commercialize Lead Compounds and Products in the Field in the Territory.

2.2 **Sublicenses.** Exelixis shall have the right to grant sublicenses, through multiple tiers of sublicensees, under the license granted in Section 2.1 to its Affiliates and other Third Parties, provided that such sublicenses shall be consistent with the terms and conditions of this Agreement and Exelixis remains responsible for the actions or omissions of its Sublicensees. Exelixis shall provide Adagene with written notice of any sublicense granted in accordance with this Section 2.2 within [***] after its execution, including the identity of the Sublicensee and the scope of the sublicense granted. If Exelixis sublicenses its rights under Section 2.1 to [***], Exelixis shall [***].

2.3 Adagene Reserved Rights. Adagene hereby expressly reserves (a) the right under the Adagene Technology to exercise its rights and perform its obligations under this Agreement, whether directly or through one or more permitted subcontractors, including the right to perform the applicable Compound Research Activities in accordance with each Target Program Plan; and (b) subject to Section 2.8, the right to practice, and to grant licenses under, the Adagene Technology outside of the scope of the rights granted in Section 2.1.

2.4 Adagene Research License. Subject to the terms and conditions of this Agreement, Exelixis hereby grants to Adagene, for the duration of the Collaboration Program, a nonexclusive, royalty-free license, with the right to grant sublicenses to permitted subcontractors for the performance of Compound Research Activities, under the Exelixis Technology solely to the extent necessary for Adagene to perform the Compound Research Activities allocated to it under each Target Program Plan.

2.5 No Implied Licenses. Except as set forth in this Agreement, neither Party shall acquire any license or other intellectual property interest, by implication or otherwise, under or to any Patents, Know-How, or other intellectual property owned or controlled by the other Party.

2.6 Technology Transfer. As reasonably required under Section 4.4, and in any event no later than the initial designation of the Compounds pursuant to Part 1 of the Target Program Plan, and otherwise as and when required under a Target Program Plan, as reasonably requested or required support Exelixis' IND filing, and as otherwise reasonably requested by Exelixis, Adagene shall disclose and make available to Exelixis all Adagene Know-How existing as of the Effective Date. Thereafter, on a Calendar Quarter basis during the performance of each Target Program Plan, or more often as reasonably requested by Exelixis, Adagene shall provide to Exelixis, at Exelixis' reasonable expense, any and all Adagene Know-How, Compound Inventions, and Data not previously disclosed to Exelixis pursuant to this Section 2.6. Adagene shall reasonably cooperate with Exelixis to provide a smooth and prompt transfer of all Adagene Know-How, Compound Inventions, and Data. Adagene shall reasonably assist Exelixis, at Exelixis' reasonable cost, in the use and understanding of such Data, Compound Inventions, and Adagene Know-How and shall provide reasonable technical assistance and make its technical personnel reasonably available to Exelixis as necessary or reasonably helpful for Exelixis to Develop and manufacture the Lead Compounds and Products. In addition to the above, Exelixis shall have the right to select [***] Compound sequences in association with each Target Program Plan at the end of Phase 2 of such Target Program Plan (in addition to the Lead Compound sequences) in consultation with Adagene. Adagene shall promptly disclose such selected Compound sequences in their entirety to Exelixis. Such Compound sequences shall be the Confidential Information of Exelixis hereunder, and Exelixis shall have the right to use and disclose such sequences as otherwise permitted under this Agreement. For the avoidance of doubt, (i) [***], and (ii) [***] under the Agreement.

2.7 Third Party Licenses. Adagene shall promptly notify Exelixis if it becomes aware of any Third Party Know-How or Patents that are necessary to Develop, make, have made, use, sell, offer for sale, or import any Lead Compound or Products. In such event, Exelixis shall have the right to negotiate and obtain a license from such Third Party under such Know-How and Patents at Exelixis' cost. Adagene shall not obtain any such license in a manner that would prohibit

Exelixis from obtaining a license under such Third Party Know-How and Patents to Develop, make, have made, use, sell, offer for sale, or import Lead Compound and Products.

2.8 Adagene Exclusivity.

(a) During the Collaboration Program. During the term of the Collaboration Program, Adagene shall not, and shall cause its Affiliates to not, directly or indirectly (including through a Third Party), research, develop, make, collaborate or consult with Third Parties, or commercialize any pharmaceutical product directed to a Collaboration Target other than Compounds and Products under this Agreement; provided, however, that Adagene shall be permitted to (i) conduct discovery activities for the discovery of novel antibodies that bind to an Exelixis Target and (ii) produce Masks that have less than a [***] sequence identity to any Mask provided by Adagene to Exelixis under this Agreement, in each case ((i) and (ii)) so long as no Compounds, Exelixis mABs, or other Exelixis Confidential Information is used, referenced, or relied upon in performing such activities.

(b) After the Collaboration Program. After the term of the Collaboration Program during the Term, Adagene shall not, and shall cause its Affiliates to not, directly or indirectly (including through a Third Party), (i) research, develop, make, collaborate or consult with Third Parties, or commercialize any product directed to any Collaboration Target to the extent that such product contains or comprises a mAB and the variable regions of such mAB have a [***] or greater sequence identity to any Exelixis mAB or (ii) produce any Mask that has a [***] or greater sequence identity to any Mask provided by Adagene to Exelixis under this Agreement for a Collaboration Target.

(c) Limitations. This Section 2.8 and the covenants set forth herein shall not apply to activities of any Third Party (or such Third Party's Affiliates) that becomes an Affiliate of Adagene after the Effective Date as a result of such Party (a) acquiring such Third Party or a portion of the business of such Third Party or (b) being acquired by such Third Party (in each case, whether by merger, stock purchase or purchase of assets) (such Third Party and its Affiliates immediately prior to the close of such transaction, the "**Adagene Acquiror**"), provided that (i) such Third Party activities are performed without using the Adagene SAFEbody™ technology and (ii) Adagene establishes reasonable firewall protections and safeguards designed to ensure the activities of its personnel under this Agreement are segregated from all activities of the Adagene Acquiror that would otherwise constitute a breach of Section 2.8(a) or 2.8(b), including without limitation ensuring that none of any Adagene Acquiror's personnel have access to Exelixis' Confidential Information, including non-public plans or information relating to the Development or Commercialization of Compounds or Products under this Agreement.

2.9 Exelixis Exclusivity. Except for Compounds and Products under this Agreement, during the Term Exelixis shall not, and shall cause its Affiliates to not, directly or indirectly (including through a Third Party), research, develop, make, collaborate or consult with Third Parties, or commercialize any product directed to a Collaboration Target to the extent that such product contains or comprises (a) an Exelixis mAB used in a Compound being Developed by or on behalf of Exelixis under this Agreement, or (b) an antibody the variable regions of which have a [***] or greater sequence identity to an Exelixis mAB used in a Compound being Developed by

or on behalf of Exelixis under this Agreement, in each case ((a) and (b)) that utilizes a protease-based masking technology that uses a peptide or small protein based domain (and in any case, [***] amino acids in length) as the blocking moiety.

3. GOVERNANCE

3.1 Joint Steering Committee. Within [***] after the Effective Date, the Parties shall establish a Joint Steering Committee (the “**Joint Steering Committee**” or the “**JSC**”), composed of an equal number of [***] of each Party, to oversee and guide the strategic direction of the collaboration of the Parties during the Collaboration Program. The JSC shall act as a joint consultative body during the Collaboration Program. The JSC in particular shall:

(a) oversee and monitor all Compound Research Activities under each Target Program Plan;

(b) discuss, review, comment on, and approve the Target Program Plan for the Collaboration Target selected pursuant to Section 4.2(b), including protocols for non-clinical and preclinical studies outlined in each Target Program Plan;

(c) discuss and review the use of any Exelixis mAB with respect to a Collaboration Target;

(d) review and discuss the status of each Target Program Plan and the Data generated during the performance thereof;

(e) discuss, review, comment on, and approve (subject to Section 3.3) any proposed amendments to a Target Program Plan, including any additional non-clinical and preclinical research, and protocols for additional non-clinical and preclinical studies;

(f) discuss and approve a transfer process and format for disclosure of Data, Compound Inventions, and Adagene Know-How to Exelixis under Section 2.6;

(g) identify the need for and create subcommittees of the JSC, which shall be subject to and report to the JSC, and assign responsibilities and authorities to such subcommittees, as the JSC determines necessary;

(h) discuss any Third Party Know-How or Patents that come to the attention of the Parties as set forth in Section 2.7 and determine whether it is advisable for either Party to negotiate with such Third Party for a license under any of such Know-How or Patents; and

(i) perform such other functions as appropriate to further the purposes of the Research Program, as expressly set forth in this Agreement or allocated to it by the Parties’ written agreement.

3.2 JSC Membership and Meetings.

(a) **JSC Members.** Each JSC representative shall have appropriate knowledge and expertise and sufficient seniority within the applicable Party to make decisions arising within the scope of the JSC's responsibilities. Each Party may replace its representatives on the JSC upon written notice to the other Party. Exelixis shall appoint the chairperson of the JSC. The chairperson shall oversee preparation and circulation of agendas to JSC members at least [***] before each JSC meeting and shall direct the preparation of reasonably detailed minutes for each JSC meeting, which shall be approved by the chairperson and circulated to JSC members within [***] after such meeting. The initial members of each of the JSC shall be determined by the Parties within [***] following the Effective Date.

(b) **Meetings.** The JSC shall hold meetings at such times as it elects to do so, but in no event less frequently than [***]. Meetings may be by telephone or video conference or in person. The first JSC meeting shall be held promptly after formation, but in any event no later than within [***] after the Effective Date. In-person meetings shall be held at locations alternately selected by the Parties. Each Party shall be responsible for all of its own expenses of participating in any JSC meeting. No action taken at any meeting of the JSC during the Collaboration Program shall be effective unless at least one (1) representative of each Party is participating. In addition, upon written notice to the other Party, either Party may request that a special *ad hoc* meeting of the JSC be convened for the purpose of resolving disputes or for the purpose of reviewing or making decisions pertaining to material subject-matter, the review or resolution of which cannot be reasonably postponed until the following scheduled JSC meeting. Such *ad hoc* meeting shall be convened at such time as may be mutually agreed by the Parties, but no later than [***] following the notification date of request that such meeting be held.

(c) **Non-Member Attendance.** Each Party may from time to time invite a reasonable number of participants, in addition to its representatives, to attend the JSC meetings in a non-voting capacity; provided that if either Party intends to have any Third Party (including any consultant) attend such a meeting, such Party shall provide reasonable prior written notice to the other Party and obtain the other Party's approval for such Third Party to attend such meeting, which approval shall not be unreasonably withheld or delayed. Such Party shall ensure that such Third Party is bound by written confidentiality and non-use obligations consistent with the terms of this Agreement.

3.3 Decision-Making.

(a) All JSC decisions shall be made by unanimous vote, with each Party's representatives collectively having one (1) vote. If after reasonable discussion and good faith consideration of each Party's view on a particular matter, the representatives of the Parties cannot reach an agreement as to such matter within [***], then either Party at any time may refer such issue to the Executive Officers for resolution.

(b) If the Executive Officers cannot resolve such matter within [***] after such matter has been referred to them, then (i) Adagene shall have final decision making authority, which shall be exercised in its sole reasonable discretion, with respect to the application of the Adagene Platform Technology in the conduct of the Compound Research Activities as described in Section 1.30(a) (i.e., in connection with the initial assembly and primary screen of Compounds, but not with respect to any *in vitro* or *in vivo* studies of such Compounds) and (ii) Exelixis shall

have final decision making authority, which shall be exercised in its sole reasonable discretion, with respect to all other Development, regulatory, and manufacturing matters with respect to any Target Program Plan, including, for clarity, the selection of the Lead Compound(s), preclinical and clinical Development work conducted solely by Exelixis (alone or in collaboration with Third Parties), regulatory activities, manufacturing and commercial matters with respect to the Lead Compound and the Product. Notwithstanding the foregoing subsection (ii), [***].

3.4 Limitations on Authority. The JSC shall have only such powers as are expressly assigned to it in this Agreement, and such powers shall be subject to the terms and conditions of this Agreement. Without limiting the generality of the foregoing, the JSC shall not have the power to amend, interpret, or waive compliance with this Agreement, and no JSC decision may be in contravention of any terms or conditions of this Agreement.

3.5 Discontinuation of the JSC. The activities to be performed by the JSC shall solely relate to governance under this Agreement, and are not intended to be or involve the delivery of services. The JSC shall continue to exist, on a Collaboration Target-by-Collaboration Target basis, until the first to occur of: (a) the Parties mutually agree to disband the JSC or (b) the completion of all Compound Research Activities under the Target Program Plan for such Collaboration Target. Upon the first to occur of the foregoing (a) or (b), the JSC shall automatically dissolve and, thereafter, each Party shall designate, to the extent necessary, a contact person for the exchange of information under this Agreement or such exchange of information shall be made through the Alliance Managers, and decisions of the JSC, if any, shall be decisions as between the Parties, subject to the other terms and conditions of this Agreement.

3.6 Alliance Managers. Promptly after the Effective Date, each Party shall appoint an individual who shall be an employee of such Party having appropriate qualification and experience to act as the alliance manager for such Party (the “**Alliance Manager**”). Each Alliance Manager shall be responsible for coordinating and managing processes and interfacing between the Parties on a day-to-day basis throughout the Collaboration Program and Term. If not already a member of the JSC, each Alliance Manager shall be permitted to attend JSC meetings as appropriate as non-voting participants. The Alliance Managers shall be the primary contact for the Parties regarding the activities contemplated by this Agreement and shall facilitate communication between the Parties with respect to all such activities hereunder. Each Party may replace its Alliance Manager with an alternative representative at any time with prior written notice to the other Party. Any Alliance Manager may designate a substitute to temporarily perform the functions of that Alliance Manager. Each Party shall bear its own costs of its Alliance Manager.

4. COMPOUND RESEARCH ACTIVITIES

4.1 Overview; Timelines. Subject to the terms and conditions of this Agreement, during the Collaboration Program the Parties will collaborate with respect to the performance of each Target Program Plan. Following the completion of a Target Program Plan, subject to Exelixis’ diligence obligations under Section 4.7, Exelixis shall have the sole and exclusive right and responsibility, at its own expense, for the Development of the Lead Compounds and Products corresponding to such Collaboration Target in the Territory.

4.2 Target Selection.

(a) **Target Nomination.** The Parties shall collaborate with respect to the performance of two (2) Target Program Plans. Subject to the remainder of this Section 4.2(a), during the Target Selection Term, Exelixis shall nominate Targets as Collaboration Targets as described in this Section 4.2 until one (1) further Target in addition to the Initial Target has been designated as a Collaboration Target under this Agreement.

(b) **Gatekeeper.** Exelixis shall nominate a Target as a Collaboration Target by providing written notice thereof (a “**Target Nomination Notice**”) to a Third Party gatekeeper mutually agreed upon by the Parties (“**Gatekeeper**”). Within [***] after the Gatekeeper’s selection, Adagene shall provide to the Gatekeeper a list of Targets that are Unavailable to enable him/her to make an accurate determination as to whether such nominated Target is Available or Unavailable. Upon the request of the Gatekeeper, Adagene shall provide supporting documentation to the Gatekeeper that is reasonably required by the Gatekeeper to verify that a Target is Unavailable. The Gatekeeper shall notify Exelixis of his/her determination within [***] after his/her receipt of Adagene’s Target list and supporting documentation (if requested), and the Gatekeeper’s determination shall be binding upon the Parties. [***]

(c) **Available.** If the Gatekeeper notifies Exelixis that a Target nominated by Exelixis in a Target Nomination Notice is Available, then, if Exelixis wishes to designate such Target as a Collaboration Target under this Agreement, Exelixis shall provide Adagene written notice confirming such nomination to Adagene (a “**Collaboration Target Designation Notice**”) within [***] after receipt of Adagene’s notice of Availability. Upon Adagene’s receipt of a Collaboration Target Designation Notice, such Target shall be automatically designated a Collaboration Target under this Agreement. As soon as reasonably practicable following such designation, the JSC shall mutually agree upon a Target Program Plan with respect to such Collaboration Target, which Target Program Plan shall be substantially similar in form and content to the Target Program Plan for the Initial Target, but taking into consideration any differences in research activities required to achieve the applicable criteria for such Collaboration Target.

(d) **Unavailable.** If the Gatekeeper notifies Exelixis that the Target is Unavailable, such Target shall not be designated a Collaboration Target and Exelixis shall have the right to nominate another Target as a Collaboration Target at any time prior to the later of (i) the expiration of the Target Selection Term and (ii) [***] after receipt of Adagene’s written notice of such Unavailability.

4.3 Target Program Plan.

(a) **General.** All Compound Research Activities for a Collaboration Target shall be conducted pursuant to a written plan (the “**Target Program Plan**”), which shall set forth the timeline and details for such Compound Research Activities. Each Target Program Plan shall also include a reasonably detailed budget of the costs by Calendar Quarter for the Compound Research Activities (the “**Target Program Budget**”) on an activity-by-activity or study-by-study basis, as appropriate. [***]. If the terms of any Target Program Plan contradict, or create inconsistencies or ambiguities with, the terms of this Agreement, then the terms of this Agreement shall govern. From time to time during the performance of each Target Program Plan, the Parties,

via the JSC, may prepare updates and amendments, as appropriate, to the then-current applicable Target Program Plan.

(b) Revisions. The JSC shall review each Target Program Plan and Target Program Budget on a Calendar Quarter basis for potential amendment in accordance with this Section 4.3(b). In addition, from time to time, either Party may propose updates and amendments, as appropriate, to a Target Program Plan to add, remove, or modify the Compound Research Activities under such Target Program Plan (including any consequential changes to the Target Program Budget) (“**Plan Revisions**”). The Party wishing to make such a revision shall provide the other Party, via the JSC, with a copy of the proposed Plan Revisions and allow the other Party a reasonable opportunity (not to exceed [***]) to review and comment thereon. Any Plan Revision must be approved by the JSC in accordance with Article 3.

4.4 Performance of Compound Research Activities. With respect to each Collaboration Target, Adagene shall use Commercially Reasonable Efforts to perform the Compound Research Activities set forth in the applicable Target Program Plan in accordance with the terms and timeline set forth therein and the terms of this Agreement to generate up to [***] Compounds that meet the applicable Development Criteria set forth in such Target Program Plan to be selected as Lead Compounds.

(a) Binders and mABs. With respect to each Collaboration Target, Exelixis will provide to Adagene an Exelixis mAB directed to the applicable Collaboration Target for use in the generation of Compounds pursuant to the following Section 4.4(b). Each Binder and mAB to be used in such activities shall be specified in the Target Program Plan for such Collaboration Target.

(b) Part 1 — Compound Generation. During Part 1 of each Target Program Plan, Adagene shall generate a series of alternative Compounds that are directed to the applicable Collaboration Target and are considered suitable for further evaluation. Prior to each regularly scheduled JSC meeting, Adagene shall provide Exelixis with all Data generated to date for a Target Program since the last such provision of Data pursuant to this Section 4.4(b). At each such meeting, the Parties shall review and discuss the Compounds Developed to date, and any Data generated by the Parties with respect thereto, and whether any such Compound is determined by Exelixis, in consultation with Adagene, to have satisfied the applicable Development Criteria and to be suitable to advance to Part 2. On a Collaboration Target-by-Collaboration Target basis, following completion of the Part 1 Compound Research Activities, Adagene shall prepare and deliver to the JSC a data package that includes the Data generated by Adagene in the performance of Part 1 of the respective Target Program Plan (the “**Data Package Part 1**”). Within [***] upon receipt of the Data Package Part 1, Exelixis may select [***] Compounds from the Data Package Part 1 to advance to Part 2. For clarity, Part 1 of the Target Program Plan corresponds to Stages 1-3 of the study plan agreed between the Parties.

(c) Part 2 — Assays and Studies. Following Exelixis’ designation of one or more Compounds pursuant to Section 4.4(b) for a Collaboration Target, Adagene shall perform the Compound Research Activities under Part 2 of the Target Program Plan for each such Compound. Exelixis shall have the right to audit Adagene’s

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performance of any Part 2 Compound Research Activities at Exelixis’ cost, and any Third Party vendors conducting such activities on Adagene’s behalf, in each case upon written request, and in each case only one time (in the absence of cause); provided, however, that if an audit of any Third Party vendor by Exelixis is not reasonably possible under Adagene’s written agreement with such vendor, Exelixis shall have the right, upon request, to review Adagene’s audit reports of such vendor. Prior to each regularly scheduled JSC meeting during the performance of the Part 2 activities, Adagene shall provide Exelixis with all Data generated under such Target Program Plan since the last such provision of Data pursuant to this Section 4.4(c). Following completion of the Part 2 Compound Research Activities for each Compound, Adagene shall prepare and deliver to the JSC a data package that includes all Data generated by Adagene in the performance of Part 2 of such Target Program Plan (the “**Data Package Part 2**”). The JSC shall review the Data Package Part 2 and determine whether any data or information is missing from such Data Package Part 2. With respect to any data or information identified as missing from such Data Package Part 2, the Parties shall cooperate to promptly update such Data Package to include such missing information and data, provided that Adagene shall not be obliged to perform further research activities, including any further *in vivo* studies other than the studies described in the applicable Target Program Plan, unless otherwise agreed by the parties pursuant to Article 3. Following the provision of such missing information to the Data Package Part 2, or if the JSC makes no requests for additional information, the Data Package, and corresponding Target Program Plan, shall be deemed complete. Upon the Data Package Part 2 being deemed complete, Exelixis shall have the right to select [***] Compounds from the Data Package to be lead compounds (each such Compound so selected, a “**Lead Compound**”). In the event that no Lead Compound for a Collaboration Target is designated, Exelixis shall have the right to terminate such Collaboration Target upon written notice to Adagene in accordance with Section 13.3. For clarity, Part 2 of the Target Program Plan corresponds to Stage 4 of the study plan agreed between the Parties.

(d) Part 3 — In Vivo Studies and Manufacturing. Following Exelixis’ designation [***] Lead Compounds pursuant to Section 4.4(c) for a Collaboration Target, Adagene shall perform the Compound Research Activities under Part 3 of the Target Program Plan for each such Lead Compound. On a Collaboration Target-by-Collaboration Target basis, following completion of the Part 3 Compound Research Activities, Adagene shall prepare and deliver to the JSC a data package that includes the Data generated by Adagene in the performance of Part 3 of the respective Target Program Plan. For clarity, Part 3 of the Target Program Plan corresponds to Stages 5 and 6 of the study plan agreed between the Parties.

4.5 Development Costs. Adagene shall be responsible for the Development Costs incurred by Adagene in the performance of the Compound Research Activities in accordance with each Target Program Plan and associated Target Program Budget. Exelixis shall reimburse Adagene for Development Costs associated with additional Compound Research Activities added to a Target Program Plan as and to the extent set forth in Section 7.2(a).

4.6 Diligence.

(a) General.

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(i) During the Collaboration Program, Adagene shall use Commercially Reasonable Efforts to perform the Compound Research Activities under and in accordance with each Target Program Plan.

(ii) Following the delivery to Exelixis of (A) a complete Data Package Part 2 in accordance with Section 4.4(c) and (B) the material specified in Section 6.1, in each case ((A) and (B)) with respect to a Collaboration Target, Exelixis shall use Commercially Reasonable Efforts to Develop and obtain Regulatory Approval for one (1) Product corresponding to such Collaboration Target in the Territory.

(b) Development Updates.

(i) **Adagene's Obligations.** At each regularly scheduled JSC meeting during the Collaboration Program, Adagene shall provide the JSC with regular reports detailing its activities under each Target Program Plan, in each case to the extent then-ongoing, and the results of such activities. The Parties shall discuss the status, progress, and results of such activities at such JSC meetings.

(ii) **Exelixis' Obligations.** Following the discontinuation of the JSC pursuant to Section 3.5, Exelixis shall provide to Adagene written reports on [***] basis regarding its significant Development (including receipts of Regulatory Approval) activities with respect to Products for each Collaboration Target in the Territory. Each such written report shall summarize in reasonable detail Exelixis', its Affiliates', and Sublicensees' significant Development activities, and the progress made with respect to the Development and Regulatory Approval of Products in the Territory, and shall contain relevant information at a level of detail reasonably required by Adagene to determine whether Exelixis is in compliance with its diligence obligations set forth herein.

4.7 Compliance. Each Party shall perform the Development activities assigned to it under each Target Program Plan in compliance with all Applicable Laws, including GLP and other good scientific and clinical practices under the Applicable Laws of the country in which such activities are conducted.

4.8 Development Records. Each Party shall maintain complete, current, and accurate records of all Development activities conducted by it hereunder, and all data and other information resulting from such activities. Such records shall fully and properly reflect all work done and results achieved in the performance of the Development activities in good scientific manner appropriate for regulatory and patent purposes. Each Party shall document all non-clinical studies and Clinical Trials in formal written study reports according to Applicable Laws. Adagene shall provide Exelixis, during regular business hours and upon reasonable notice, the opportunity to inspect and copy all such Development records created by Adagene in the performance of any Target Program Plan.

4.9 Use of Subcontractors. Adagene may perform its Development activities under this Agreement through one or more subcontractors with Exelixis' prior written consent, provided that (a) Adagene will remain responsible for the work allocated to, and payment to, such

subcontractors to the same extent it would if it had done such work itself; (b) each subcontractor undertakes in writing obligations of confidentiality and non-use regarding Confidential Information that are substantially the same as those undertaken by the Parties pursuant to Article 13; and (c) each subcontractor agrees in writing to assign all intellectual property developed in the course of performing any such work to Adagene. For clarity, Exelixis may subcontract any of its Development activities under this Agreement in its sole discretion, provided that any such subcontracting shall be consistent with the terms and conditions of this Agreement, including the confidentiality obligations and the intellectual property provisions.

4.10 Regulatory Responsibilities. Exelixis shall own all Regulatory Filings and Regulatory Approvals for the Products in the Field in the Territory, and shall be solely responsible for preparing any and all Regulatory Filings for the Product in the Field in the Territory at its expense. Upon Exelixis' request and at its expense, Adagene shall reasonably assist Exelixis in connection with the preparation and filing of such Regulatory Filings, including by providing any and all documentation, Data, and other Adagene Know-How reasonably requested by Exelixis in support of such Regulatory Filings, in each case to the extent not previously provided pursuant to Section 2.6.

5. COMMERCIALIZATION

5.1 General. Subject to the terms and conditions of this Article 5, Exelixis shall have the sole and exclusive right and responsibility, at its expense, for all aspects of the Commercialization of Products corresponding to Collaboration Targets in the Territory.

5.2 Diligence.

(a) General. During the Term, with respect to each Collaboration Target, Exelixis, either itself or through its Affiliates or Sublicensees, shall use Commercially Reasonable Efforts to Commercialize one Product for which Regulatory Approval has been obtained in the Territory. For clarity, as between the Parties and subject to the foregoing sentence, Exelixis shall retain at all times sole and complete decision-making authority regarding whether to launch any particular Product in any particular country, and whether to launch any particular Product itself or through its Affiliates or Sublicensees anywhere in the Territory.

(b) Commercial Updates. Exelixis shall provide to Adagene written reports on [***] basis regarding its significant Commercialization activities with respect to Products in the Territory. Each such written report shall summarize in reasonable detail Exelixis', its Affiliates', and Sublicensees' significant Commercialization activities that have been conducted since the last reporting date and that are planned for the current year with respect to the Commercialization of each Product in the Territory.

6. MANUFACTURE AND SUPPLY

6.1 Adagene Manufacture and Supply. Adagene, itself or through one or more Third Party contract manufacturers, shall be responsible, at its own expense, for all manufacturing activities necessary to perform any of its obligations under each Target Program Plan. In addition, Adagene shall provide, at its own expense, Exelixis with the sequence of each Lead Compound,

as well as 10ug of cDNA for each Lead Compound for a Collaboration Target, in each case promptly after the designation thereof pursuant to Section 4.4(c).

6.2 Exelixis Supply. Following the completion of a Target Program Plan in respect of a Collaboration Target, Exelixis, either directly or through one or more Third Party contract manufacturers, shall provide all Lead Compounds and Products for use in the Development and Commercialization of such Products under this Agreement.

7. FINANCIAL PROVISIONS

7.1 Upfront Payment. In partial consideration of the rights granted to Exelixis under this Agreement, within [***] after the Effective Date, Exelixis shall pay to Adagene a one-time, non-refundable, non-creditable upfront payment of eleven million dollars (\$11,000,000).

7.2 Reimbursement of Development Costs.

(a) Development Costs. Adagene shall be responsible for the costs incurred in performing the Compound Research Activities specified under the initial Target Program Plan for each Collaboration Target (including FTE Costs and amounts that Adagene pays to Third Parties subcontracted to perform certain of Adagene's obligations under a Target Program Plan) (collectively, "**Development Costs**"). In addition, Adagene shall be responsible for the Development Costs incurred in performing additional activities or studies added to the initial Target Program Plan pursuant to Section 4.3(b) to the extent that such Development Costs do not exceed, in the aggregate, [***] more than the total amount budgeted under the initial Target Program Budget for such Target Program Plan (the "**R&D Cap**"). Exelixis shall reimburse Adagene for any Development Costs incurred by Adagene in performing additional activities or studies added to the initial Target Program Plan pursuant to Section 4.3(b) in excess of the R&D Cap to the extent such costs are incurred in accordance with the applicable Target Program Budget (as revised pursuant to Section 4.3(b)).

(b) Estimates and Invoices. Within [***] after the end of each month in which Adagene incurs any Development Costs under a Target Program Plan in excess of the R&D Cap, Adagene shall submit to Exelixis (i) an invoice setting forth the total Development Costs incurred by Adagene in accordance with the applicable Target Program Budget in such month that exceed the R&D Cap, (ii) monthly timesheets indicating the percentage of time and the hour equivalent of such percentage Adagene personnel allocated to such Compound Research Activities during such month, as reasonably necessary to confirm the manner and circumstances in which such costs were incurred, and (iii) documentation confirming any out-of-pocket expenses paid by Adagene for such Compound Research Activities during such month. Exelixis shall pay the undisputed amounts invoiced within [***] after receipt of the invoice. Adagene shall notify Exelixis as soon as reasonably practicable in the event that it becomes aware that Development Costs are expected to deviate from the Target Program Budget, whereupon the Parties shall discuss the causes of such deviation, and an appropriate adjustment to the Target Program Budget. The Parties shall refer any Target Program Budget increase amounting to greater than [***] of the previously approved amount to the JSC for prior approval. Unless such budget increase is approved by the JSC, Exelixis

shall have no responsibility for, nor any liability in respect of, any Development Costs in excess of [***] of the amount set out in the Target Program Budget (as revised from time to time).

7.3 Development Milestone Payments.

(a) **Development Milestones.** Subject to the remainder of this Section 7.3(a), on a Collaboration Target-by-Collaboration Target basis, Exelixis shall pay to Adagene the one-time, non-refundable, non-creditable payment set forth in the table below upon the first achievement of the respective applicable milestone event by a Product corresponding to such Collaboration Target (whether by or on behalf of Exelixis or its Affiliates or Sublicensees):

Milestone Event	Milestone Payment
1. [***]	[***]
2. [***]	[***]
3. [***]	[***]
4. [***]	[***]
5. [***]	[***]
6. [***]	[***]
7. [***]	[***]
8. [***]	[***]

For clarity, the foregoing milestone payments shall be payable one-time only with respect to each Collaboration Target, regardless of the number of Products corresponding to such Collaboration Target to achieve such milestone event. With respect to the milestone events and milestone payments listed under number 1 through number 5 above, if any such milestone event has not occurred until the achievement of the subsequent milestone event for a Product directed to the same Collaboration Target, the milestone payment for such milestone event shall become due upon the achievement of the subsequent milestone event by a Product directed to the same Collaboration Target.

(b) **Notice and Payment.** Exelixis shall notify Adagene in writing within [***] after the achievement of any milestone set forth in this Section 7.3 by Exelixis, its Affiliates, or Sublicensee(s). Exelixis shall pay to Adagene the applicable development milestone payments within [***] after the achievement of the applicable milestone event.

7.4 Sales-Based Milestones Payments.

(a) **Sales Milestones.** Subject to the remainder of this Section 7.4(a), on a Collaboration Target-by-Collaboration Target basis, Exelixis shall pay to Adagene the non-refundable, non-creditable payments set forth in the table below within [***] after the end of the Calendar Year in which the corresponding milestone event is first achieved by all Products corresponding to a Collaboration Target (whether by or on behalf of Exelixis or its Affiliates or Sublicensees).

Annual Net Sales of all Products in the Territory directed to a particular Collaboration Target	Sales Milestone Payments
Exceed [***]	[***]
Exceed [***]	[***]
Exceed [***]	[***]

For clarity, the foregoing milestone payments shall be payable one-time only with respect to each Collaboration Target, regardless of the number of Products corresponding to such Collaboration Target to achieve such milestone event. For the avoidance of doubt, from and after the expiration of the Royalty Term for a Product in a country, Net Sales of such Product in such country shall be excluded for purposes of calculating the Net Sales thresholds set forth in this Section 7.4(a). If more than one sales milestone event is achieved in a single Calendar Year, then the sales milestone payments for all sales milestone events achieved in such Calendar Year shall be due.

(b) **Notice and Payment.** As part of the report provided pursuant to Section 8.1, on a Collaboration Target-by-Collaboration Target basis, Exelixis shall provide written notice to Adagene upon the annual Net Sales of the Products for a Collaboration Target in the Territory first reaching each of the values set forth in Section 7.4(a) above. Each milestone payment set forth in the table above shall be paid as set forth in Section 7.4(a).

7.5 Royalty Payments.

(a) **Royalty Rate.** On a Collaboration Target-by-Collaboration Target basis, subject to the remainder of this Section 7.5, during the Royalty Term for a Product, Exelixis shall pay to Adagene a non-refundable, non-creditable [***] royalty on the aggregate annual Net Sales of such Product in the Territory, as calculated by multiplying the foregoing royalty rate by the amount of Net Sales of such Product in the Territory in the applicable Calendar Year.

(b) **Royalty Term.** Royalties shall be paid on a Product-by-Product and country-by-country basis from the First Commercial Sale of a Product in a country by or on behalf of Exelixis, its Affiliates, or Sublicensees until the latest of (i) ten (10) years after the First Commercial Sale of such Product in such country, (ii) expiration of the last-to-expire Valid Claim of the Adagene Patents, Adagene Platform Patents, and Product Patents in such country covering such Product's composition of matter or method of use as such Product is sold in such country, or

(iii) the expiration of Regulatory Exclusivity for such Product in such country (the “**Royalty Term**”).

(c) Royalty Reductions.

(i) Biosimilar Entry. If one or more Biosimilar Products to a Product is sold in a country in the Territory in any Calendar Quarter during the Royalty Term for such Product in such country, and during such Calendar Quarter [***], then the royalty rate provided in Section 7.5(a) for such Product in such country shall be reduced [***] for such Calendar Quarter and each subsequent Calendar Quarter during such Royalty Term.

(ii) Patent Expiry. If during any Calendar Quarter of the Royalty Term for a particular Product in a country no Valid Claim of any Adagene Patent, Adagene Platform Patent, or Product Patent in such country covers such Product’s composition of matter or method of use as such Product is sold in such country, then the royalty rate provided in Section 7.5(a) shall be reduced by [***] for such Product in such country for such Calendar Quarter.

(iii) Royalty Floor. Notwithstanding the foregoing Sections 7.5(c)(i) and 7.5(c)(ii) with respect to any Product in any Calendar Quarter, the operation of Sections 7.5(c)(i) and 7.5(c)(ii), individually or in combination, shall not reduce by more than [***] the royalties that would otherwise have been due under Section 7.5(a) with respect to Net Sales of such Product in the applicable country(ies) during such Calendar Quarter.

(d) Compulsory Licenses. If Exelixis or its Affiliate or Sublicensee is required in a given country to issue a Compulsory Sublicense to one or more Compulsory Licensees for the sale of Product(s) in such country, then the sales of such Compulsory Sublicensee shall be excluded from Net Sales, and Exelixis shall pay to Adagene [***] of any royalty payment received by Exelixis from the compulsory sublicensee for the sale of such Product(s) in such country.

8. PAYMENT; RECORDS; AUDITS

8.1 Payment; Reports. Royalty payments due by Exelixis to Adagene under Section 7.5 shall be calculated and reported for each Calendar Quarter during the Royalty Term. Royalty payments due under Section 7.5 shall be paid within [***] after the end of each Calendar Quarter and shall be accompanied by a report setting forth the Net Sales of the Products by Exelixis and its Affiliates and Sublicensees in the Territory in sufficient detail to permit confirmation of the accuracy of the royalty payment made, including the number of Products sold, the gross sales and Net Sales of Products ([***]), the royalties payable, the method used to calculate the royalties, the exchange rates used, and whether any sales milestone under Section 7.4 has been achieved.

8.2 Exchange Rate; Manner and Place of Payment. All references to dollars and “\$” herein shall refer to U.S. dollars. All payments hereunder shall be payable in U.S. dollars.

With respect to conversion of Net Sales in any currency other than U.S. dollars to U.S. dollars, Exelixis or its Affiliates or Sublicensees shall convert the Net Sales for the Products using its then-current standard worldwide currency conversion methodology applied to its external reporting. All payments owed to Adagene under this Agreement shall be made by wire transfer in immediately available funds to a bank and account designated in writing by Adagene, unless otherwise specified in writing by Adagene.

8.3 Adagene Reporting.

(a) As of the Effective Date, Adagene shall provide Exelixis with a copy of Adagene's most recent unaudited financial statements prepared in accordance with Accounting Standards aggregated summary of programs by development stage (i.e. non-clinical vs clinical) and aggregated summary of resources (FTE allocation) ("**Initial Financial Information**") as required to make a determination if entering into this Agreement imposes on Exelixis the requirements of the variable interest entity consolidation model as described in Accounting Standards Codification 810 ("**VIE Accounting**"). During the Term, Adagene shall provide Exelixis its annual audited financial statements in accordance with Accounting Standards within four (4) months of the end of each fiscal year of Adagene. In addition, Adagene shall provide Exelixis updates to the Initial Financial Information and information pertaining to other material changes to its business as reasonably necessary for such determination on a quarterly basis, as set forth in **Exhibit D** by the end of the second month of each Calendar Quarter. If Exelixis determines in its reasonable judgment that VIE Accounting is required in any Calendar Quarter (the "**VIE Determination**") then the provisions of Sections 8.3(b) and 8.3(c) below will apply until such time as VIE Accounting is no longer required. In the event that a VIE Determination is not made, or a VIE Determination is removed, then the provisions of Sections 8.3(b) and 8.3(c) below will not apply, but Adagene will continue to provide quarterly updates in accordance with this Section 8.3(a).

(b) Subject to Section 8.3(a), Adagene shall continue to provide updates to the Initial Financial Information on a quarterly basis, and annual audited financial statements on an annual basis. In addition, Adagene will provide information reasonably required to prepare consolidated financial statements in accordance with US GAAP. Adagene may choose to remove any confidential information that is not required for Exelixis consolidation reporting.

(c) Subject to Section 8.3(a), should Adagene enter into a Relevant Transaction, Adagene shall disclose the occurrence of such Relevant Transaction within thirty (30) days of such occurrence to Exelixis in writing, and, upon Exelixis' request, shall provide to Exelixis additional information about such Relevant Transaction solely to the extent reasonably necessary for Exelixis to confirm the VIE Determination in accordance with Accounting Standards, and subject to any confidentiality obligations of Adagene to any Third Party. For purposes of the above, "**Relevant Transaction**" means any of the following: (i) a bona fide equity financing which results in a material change to Adagene's capitalization, or (ii) any material purchase or redemption by Adagene of any shares of Adagene's capital stock other than (A) repurchases or redemptions of stock from former employees, officers, directors, consultants or other persons who performed services for Adagene or any subsidiary in connection with the

cessation of such employment or service, or (B) the exercise of a contractual right of first refusal by Adagene, or (iii) any material contractual arrangement under which Adagene licenses to Third Parties any Adagene's technology or platform, any divestitures, reorganization of the entity.

8.4 The Initial Financial Statements and any other information provided to Exelixis under this Section 8.4: (i) shall be deemed Licensor's Confidential Information under Article 12, (ii) without limiting the foregoing, may only be used by Exelixis' Chief Financial Officer, controller or similarly situated employee (and personnel working under such controller, on a need-to-know basis) and Exelixis' independent auditors solely for the purpose of confirming VIE Determination in accordance to U.S. GAAP, and (iii) may not be shared with any other Exelixis employee.

8.5 Taxes.

(a) Taxes on Income. Except as otherwise provided in this Section 8.5, each Party shall be solely responsible for the payment of all taxes imposed on its share of income arising directly or indirectly from the activities of the Parties under this Agreement.

(b) Tax Cooperation. The Parties shall cooperate with one another and use reasonable efforts to avoid or reduce tax withholding (e.g., withholding imposed under any U.S. federal, state or local statute, regulation or code) or similar obligations in respect of the milestone payments, royalty payments, and other payments made by Exelixis under this Agreement. To the extent Exelixis is required by Applicable Laws to deduct and withhold taxes on any payment to Adagene, Exelixis shall pay the amounts of such taxes to the proper Governmental Authority in a timely manner and promptly transmit to Adagene an official tax certificate or other evidence of such payment sufficient to enable Adagene to claim such payment of taxes. Adagene shall provide Exelixis any tax forms that may be reasonably necessary in order for Exelixis to not withhold tax or to withhold tax at a reduced rate under an applicable bilateral income tax treaty, to the extent legally able to do so. Adagene shall use reasonable efforts to provide any such tax forms to Exelixis in advance of the due date. Exelixis shall provide Adagene with reasonable assistance to enable the recovery, as permitted by Applicable Laws, of withholding taxes or similar obligations resulting from payments made under this Agreement, such recovery to be for the benefit of Adagene. Exelixis shall have the right to deduct any such tax, levy, or charge actually paid from payment due to Adagene. Each Party agrees to assist the other Party in claiming exemption from such deductions or withholdings under double taxation or similar agreement or treaty from time to time in force and in minimizing the amount required to be so withheld or deducted.

(c) Other Tax Liability. All amounts payable under this Agreement are exclusive of value added or similar taxes (the "VAT"). In the event of value added or similar taxes incurred by a Party with respect to payments made to a Party hereunder or the activities underlying such payments, each Party and its Affiliates will use reasonable efforts to secure available exemption(s) from VAT or cooperate with the other Party's efforts to obtain maximum recovery of VAT paid or incurred by such Party or any Affiliate, to the extent permitted by Applicable Law.

8.6 Records; Audit. Each Party shall keep, and shall require its Affiliates, contractors, and Sublicensees to keep, complete and accurate records pertaining to, in the case of Adagene, the performance of each Compound Research Activity and, in the case of Exelixis, the sale or other

disposition of Products, in each case in sufficient detail to permit the other Party to confirm the accuracy of any payment due hereunder. Each Party will keep such books and records for [***] following the Calendar Year to which they pertain, or such longer period of time as may be required by Applicable Laws. Upon reasonable prior notice and during regular business hours at such place or places where such records are customarily kept, a Party may inspect the other Party's records via an independent certified public accountant (the "Auditor") selected by the auditing Party and reasonably acceptable to the audited Party for the sole purpose of verifying for the auditing Party the accuracy of the financial reports furnished by the audited Party pursuant to this Agreement and of any payments made, or required to be made, to such auditing Party pursuant to this Agreement. Before beginning its audit, the Auditor shall execute an undertaking acceptable to each Party by which the Auditor agrees to keep confidential all information reviewed during the audit. Such audits shall be limited to once each Calendar Year and once with respect to records covering any specific period of time. Such auditor shall not disclose any of the audited Party's Confidential Information to the auditing Party, and shall only report whether the financial reports provided by the audited Party and the amount of payments made to or by a Party are correct or not, and the amount of any discrepancy. In the event that the final result of the inspection reveals an undisputed underpayment or overpayment, the underpaid or overpaid amount shall be settled within [***] after the Auditor's report. The auditing Party shall bear the full cost of such audit unless such audit reveals a discrepancy in the auditing Party's favor of more than [***] from the reported amounts, in which case the audited Party shall reimburse the auditing Party for the reasonable out-of-pocket costs incurred for such audit.

8.7 Late Payments. In the event that any payment due under this Agreement is not paid when due in accordance with the applicable provisions of this Agreement, the payment shall accrue interest from thirty (30) day past the date due at the interest rate of [***]; provided, however, that in no event shall such rate exceed the maximum legal annual interest rate.

9. INTELLECTUAL PROPERTY

9.1 Ownership.

(a) Data. All Data shall be the exclusive property of Exelixis. Adagene hereby assigns to Exelixis all of Adagene's right, title, and interest in and to all Data. To effectuate the foregoing assignment, Adagene shall ensure that all entities and individuals that perform any activities on behalf of Adagene hereunder are under a written or other legally enforceable obligation to assign all right, title, and interest in any intellectual property created pursuant to such activities directly to Adagene or to an entity that is obligated to assign all such right, title, and interest to Adagene.

(b) Compound Inventions. All Inventions other than Platform Inventions, whether or not patentable, that (i) constitute the composition of matter, synthesis or manufacture, use, or formulation of any Compound or (ii) relate to an Exelixis mAB, in each case ((i) and (ii)) that are generated in the performance of any Compound Research Activities or other Development, manufacturing, or regulatory activities under this Agreement solely by one or more employees or agents of a Party or its Affiliates or other persons acting on such Party's behalf, or jointly by one or more employees or agents of each Party or its Affiliates or other persons acting on their behalf, (collectively, the "Compound Inventions") shall be the exclusive property of Exelixis. Adagene

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shall notify and transfer to Exelixis each Compound Invention promptly after its conception, development, or reduction to practice by Adagene or its Affiliate or contractor. Adagene hereby assigns to Exelixis all of Adagene's right, title, and interest in and to all Compound Inventions. To effectuate the foregoing assignment, Adagene shall ensure that all entities and individuals that perform any activities on behalf of Adagene under this Agreement are under a written or other legally enforceable obligation to assign all right, title, and interest in any intellectual property created pursuant to such activities directly to Adagene or to an entity that is obligated to assign all such right, title, and interest to Adagene. Adagene shall assist Exelixis, at Exelixis' expense, in securing for Exelixis any patents, copyrights, or other proprietary rights in such Compound Inventions, and to perform all acts that may be reasonably required to vest in Exelixis all right, title, and interest in such Compound Inventions.

(c) Platform Inventions. All Inventions, whether or not patentable, that (i) pertain generally to the Adagene Platform Technology or any improvements, enhancements or modifications thereto and (ii) do not specifically describe or claim a Compound or a Product and do not use or incorporate and are not based on any Exelixis mAB, Exelixis Technology, or any of Exelixis' Confidential Information, and in each case are generated under this Agreement solely by one or more employees or agents of a Party or its Affiliates or other persons acting on a Party's behalf, or jointly by one or more employees or agents of each Party or its Affiliates or other persons acting on their behalf, (collectively, the "Platform Inventions") shall be the exclusive property of Adagene. All Platform Inventions shall be included within the Adagene Platform Technology. Exelixis hereby assigns to Adagene all of Exelixis' right, title, and interest in and to all Platform Inventions. To effectuate the foregoing assignment, Exelixis shall ensure that all entities and individuals that perform any activities on behalf of Exelixis hereunder are under a written or other legally enforceable obligation to assign all right, title, and interest in any intellectual property created pursuant to such activities directly to Exelixis or to an entity that is obligated to assign all such right, title, and interest to Exelixis. Exelixis shall assist Adagene, at Adagene's expense, in securing for Adagene any patents, copyrights, or other proprietary rights in such Platform Inventions, and to perform all acts that may be reasonably required to vest in Adagene all right, title, and interest in such Platform Inventions.

(d) Other Inventions. Except for Compound Inventions and Platform Inventions, each Party shall solely own any discovery or invention made solely by it and its Affiliates' employees, agents, or independent contractors (the "Sole Inventions"), and the Parties shall jointly own any discovery or invention made jointly by employees, agents, or independent contractors of one Party or its Affiliates together with employees, agents, or independent contractors of the other Party or its Affiliates (the "Joint Inventions"). All Patents claiming patentable Joint Inventions shall be referred to herein as "Joint Patents". Except to the extent either Party is restricted by the licenses granted to the other Party under this Agreement, each Party shall be entitled to practice, license, assign, and otherwise exploit its interest under the Joint Inventions and Joint Patents without the duty of accounting or seeking consent from the other Party.

9.2 Patent Prosecution and Maintenance.

(a) Exelixis Patents. Exelixis shall have the sole right, but not the obligation, to control the preparation, filing, prosecution, and maintenance (including any interferences,

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reissue proceedings, reexaminations, inter partes review, patent term extensions, applications for supplementary protection certificates, oppositions, invalidation proceedings, and defense of validity or enforceability challenges) of the Exelixis Patents (including Product Patents, but excluding Joint Patents) worldwide, using counsel of its own choice.

(b) Adagene Patents and Adagene Platform Patents. Adagene shall have the sole right, but not the obligation, to control the preparation, filing, prosecution, and maintenance (including any interferences, reissue proceedings, reexaminations, patent term extensions, applications for supplementary protection certificates, oppositions, invalidation proceedings and defense of validity or enforceability challenges) of all Adagene Patents (other than Joint Patents) and Adagene Platform Patents worldwide, at its sole cost and expense and by counsel of its own choice. Adagene shall keep Exelixis reasonably informed of the status of filing, prosecution, maintenance, and defense, if any, of the Adagene Patents that are listed in Exhibit B (but not the Adagene Platform Patents), and Adagene shall consult with, and consider in good faith the requests and suggestions of, Exelixis with respect to strategies for filing, prosecuting, and defending the Adagene Patents listed in Exhibit B (but not the Adagene Platform Patents).

(c) Joint Patents.

(i) Subject to this Section 9.2(c), Exelixis shall have the first right, but not the obligation, to prepare, file, prosecute, and maintain (including any interferences, reissue proceedings, reexaminations, patent term extensions, applications for supplementary protection certificates, oppositions, invalidation proceedings and defense of validity or enforceability challenges) the Joint Patents using a patent counsel selected by Exelixis and reasonably acceptable to Adagene. Adagene shall reimburse Exelixis for [***] of the costs and expenses incurred with respect to the preparation, filing, prosecution, and maintenance of the Joint Patents in the Territory, within [***] after receipt of an invoice for such costs and expenses together with reasonable documentation evidencing such costs and expenses provided by Exelixis. In the event that Adagene does not reimburse Exelixis for such costs and expenses for any Joint Patent or notifies Exelixis in writing that it elects to cease reimbursing Exelixis for such costs and expense for any Joint Patent, Adagene shall execute such documents and perform such acts, at Adagene's expense, as may be reasonably necessary to effect an assignment of Adagene's entire right, title, and interest in and to such Joint Patent to Exelixis, and such Patent shall cease to be a Joint Patent and shall no longer be subject to the licenses and other rights granted by Exelixis to Adagene under this Agreement. Exelixis shall keep Adagene informed of material progress with regard to the preparation, filing, prosecution, maintenance, and defense, if any, of the Joint Patents, including content, timing, and jurisdiction of the filing of such Joint Patents, and Exelixis shall consult with, and consider in good faith the reasonable comments of Adagene with respect to filing, prosecuting, and defending the Joint Patents in the Territory.

(ii) If Exelixis desires to abandon or cease prosecution or maintenance of any Joint Patent in the Territory, Exelixis shall provide reasonable prior written notice to Adagene of such intention to abandon (which notice shall, to the extent possible, be given no later than [***] prior to the next deadline for any action that must be taken with respect to any such Joint Patent in the relevant patent office). In such case, at Adagene's sole discretion, upon written notice from Adagene to Exelixis, Adagene may elect to continue prosecution or maintenance of any such Joint Patent at its own expense, and Exelixis shall execute such documents and perform

such acts, at Adagene's expense, as may be reasonably necessary to assign to Adagene Exelixis' ownership interest therein and to allow Adagene to continue the prosecution and maintenance of such Joint Patent in the Territory. Any patents so assigned shall cease to be a Joint Patent and shall thereafter be Adagene Patents subject to the licenses herein.

(d) Patent Term Extensions. Exelixis shall have the right to apply, in its sole discretion and at its expense, for any patent term extension for any Exelixis Patent or Joint Patent in the Territory. Adagene shall have the right to apply, in its sole discretion and at its expense, for any patent term extension for all Adagene Patents (other than Joint Patents) and Adagene Platform Patents in the Territory to the extent that such Patent Term Extensions are not based on Products approval.

(e) Cooperation. Each Party agrees to cooperate fully in the preparation, filing, prosecution, maintenance, and defense, if any, of Patents under Section 9.2 and in the obtaining and maintenance of any patent term extensions, supplementary protection certificates, and their equivalent with respect thereto, at its own cost (except as expressly set forth otherwise in this Section 9.2). Such cooperation includes (i) executing all papers and instruments, or requiring its employees or contractors, to execute such papers and instruments, so as to enable the other Party to apply for and to prosecute patent applications in any country as permitted by Section 9.2; and (ii) promptly informing the other Party of any matters coming to such Party's attention that may affect the preparation, filing, prosecution, or maintenance of any such patent application and the obtaining of any patent term extensions, supplementary protection certificates, and their equivalent.

(f) Separation of Patent Rights. In order to more efficiently enable the prosecution and maintenance of the Product Patents and the Adagene Platform Patents relating to the Inventions described above, the Parties will use good faith efforts to separate Compound Inventions, Platform Inventions, and Joint Inventions into separate filings to the extent possible and without adversely impacting such prosecution and maintenance. For clarity, it is also the mutual intent of the parties that Adagene Patents, if any, shall be separated and prosecuted as either Product Patents or Adagene Platform Patents to the extent possible without adversely impacting the prosecution and maintenance thereof.

9.3 Patent Enforcement.

(a) Notice. Each Party shall notify the other within [***] after becoming aware of any alleged or threatened infringement by a Third Party of any of the Adagene Patents, Adagene Platform Patents, or Product Patents in the Territory, which infringement adversely affects or could reasonably be expected to adversely affect the Development, manufacture, or Commercialization of any Product (collectively, "**Product Infringement**").

(b) Enforcement Right. Exelixis shall have the first right to bring and control any legal action in connection with such Product Infringement at its own expense as it reasonably determines appropriate; provided, however, that with respect to any action involving any Adagene Patent or Adagene Platform Patent, Exelixis shall only have the right to enforce such Adagene Patent or Adagene Platform Patent, as applicable, to the extent such Patent directly and exclusively

relates to a Compound or Product (and not to any compound or product that is not a Compound or Product). Exelixis shall not compromise or settle any such litigation in a way that could reasonably be expected to adversely affect the validity or enforceability of any Adagene Patent or Adagene Platform Patent or result in an admission of liability on the part of Adagene, without Adagene's prior written consent, such consent not to be unreasonably withheld. If Exelixis (i) elects to not bring such legal action against a Product Infringement (the decision of which Exelixis shall inform Adagene promptly) or (ii) Exelixis otherwise fails to bring such legal action against a Product Infringement within [***] after first becoming aware of such Product Infringement, then, to the extent such Product Infringement pertains solely to one or more Adagene Patents or Adagene Platform Patents and not to any Product Patents, Adagene shall have the right to bring and control any legal action in connection with such Product Infringement at its own expense as it reasonably determines appropriate after consultation with Exelixis; provided, however, that Adagene shall not have the right to bring any such action to the extent it relates to a Patent claiming the composition of matter or method of use of any Compound or Product without Exelixis' prior written consent, which Exelixis may grant in its sole discretion.

(c) **Collaboration.** Each Party shall provide to the enforcing Party reasonable assistance in such enforcement, at such enforcing Party's request and expense, including to be named in such action if required by Applicable Laws to pursue such action. The enforcing Party shall keep the other Party regularly informed of the status and progress of such enforcement efforts and shall reasonably consider the other Party's comments on any such efforts, including determination of litigation strategy and filing of material papers to the competent court. The non-enforcing Party shall be entitled to separate representation in such matter by counsel of its own choice and at its own expense, but such Party shall at all times cooperate fully with the enforcing Party.

(d) **Expense and Recovery.** Except as otherwise agreed by the Parties in connection with a cost sharing arrangement, any recovery realized as a result of any litigation described in Section 9.3 (whether by way of settlement or otherwise) shall be first allocated to reimburse the Parties for their costs and expenses in making such recovery (which amounts shall be allocated *pro rata* if insufficient to cover the totality of such expenses). Any remainder after such reimbursement is made shall be allocated as follows: [***].

(e) **Other Infringement.** Except for Product Infringement as set forth above, each Party shall have the exclusive right to enforce its own Patent against any infringement anywhere in the world. For clarity, Exelixis shall have the exclusive right to enforce the Exelixis Patents against any infringement in the Territory at its own expense as it determines appropriate.

9.4 Infringement of Third Party Rights. If any Product used or sold by Exelixis, its Affiliates, or Sublicensees becomes the subject of a Third Party's claim or assertion of infringement of intellectual property rights in a jurisdiction within the Territory ("**Third Party Infringement Claim**"), Exelixis shall promptly notify Adagene and the Parties shall promptly meet to consider the claim or assertion and the appropriate course of action and may, if appropriate, agree on and enter into a "common interest agreement" wherein the Parties agree to their shared, mutual interest in the outcome of such potential dispute. Absent any agreement to the contrary, and subject to claims for indemnification under Article 11, each

Party shall defend itself from any such Third Party Infringement at its own cost and expense, provided, however, that the provisions of Section 9.3 shall govern the right of a Party to assert a counterclaim of infringement of any Adagene Patent, Adagene Platform Patent, or Product Patent.

10. REPRESENTATIONS AND WARRANTIES

10.1 Mutual Representations and Warranties. Each Party represents and warrants to the other that, as of the Effective Date:

- (a) it is duly organized and validly existing under the laws of its jurisdiction of incorporation or formation, and has full corporate or other power and authority to enter into this Agreement and to carry out the provisions hereof;
- (b) it is duly authorized to execute and deliver this Agreement and to perform its obligations hereunder, and the person or persons executing this Agreement on its behalf has been duly authorized to do so by all requisite corporate or partnership action;
- (c) this Agreement is legally binding upon it, enforceable in accordance with its terms, and does not conflict with any agreement, instrument, or understanding, oral or written, to which it is a party or by which it may be bound, nor violate any material law or regulation of any court, governmental body, or administrative or other agency having jurisdiction over it; and
- (d) it has the right to grant the licenses granted by it under this Agreement.

10.2 Covenants.

(a) **Employees, Consultants, and Contractors.** Each Party covenants that it has obtained or will obtain written agreements from each of its employees, consultants, and contractors who perform Development activities pursuant to this Agreement, which agreements will obligate such persons to obligations of confidentiality and non-use and to assign (or, in the case of contractor, grant a license under) Inventions in a manner consistent with the provisions of this Agreement.

(b) **Debarment.** Each Party represents, warrants, and covenants to the other Party that it is not debarred or disqualified under the U.S. Federal Food, Drug and Cosmetic Act, as may be amended, or comparable laws in any country or jurisdiction other than the U.S., and it does not, and will not during the Term, employ or use the services of any person who is debarred or disqualified, in connection with activities relating to any Product. In the event that either Party becomes aware of the debarment or disqualification or threatened debarment or disqualification of any person providing services to such Party, including the Party itself or its Affiliates or Sublicensees, that directly or indirectly relate to activities contemplated by this Agreement, such Party shall immediately notify the other Party in writing and such Party shall cease employing, contracting with, or retaining any such person to perform any such services.

(c) **Compliance.** In the performance of its obligations under this Agreement, such Party shall comply and shall cause its and its Affiliates' employees and contractors to comply with all Applicable Laws.

10.3 Additional Adagene Representations and Warranties. Adagene represents and warrants as of the Effective Date:

(a) Exhibit B lists all Adagene Patents and Adagene Platform Patents as of the Effective Date and all such Patents listed on Exhibit B are (i) to Adagene's knowledge, valid and enforceable, (ii) solely and exclusively owned by Adagene, free of any encumbrance, lien, or claim of ownership by any Third Party, and (iii) filed and maintained properly and correctly and all applicable application and maintenance fees have been paid on or before the due date for payment or during any extension thereof.

(b) Each of the Adagene Patents and Adagene Platform Patents properly identifies each and every inventor of the claims thereof as determined in accordance with the laws of the jurisdiction in which such Patent is issued or such application is pending; all such inventors have assigned their entire right, title, and interest in and to such inventions to Adagene and all such assignments have been duly recorded and are enforceable in accordance with Applicable Law; and there are no claims or assertions in writing received by Adagene regarding the inventorship of any Adagene Patent alleging that additional or alternative inventors should be listed.

(c) Adagene has the right to grant all rights and licenses it purports to grant to Exelixis with respect to the Adagene Technology and Adagene Platform Technology under this Agreement, free and clear of any rights therein granted to any Third Party.

(d) All Adagene Technology and Adagene Platform Technology is owned by Adagene or its Affiliates and is not held under license from any Third Party.

(e) Adagene has not granted any liens or security interests on any of the Adagene Technology or Adagene Platform Technology.

(f) Adagene has not as of the Effective Date, granted to any Third Party any right under the Adagene Technology or Adagene Platform Technology that would conflict with the rights granted to Exelixis hereunder.

(g) (i) No claim or action has been brought or, to Adagene's knowledge, threatened in writing, by any Third Party alleging that (A) the Adagene Patents or Adagene Platform Patents are invalid or unenforceable, or (B) the use of the Adagene Know-How or Adagene Platform Technology infringes or misappropriates, or would infringe or misappropriate, any intellectual property right of any Third Party, and (ii) to Adagene's knowledge, no facts or circumstances exist that would reasonably be expected to give rise to any such claims described in subsection (i).

(h) There are no pending or, to Adagene's knowledge, alleged or threatened, (i) inter partes reviews, post-grant reviews, interferences, re-examinations, or oppositions involving the Adagene Patents or Adagene Platform Patents that are in or before any patent authority (or

other governmental authority performing similar functions) or (ii) inventorship challenges involving the Adagene Patents or Adagene Platform Patents that are in or before any patent or other Governmental Authority.

(i) To Adagene's knowledge, no Third Party is infringing or misappropriating or has infringed or misappropriated or is threatening to infringe or misappropriate the Adagene Technology or the Adagene Platform Technology.

(j) The inventions claimed by the Adagene Patents and the Adagene Platform Patents (i) were not conceived, discovered, developed, or otherwise made in connection with any research activities funded, in whole or in part, by the federal government of the United States or any agency thereof and (ii) are not a "subject invention" as that term is described in 35 U.S.C. Section 201(e), and (iii) are not otherwise subject to the provisions of the Patent and Trademark Law Amendments Act of 1980, as amended, codified at 35 U.S.C. §§ 200-212, as amended, as well as any regulations promulgated pursuant thereto, including in 37 C.F.R. part 401.

(k) To Adagene's knowledge, there are no issues or information related to the Adagene Technology or Adagene Platform Technology, which in Adagene's reasonable opinion, are reasonably likely to have a material impact on the Development or Commercialization of the Compounds (including any infringement of the Adagene Patents or the Adagene Platform Patents of Third Party patent rights) that have not been fully disclosed to Exelixis in the course of Exelixis' due diligence.

10.4 Adagene Covenants. Adagene covenants to Exelixis:

(a) Adagene will not during the Term grant to any Third Party any right under the Adagene Technology or Adagene Platform Technology that would conflict with the rights granted to Exelixis hereunder.

(a) Adagene shall perform the Compound Research Activities allocated to it under each Target Program Plan in accordance with the same level of care and diligence practiced by recognized firms in performing activities of a similar nature and the Compound Research Activities it performs will conform to the requirements and other terms in the applicable Target Program Plan and this Agreement.

10.5 Disclaimer. EXCEPT FOR THE EXPRESS WARRANTIES SET FORTH HEREIN, EACH PARTY EXPRESSLY DISCLAIMS ANY AND ALL WARRANTIES OF ANY KIND, WHETHER EXPRESS, IMPLIED, OR STATUTORY, INCLUDING WARRANTIES OF DESIGN, MERCHANTABILITY, FITNESS FOR A PARTICULAR PURPOSE, NONINFRINGEMENT OF THE INTELLECTUAL PROPERTY RIGHTS OF THIRD PARTIES, OR ARISING FROM A COURSE OF DEALING, USAGE, OR TRADE PRACTICES, IN ALL CASES WITH RESPECT THERETO. NEITHER PARTY MAKES ANY REPRESENTATION OR WARRANTY, EITHER EXPRESS OR IMPLIED, THAT ANY OF THE DEVELOPMENT, MANUFACTURING, OR COMMERCIALIZATION EFFORTS WITH REGARD TO ANY PRODUCT WILL BE SUCCESSFUL.

11. INDEMNIFICATION

11.1 Indemnification by Exelixis. Exelixis hereby agrees to defend, indemnify, and hold harmless Adagene and its Affiliates and their respective directors, officers, employees, and agents (each, an “**Adagene Indemnatee**”) from and against any and all liabilities, expenses, and losses, including reasonable legal expenses and attorneys’ fees (collectively, “**Losses**”), to which any Adagene Indemnatee may become subject as a result of any claim, demand, action, or other proceeding (each, a “**Claim**”) by any Third Party to the extent such Losses arise out of:

- (a) the Development, use, manufacture, or Commercialization of any Compound or Product by Exelixis or its Affiliates or Sublicensees or the contractors of any of them (excluding any activities by or on behalf of Adagene or its Affiliates);
- (b) the negligence or willful misconduct of any Exelixis Indemnatee; or
- (c) the breach by Exelixis of any warranty, representation, covenant, or agreement made by Exelixis in this Agreement;

except, in each case (a)-(c), to the extent such Losses arise out of any activities set forth in Sections 11.1(a)-(d) for which Adagene is obligated to indemnify any Exelixis Indemnatee under Section 11.2.

11.2 Indemnification by Adagene. Adagene hereby agrees to defend, indemnify, and hold harmless Exelixis, its Affiliates, and Sublicensees and their respective directors, officers, employees, and agents (each, an “**Exelixis Indemnatee**”) from and against any and all Losses to which any Exelixis Indemnatee may become subject as a result of any Claim by any Third Party to the extent such Losses arise out of:

- (a) the Development, use, or manufacture of any Compound or Product by Adagene, its Affiliates, or licensees (other than Exelixis) or the contractor of any of them;
- (b) the negligence or willful misconduct of any Adagene Indemnatee; or
- (c) the infringement or misappropriation by Adagene or any of its Affiliates of any Third Party intellectual property rights resulting from the use or practice of any Adagene Technology or the Adagene Platform Technology, except to the extent such infringement or misappropriation is caused directly by the use of an Exelixis mAB, including the Collaboration Target to which such Exelixis mAB is directed; or
- (d) the breach by Adagene of any warranty, representation, covenant, or agreement made by Adagene in this Agreement;

except, in the case of (a), (b), or (d), to the extent such Losses arise out of any activities set forth in Sections 11.1(a)-(c) for which Exelixis is obligated to indemnify any Adagene Indemnatee under Section 11.1.

11.3 Indemnification Conditions and Procedure.

(a) **Notice.** The Party claiming indemnity under this Article 11 (the “**Indemnified Party**”) shall give written notice to the Party from whom indemnity is being sought (the “**Indemnifying Party**”) promptly after learning of such Claim.

(b) **Control.** The Indemnified Party shall provide the Indemnifying Party with reasonable assistance, at the Indemnifying Party’s expense, in connection with the defense of the Claim for which indemnity is being sought. The Indemnified Party may participate in and monitor such defense with counsel of its own choice at its own expense; provided, however, that the Indemnifying Party shall have the right to assume and conduct the defense of the Claim with counsel of its choice.

(c) **Settlement Rights.** The Indemnifying Party shall not settle any Claim without the prior written consent of the Indemnified Party, not to be unreasonably withheld, conditioned, or delayed, unless the settlement involves only the payment of money, no admission of wrong-doing or fault by the Indemnified Party, and no restriction on the future actions or activities of the Indemnified Party. So long as the Indemnifying Party is actively defending the Claim in good faith, the Indemnified Party shall not settle such Claim without the prior written consent of the Indemnifying Party.

(d) **Backup Right.** If the Indemnifying Party does not assume and conduct the defense of the Claim as provided above, (i) the Indemnified Party may defend against and consent to the entry of any judgment, or enter into any settlement with respect to, the Claim in any manner the Indemnified Party may deem reasonably appropriate (and the Indemnified Party need not consult with, or obtain any consent from, the Indemnifying Party in connection therewith so long as the settlement involves only the payment of money, no admission of wrong-doing or fault by the Indemnifying Party, and no restriction on the future actions or activities of the Indemnifying Party), and (ii) the Indemnifying Party will remain responsible to indemnify the Indemnified Party as provided in this Article 11.

11.4 Insurance. During the Term and for a commercially reasonable period of time thereafter, each Party, at its own expense, shall maintain commercial general liability insurance, product liability and other appropriate insurance (or, in the case of Exelixis, self-insure) in an amount consistent with sound business practice and reasonable in light of its obligations under this Agreement. Without limiting the generality of the foregoing, Adagene represents, warrants, and covenants that it possesses and shall maintain during the Term and for [***] thereafter, at its own expense, comprehensive general liability insurance [***]. Each Party shall provide a certificate of insurance (or evidence of self-insurance) evidencing such coverage to the other Party upon request.

11.5 Limitation of Liability. EXCEPT FOR DAMAGES THAT (A) ARISE IN CONNECTION WITH A PARTY’S (I) WILLFUL MISCONDUCT OR FRAUD OR (II) BREACH OF ITS OBLIGATIONS UNDER ARTICLE 12, OR (B) ARE SUBJECT TO INDEMNIFICATION UNDER SECTION 11.1 OR 11.2, NEITHER PARTY SHALL BE LIABLE IN CONTRACT, TORT, NEGLIGENCE, BREACH OF STATUTORY DUTY, OR OTHERWISE FOR ANY SPECIAL, PUNITIVE, OR CONSEQUENTIAL DAMAGES OR FOR LOSS OF PROFITS SUFFERED BY THE OTHER PARTY (OR ITS AFFILIATES OR

(SUB)LICENSEES), REGARDLESS OF ANY NOTICE OF THE POSSIBILITY OF SUCH DAMAGES.

12. CONFIDENTIALITY

12.1 Confidential Information. Except to the extent expressly authorized by this Agreement or otherwise agreed in writing by the Parties, the Parties agree that, during the Term and for [***] thereafter, the receiving Party shall keep confidential and shall not publish or otherwise disclose, and shall not use for any purpose other than as expressly provided for in this Agreement, any Confidential Information of the other Party, and both Parties shall keep confidential and shall not publish or otherwise disclose the terms of this Agreement except as permitted herein. Each Party may use the other Party's Confidential Information only to the extent required to accomplish the purposes of this Agreement, including exercising its rights and performing its obligations under this Agreement. Each Party will use at least the same standard of care as it uses to protect proprietary or confidential information of its own (but no less than reasonable care) to ensure that its employees, agents, consultants, contractors, and other representatives do not disclose or make any unauthorized use of the other Party's Confidential Information. Each Party will promptly notify the other upon discovery of any loss or unauthorized use or disclosure of the other Party's Confidential Information.

12.2 Exceptions. The obligations of confidentiality and restriction on use under Section 12.1 will not apply to any information that the receiving Party can prove by competent written evidence: (a) is now, or hereafter becomes, through no act or failure to act on the part of the receiving Party, generally known or available to the public; (b) is known by the receiving Party at the time of receiving such information, other than by previous disclosure of the disclosing Party, or its Affiliates, employees, agents, consultants, or contractors; (c) is hereafter furnished to the receiving Party without restriction by a Third Party who has no obligation of confidentiality or limitations on use with respect thereto, as a matter of right; or (d) is independently discovered or developed by the receiving Party without the use of the disclosing Party's Confidential Information.

12.3 Authorized Disclosure. Each Party may disclose Confidential Information belonging to the other Party as expressly permitted by this Agreement or if and to the extent such disclosure is reasonably necessary in the following instances:

12.4 filing, prosecuting, or maintaining Patents as permitted by this Agreement;

12.5 filing Regulatory Filings for Products that such Party has a license or right to Develop, manufacture, and Commercialize hereunder in a given country or jurisdiction;

12.6 prosecuting or defending litigation as permitted by this Agreement;

(a) complying with Applicable Laws or regulations (including regulations promulgated by securities exchanges, subject to Section 12.5) or court or administrative orders;

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(b) disclosure to actual and bona fide potential investors, acquirors, licensees, and other financial or commercial partners solely for the purpose of evaluating or carrying out an actual or potential investment, acquisition, or collaboration, in each case under a written agreement containing obligations of confidentiality and non-use at least as stringent as those herein; and

(c) disclosure to its and its Affiliates' employees, consultants, contractors, agents, licensees, and sublicensees, in each case on a need-to-know basis in connection with the Development, manufacture, or Commercialization of the Products in accordance with the terms of this Agreement, in each case under a written agreement containing obligations of confidentiality and non-use at least as stringent as those herein.

Notwithstanding the foregoing, the Parties shall use good faith efforts to agree on a redacted version of this Agreement within [***] following the Effective Date, it being understood that no consent shall be required for a disclosure of such agreed redacted version pursuant to Section 12.3(b) or 12.3(c).

Notwithstanding the foregoing, in the event that a Party is required to make a disclosure of the other Party's Confidential Information pursuant to Section 12.3(a) or 12.3(b), it will, except where impracticable, give reasonable advance notice to the other Party of such disclosure and use efforts to secure confidential treatment of such Confidential Information at least as diligent as such Party would use to protect its own confidential information, but in no event less than reasonable efforts. In any event, the Parties agree to take all reasonable action to avoid disclosure of Confidential Information hereunder. Any information disclosed pursuant to Section 12.3 shall remain Confidential Information and subject to the restrictions set forth in this Agreement, including the foregoing provisions of this Article 12.

Without limiting a Party's obligations under this Article 12, a Party shall be liable for any damages incurred by the other Party to the extent arising from any disclosure by such Party of the terms of this Agreement to any Third Party in breach of this Article 12. Each Party acknowledges that money damages alone may not adequately compensate the other Party in the event of a breach or threatened breach of this paragraph or the other provisions of this Article 12, and that, in addition to all other remedies available to the other Party at law or in equity, the other Party shall be entitled to seek injunctive relief for the enforcement of its rights, including requiring such Party to enforce the confidentiality and non-use provisions of any agreement with a Third Party in accordance with the terms of this Agreement. For the avoidance of doubt, the provisions of this paragraph are in addition to the other provisions contained in this Article 12.

12.7 Publications.

(a) **Regarding Adagene Platform Technology.** Adagene shall have the sole right, in its sole discretion, to publish or otherwise disclose any information regarding the Adagene Platform Technology or any improvements thereto; provided that in no event shall Adagene disclose any Confidential Information of Exelixis in violation of Section 12.1.

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(b) **Collaboration Targets.** Exelixis shall have the sole right, in its sole discretion, to publish or otherwise disclose the results of or other information regarding any and all research and Development activities performed hereunder with respect to the Products, including all Compound Research Activities for each Collaboration Target; provided that in no event shall Exelixis disclose any Confidential Information of Adagene in violation of Section 12.1.

12.8 Publicity; Public Disclosures.

(a) **Initial Press Release.** The Parties agree to issue a joint press release substantially in a form agreed by the Parties and attached to this Agreement as Exhibit C announcing the signature of this Agreement at or shortly after the Effective Date within the time-period required by applicable securities laws.

(b) **Subsequent Releases.** If a Party wishes to release any subsequent press release or other public statement or announcement about the terms of this Agreement, the Parties' relationship under this Agreement, performance of this Agreement, or activities conducted under this Agreement (each, a "**Release**"), such Party shall first obtain the other Party's written approval of the proposed Release. The reviewing Party will have [***], or such shorter period of time as agreed by the Parties or necessary for the releasing Party to comply with Applicable Law, to review and provide comments to such Release and the reviewing Party's consent of such approval shall not be unreasonably withheld to the extent such information has been previously disclosed. Notwithstanding the foregoing, a Party need not obtain the other Party's approval for a Release to the extent such Release (a) discusses a Product which such Party has the right to Commercialize under this Agreement and such Release does not mention the other Party hereto or the relationship of the Parties under this Agreement, or (b) discloses text previously approved in any filing with the SEC or in an offering circular for an unregistered securities offering, but only to the extent the underlying facts disclosed in such previously approved text are still true and accurate, and where the circumstances surrounding such disclosure have not changed.

(c) **SEC Disclosures.** With respect to complying with disclosure requirements of the SEC or other stock exchange on which a Party's securities are publicly traded (or to which an application for listing has been submitted) ("**Exchange**") in connection with any required filing of this Agreement, the filing Party will consult with the other Party on the provisions of this Agreement to be redacted in any filings made by such Party with the SEC or other Exchange; provided that each Party shall have the right to make any such filing as it reasonably determines necessary under Applicable Laws and will not rely on any statements made by the other Party related to securities laws or regulations. If Adagene reasonably concludes that it must make a Release, or that any portion of this Agreement must be disclosed, in each case pursuant to the requirements of the SEC or other Exchange, and Exelixis would prefer that such Release not be made, that the information so disclosed within such Release be modified or limited, or that disclosure of the Agreement be limited, then Adagene shall modify or limit such Release to address Exelixis' concerns, including by incorporating statements suggested by Exelixis (to the extent any such statement is true and accurate) and seeking confidential treatment of the information included in such Release; provided that Adagene may provide to Exelixis a written statement of why any particular modification or limitation proposed by Exelixis would be contrary to requirements of Applicable Law or the rules of the SEC or applicable Exchange or are otherwise necessary for

appropriate market disclosure; provided further that any statement proposed by Exelixis which does reduce the information disclosed in such Release shall be included by Adagene in such Release (to the extent such statement is true and accurate).

12.9 Equitable Relief. Given the nature of the Confidential Information and the competitive damage that a Party would suffer upon unauthorized disclosure, use, or transfer of its Confidential Information to any Third Party, the Parties agree that monetary damages may not be a sufficient remedy for any breach of this Article 12. In addition to all other remedies, a Party shall be entitled to seek specific performance and injunctive and other equitable relief as a remedy for any breach or threatened breach of this Article 12.

13. TERM AND TERMINATION

13.1 Term. This Agreement shall commence on the Effective Date and, unless terminated earlier as provided in this Article 13 or by mutual written agreement of the Parties, shall continue in effect, on a Product-by-Product and country-by-country basis, until the expiration of the Royalty Term for such Product in such country (the “**Term**”). Upon the expiration of the Royalty Term for a particular Product in a particular country, the license granted to Exelixis in Section 2.1 with respect to such Product in such country shall be deemed fully-paid, royalty-free, perpetual, and irrevocable.

13.2 Termination for Cause.

(a) Material Breach. Each Party shall have the right to terminate this Agreement immediately in its entirety upon written notice to the other Party if such other Party materially breaches this Agreement, such material breach to include a failure of Exelixis to comply with its diligence obligations under this Agreement and Adagene’s failure to comply with its exclusivity obligations hereunder, and has not cured such breach to the reasonable satisfaction of the other Party within [***] ([***] with respect to any payment breach) after notice of such breach from the non-breaching Party. If the breach is capable of being cured, but cure of such breach (other than non-payment) cannot reasonably be effected within such [***] period, the breaching Party shall promptly notify the non-breaching Party thereof, and the Parties shall agree on a plan to cure such breach promptly, but in any event within an additional [***]. So long as the breaching Party is diligently carrying out such plan, the non-breaching Party shall not have the right to terminate this Agreement. If the breaching Party fails to diligently carry out such plan and cure such breach within the additional cure period as provided above, then the non-breaching Party may terminate this Agreement upon written notice to the breaching Party; provided that if such material breach relates solely to one Collaboration Target under the Agreement (but not to both Collaboration Targets), then the terminating Party shall have the right to terminate this Agreement pursuant to this Section 13.2(a) solely with respect to the Collaboration Target as to which such material breach relates.

(b) Disputed Breach. If the alleged breaching Party disputes in good faith the existence or materiality of a breach specified in a notice provided by the other Party, and such alleged breaching Party provides the other Party notice of such dispute within [***], then the other Party shall not have the right to terminate this Agreement under Section 13.2(a) unless and until

a panel of arbitrators, in accordance with Article 14, has determined that the alleged breaching Party has materially breached the Agreement and, if the breach is then curable, such Party fails to cure such breach within the applicable cure period set forth above following such decision.

(c) **Bankruptcy.** Each Party shall have the right to terminate this Agreement immediately in its entirety upon written notice to the other Party if such other Party makes a general assignment for the benefit of creditors, files an insolvency petition in bankruptcy, petitions for or acquiesces in the appointment of any receiver, trustee, or similar officer to liquidate or conserve its business or any substantial part of its assets, commences under the laws of any jurisdiction any proceeding involving its insolvency, bankruptcy, reorganization, adjustment of debt, dissolution, liquidation, or any other similar proceeding for the release of financially distressed debtors or becomes a party to any proceeding or action of the type described above and such proceeding is not dismissed within [***] after the commencement thereof.

(d) **Safety Reasons.** Exelixis shall have the right to terminate this Agreement on a Collaboration Target-by-Collaboration Target basis immediately upon written notice to Adagene if Exelixis reasonably determines, based upon additional information that becomes available or an analysis of the existing information at any time, that the medical risk/benefit of a Product directed to the applicable Collaboration Target is so unfavorable that it would be incompatible with the welfare of patients to Develop or Commercialize or to continue to Develop or Commercialize such Product.

13.3 Termination without Cause by Exelixis. Exelixis shall have the right to terminate this Agreement in its entirety or on Collaboration Target-by-Collaboration Target basis at any time, for any or no reason, upon [***] written notice to Adagene.

13.4 Effects of Termination. Upon any termination of this Agreement in its entirety by either Party, the following terms will apply. Upon any termination of this Agreement with respect to a particular Collaboration Target, the following terms will apply solely with respect to such terminated Collaboration Target. (For clarity, during the pendency of any dispute regarding material breach and/or any termination notice period, all of the terms and conditions of this Agreement shall remain in effect and the Parties shall continue to perform all of their respective obligations hereunder.)

(a) **Licenses.** All licenses granted by either Party will automatically terminate.

(b) **Return of Exelixis Property.** Adagene shall return to Exelixis all of Exelixis' Confidential Information and shall deliver to Exelixis, or destroy at Exelixis' request, the Compound Inventions (in each case to the extent not already provided) and any other deliverables specified in the Target Program Plan (in whatever stage of development or completion).

(c) **Non-use Covenant for Masks.** Exelixis shall not, and shall procure that its Affiliates and Sublicensees will not, use, commercialize or otherwise exploit in any way, assign, transfer, license or otherwise grant any rights to any Third Party under the Masks Developed and assigned by Adagene to Exelixis under this Agreement. Provided, however, that in the event that Exelixis, its Affiliates and/or its Sublicensees do use, commercialize or otherwise exploit such

Masks, such use or commercialization shall be subject to all of the milestone and royalty payment terms and all of the associated reporting terms hereunder.

(d) License to Adagene. Upon Adagene's written request to Exelixis within thirty (30) days following the effective date of termination of a Collaboration Target by Exelixis pursuant to Section 13.3, and subject to Exelixis' agreement to such request in its sole discretion, the Parties shall negotiate in good faith, for a period of [***] (or such longer period agreed by the Parties), a non-exclusive or exclusive royalty-bearing license under the Exelixis Technology for such terminated Collaboration Target to research, Develop, use, make, have made, sell, offer for sale, import, and otherwise Commercialize the Compounds and Products directed to such Collaboration Target, on commercially reasonable terms taking into account the then current stage of the Product(s) for the applicable terminated Collaboration Target and the license rights requested by Adagene. Following the expiration of such [***] period (or such longer period agreed by the Parties), Exelixis shall have no further obligation to Adagene under this Section.

13.5 Right to Maintain License. Notwithstanding the foregoing, if Exelixis would otherwise have the right to terminate this Agreement pursuant to Section 13.2(a) (Material Breach), in lieu of such termination Exelixis may elect to continue this Agreement in full force and effect, except that, upon written notice to Adagene, Exelixis' diligence obligations under Sections 4.6 and 5.2 shall be deemed fully satisfied, and any amounts that would have been due to Adagene by Exelixis accruing after such termination with respect to any Product under Sections 7.3, 7.4, and 7.5 shall each be reduced by [***]. Notwithstanding the above, if Adagene disputes in good faith the existence or materiality of such breach and provides Exelixis notice of such dispute within [***] after receipt of Exelixis' notice, then the preceding subclauses (a) and (b) will not apply unless and until a panel of arbitrators, in accordance with Article 14, has determined that Adagene has committed such breach failed to cure such breach within the cure period. For clarity, during the pendency of any dispute regarding material breach and/or any cure period, all of the terms and conditions of this Agreement shall remain in effect and the Parties shall continue to perform all of their respective obligations hereunder.

13.6 Confidential Information. Upon expiration or termination of this Agreement in its entirety, each Party shall promptly return to the other Party, or delete or destroy, all relevant records and materials in such Party's possession or control containing Confidential Information of the other Party; provided that a Party may keep one copy of such materials for legal archival purposes subject to continuing confidentiality obligations.

13.7 Rights in Bankruptcy. All rights and licenses granted under or pursuant to this Agreement by one Party to the other Party are, and otherwise will be deemed to be, for purposes of Section 365(n) of the U.S. Bankruptcy Code or comparable provision of applicable bankruptcy or insolvency laws, licenses of right to "intellectual property" as defined under Section 101 of the U.S. Bankruptcy Code or comparable provision of applicable bankruptcy or insolvency laws. The Parties agree that a Party that is a licensee of such rights under this Agreement will retain and may fully exercise all of its rights and elections under the U.S. Bankruptcy Code or comparable provision of applicable bankruptcy or insolvency laws. The Parties further agree that, in the event of the commencement of a bankruptcy proceeding by or against a Party to this Agreement under the U.S. Bankruptcy Code or comparable provision of applicable bankruptcy or insolvency laws,

the other Party will be entitled to a complete duplicate of (or complete access to, as appropriate) any such intellectual property and all embodiments of such intellectual property, and same, if not already in its possession, will be promptly delivered to it (a) upon any such commencement of a bankruptcy or insolvency proceeding upon its written request therefor, unless the bankrupt Party elects to continue to perform all of its obligations under this Agreement, or (b) if not delivered under (a) above, following the rejection of this Agreement by or on behalf of the bankrupt Party upon written request therefor by the other Party.

13.8 Survival. Expiration or termination of this Agreement shall not relieve the Parties of any obligation or right accruing prior to such expiration or termination. Except as set forth below or elsewhere in this Agreement, the obligations and rights of the Parties under the following provisions of this Agreement shall survive expiration or termination of this Agreement: Article 1 (as applicable), Section 2.2 (with respect to liability for breaches by Sublicensees solely to the extent such breach occurred prior to the expiration or termination of this Agreement), Section 2.5, Sections 7.3, 7.4, 7.5, 8.1, 8.2, and 8.7 (solely in connection with the payment of amounts accrued by not paid as of the termination or expiration of this Agreement, and as applicable under Section 13.4(c)), Section 8.6, Section 9.1, Article 11 (solely to the extent the applicable indemnification claim occurred prior to the expiration or termination of this Agreement), Article 12, Section 13.1, Section 13.4, Section 13.6, Section 13.7, Section 13.8, Article 14, and Article 15 (as applicable).

14. DISPUTE RESOLUTION

14.1 Objective. It is the objective of the Parties to establish procedures to facilitate the resolution of any disputes arising under or related to this Agreement in an expedient manner by mutual cooperation and without resort to litigation. To accomplish this objective, the Parties agree to follow the procedures set forth in this Article 14 to resolve any such dispute if and when it arises.

14.2 Executive Negotiations. The Parties will try to settle any dispute, controversy, or claim that arises out of, or relates to, any provision of the Agreement (“**Disputed Matter**”) by first referring the Disputed Matter to the Parties’ Executive Officers. Either Party may initiate such informal dispute resolution by sending written notice of the Disputed Matter to the other Party, and, within [***] after such notice, the Executive Officers (or their respective designees having the authority to settle such Disputed Matter) of the Parties will meet for attempted resolution by good faith negotiations. If the Executive Officers (or their respective designees) are unable to resolve such dispute within [***] after their first meeting for such negotiations, either Party may seek to have such dispute resolved in accordance with Section 14.3 below.

14.3 Dispute Resolution.

(a) If the Parties are unable to resolve a Disputed Matter using the process described in Section 14.2, then a Party seeking further resolution of the Disputed Matter (other than an Excluded Claim) shall submit the Disputed Matter to resolution by final and binding arbitration. Whenever a Party determines to institute arbitration proceedings, it will give written notice to that effect to the other Party. The legal seat of arbitration will be San Francisco, California, U.S., and the arbitration will be administered by JAMS pursuant

to its Streamlined Arbitration Rules then in effect (the “**Rules**”), except as otherwise provided herein and applying the substantive law specified in Section 15.1. The arbitration will be conducted by three arbitrators appointed in accordance with the Rules, provided that such arbitrators must have significant business or legal experience in the pharmaceutical business. After conducting any hearing and taking any evidence deemed appropriate for consideration, the arbitrators will render their opinion within thirty (30) days after the final arbitration hearing. The arbitrators will not have the power to award damages excluded pursuant to Section 11.5 under this Agreement and any arbitral award that purports to award such damages is expressly prohibited and void *ab initio*. Decisions of the arbitrators that conform to the terms of this Section 14.3 will be final and binding on the Parties and judgment on the award so rendered may be entered in any court of competent jurisdiction and the Parties undertake to carry out any award without delay. The losing Party, as determined by the arbitrators, will pay all of the JAMS administrative costs and fees of the arbitration and the fees and costs of the arbitrators, and the arbitrators will be directed to provide for payment or reimbursement of such fees and costs by the losing Party. If the arbitrators determine that there is no losing Party, the Parties will each be responsible for one-half of those costs and fees and the arbitrator’s award will so provide. Notwithstanding the foregoing, each Party shall be responsible for its own attorneys’ fees, expert or witness fees, and any other fees and costs, and no such fees or costs will be shifted to the other Party.

(b) Notwithstanding the terms of and procedures set forth in Section 14.2 or 14.3, any applications, motions, or orders to show cause seeking temporary restraining orders, preliminary injunctions or other similar preliminary or temporary legal or equitable relief (“**Injunctive Relief**”) concerning a Disputed Matter (including, but not limited to, Disputed Matters arising out of a potential or actual breach of the confidentiality and non-use provisions in Article 12) may immediately be brought in the first instance and without invocation or exhaustion of the procedures set forth in Section 14.2 or 14.3(a) for hearing and resolution in and by a court of competent jurisdiction. Alternatively, a party seeking Injunctive Relief may immediately institute arbitral proceedings without invocation or exhaustion of the procedures set forth in Section 14.2 or 14.3(a), and any such Injunctive Relief proceedings will be administered by the ICC pursuant to its emergency arbitration procedures then in effect and applying the substantive law specified in Section 15.1. In either event, once the Injunctive Relief proceedings have been conducted and a decision rendered thereon by the court or arbitral forum, the Parties will, if the Disputed Matter is not finally resolved by the Injunctive Relief, proceed to resolve the Disputed Matter in accordance with the terms of Section 14.2 and 14.3(a). This Section shall be specifically enforceable.

(c) Except to the extent necessary to confirm, enforce, or challenge an award of the arbitration, to protect or pursue a legal right, or as otherwise required by Applicable Law, neither Party nor the arbitrator may disclose the existence, content, or results of an arbitration under this Section 14.3 without the express, prior written consent of both Parties. In no event shall an arbitration be initiated after the date when commencement of a legal or equitable proceeding based on the dispute, controversy, or claim would be barred by the

applicable California statute of limitations. Any disputes concerning the propriety of the commencement of the arbitration, or the validity, or application of this Section 14.3 shall be finally settled by the arbitrator.

15. GENERAL PROVISIONS

15.1 Governing Law. This Agreement, and all questions regarding the existence, validity, interpretation, breach, or performance of this Agreement, shall be governed by, and construed and enforced in accordance with, the laws of the State of California, United States, without reference to its conflicts of law principles.

15.2 Entire Agreement; Modification. This Agreement, including the Exhibits, is both a final expression of the Parties' agreement and a complete and exclusive statement with respect to all of its terms. This Agreement supersedes all prior and contemporaneous agreements and communications, whether oral, written, or otherwise, concerning any and all matters contained herein, including the Confidentiality Agreement. This Agreement may only be modified or supplemented in a writing expressly stated for such purpose and signed by an authorized representative of each Party.

15.3 Relationship Between the Parties. The Parties' relationship, as established by this Agreement, is solely that of independent contractors. This Agreement does not create any partnership, joint venture, or similar business relationship between the Parties. Neither Party is a legal representative of the other Party, and neither Party can assume or create any obligation, representation, warranty, or guarantee, express or implied, on behalf of the other Party for any purpose whatsoever.

15.4 Performance by Affiliates. Exelixis may discharge any obligations and exercise any right hereunder through any of its Affiliates. Exelixis hereby guarantees the performance by its Affiliates of its obligations under this Agreement, and shall cause its Affiliates to comply with the provisions of this Agreement in connection with such performance.

15.5 Waiver. The waiver by either Party of any right hereunder or of the failure to perform or of a breach by the other Party shall not be deemed a waiver of any other right hereunder or of any other breach or failure by such other Party whether of a similar nature or otherwise. Any waiver by a Party of a particular term or condition will be effective only if set forth in a written instrument duly executed by or on behalf of the Party waiving such term or condition.

15.6 Assignment. Except as expressly provided hereunder, neither this Agreement nor any rights or obligations hereunder may be assigned or otherwise transferred by either Party without the prior written consent of the other Party (which consent shall not be unreasonably withheld). The Parties may assign or otherwise transfer this Agreement and its rights and obligations hereunder without the other Party's consent as follows:

(a) by either Party in connection with the transfer or sale of all or substantially all of the business or assets relating to this Agreement to a Third Party, whether by merger, consolidation, divestiture, restructure, sale of stock, sale of assets, or otherwise, provided that in the event of any such transaction (whether this Agreement is actually assigned or is assumed by the

acquiring party by operation of law (e.g., in the context of a reverse triangular merger)), intellectual property rights of the acquiring party to such transaction shall not be included in the technology licensed or assigned hereunder unless used by such acquiring party to Develop, use, make, or sell any Compound or Product; or

(b) by either Party, to an Affiliate, provided that if the entity to which this Agreement is assigned ceases to be an Affiliate of the assigning Party, the Agreement shall be automatically assigned back to the assigning Party or its successor.

The rights and obligations of the Parties under this Agreement shall be binding upon and inure to the benefit of the successors and permitted assigns of the Parties specified above, and the name of a Party appearing herein will be deemed to include the name of such Party's successors and permitted assigns to the extent necessary to carry out the intent of this Section 15.6. Any assignment not in accordance with this Section 15.6 shall be null and void and of no legal force or effect.

15.7 Severability. If, for any reason, any part of this Agreement is adjudicated invalid, unenforceable, or illegal by a court of competent jurisdiction, such adjudication shall not affect or impair, in whole or in part, the validity, enforceability, or legality of any remaining portions of this Agreement. The Parties will in such an instance use their best efforts to replace the invalid, unenforceable, or illegal provision(s) with a valid, enforceable, and legal provision(s) that best implements the original intent of the Parties and purposes of this Agreement.

15.8 Notices. Any notice to be given under this Agreement must be in writing and delivered either (a) in person or (b) by (i) air mail (postage prepaid) requiring return receipt, (ii) recognized overnight courier, or (iii) facsimile, in each case to the Party to be notified at its address given below, or at any other address such Party may designate by prior written notice to the other in accordance with this Section 15.8. Notice shall be deemed sufficiently given for all purposes upon the earliest of: (w) if personally delivered, the date of actual receipt; (x) if air mailed, five (5) days after the date of postmark; (y) if delivered by overnight courier, the next day the overnight courier regularly makes deliveries; or (z) if sent by facsimile, the date of confirmation of receipt if during the recipient's normal business hours, otherwise the next business day.

If to Adagene, notices must be addressed to:

Adagene, Inc.
315 Montgomery Street, Suite 900
San Francisco, CA 94104
Attn: General Counsel

If to Exelixis, notices must be addressed to:

Exelixis, Inc.
1851 Harbor Bay Parkway
Alameda, CA 94502
Attention: EVP and General Counsel

15.9 Force Majeure. Each Party shall be excused from liability for the failure or delay in performance of any obligation under this Agreement by reason of any event beyond such Party’s reasonable control, including Acts of God, fire, flood, explosion, earthquake, pandemic, or other natural forces, war, civil unrest, acts of terrorism, accident, destruction, or other casualty, any lack or failure of transportation facilities, any lack or failure of supply of raw materials, or any other event similar to those enumerated above. Such excuse from liability shall be effective to the extent and duration of the event(s) causing the failure or delay in performance and provided that the Party has not caused such event(s) to occur. Notice of a Party’s failure or delay in performance due to force majeure must be given to the other Party as soon as reasonably practicable after its occurrence. All delivery dates under this Agreement that have been affected by force majeure shall be tolled for the duration of such force majeure.

15.10 Interpretation. The headings of clauses contained in this Agreement preceding the text of the sections, subsections, and paragraphs hereof are inserted solely for convenience and ease of reference only and shall not constitute any part of this Agreement, or have any effect on its interpretation or construction. All references in this Agreement to the singular shall include the plural where applicable. Unless otherwise specified, references in this Agreement to any Article shall include all Sections, subsections, and paragraphs in such Article, references to any Section shall include all subsections and paragraphs in such Section, and references in this Agreement to any subsection shall include all paragraphs in such subsection. The word “including” and similar words means including without limitation. The word “or” means “and/or” unless the context dictates otherwise because the subjects of the conjunction are or intended to be mutually exclusive. The words “herein”, “hereof”, and “hereunder” and other words of similar import refer to this Agreement as a whole and not to any particular Section or other subdivision. All references to days in this Agreement mean calendar days, unless otherwise specified. Ambiguities and uncertainties in this Agreement, if any, shall not be interpreted against either Party, irrespective of which Party may be deemed to have caused the ambiguity or uncertainty to exist. This Agreement has been prepared in the English language and the English language shall control its interpretation. In addition, all notices required or permitted to be given hereunder, and all written, electronic, oral, or other communications between the Parties regarding this Agreement shall be in the English language.

15.11 Counterparts; Electronic or Facsimile Signatures. This Agreement may be executed in any number of counterparts, each of which shall be an original, but all of which together shall constitute one instrument. This Agreement may be executed and delivered electronically or by PDF or facsimile and, upon such delivery, such electronic or PDF or facsimile signature will be deemed to have the same effect as if the original signature had been delivered to the other Party.

{SIGNATURE PAGE FOLLOWS}

IN WITNESS WHEREOF, the Parties hereto have caused this **COLLABORATION AND LICENSE AGREEMENT** to be executed and entered into by their duly authorized representatives as of the Effective Date.

EXELIXIS, INC.

ADAGENE INCORPORATED

By: /s/ Michael M. Morrissey

By: /s/ Peter Luo

Name: Michael M. Morrissey

Name: Peter Luo

Title: President & Chief Executive Officer

Title: Chief Executive Officer & Chairman

List of Exhibits:

Exhibit A: Adagene Know-How

Exhibit B: Adagene Patents and Adagene Platform Patents

Exhibit C: Press Release

Exhibit D: VIE Reporting

Exhibit A

Adagene Know-How

Exhibit B

Adagene Patents

Exhibit C

Form of Press Release

Exhibit D

Quarterly VIE assessment

CONSENT OF INDEPENDENT REGISTERED PUBLIC ACCOUNTING FIRM

We hereby consent to the use in this Registration Statement on Form F-1 of Adagene Inc. of our report dated September 22, 2020 relating to the financial statements of Adagene Inc., which appears in this Registration Statement. We also consent to the reference to us under the heading “Experts” in such Registration Statement.

/s/ PricewaterhouseCoopers Zhong Tian LLP

Shanghai, the People’s Republic of China

February 1, 2021
