
REGULATORY OVERVIEW

OVERVIEW OF LAWS AND REGULATIONS IN THE PRC

This section summarizes the principal PRC laws, rules and regulations that are relevant to our business:

Regulatory Authorities

The regulatory authorities of the drug industry in the PRC include: the NMPA, the National Health Commission of the PRC (中華人民共和國國家衛生健康委員會) (the “NHC”) and the National Healthcare Security Administration (國家醫療保障局) (the “NHSA”).

The NMPA is an authority under the State Administration for Market Regulation (國家市場監督管理總局) (the “SAMR”) and is the primary regulator for medical products. It is primarily responsible for supervising and managing drugs, medical devices and cosmetics, including drafting of relevant regulations and policies; undertaking standard management, registration regulation, quality management and post-market risk management for drugs, medical devices and cosmetics; organizing and guiding the supervision and inspection of drugs, medical devices and cosmetics; and undertaking management of qualifications for licensed pharmacists.

The NHC is the primary national regulator for public health. It is primarily responsible for drafting national health policies, supervising and regulating public health, healthcare services and health emergency systems, coordinating the reform of medical and health system, organizing the formulation of national drug policies and national essential medicine system, launching an early warning mechanism for the monitoring of the use and clinical comprehensive evaluation of medicine as well as the drug shortage, giving suggestions on the pricing policy of national essential medicine, and regulating the operation of medical institutions and practicing of medical personnel.

The NHSA is an authority directly under the State Council responsible for the management of the healthcare security system. It is primarily responsible for drafting and implementing policies and standards on medical insurance, maternity insurance and medical assistance; supervising and administering the healthcare security funds; organizing the formulation of a uniform medical insurance catalog and payment standards on drugs, medical disposables and healthcare services; and formulating and supervising the implementation of the bidding and tendering policies for drugs and medical disposables.

Laws and Regulations in Relation to New Drugs

Application for New Drug Registration

Drug registration refers to an approval process where the NMPA conducts review of the safety, efficacy and quality controllability of the drugs intended for marketing according to the application for drug registration made by an applicant, and decides whether to approve the application. Pursuant to the provisions of the Measures for the Administration of Drug Registration (《藥品註冊管理辦法》), promulgated by the SAMR on January 22, 2020 and came into effect on July 1, 2020, the Measures for the Administration of Drug Registration (2020) shall apply to the development, registration, supervision and management activities carried out in the territory of the PRC for marketing of drugs. In accordance with the Measures for the Administration of Drug Registration (2020), drugs registration refers to activities that a drug registration applicant files an application and other supplementary applications for clinical drug trial, approval for drug marketing and reregistration, among others, under the legal procedures and according to the relevant requirements, and that the medical products administrative department examines the safety, effectiveness and quality controllability based on the laws and regulations, and the existing scientific cognitions, to decide whether to agree with the activities applied for. A drug registration certificate shall be valid for five years. During the validity period, a holder of a drug registration certificate shall continue to ensure the safety, effectiveness and quality controllability of the marketed drug, and apply for re-registration of the drug six months prior to the expiry of the validity period.

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Non-clinical Research and Animal Testing

The non-clinical safety assessment of drugs for marketing approval shall be conducted in accordance with the Good Laboratory Practices for Non-clinical Laboratory Studies (《藥物非臨床研究質量管理規範》) promulgated by the State Food and Drug Administration (the “SFDA”) in August 2003 and latest amended by the China Food and Drug Administration (國家食品藥品監督管理總局) (the “CFDA”, revoked on March 18, 2018) in July 2017 and came into effect on September 1, 2017. The SFDA promulgated the Administrative Measures for the Certification of Good Laboratory Practices for Non-clinical Laboratory Studies (《藥物非臨床研究質量管理規範認證管理辦法》) in April 2007, which specifies the requirements for institutions applying for Good Laboratory Practices (GLP) certification of non-clinical laboratory studies. On January 19, 2023, the NMPA amended the Administrative Measures for the Certification of Good Laboratory Practices for Non-clinical Laboratory Studies (《藥物非臨床研究質量管理規範認證管理辦法》), which came into effect on July 1, 2023.

According to the Regulations for the Administration of Affairs Concerning Experimental Animals (《實驗動物管理條例》) promulgated by the State Science and Technology Commission in November 1988 and lastly amended in March 2017 by the State Council, the Administrative Measures on Good Practice of Experimental Animals (《實驗動物質量管理辦法》) jointly promulgated by the State Science and Technology Commission and the State Bureau of Quality and Technical Supervision in December 1997, and the Administrative Measures on the Certificate for Experimental Animals (Trial) (《實驗動物許可證管理辦法(試行)》) promulgated by the Ministry of Science and Technology and other regulatory authorities in December 2001 and came into effect in January 2002, using experimental animals and related products requires a Certificate for Utilization of Laboratory Animals. A Certificate for Utilization of Laboratory Animals shall be valid for five years, and the holder shall apply for renewal six months prior to the expiry of the validity period. A Certificate for Utilization of Laboratory Animals shall be inspected annually by the local Science and Technology Bureau.

Application for Clinical Trial

After completing the pre-clinical studies, the applicant must obtain approval for clinical trials of drugs from the NMPA before the conduction of new clinical drug trials. According to the Decision on Adjusting the Approval Procedures of Certain Administrative Approval Items for Drugs (《關於調整部分藥品行政審批事項審批程序的決定》) promulgated by the CFDA on March 17, 2017 and came into effect on May 1, 2017, the decision on the approval of clinical trials of drugs enacted by the CFDA can be made by the CDE from May 1, 2017. Pursuant to the Drug Administration Law of the PRC (《中華人民共和國藥品管理法》) (the “Drug Administration Law”) promulgated by the Standing Committee of the National People’s Congress (the “SCNPC”) in September 1984 and lastly amended in August 2019 and came into effect in December 2019, the dossier on a new drug research and development, including the manufacturing method, quality specifications, results of pharmacological and toxicological tests and the relevant data, files and samples, shall, in accordance with the regulations of the drug regulatory authority under the State Council be truthfully submitted to the said department for approval before clinical drug trial is conducted.

The Drug Administration Law further stipulates that the drug regulatory authority under the State Council shall decide whether to approve the clinical trial application and notify the decision to the clinical trial applicant within sixty (60) business days from the date of accepting the clinical trial application. If the drug regulatory authority under the State Council fails to do so, the clinical trial application shall be deemed as approved, and if the bioequivalence test is conducted, it is required to report it to the drug regulatory authority under the State Council for filing. According to the Announcement of the Adjustment of Procedures of the Evaluation and Examination for Drug Clinical Trial (《關於調整藥物臨床試驗審評審批程序的公告》), both promulgated by NMPA on

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and effective from July 24, 2018, applicants could proceed with their clinical trials if they have not received any denial or query from the CDE within 60 business days after the application has been accepted and the relevant application fees have been paid.

Before conducting the clinical trial, the applicant shall file a series of detailed documents with the NMPA. According to the Announcement on Drug Clinical Trial Information Platform (《關於藥物臨床試驗信息平台的公告》), which came into effect in September 2013, and the Standard for the Management of Drug Clinical Trial Registration and Information Disclosure (Trial) (《藥物臨床試驗登記與信息公示管理規範(試行)》), which came into effect in July 2020, all clinical trials approved by the CFDA and conducted in the PRC shall complete the clinical trial registration and information disclosure on the drug clinical trial information platform. The applicant must complete the initial registration of the trial within one month after obtaining the approval of the clinical trial to obtain the unique registration number of the trial; and complete the subsequent data registration before the first patient is enrolled and submit it for the first time for disclosure. If the aforementioned disclosure has not been submitted within one year after applicant's obtaining the clinical trial approval, the applicant shall submit an explanation, and if the procedure is not completed within three years, the clinical trial approval shall automatically be annulled.

After obtaining clinical trial approval, the applicant shall choose institutions qualified for clinical trials of the drug to conduct clinical trials. Pursuant to the Administrative Regulations for Drug Clinical Trial Institutions (《藥物臨床試驗機構管理規定》), which came into effect in December 2019, drug development activities and clinical trials of drugs (including bioequivalence test conducted after filing) approved by the NMPA within the territory of the PRC, shall be conducted in the drug clinical trial institutions. Drug clinical trial institutions shall be subject to filing administration. Institutions that only engage in analysis of biological samples related to drug clinical trials shall not be subject to filing. The national drug regulatory authority is responsible for setting up a filing management information platform for the registration, filing and operation management of such institutions, as well as for the entry, sharing and disclosure of information on the supervision and inspection conducted by the drug regulatory authority and competent healthcare authority.

Conduct of Clinical Trial

According to the Announcement of Several Policies on the Evaluation and Examination for Drug Registration (《關於藥品註冊審評審批若干政策的公告》) promulgated by the CFDA on November 11, 2015, the INDs of new drugs are subject to one-off umbrella approval instead of declaration, evaluation and approval by stages. The Measures for the Administration of Drug Registration (《藥品註冊管理辦法》) further confirms that the CDE shall, within 60 working days from the date on which the application for a clinical trial is accepted, decide on whether to approve it and then notify the clinical trial applicant. In the case of failure to notify the applicant within the prescribed time limit, it shall be deemed as approved.

In compliance with the Measures for the Administration of Drug Registration (《藥品註冊管理辦法》), clinical trials are divided into Phase I, Phase II, Phase III, Phase IV and bioequivalence trial:

A clinical drug trial to be carried out shall be examined and approved by the ethics committee. The management of drugs used in a clinical drug trial shall satisfy the relevant requirements of the GCP. A sponsor approved to carry out clinical drug trial shall, before carrying out subsequent clinical drug trial by stages, develop corresponding plan for clinical drug trial, carry out clinical drug trial upon examination and with consent of the ethics committee, and submit corresponding plan for clinical drug trial and supporting materials on the website of the CDE.

Clinical trials shall be conducted for the application of new drug registration and shall be implemented in accordance with the GCP. The GCP stipulates the criteria for the entire procedure of the clinical trial including pre-clinical trial preparation and the necessary conditions, protection

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of testees' rights and interests, trial protocols, duties of researchers, duties of sponsors, duties of monitors, trial record and report, data management and statistical analysis, administration of drug products for trial, guarantee for quality, polycentric trials, with reference to the internationally recognized principles.

According to the Announcement of the Adjustment of Procedures of the Evaluation and Examination for Drug Clinical Trial (《關於調整藥物臨床試驗審評審批程序的公告》), if a new drug clinical trial has been approved to be carried out, after the completion of Phase I and Phase II clinical trials and before the implementation of Phase III clinical trials, the applicant shall submit an application for a communication meeting to the CDE to discuss with the CDE on key technical issues including the design of the Phase III clinical trial. The applicant can also apply for communication meeting on key technical issues at different stages of clinical research and development.

According to the Measures for the Administration of Drug Registration (《藥品註冊管理辦法》), applicants may communicate with the CDE on major issues at critical stages such as prior to application for clinical trial of a drug, during the process of clinical trial of a drug, and prior to application for marketing authorization of a drug. According to the Measures for the Administration of Communication and Exchange in Drug Development and Technology Review (《藥物研發與技術審評溝通交流管理辦法》) promulgated by the CDE on December 10, 2020, an applicant may propose to convene a communication meeting with the CDE during the process of drug research and development and registration application. There are three types of communication and exchange meetings: Type I meetings are held to resolve major safety issues encountered in the course of clinical trials of drugs and major technical issues in the course of R&D of breakthrough therapeutic drugs; Type II meetings are held for drugs at critical stages of R&D, which mainly include pre-application meetings for new drugs, meetings after the conclusion of Phase II clinical trials and before the commencement of Phase III clinical trials, meetings before application for marketing authorization of new drugs, and meetings for risk assessment and evaluation of new drugs. Type III meetings shall refer to meetings other than Type I and Type II meetings.

According to the Announcement of the Opinions on Handling Issues Related to Verification of Drug Clinical Trial Data (《關於藥物臨床試驗數據核查有關問題處理意見的公告》), both promulgated by CFDA on and effective from May 22, 2017, if the clinical trial data is incomplete, ill-formed or insufficient to prove the safety and efficacy of the drug, the registration application of the drug will be rejected.

New Drug Application

Pursuant to the Measures for the Administration of Drug Registration (《藥品註冊管理辦法》), after completing the pharmaceutical research, pharmacological and toxicological research, clinical drug trial, and other researches supporting the marketing registration of a drug, determining the quality standards, completing the verification of commercial large-scale production process, and making sound preparation for the acceptance of drug registration inspection and examination, an applicant shall file an application for drug marketing authorization, and submit relevant research materials in accordance with the requirements of the application materials. After the formal examination of the application materials, an application that satisfies the requirements shall be accepted. Where a generic drug, in vitro diagnostic reagent managed as a drug, or any other eligible circumstance assessed by an applicant to be unnecessary or impossible for conducting clinical drug trial and meeting the conditions for exempting clinical drug trial, the applicant may directly file an application for drug marketing authorization. The technical guiding principles and relevant specific requirements for exempting clinical drug trial shall be developed and announced by the CDE.

The CDE shall organize pharmaceutical, medical and other technical personnel to evaluate the accepted applications for drug marketing authorization. Where the comprehensive evaluation conclusion is adopted, the drug shall be approved for marketing, and a drug registration certificate

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shall be issued. If the comprehensive evaluation conclusion is not adopted, a disapproval decision shall be made. A drug registration certificate shall specify the drug approval number, holder, manufacturer and other information.

Drug registration inspection means the inspection activities carried out for the development sites and production sites to verify the authenticity and consistency of the application materials and the commercial production conditions for marketing of drugs, and to examine, among other things, the compliance of drug development and data reliability. Where necessary, it also includes the extended examination activities carried out for manufacturers, suppliers, or other entrusted institutions of chemical active pharmaceutical ingredients, auxiliary materials, and packaging materials and containers in direct contact with drugs involved in the application for drug registration.

The CDE shall decide whether to carry out on-site inspection of drug registration development based on risks, according to the degree of drug innovation and the previous acceptance of inspection by drug research institutions.

The CDE shall decide whether to launch production site inspection for drug registration based on risks according to factors such as variety, process, facility and previous acceptance of inspection for which an application is filed for registration. For innovative drugs, new modified drugs and biological products, production site inspection for drug registration and pre-marketing examination for management standards for drug production quality shall be conducted. For generic drugs, production site inspection for drug registration and pre-marketing examination for management standards for drug production quality shall be conducted based on the risks, according to whether a drug production license for the corresponding production scope has been obtained and whether a variety of the same dosage form has been marketed.

After an application for drug registration is accepted, the CDE shall conduct preliminary examination within forty (40) business days of acceptance, notify the Center for Food and Drug Inspection of NMPA (the "Center for Inspection") inspection and provide the relevant materials required for inspection, where production site inspection for drug registration is required, and concurrently notify the applicant and the medical products administrative department of the province, autonomous region or municipality in the place where the applicant or production enterprise is located. In principle, the Center for Inspection shall complete the inspection work forty (40) business days prior to the expiry of the time limit for inspection, and report the inspection information, inspection results and other relevant materials to the CDE.

Drug registration examination shall include standard review and sample examination. Standard review means the laboratory assessment of the scientificity of the items specified in the standards for the drug for which the applicant applies, the feasibility of the test methods and the rationality of quality control indicators, among others. Sample examination means the laboratory examination carried out for samples according to the application of the applicant or the drug quality standards verified by the CDE.

The review period for an application for drug marketing authorization shall be two hundred (200) business days. The review period for the procedures for prioritized review and approval shall be one hundred and thirty (130) business days, and the review period for the procedures for prioritized review and approval for clinically and urgently needed overseas-marketed drug for a rare disease shall be seventy (70) business days.

The following duration shall be excluded from the relevant work period: (i) time taken for the applicant to provide supplementary materials, to make corrections upon examination as well as to verify manufacturing process, quality standards and literature in accordance with the requirements; (ii) delay in examination or inspection due to reason of the applicant, time taken for organizing

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expert advisory meetings; (iii) the suspended duration in the event of suspension of review and approval procedures pursuant to the provisions of laws and regulations; and (iv) time taken for overseas examination where such overseas examination is activated.

Reform of Evaluation and Approval System for Drugs

In August 2015, the State Council promulgated the Opinions on the Reform of Evaluation and Approval System for Drugs and Medical Devices and Equipment (《關於改革藥品醫療器械審評審批制度的意見》) (the “Reform Opinions”), which provide a framework for reforming the evaluation and approval system for drugs and indicate the enhancement of the standard of approval for drug registration and the acceleration of the evaluation and approval process for innovative drugs.

In November 2015, the CFDA promulgated the Announcement on Certain Policies for Drug Registration, Evaluation and Approval (《關於藥品註冊審評審批若干政策的公告》) (the “Certain Policies Announcement”), which further clarifies the measures and policies on simplifying and accelerating the approval process on the basis of the Reform Opinions.

Pursuant to the Decision on Adjusting the Approval Procedures of Certain Administrative Approval Items for Drugs (《關於調整部分藥品行政審批事項審批程序的決定》) promulgated by the CFDA in March 2017 and came into effect in May 2017, the clinical trial approval decisions on drugs (including domestic and imported) can be directly made by the CDE in the name of the CFDA; decisions on approval of drug supplementary applications (including domestic and imported); decisions on approval of re-registration of imported drugs. In addition, the Circular on Issues Concerning Optimizing Drug Registration Review and Approval (《關於優化藥品註冊審評審批有關事宜的公告》), both promulgated by NMPA and NHC on and effective from May 17, 2018, further simplifies and accelerates the approval process of specific kinds of drugs.

The Evaluation and Approval Procedures for Breakthrough Therapeutic Drugs (Trial) (《突破性治療藥物審評工作程序(試行)》), the Evaluation and Approval Procedures for Conditionally Approved Drugs (Trial) (《藥品附條件批准上市申請審評審批工作程序(試行)》) and The Preferential Evaluation and Approval Procedures for Drug Marketing Authorization (Trial) (《藥品上市許可優先審評審批工作程序(試行)》) promulgated by the NMPA in July 2020 and came into effect in July 2020, replace the Opinions on Implementing Priority Review and Approval to Encourage Drug Innovation (《關於鼓勵藥品創新實行優先審評審批的意見》) promulgated by the CFDA in December 2017 and came into effect in December 2017, which further clarified the Accelerating Registration Procedures for Drugs.

Administrative Protection and Monitoring Periods for New Drugs

According to the Implementing Rules for PRC Drug Administration Law (《中華人民共和國藥品管理法實施條例》) issued on March 2, 2019 latest amended and effective on January 20, 2025 and the Reform Plan for Registration Category of Chemical Drugs (《化學藥品註冊分類改革工作方案》) issued on March 4, 2016, the NMPA may, for the purpose of protecting public health, provide for an administrative monitoring period of five years for new Category 1 drugs approved to be manufactured, commencing from the date of approval, to continually monitor the safety of those new drugs. During the monitoring period of a new drug, the NMPA will not approve any other enterprises’ applications to manufacture or import the said drug.

Regulations on International Multi-Center Clinical Trials and Acceptance of Overseas Clinical Trial Data

According to the Notice on Issuing the International Multi-Center Clinical Trial Guidelines (Trial) (《關於發佈國際多中心藥物臨床試驗指南(試行)的通告》), (“the Multi-Center Clinical Trial Guidelines”), promulgated by the NMPA on January 30, 2015 and came into effect from March 1, 2015, international multi-center clinical trial applicants may simultaneously perform clinical

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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 Rules for PRC Drug Administration Law and the Measures for the Administration of Drug Registration, execute the GCP, make reference to universal international principles such as the International Council for Harmonization of Technical Requirements for Pharmaceuticals for Human Use (ICH), 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 and other related laws and regulations.

According to the Opinions on Deepening the Reform of the Evaluation and Approval Systems and Encouraging Innovation of Drugs and Medical Devices (《關於深化審評審批制度改革鼓勵藥品醫療器械創新的意見》), clinical trial data obtained in an international multi-center that conforms to China’s requirements for registration of drugs and medical devices can be used for the application for registration in China.

According to the Technical Guiding Principles for the Acceptance of Overseas Clinical Trial Data of Drugs (《接受藥品境外臨床試驗數據的技術指導原則》) promulgated by the NMPA on July 6, 2018, the basic principles for accepting overseas clinical trial data include: (i) applicants shall ensure the authenticity, integrity, accuracy and trace-ability of overseas clinical trial data; (ii) the process of generating overseas clinical trial data shall comply with the relevant requirements of the ICH-GCP; (iii) applicants shall ensure the scientific design of overseas clinical trials, the compliance of clinical trial quality management system with the requirements, and the accuracy and integrity of statistical analysis of data; and (iv) to ensure that the clinical trial design and statistical analysis of the data are scientific and reasonable, for the drugs with simultaneous R&D at home and abroad and forthcoming clinical trials in China, the applicants may, prior to implementing registrational clinical trials, contact the CDE to ensure the compliance of registrational clinical trial’s design with the essential technical requirements for drug registration in China.

Marketing Authorization Holder System

Pursuant to the Drug Administration Law and the Measures for the Administration of Drug Registration, the state implements the drug marketing authorization holder system for drug management. After obtaining a drug registration certificate, an applicant shall be the drug marketing authorization holder. During the validity period, a holder of a drug registration certificate shall continue to ensure the safety, effectiveness and quality controllability of the marketed drug, and apply for re-registration of the drug six months prior to the expiry of the validity period.

The drug marketing authorization holder shall proactively carry out post-marketing research on drugs, further confirm the safety, effectiveness and quality controllability of drugs, and strengthen the continuous management of marketed drugs. Where a drug registration certificate and its annex require the marketing authorization holder to carry out relevant research work after the drug is marketed, the marketing authorization holder shall complete the research within the prescribed time limit and file a supplementary application, undergo recordation formalities or report as required. After a drug is approved for marketing, the marketing authorization holder shall continue to conduct research on drug safety and effectiveness, undergo recordation formalities in a timely manner or file a supplementary application for revising the instructions according to the relevant data, and continuously update and improve the instructions and labels. The medical products administrative department may require the marketing authorization holder to revise the instructions and labels based on the monitoring of adverse drug reactions and the post-marketing reevaluation results of the drug.

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The marketing authorization holder shall apply for re-registration six months prior to the expiry of the validity period of the drug registration certificate. An application for re-registration of a domestically produced drug shall be filed by the marketing authorization holder with the medical products administrative department of the province, autonomous region or municipality directly under the central government, and an application for re-registration of a drug produced overseas shall be filed by the marketing authorization holder with the CDE.

National Reimbursement Drug List of China (the “NRDL”)

Participants in the National Health Insurance Scheme and their employers (if any) have to pay a monthly premium. Participants may be reimbursed for all or part of the cost of medicines included in the medical insurance catalog. The Tentative Measures for the Administration of the Scope of Medical Insurance Coverage for Pharmaceutical Products for Urban Employee (《城鎮職工基本醫療保險用藥範圍管理暫行辦法》), jointly issued by the National Development and Reform Commission (the “NDRC”), the MOF and other authorities, came into effect on May 12, 1999, stipulates that the medicines included in the medical insurance catalog must be clinically necessary, safe and effective, reasonably priced, convenient to use and the supply of which can be guaranteed by the market.

The NRDL for Basic Medical Insurance, Maternity Insurance and Work-related Injury Insurance (《國家基本醫療保險、生育保險和工傷保險藥品目錄》) sets out the standards for payment of medicines by the basic medical insurance, work-related injury insurance and maternity insurance funds. The NHSA and other governmental organizations have the authority to determine the drugs to be included in the NRDL. Drugs listed in the NRDL are divided into two parts: Class A and Class B. Class A drugs are widely used for clinical treatment, with favorable efficacy and lower prices than their counterparts, while Class B drugs are used for clinical treatment, with favorable efficacy and slightly higher prices than Class A drugs.

On December 5, 2025, the NHSA and the Ministry of Human Resources and Social Security of the PRC released the latest NRDL (effective from January 1, 2026), which has been expanded to cover a total of 3,253 drugs. Inclusion in the NRDL will generally result in increased sales volume and lower drug prices (which are determined on a case-by-case basis and negotiated based on factors such as the initial drug price).

On July 30, 2020, the NHSA issued the Provisional Measures for the Administration of Medicines for Basic Medical Insurance (《基本醫療保險用藥管理暫行辦法》) (“Measures for the Administration of the NRDL”), which came into effect on September 1, 2020. The Measures for the Administration of the NRDL provides guidance on the inclusion and adjustment of the NRDL and the payment, management and supervision of basic medical insurance. According to the Measures for the Administration of the NRDL, a dynamic adjustment mechanism shall be established for the NRDL, which shall be adjusted annually in principle.

National Essential Drug List of China (the “NEDL”)

On August 18, 2009, the MOH and eight other ministries and commissions in the PRC issued the Provisional Measures on the Administration of the NEDL (《國家基本藥物目錄管理辦法(暫行)》), which was revised on February 13, 2015 by the Notice on Issuing the Measures on the Administration of the NEDL (《關於印發國家基本藥物目錄管理辦法的通知》), and the Guidelines on the Implementation of the NEDL System (《關於建立國家基本藥物制度的實施意見》), which aims to promote essential medicines sold to consumers at fair prices in the PRC and ensure that the general public in the PRC has equal access to the drugs contained in the NEDL. On September 13, 2018, the General Office of the State Council issued the Opinions of the General Office of the State Council on Improving the National Essential Drug System (《國務院辦公廳關於完善國家基本藥物制度的意見》). The NHC and the National Administration of Traditional Chinese Medicine promulgated the NEDL (2018 Edition) (《國家基本藥物目錄(2018年版)》) on September 30, 2018, replacing the NEDL (2012 Edition) (《國家基本藥物目錄(2012年版)》) which was promulgated on

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March 13, 2013. According to these regulations, basic healthcare institutions funded by the government, which primarily include county-level hospitals, county-level Chinese medicine hospitals, rural clinics and community clinics, shall stock and use drugs listed in the NEDL. The drugs listed in NEDL shall be purchased by centralized tender process and are subject to the price control by the NDRC. Remedial drugs in the NEDL are all listed in the Medical Insurance Catalog and the full amount of the purchase price of such drugs is entitled to reimbursement.

Human Genetic Resources Approval or Filing

Pursuant to the Regulations of the PRC on the Administration of Human Genetic Resources (《中華人民共和國人類遺傳資源管理條例》) (the “Human Genetic Resource Regulation”) promulgated by the State Council on May 28, 2019 and implemented on July 1, 2019, newly amended in March 2024 and came into effect on May 1, 2024, human genetic resource includes human genetic resource materials and information. Human genetic resource materials shall refer to organs, tissues, cells and other genetic materials containing human genome, genes and other genetic materials. Human genetic resource information shall refer to information, such as data, generated by human genetic resources materials. The Human Genetic Resource Regulation formalized the specific requirements to obtain approval relating to research collaborations between Chinese and foreign entities. Utilization of China’s human genetic resources in internally cooperative clinical trial with the purpose of obtaining marketing licenses for drugs and medical devices, which involves no export or emigration of human genetic resources, is not subject to relevant approval. However, parties to such internationally cooperative clinical trial shall submit a filing about the category, quantity, usage of human genetic resources. Foreign organisations, individuals and institutions established or actually controlled by foreign organizations and individuals are not allowed to collect or preserve human genetic resources in China or provide human genetic resources abroad.

The Bio-security Law of the PRC (《中華人民共和國生物安全法》) promulgated by the SCNPC on October 17, 2020 and last amended on April 26, 2024 and effective from the same date, provides that the PRC shall have sovereignty over the human genetic resources and biological resources of China. The Bio-security Law of the PRC further stipulates that the competent health department under the State Council shall be the competent authority for the approval or filing of activities involving China’s human genetic resources.

Good Clinical Practice Certification and Compliance with the Good Clinical Practice (GCP)

To improve the quality of clinical trials, the NMPA and NHC promulgated the Good Clinical Practice for Drug Trials (《藥物臨床試驗質量管理規範》) (the “GCP”) in April 2020 and came into effect on July 1, 2020, which aims to ensure that the clinical trials of drugs are standardized and the results are scientific and reliable, protecting the rights and safety of human subjects. Pursuant to the Opinions on Deepening the Reform of the Evaluation and Approval Systems and Encouraging Innovation of Drugs and Medical Devices (《關於深化審評審批制度改革鼓勵藥品醫療器械創新的意見》) promulgated by the general offices of the Chinese Communist Party Central Committee and the General Office of the State Council in October 2017, the qualification of clinical trial institutions shall be subject to record management. Clinical trials should follow GCP and protocols approved by the ethics committee of each research center.

Laws and Regulations in Relation to Drug Manufacturer

Drug Manufacturing Permit

Pursuant to the Drug Administration Law and the Measures on Supervision of Pharmaceutical Manufacturing (《藥品生產監督管理辦法》), promulgated by the NMPA on December 11, 2002, last amended on January 22, 2020 and effective on July 1, 2020, the state adopts an industry entry permit system for drug manufacturers. The conduct of drug manufacturing activities shall be approved and granted with a Drug Manufacturing License (《藥品生產許可證》) by the drug

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regulatory authority of the people’s government at provincial, autonomous regional or municipal level. The Drug Manufacturing License shall indicate the validity period and the scope of production, and shall be reviewed for renewing upon expiration.

Good Manufacturing Practices

Prior to December 1, 2019, establishment of a new drug manufacturer, construction of new production premise for a drug manufacturer or production of new dosage form were required to submit application for good manufacturing practice certification (GMP certification) with the drug regulatory authority in accordance with relevant provisions. If the Good Manufacturing Practices are satisfied, a GMP certificate will be issued. Pursuant to the Announcement on the Relevant Issues Concerning the Implementation of the Drug Administration Law of the PRC (《關於貫徹實施〈中華人民共和國藥品管理法〉有關事項的公告》), promulgated by the NMPA on November 29, 2019, and the Drug Administration Law, the GMP and Good Supply Practice (GSP) certifications have been canceled, applications for GMP and GSP certifications are no longer accepted, and GMP and GSP certificates are no longer issued. When engaging in drug manufacturing activities, a manufacturer shall comply with the GMP and establish a sound GMP management system, to ensure that the entire process of drug manufacturing continue to meet the statutory requirements, and meet the GMP requirements enacted by the drug regulatory authority under the State Council in accordance with the law. The legal representative of and principal person in charge of a drug manufacturer are fully responsible for the drug manufacturing activities of the enterprise.

The Good Manufacturing Practices (《藥品生產質量管理規範》), promulgated by the Ministry of Health of the PRC (the “MOH”, revoked on March 10, 2013, now known as the NHC) in December 1992, newly amended in January 2011 and came into effect on March 1, 2011, provided guidance for the quality management, organization and staffing, production premises and facilities, equipments, material and products, recognition and inspection, documentation maintenance, manufacture management, quality control and quality assurance, contractual manufacture and contractual inspection for the products, product delivery and recalls of a manufacturer in a systematical manner.

Contract Manufacturing of Drug

Pursuant to the Administrative Regulations for the Contract Manufacturing of Drug (《藥品委託生產監督管理規定》), promulgated by CFDA on August 14, 2014 and effective from October 1, 2014, in the event that a drug manufacturer in China that has obtained a drug marketing authorization temporarily lacks manufacturing conditions as a result of technology upgrade or is unable to ensure market supply due to insufficient manufacturing capabilities, it can entrust the manufacturing of that drug to another domestic drug manufacturer. Such contract manufacturing arrangements need to be approved by the provincial branch of NMPA. The Regulations further prohibit the contract manufacturing arrangement of certain special drugs, including narcotic drugs, psychoactive drugs, biochemical drugs and active pharmaceutical ingredients.

Pursuant to the Drug Administration Law, a drug marketing authorization holder may produce drugs by itself or may entrust other drug manufacturers with production. A drug marketing authorization holder that intends to manufacture drugs on its own shall obtain a drug manufacturing permit. If the holder intends to entrust a third-party to manufacture, it shall entrust a qualified drug manufacturer. The drug marketing authorization holder and the commissioned manufacturer shall enter into an entrustment agreement and a quality agreement, and strictly perform the obligations pursuant to such agreements. Blood products, anesthetics, psychotropic pharmaceuticals, toxic pharmaceuticals for medical treatment and pharmaceutical precursor chemicals may not be produced through entrustment, except as otherwise prescribed by the drug administrative department of the State Council.

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Export of Drug

Pursuant to the Reply by NMPA on Certain Issues of Pharmaceutical Products Export (《國家藥品監督管理局關於藥品出口有關問題的批覆》), both promulgated on and effective from September 20, 1999, enterprises have the right to operate import and export of pharmaceutical products and the qualification shall be decided by the foreign trade authority. Export of pharmaceutical products shall mainly comply with the requirements of the importing country, so long as there is no special requirement by the importation country, the NMPA would, in principle, support the export based on the national policy of encouraging exports. However, pursuant to the Drug Administration Law, the export of narcotics and psychotropic drugs prescribed by the PRC requires an export license issued by the NMPA.

Other Laws and Regulations in Relation to Medical Industry

Basic Medical Insurance Policy

Pursuant to the Decision on the Establishment of the Urban Employee Basic Medical Insurance Program (《關於建立城鎮職工基本醫療保險制度的決定》) promulgated by the State Council on December 14, 1998 and the Tentative Measures for the Administration of the Scope of Medical Insurance Coverage for Pharmaceutical Products for Urban Employee (《城鎮職工基本醫療保險用藥範圍管理暫行辦法》), all employers in cities and towns, including enterprises (state-owned enterprises, collective enterprises, foreign-invested enterprises, private enterprises, etc.), institutions, public institutions, social organizations, private non-enterprise units and their employees are required to participate in basic medical insurance. Pursuant to the Guiding Opinions on the Pilot of Basic Medical Insurance for Urban Residents (《關於開展城鎮居民基本醫療保險試點的指導意見》) promulgated by the State Council on July 10, 2007, urban residents (not urban employees) in the pilot areas can voluntarily participate in the basic medical insurance for urban residents. Pursuant to the Opinions of the State Council on the Integration of the Basic Medical Insurance System for Urban and Rural Residents (《國務院關於整合城鄉居民基本醫療保險制度的意見》) promulgated by the State Council on January 3, 2016, a unified basic medical insurance system for urban and rural residents was established, integrating the existing urban residents' medical insurance and the new rural cooperative medical system, covering all urban and rural residents except those who should be covered by the employee's basic medical insurance.

Medical Insurance Catalog

Pursuant to the Tentative Measures for the Administration of the Scope of Medical Insurance Coverage for Pharmaceutical Products for Urban Employee (《城鎮職工基本醫療保險用藥範圍管理暫行辦法》), the scope of medical insurance coverage for pharmaceutical products needs to be managed through the formulation of the Medical Insurance Catalog. A pharmaceutical product listed in the Medical Insurance Catalog must be clinically needed, safe, effective, reasonably priced, easy to use, available in sufficient quantity, and must meet the following requirements: it is set forth in the Pharmacopoeia of the PRC (current edition) (《中華人民共和國藥典》(現行版)); it meets the standards promulgated by the NMPA; and if imported, it is approved by the NMPA for import. According to the Opinions of the NHSA and the Ministry of Finance on Establishing a List-Based System for Healthcare Security Benefits (《國家醫保局、財政部關於建立醫療保障待遇清單制度的意見》), which came into effect in January 2021, all provinces shall implement the NRDL in a strict manner, and shall not have the discretion to formulate the catalog, add drugs to the catalogue, or adjust the scope of limited payment unless explicitly stipulated. After several adjustments, the currently effective one is the National Insurance Drug List for Basic Medical Insurance, Maternity Insurance and Work-related Injury Insurance (2025) (《國家基本醫療保險、生育保險和工傷保險藥品目錄(2025年)》) came into effect since January 1, 2026.

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Drug Price

Pursuant to the Drug Administration Law, for drug products with market-regulated prices in accordance with the law, the drug marketing authorization holder, the drug manufacturer, the drug distributor and medical institution shall determine the price pursuant to the principles of fairness, reasonableness, integrity and trustworthiness as well as quality for value in order to supply drug users with reasonably priced drug products; and shall comply with the requirements relating to drug price administration promulgated by the State Council’s pricing authorities, determine and clearly mark the retail prices of drug products. Pursuant to the Notice on Issuing Opinions on Promoting Drug Price Reform (《關於印發<推進藥品價格改革意見>的通知》) jointly promulgated by the NDRC, the NHC, the Ministry of Human Resources and Social Security, Ministry of Industry and Information Technology, the Ministry of Finance of the PRC (中華人民共和國財政部) (the “MOF”), the Ministry of Commerce of the PRC (中華人民共和國商務部) (the “MOFCOM”) and the CFDA on May 4, 2015 and came into effect on June 1, 2015, government-set drug prices were abolished except for narcotic drugs and first-class psychotropic drugs.

Advertising of Pharmaceutical Products

The Advertising Law of the PRC (《中華人民共和國廣告法》), as amended and effective on April 29, 2021, outlines the regulatory framework for the advertising industry. Advertisers, advertising service providers and advertising publishers are required to ensure that the contents of the advertisements they prepare or distribute are true and in full compliance with applicable laws and regulations. For advertisement of drugs, the advertisement contents shall be examined by the relevant authorities prior to their publishing. Pursuant to the Interim Administrative Measures for the Review of Advertisements for Drugs, Medical Devices, Health Food and Formula Food for Special Medical Purposes (《藥品、醫療器械、保健食品、特殊醫學用途配方食品廣告審查管理暫行辦法》) promulgated by SAMR in December 2019 and came into effect on March 1, 2020, advertisements for drugs, medical devices, health food and formula food for special medical purposes shall be true and legitimate, and shall not contain any false or misleading contents.

Insert Sheet, Labels and Packaging of Pharmaceutical Products

Pursuant to the Measures for the Administration of the Insert Sheets and Labels of Drugs (《藥品說明書和標籤管理規定》), promulgated by the SFDA and came effective on June 1, 2006, the insert sheets and labels of drugs should be reviewed and approved by the SFDA. A drug insert sheet should include the important scientific data, conclusions and information concerning drug safety and efficacy in order to direct the safe and rational use of drugs. The inner label of a drug should bear such information as the drug’s name, indication or function, strength, dose and usage, production date, batch number, expiry date and drug manufacturer. The outer label of a drug should indicate such information as the drug’s name, ingredients, description, indication or function, strength, dose and usage, adverse reaction, contraindications, precautions, storage, production date, batch number, expiry date, approval number and the name of the drug manufacturer. Pursuant to the Measures for The Administration of Pharmaceutical Packaging (《藥品包裝管理辦法》) which came effective on September 1, 1988, pharmaceutical packaging must comply with national and professional standards. If no national or professional standards are available, the enterprise can formulate and implement its standards after obtaining the approval of the food and drug administration and bureau of standards at provincial level. The enterprise shall reapply with the relevant authorities if it needs to change its packaging standard. Drugs without packaging standards must not be sold or traded (except for drugs for the military).

Provisions in Relation to the Company’s Principal Activities

The “Healthy China 2030” Planning Outline (《“健康中國2030”規劃綱要》) jointly issued and implemented by the State Council and the general offices of the Chinese Communist Party Central Committee on October 25, 2016 proposes the establishment of a comprehensive clinical evaluation system focusing on essential medicines, aiming to complete the drug price formation

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mechanism in accordance with the principle of combining government regulation with market adjustment, strengthen the linkage between the pricing, health insurance, procurement policies while insisting on categorized management, strengthen supervision of prices of medicines with insufficient market competition and high-value medical consumables, establish a system for monitoring and disclosing information on drug prices, and formulate and improve policies on and standards for payment for drugs covered by medical insurance.

The 14th Five-Year Plan for National Health (《“十四五”國民健康規劃》) issued and implemented by the General Office of the State Council on April 27, 2022 proposes to encourage the research and development, innovation and application of new medicines, accelerate the research, development and industrialization of urgently-needed medicines for the clinical treatment of major diseases, and support the research and development of high-quality generic medicines, deepen the reform of the review and approval procedures for medicines and medical devices, and expedite the review and approval process for innovative, clinically-needed drugs and medical devices, and drugs for the treatment of rare diseases.

The Several Opinions of the General Office of the Shanghai Municipal People’s Government on Supporting the Whole-Chain Innovation and Development of the Biopharmaceutical Industry (《上海市人民政府辦公廳關於支持生物醫藥產業全鏈條創新發展的若干意見》) issued by the General Office of the Shanghai Municipal People’s Government on July 15, 2024 and implemented on August 1, 2024 (valid until July 31, 2029) proposes to support the innovation and development of the whole chain of the biopharmaceutical industry, including accelerating the transition from the R&D achievements to pre-clinical research, continuously increasing the support for R&D of innovative drugs, optimizing the mechanism for the transformation of clinical research achievements, shortening the start-up time of clinical trials, etc.

The Several Measures for the Whole-Chain Support of the High-Quality Development of Innovative Drugs and Medical Devices (《浙江省人民政府辦公廳關於印發浙江省全鏈條支持創新藥械高質量發展若干舉措的通知》) issued by the General Office of the People’s Government of Zhejiang Province on December 30, 2024 and implemented on February 15, 2025, proposes to support the whole-chain development of innovative drugs and medical devices through initiatives including enhancing R&D capabilities, optimizing clinical research and review processes, promoting application, expanding payment channels, and strengthening data/industrial factor support, etc.

Laws and Regulations in Relation to Intellectual Property

Patent

Patents in the PRC are mainly protected by the Patent Law of the PRC (《中華人民共和國專利法》) (the “Patent Law”), which was promulgated by the SCNPC on March 12, 1984 and latest amended on October 17, 2020 and came into effect on June 1, 2021, and the Implementation Rules of the Patent Law of the PRC (《中華人民共和國專利法實施細則》) (the “Implementation Rules”), promulgated by the State Council on June 15, 2001 and latest amended on December 11, 2023 and came into effect on January 20, 2024. The Patent Law and the Implementation Rules provide for three types of patents, namely “invention”, “utility model” and “design”. “Invention” refers to any new technical solution relating to a product, a process or improvement thereof; “utility model” refers to any new technical solution relating to the shape, structure, or their combination, of a product, which is suitable for practical use; and “design” refers to any new design of the shape, pattern, color or the combination of any two of them, of a product, which creates an aesthetic feeling and is suitable for industrial application. The duration of a patent right for “invention” is twenty (20) years; the duration of a patent right for “utility model” is ten (10) years; and the duration of a patent right for “design” is fifteen (15) years, all of which duration are from the date of application. According to the Patent Law, for the purpose of public health, the patent

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administration department of the State Council may grant mandatory licensing for patented drugs manufactured and exported to countries or regions which comply with the provisions of the relevant international treaty participated by the PRC.

The newly amended Patent Law introduces patent extensions to patents of new drugs launched in the PRC, and stipulates that the patent administration department under the State Council shall, upon request of the patentee, extend the patent term of relevant invention patents of the new drug that is approved to be listed on the market in China, to compensate for the time spent for the review, examination and approval of the listing of a new drug on the market.

The compensated extension shall not exceed five (5) years, and the total valid patent term after the new drug is approved for the market shall not exceed fourteen (14) years. Such newly adopted patent term extension rule benefits the Company through longer protection terms of patents applied or registered in the PRC and related to our product candidates. During the compensated extension period of the patent term of the patent for invention related to a new drug, the scope of protection of the patent is limited to the new drug and the technical solutions related to the approved indications of the new drug. Within the scope of protection, the rights and obligations of the patentee remain the same as before the compensated extension period.

Trademarks

Registered trademarks in the PRC are mainly protected by the Trademark Law of the PRC (《中華人民共和國商標法》), which was promulgated by the SCNPC on August 23, 1982 and latest amended on April 23, 2019 and came into effect on November 1, 2019, and the Implementation Rules of the Trademark Law of the PRC (《中華人民共和國商標法實施條例》), which were promulgated by the State Council on August 3, 2002 and latest amended on April 29, 2014 and came into effect on May 1, 2014. The Trademark Office is responsible for the registration and administration of trademarks throughout the PRC, and grants a term of ten (10) years to registered trademarks. When it is necessary to continue using the registered trademark upon expiration of period of validity, a trademark registrant shall make an application for renewal within twelve (12) months before the expiration in accordance with the requirements. If such an application cannot be filed within that period, an extension period of six months may be granted. The period of validity for each renewal of registration shall be ten (10) years as of the next day of the previous period of validity. If the formalities for renewal have not been handled upon expiration of period of validity, the registered trademarks will be deregistered.

Domain Names

Domain names are regulated under the Administrative Measures on the Internet Domain Names (《互聯網域名管理辦法》) issued by the Ministry of Industry and Information Technology (the “MIIT”), on August 24, 2017 and effective from November 1, 2017. The MIIT is the main regulatory authority responsible for the administration of the PRC internet domain names. Domain names registrations are handled through domain name service agencies established under the relevant regulations, and the applicants become domain name holders upon successful registration.

Trade Secret

According to the Anti-Unfair Competition Law of the PRC (《中華人民共和國反不正當競爭法》) promulgated by SCNPC, latest amended on June 27, 2025 and came into effect on October 15, 2025, 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 Anti-Unfair Competition Law of the PRC, business persons are prohibited from infringing others’ trade secrets by: (i) acquiring a trade secret from the right holder by theft, bribery, fraud, coercion, electronic intrusion, or any other means; (ii) disclosing, using, or allowing another person to use a trade secret acquired from the right holder by any means as specified in the item (i) above; (iii) disclosing, using, or allowing another

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person use a trade secret in its possession, in violation of its confidentiality obligation or the requirements of the right holder for keeping the trade secret confidential; (iv) abetting a person, or tempting another person into or in acquiring, disclosing, using, or allowing another person to use the trade secret of the right holder in violation of his or her non-disclosure obligation or the requirements of the right holder for keeping the trade secret confidential. 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 impose fine on the infringing parties.

The Company Law and Regulations

The Company Law of the PRC (《中華人民共和國公司法》), which was amended by the SCNPC on December 29, 2023 and became effective on July 1, 2024, provides for the establishment, corporate structure and corporate management of companies, which also applies to foreign-invested enterprises in PRC.

Regulations in Relation to Foreign Direct Investment

Since January 1, 2020, the Foreign Investment Law of the PRC (《中華人民共和國外商投資法》) (the “Foreign Investment Law”) promulgated by the National People’s Congress (the “NPC”) and the Regulations for Implementation of the PRC Foreign Investment Law (《中華人民共和國外商投資法實施條例》) promulgated by the State Council came into effect and became the principal laws and regulations governing foreign investment in the PRC. The Law of the PRC on Sino-Foreign Equity Joint Ventures (《中華人民共和國中外合資經營企業法》), the Law of the PRC on Wholly Foreign-Owned Enterprises (《中華人民共和國中外合作經營企業法》) and the Law of the PRC on Sino-Foreign Cooperative Joint Ventures (《中華人民共和國中外合作經營企業法》), together with their implementation rules and ancillary regulations were abolished at the same time. Pursuant to the Foreign Investment Law, “foreign investment” shall refer to the investment activities conducted directly or indirectly by foreign individuals, enterprises or other entities in the PRC, including the following circumstances: (i) the establishment of foreign-invested enterprises in the PRC by foreign investors solely or jointly with other investors; (ii) a foreign investor’s acquisition of shares, equity interests, property portions or other similar rights and interests of enterprises in the PRC; (iii) investment in new projects in the PRC by foreign investors solely or jointly with other investors; and (iv) investments made by foreign investors through means provided in laws, administrative regulations, or other methods prescribed by the State Council. The organization form, institutional framework and standard of conduct of foreign-invested enterprises shall be subject to the provisions of the Company Law of the PRC and other laws.

The PRC government will implement the management system of pre-entry national treatment and the Negative List for foreign investment abolished the original approval and filing administration system for the establishment and change of foreign-invested enterprises. Pre-entry national treatment refers to the treatment accorded to foreign investors and their investments at the stage of investment entry which is no less favorable than the treatment accorded to domestic investors and their investments. Negative List refers to a special administrative measure for the entry of foreign investment in specific sectors as imposed by the PRC. The PRC accords national treatment to foreign investment outside of the Negative List. The current Negative List is the Special Management Measures (Negative List) for the Access of Foreign Investment (2024 Revision) (《外商投資准入特別管理措施(負面清單)(2024年版)》) issued by the NDRC and the MOFCOM on September 6, 2024 and came into effect on November 1, 2024, which lists the special management measures for foreign investment access for industries regulated by the Negative List, such as equity requirements and senior management requirements. While strengthening investment promotion and protection, the Foreign Investment Law further regulates foreign investment management and proposes the establishment of a foreign investment information reporting system that replaces the original foreign investment enterprise approval and filing system of the MOFCOM.

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The foreign investment information reporting is subject to the Foreign Investment Information Reporting Method (《外商投資信息報告辦法》) jointly developed by the MOFCOM and the SAMR, which came into effect on January 1, 2020. According to the Foreign Investment Information Reporting Method, the MOFCOM is responsible for coordinating and guiding the reporting of foreign investment information nationwide. The competent commercial department of the local people’s government at or above the county level, as well as the relevant agencies of the pilot free trade zone and the national economic and technological development zone, are responsible for reporting information on foreign investment in the region. Foreign investors who directly or indirectly carry out investment activities in China shall submit investment information to the competent commercial department through the enterprise registration system and the national enterprise credit information publicity system and the reporting methods include initial reports, change reports, cancellation reports, and annual reports. Foreign investors who establish foreign invested enterprises in China or acquire domestic non-foreign-invested enterprises through equity merger and acquisition shall submit initial reports through the enterprise registration system when applying for the registration of the establishment of foreign-invested enterprises or applying for the registration of the change of the acquired enterprises. If the change in the information of initial reports involves registration or filing of the change of enterprises, foreign-invested enterprises shall submit change reports through the enterprise registration system when applying for the registration or filing of change of enterprises. If the change in the information of initial reports does not involve registration or filing of the change of enterprises, foreign-invested enterprises shall submit change reports through the enterprise registration system within twenty (20) business days after the change. Foreign-invested listed companies may report information on changes in investors and their shareholdings only when the cumulative change in the foreign investors’ shareholding ratio exceeds 5% or the foreign parties’ shareholding or relative holding status have changed.

Regulations on The Security Review of Foreign Investment

On December 19, 2020, the NDRC and the MOFCOM jointly promulgated the Measures on the Security Review of Foreign Investment (《外商投資安全審查辦法》), effective on January 18, 2021, setting forth provisions concerning the security review mechanism on foreign investment, including the types of investments subject to review, the scopes of review and procedures to review, among others.

Regulations in Relation to Product Liability

The Product Quality Law of the PRC (《中華人民共和國產品質量法》), promulgated by the SCNPC on February 22, 1993 and latest amended on December 29, 2018 (the “Product Quality Law”), is the principal governing law relating to the supervision and administration of product quality. According to the Product Quality Law, manufacturers shall be liable for the quality of products produced by them and sellers shall take measures to ensure the quality of the products sold by them. A manufacturer shall be liable to compensate for any bodily injuries or damage to property other than the defective product itself resulting from the defects in the product, unless the manufacturer is able to prove that: (1) the product has never been circulated; (2) the defects causing injuries or damage did not exist at the time when the product was circulated; or (3) the science and technology at the time when the product was circulated were at a level incapable of detecting the defects. A seller shall be liable to compensate for any bodily injuries or damage to property of others caused by the defects in the product if such defects are attributable to the seller. A seller shall pay compensation if it fails to indicate neither the manufacturer nor the supplier of the defective product. A person who is injured or whose property is damaged by the defects in the product may claim for compensation from the manufacturer or the seller.

Pursuant to the PRC Civil Code (《中華人民共和國民法典》) promulgated by the NPC on May 28, 2020 and coming into effect on January 1, 2021, where a patient suffers damage due to defects in drugs, he may seek compensation from the drug marketing authorization holder, producer

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or also from the medical institution. Where the patient seeks compensation from the medical institution, the medical institution, after it has made the compensation, shall have the right to recover the compensation from the liable drug marketing authorization holder or producer.

The Law of the PRC on the Protection of the Rights and Interests of Consumers (《中華人民共和國消費者權益保護法》) was promulgated on October 31, 1993 and latest amended on October 25, 2013 and came into effect on March 15, 2014 to protect consumers’ rights when they purchase or use goods and accept services. All business operators must comply with this law when they manufacture or sell goods and/or provide services to customers. All business operators must pay high attention to protecting customers’ privacy and must strictly keep confidential any consumer information they obtain during their business operations.

Regulations in Relation to Production Safety

The Production Safety Law of the PRC (《中華人民共和國安全生產法》), promulgated by the SCNPC on June 29, 2002 and latest amended on June 10, 2021 and came into effect on September 1, 2021, is the basic law for governing production safety. It provides that, any entity whose production safety conditions do not meet the requirements may not engage in production and business operation activities. The production and business operation entities shall educate and train employees regarding production safety so as to ensure that the employees have the necessary knowledge of production safety, are familiar with the relevant regulations and rules for safe production and the rules for safe operation, master the skills of safe operation in their own positions, understand the emergency measures, and know their own rights and duties in terms of production safety. Employees who fail the education and training programs on production safety may not commence working in their positions. Safety facilities of new building, rebuilding or expanding project (the “construction project”) shall be designed, constructed and put into operation simultaneously with the main body of the project. Investment in safety facilities shall be included in the budget of the construction project.

Regulations in Relation to Environmental Protection and Fire Safety

According to the Environmental Protection Law of the PRC (《中華人民共和國環境保護法》), promulgated by the SCNPC on December 26, 1989 and latest amended on April 24, 2014 and came into effect on January 1, 2015, the Environmental Impact Assessment Law of the PRC (《中華人民共和國環境影響評價法》), promulgated by the SCNPC on October 28, 2002 and latest amended on December 29, 2018, and the Administrative Regulations on the Environmental Protection of Construction Project (《建設項目環境保護管理條例》), promulgated by the State Council on November 29, 1998 and latest amended on July 16, 2017 and came into effect on October 1, 2017, enterprises which plan to construct projects shall engage qualified professionals to provide the assessment reports, assessment form, or registration form on the environmental impact of such projects. The assessment reports, assessment form, or registration form shall be filed with or approved by the relevant environmental protection bureau prior to the commencement of any construction work.

Enterprises generating environmental pollution in the PRC must comply with the PRC Law on the Prevention from Water Pollution (《中華人民共和國水污染防治法》) promulgated by the SCNPC on May 11, 1984, and last amended on June 27, 2017, the PRC Law on the Prevention from Atmospheric Pollution (《中華人民共和國大氣污染防治法》) promulgated by the SCNPC on September 5, 1987, and last amended on October 26, 2018, the PRC Law on the Prevention from Noise Pollution (《中華人民共和國噪聲污染防治法》) promulgated by the SCNPC on December 24, 2021 and effective from June 5, 2022, and the PRC Law on the Prevention from Environmental Pollution of Solid Waste (《中華人民共和國固體廢物污染環境防治法》) promulgated by the SCNPC on October 30, 1995, and last amended on April 29, 2020. The abovementioned laws regulate extensive issues in relation to the environment protection including waste water discharge, air pollution control, noise emission and solid waste pollution control. Pursuant to these laws, all

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the enterprises that may cause environmental pollution in the course of their production and business operation shall introduce environmental protection measures in their plants and establish a reliable system for environmental protection.

Enterprises that engage in the activities of industry, construction, catering, and medical treatment, etc. that discharges sewage into urban drainage facilities shall apply to the relevant competent urban drainage department for the permit for discharging sewage into drainage pipelines under relevant laws and regulations, including the Regulations on Urban Drainage and Sewage Disposal (《城鎮排水與污水處理條例》), which was promulgated on October 2, 2013 and came into force on January 1, 2014, and the Measures for the Administration of Permits for the Discharge of Urban Sewage into the Drainage Network (《城鎮污水排入排水管網許可管理辦法》), which was promulgated on January 22, 2015 and last amended on December 1, 2022 and took effect on February 1, 2023. 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.

According to the Administrative Measures on Pollutant Discharge Permit (《排污許可管理辦法》) issued by the Ministry of Ecology and Environment on April 1, 2024 and came into effect on July 1, 2024, enterprises, public institutions and other producers and operators that are subject to the administration of pollutant discharge permits shall apply for pollutant discharge permit and discharge pollutants in accordance with the requirements of the pollutant discharge permit; and those who have not obtained the pollutant discharge permits shall not discharge pollutants. According to the Classification Management List for Fixed Source Pollution Permits (2019 Edition) (《固定污染源排污許可分類管理名錄(2019年版)》), the manufacturing of biological drugs and products falls into the classification management scope for fixed source pollution permits.

According to the Opinions of the general offices of the Chinese Communist Party Central Committee and the State Council on Fully and Accurately Implementing the New Development Concept and Doing a Good Job in Carbon Peak and Carbon Neutrality, (《中共中央、國務院關於完整準確全面貫徹新發展理念做好碳達峰碳中和工作的意見》) promulgated and came into effect on September 22, 2021, significant progress will be made in green transformation by 2030, with key industries reaching advanced international energy efficiency levels. By 2060, a green, low-carbon, circular economy and a clean, safe, and efficient energy system will be fully established, achieving the carbon neutrality goal.

According to the Fire Safety Law of the PRC (《中華人民共和國消防法》) promulgated by the SCNPC in April 1998, last amended and effective on April 29, 2021, and the Interim Provisions on Administration of Fire Protection Design Review and Acceptance of Construction Projects (《建設工程消防設計審查驗收管理暫行規定》) (the “Interim Provisions”) promulgated by the Ministry of Housing and Urban-Rural Development on April 1, 2020, and last amended on August 21, 2023, the fire protection design or construction of a construction project must conform to the national fire protection technical standards for project construction and construction projects shall undergo the fire protection design review and acceptance system. The special construction projects as defined in the Interim Provisions must apply to the fire control department for fire protection design review, and complete the fire protection acceptance procedures after the completion of the construction project. The construction unit of other construction projects must complete the fire protection filing of the fire protection design and the completion acceptance within five (5) business days after the completion acceptance of the construction project. If a construction project fails to pass the fire safety inspection before it is put into use, or does not meet the fire safety requirements after the inspection, it will be ordered to suspend the construction and use of such project, or suspend production and business, and be imposed a fine.

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Regulations in Relation to Prevention and Control of Occupational Diseases

The Prevention and Control of Occupational Diseases Law of the PRC (《中華人民共和國職業病防治法》), which was promulgated by the SCNPC on October 27, 2001 and latest amended on December 29, 2018 (the “Prevention and Control of Occupational Diseases Law”), is the basic law for the prevention and control of occupational diseases. According to the Prevention and Control of Occupational Diseases Law, budget for facilities for the prevention and control of occupational diseases of a construction project shall be included in the budget of the project and those facilities shall be designed, constructed and put into operation simultaneously with the main body of the project. The entity that takes charge of the project should carry out the assessment of the effectiveness of measures for the prevention and control of occupational diseases before the final acceptance of the construction project. In addition, employers shall take required administrative measures to prevent and control occupational diseases in work.

Employee Stock Option Plans

On February 15, 2012, the SAFE issued the Circular on Issues concerning the Foreign Exchange Administration for Domestic Individuals Participating in Share Incentive Plans of Overseas Publicly-Listed Companies (《關於境內個人參與境外上市公司股權激勵計劃外匯管理有關問題的通知》) (the “Share Option Rules”). Under the Share Option Rules, the PRC citizens or residents habitually residing in the PRC continuously for over one year, with a few exceptions, and who have been granted, restricted shares or share options by an overseas listed company according to its employee share option or share incentive plan, are required to appoint a qualified PRC agent, register with the SAFE or its local counterparts and complete certain other procedures related to the shareholding plan, share option plan or other similar share incentive plans. Concurrent with registration with the SAFE or its local counterparts, the qualified PRC agent is required to obtain an approval from the SAFE for an annual allowance for the foreign exchanges in connection with shareholding or the exercise of a share option, and an approval for opening a special foreign exchange account at a PRC domestic bank to hold the funds required in connection with share purchases or share option exercises, returned principals or profits upon sale of shares, dividends issued on the stock and any other income or expenditures approved by the SAFE. Currently, foreign exchange income of the participating PRC residents received from the sale of share and dividends distributed by the overseas listed company are required to be fully remitted into such special domestic foreign currency account before distribution to such participants. In addition, the PRC agents are required to amend or deregister the registrations with the SAFE or its local counterparts in case of any material change in, or termination of, the share incentive plans within the time periods provided by the Share Option Rules.

Regulations in Relation to Employment and Social Securities

Pursuant to the Labor Law of the PRC (《中華人民共和國勞動法》), promulgated by the SCNPC on July 5, 1994 and latest amended on December 29, 2018 and the Labor Contract Law of the PRC (《中華人民共和國勞動合同法》), promulgated by the SCNPC on June 29, 2007 and latest amended on December 28, 2012 and came into effect on July 1, 2013, employers shall execute written labor contracts with full-time employees. All employers shall comply with local minimum wage standards. Employers shall establish a comprehensive management system to protect the rights of their employees, including a system governing occupational health and safety to provide employees with occupational training to prevent occupational injury, and employers are required to truthfully inform prospective employees of the job description, working conditions, working location, occupational hazards, and status of safe production as well as remuneration and other conditions.

According to Social Security Law of the PRC (《中華人民共和國社會保險法》), which was promulgated on October 28, 2010 and amended on December 29, 2018, an employer is required to make contributions to social insurance schemes for its employees, including basic pension insurance, basic medical insurance, unemployment insurance, maternity insurance and work-related

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injury insurance. If the employer fails to make social insurance contributions in full and on time, the social insurance authorities may demand the employer to make payments or supplementary payments for the unpaid social insurance premium within a prescribed time limit together with a 0.05% surcharge of the unpaid social insurance premium from the due date. If the payment is not made within such time limit, the relevant administrative authorities will impose a fine ranging from one to three times the total outstanding amount.

According to the Interim Measures for the Participation in Social Insurance of Foreigners Employed in China (《在中國境內就業的外國人參加社會保險暫行辦法》), which was promulgated on September 6, 2011, amended on December 23, 2024 and became effective on the same date, employers who hire foreigners shall register them for social insurance within 30 days from the date of obtaining employment certificates. Foreigners who participate in social insurance and meet the requirements shall enjoy social insurance benefits in accordance with the law.

According to the Reform Plan of the State Tax and Local Tax Collection Administration System (《國稅地稅徵管體制改革方案》), which was promulgated on July 20, 2018, commencing from January 1, 2019, all the social insurance premiums including the premiums of the basic pension insurance, unemployment insurance, maternity insurance, work-related injury insurance and basic medical insurance shall be collected by the tax authorities. According to the Notice on Conducting the Relevant Work Concerning the Administration of Collection of Social Insurance Premiums in a Steady, Orderly and Effective Manner (《關於穩妥有序做好社會保險費徵管有關工作的通知》) promulgated by the General Office of the STA on September 13, 2018, all the local authorities responsible for the collection of social insurance are strictly forbidden to conduct self-collection of historical unpaid social insurance contributions from enterprises. The Notice on Implementing Measures to Further Support and Serve the Development of Private Economy (《關於實施進一步支持和服務民營經濟發展若干措施的通知》), promulgated by the STA on November 16, 2018, repeats that tax authorities at all levels may not organize self-collection of arrears of taxpayers including private enterprises from the previous years. The Notice of General Office of the State Council on Promulgation of the Comprehensive Plan for the Reduction of Social Insurance Premium Rate (《國務院辦公廳關於印發降低社會保險費率綜合方案的通知》), promulgated on April 1, 2019, requires steady advancement of the reform of the system of social security collection. In principle, the basic pension insurance for enterprise employees and other insurance types for enterprise employees shall be collected temporarily according to the existing collection system to stabilize the payment method. It also emphasizes that the historical unpaid arrears of the enterprise shall be properly treated. In the process of reformation of the collection system, it is not allowed to conduct self-collection of historical unpaid arrears from enterprises, and it is not allowed to adopt any method of increasing the actual payment burden of small and micro enterprises to avoid causing difficulties in the production and operation of the enterprises.

According to the Administrative Regulations on Housing Provident Funds (《住房公積金管理條例》), which was promulgated on April 3, 1999 and latest amended on March 24, 2019, employers are required to make contribution to housing provident funds for their employees. Where an employer fails to pay up housing provident funds, the housing provident fund administration center may order it to make payment within a prescribed time limit. If the employer still fails to do so, the housing provident fund administration center may apply to the court for compulsory enforcement of the unpaid amount.

Regulations in Relation to Information Security and Data Privacy

Data Security and Export

The SCNPC promulgated the Data Security Law of the PRC (《中華人民共和國數據安全法》), on June 10, 2021 (effective from September 1, 2021), for the establishment of a data classification and grading protection system to conduct classified and hierarchical protection of

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data. Entities engaged in data processing activities shall, in accordance with laws and regulations, establish a sound full-process data security management system, organize data security education and training, and take corresponding technical measures and other necessary measures to ensure data security.

The Measures for Cybersecurity Review (《網絡安全審查辦法》) (the “MCR”) were promulgated jointly by the Cyberspace Administration of China and the other 12 governmental authorities, and became effective on February 15, 2022. The MCR stipulates that the operators carrying out data processing activities that affect or may affect national security, shall conduct cybersecurity review. Pursuant to Article 2 of the MCR, besides the procurement of network products and services by critical information infrastructure operators, any data processing activity by network platform operators that affects or may affect national security shall be subject to the cybersecurity review. In accordance with Article 7 of the MCR, network platform operators mastering personal information of more than one million users must apply to the Cybersecurity Review Office for cybersecurity review when listing abroad.

According to the Measures for Security Assessment of Data Export (《數據出境安全評估辦法》) issued by the Cyberspace Administration of China on July 7, 2022 and came into effect on September 1, 2022, a data processor that provides data overseas under any of the following circumstances shall apply to the national cyberspace administration for the security assessment of the outbound data transfer through local provincial cyberspace administration: (i) a data processor provides important data abroad; (ii) the critical information infrastructure operator or the data processor that has processed the personal information of more than 1 million people provides personal information abroad; (iii) the data processor that has provided the personal information of over 100,000 people or the sensitive personal information of over 10,000 people cumulatively since January 1 of the previous year provides personal information abroad; and (iv) any other circumstance where an application for the security assessment of outbound data transfer is required by the national cyberspace administration.

According to the Measures for Standard Contract for Outbound Transfer of Personal Information (《個人信息出境標準合同辦法》) issued by the Cyberspace Administration of China on February 22, 2023 and effective from June 1, 2023, to provide personal information to an overseas recipient through the conclusion of the standard contract, a personal information processor shall meet all of the following circumstances: (i) it is not a critical information infrastructure operator; (ii) it has processed the personal information of less than one million individuals; (iii) it has cumulatively provided the personal information of less than 100,000 individuals to overseas recipients since January 1 of the previous year; and (iv) it has cumulatively provided the sensitive personal information of less than 10,000 individuals since January 1 of the previous year.

According to the Provisions on Facilitating and Regulating Cross-border Data Flows (《促進和規範數據跨境流動規定》), both issued by the Cyberspace Administration of China on and effective from March 22, 2024, a data handler that is not a critical information infrastructure operator, will be exempted from declaring for security assessment for outbound data transfer, signing a standard contract with overseas recipient or passing the personal protection certification, if such data handler accumulatively transfers overseas ordinary personal information of less than 100,000 individuals since the January 1 of the current year.

Personal Information Protection

According to the Civil Code (《民法典》), personal information of natural persons is protected by law. If any organization or individual needs to obtain other people’s personal information, they should obtain it in accordance with the law and ensure the security of the information. They must not illegally collect, use, process, or transmit other people’s personal information, and must not illegally buy, sell, provide, or disclose the information. The Personal Information Protection Law of the PRC (《中華人民共和國個人信息保護法》) promulgated by the

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SCNPC on August 20, 2021 and implemented on November 1, 2021, further emphasizes the obligations and responsibilities of processors for the protection of personal information, and requests higher level of protective measures on the processing of sensitive personal information.

According to the Cybersecurity Law of the PRC (《中華人民共和國網絡安全法》), promulgated by the SCNPC on November 7, 2016 and effective on June 1, 2017, as amended on October 28, 2025 and came into effect on January 1, 2026, network operators must follow the principles of legality, legitimacy and necessity when collecting and using personal information, and publicly disclose the rules for collection and use, clearly state the purpose, method and scope of collecting and using information, and obtain the consent of the person whose data is being collected. Network operators shall not collect personal information unrelated to the services they provide. Network operators are not allowed to leak, tamper with, or damage the personal information they collect; they are not allowed to provide personal information to others without the consent of the person whose data is being collected. However, this does not apply to cases where a specific individual cannot be identified and the identity cannot be recovered after processing. Network operators should take technical measures and other necessary measures to ensure the security of the personal information they collect and prevent leakage, damage and loss of information.

Laws and Regulations in Relation to Anti-Bribery

According to the Anti-Unfair Competition Law of the PRC (《中華人民共和國反不正當競爭法》), and the Interim Provisions on the Prohibition of Commercial Bribery (《關於禁止商業賄賂行為的暫行規定》) promulgated by the State Administration of Industry and Commerce of the PRC (中華人民共和國國家工商行政管理總局, the predecessor of the SAMR) on November 15, 1996, any business operator shall not provide or promise to provide economic benefits (including cash, other property or by other means) to a counter-party in a transaction or a third party that may be able to influence the transaction, in order to entice such party to secure a transactional opportunity or competitive advantages for the business operator. Any business operator breaching the relevant anti-bribery rules above-mentioned may be subject to administrative punishment or criminal liability depending on the seriousness of the cases.

Pursuant to the Provisions on the Establishment of Adverse Records of Commercial Briberies in the Medicine Purchase and Sales Industry (《關於建立醫藥購銷領域商業賄賂不良記錄的規定》), promulgated by the National Health and Family Planning Commission (currently the NHC) and came into effect on March 1, 2014, any medicine production and operation enterprises or agents that are involved in criminal, investigational or administrative procedures for commercial bribery will be listed in the adverse records of commercial bribes by the relevant government authorities, as a result of which, for two years from the date the list of adverse records of commercial bribes is published, (i) their products cannot be purchased by public medical institutions or medical and health institutions receiving financial subsidies within the relevant provinces, and (ii) the scores of their products in the centralized tender processes of public medical institutions or medical and health institutions receiving financial subsidies in other provinces will be reduced. As for those enterprises or agents listed in adverse records twice within five years, their products cannot be purchased by public medical institutions or medical and health institutions receiving financial subsidies throughout China for two years from the date the list of adverse records of commercial bribes is published.

REGULATIONS ON TAXATION

Enterprise Income Tax

The PRC Enterprise Income Tax Law (《中華人民共和國企業所得稅法》) and Regulations for the Implementation of the PRC Enterprise Income Tax (《中華人民共和國企業所得稅法實施條例》) (collectively, the “EIT Laws”) were promulgated on March 16, 2007 and December 6, 2007, respectively, and were amended on December 29, 2018 and April 23, 2019, respectively. Pursuant to the EIT Laws, taxpayers consist of resident enterprises and non-resident enterprises. Resident

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enterprises are defined as enterprises that are established in China in accordance with PRC laws, or that are established in accordance with the laws of foreign countries but whose actual or de facto control is administered within China. Non-resident enterprises are defined as enterprises that are set up in accordance with the laws of foreign countries and whose actual administration is conducted outside China, but have established institutions or premises in China, or have no such established institutions or premises but have income generated from inside China. Pursuant to the EIT Laws and relevant implementing regulations, a uniform 25% enterprise income tax rate is imposed to both foreign invested enterprises and domestic enterprises. However, if non-resident enterprises have not formed permanent establishments or premises in China, or if they have formed permanent establishment institutions or premises in China but there is no actual relationship between the relevant income derived in China and the established institutions or premises set up by them, the enterprise income tax is, in that case, set at the rate of 10% for their income sourced from inside China. The enterprise income tax rate is reduced to 20% for qualifying small low-profit enterprises. The high-tech enterprises that need full support from the PRC’s government will enjoy a reduced tax rate of 15% for enterprise income tax.

Value-added Tax

Pursuant to the Value-Added Tax Law of the PRC (《中華人民共和國增值稅法》), which was promulgated by the SCNPC on December 25, 2024 and effective from January 1, 2026, entities and individuals engaging in selling goods, services, intangible assets, or immovables within the territory of the PRC are taxpayers of the value-added tax (“VAT”).

According to the Notice of the Ministry of Finance and the State Taxation Administration on the Adjusting Value-added Tax Rates (《財政部、稅務總局關於調整增值稅稅率的通知》) effective in May 2018, the VAT rates of 17% and 11% on sales, imported goods shall be adjusted to 16% and 10%, respectively.

According to the Announcement of the Ministry of Finance, the State Taxation Administration and the General Administration of Customs on Relevant Policies for Deepening the Value-Added Tax Reform (《財政部、稅務總局、海關總署關於深化增值稅改革有關政策的公告》) promulgated on March 20, 2019 and effective from April 1, 2019, the VAT rates of 16% and 10% on sales, imported goods shall be adjusted to 13% and 9%, respectively.

REGULATIONS ON FOREIGN EXCHANGE

Foreign Exchange Regulation

On January 29, 1996, the State Council promulgated the Administrative Regulations on Foreign Exchange of the PRC (《中華人民共和國外匯管理條例》) which became effective on April 1, 1996 and was amended on January 14, 1997 and August 5, 2008. Foreign exchange payments under current account items shall, pursuant to the administrative provisions of the foreign exchange control department of the State Council on payments of foreign currencies and purchase of foreign currencies, be made using self-owned foreign currency or foreign currency purchased from financial institutions engaging in conversion and sale of foreign currencies by presenting the valid document. Domestic entities and domestic individuals making overseas direct investments or engaging in issuance and trading of overseas securities and derivatives shall process registration formalities pursuant to the provisions of the foreign exchange control department of the State Council.

On November 19, 2012, the SAFE issued the Circular of Further Improving and Adjusting Foreign Exchange Administration Policies on Foreign Direct Investment (《關於進一步改進和調整直接投資外匯管理政策的通知》), or the SAFE Circular 59, which came into effect on December 17, 2012 and was revised on May 4, 2015, October 10, 2018 and partially abolished on December 30, 2019. The SAFE Circular 59 aims to simplify the foreign exchange procedure and promote the facilitation of investment and trade. According to the SAFE Circular 59, the opening of various

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special purpose foreign exchange accounts, such as pre-establishment expenses accounts, foreign exchange capital accounts and guarantee accounts, the reinvestment of RMB proceeds derived by foreign investors in the PRC, and remittance of foreign exchange profits and dividends by a foreign-invested enterprise to its foreign shareholders no longer require the approval or verification of the SAFE, multiple capital accounts for the same entity may be opened in different provinces as well. Later, the SAFE promulgated the Circular on Further Simplifying and Improving Foreign Exchange Administration Policies in Respect of Direct Investment (《關於進一步簡化和改進直接投資外匯管理政策的通知》) in February 2015, which was partially abolished in December 2019, prescribed that the bank instead of the SAFE can directly handle the foreign exchange registration and approval under foreign direct investment while SAFE and its branches indirectly supervise the foreign exchange registration and approval under foreign direct investment through the bank.

On May 10, 2013, the SAFE issued the Administrative Provisions on Foreign Exchange in Domestic Direct Investment by Foreign Investors (《外國投資者境內直接投資外匯管理規定》), or the SAFE Circular 21, which became effective on May 13, 2013, amended on October 10, 2018 and partially abolished on December 30, 2019. The SAFE Circular 21 specifies that the administration by the SAFE or its local branches over direct investment by foreign investors in the PRC must be conducted by way of registration and banks must process foreign exchange business relating to the direct investment in the PRC based on the registration information provided by the SAFE and its branches.

According to the Notice on Relevant Issues Concerning the Administration of Foreign Exchange for Overseas Listing (《關於境外上市外匯管理有關問題的通知》) issued by the SAFE on December 26, 2014, the domestic companies shall register the overseas listed with the foreign exchange control bureau located at its registered address in 15 working days after completion of the overseas listing and issuance. The funds raised by the domestic companies through overseas listing may be repatriated to China or deposited overseas, provided that the intended use of the fund shall be consistent with the contents of the document and other public disclosure documents.

According to the Notice of the State Administration of Foreign Exchange on Reforming the Management Mode of Foreign Exchange Capital Settlement of Foreign Investment Enterprises (《國家外匯管理局關於改革外商投資企業外匯資本金結匯管理方式的通知》), or the SAFE Circular 19 promulgated on March 30, 2015, coming effective on June 1, 2015 and partially abolished on December 30, 2019 and March 23, 2023, foreign-invested enterprises could settle their foreign exchange capital on a discretionary basis according to the actual needs of their business operations. Whilst, foreign-invested enterprises are prohibited to use the foreign exchange capital settled in RMB (a) for any expenditures beyond the business scope of the foreign-invested enterprises or forbidden by laws and regulations; (b) for direct or indirect securities investment; (c) to provide entrusted loans (unless permitted in the business scope), repay loans between enterprises (including advances by third parties) or repay RMB bank loans that have been on-lent to a third party; and (d) to purchase real estates not for self-use purposes (save for real estate enterprises).

On June 9, 2016, the SAFE issued the Notice of the State Administration of Foreign Exchange on Reforming and Standardizing the Foreign Exchange Settlement Management Policy of Capital Account (《國家外匯管理局關於改革和規範資本項目結匯管理政策的通知》), or the SAFE Circular 16, which came into effect on the same day and partially amended on December 4, 2023. The SAFE Circular 16 provides that discretionary foreign exchange settlement applies to foreign exchange capital, foreign debt offering proceeds and remitted foreign listing proceeds, and the corresponding RMB capital converted from foreign exchange may be used to extend loans to related parties or repay inter-company loans (including advances by third parties). However, there remain substantial uncertainties with respect to the SAFE Circular 16's interpretation and implementation in practice.

On October 23, 2019, the SAFE promulgated the Notice on Further Facilitating Cross-Board Trade and Investment (《國家外匯管理局關於進一步促進跨境貿易投資便利化的通知》), which became effective on the same date (except for Article 8.2, which became effective on January 1,

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2020), and partially amended on December 4, 2023. The notice canceled restrictions on domestic equity investments made with capital funds by non-investing foreign-funded enterprises. In addition, restrictions on the use of funds for foreign exchange settlement of domestic accounts for the realization of assets have been removed and restrictions on the use and foreign exchange settlement of foreign investors' security deposits have been relaxed. Eligible enterprises in the pilot area are also allowed to use revenues under capital accounts, such as capital funds, foreign debts and overseas listing revenues for domestic payments without providing materials to the bank in advance for authenticity verification on an item-by-item basis, while the use of funds should be true, in compliance with applicable rules and conforming to the current capital revenue management regulations.

According to the Circular on Optimizing Administration of Foreign Exchange to Support the Development of Foreign-related Business (《關於優化外匯管理支持涉外業務發展的通知》) issued by the SAFE on April 10, 2020, eligible enterprises are allowed to make domestic payments by using their capital funds, foreign credits and the income under capital accounts of overseas listing, without submitting the evidentiary materials concerning authenticity of such capital for banks in advance, provided that their capital use is authentic and in compliance with administrative regulations on the use of income under capital accounts. The bank in charge shall conduct post spot checking in accordance with the relevant requirements.

Regulations in Relation to Overseas Securities Offering and Listing by Domestic Companies

On July 6, 2021, the general offices of the Chinese Communist Party Central Committee and the General Office of the State Council jointly promulgated the Opinions on Strictly Cracking Down on Illegal Securities Activities in accordance with the Law (《關於依法從嚴打擊證券違法活動的意見》) (the “Opinions on Securities Activities”). The Opinions on Securities Activities emphasize the need to strengthen the administration over illegal securities activities and the supervision on overseas listings by China-based companies and proposed to take effective measures, such as promoting the construction of relevant regulatory systems to deal with the risks and incidents faced by China-based overseas-listed companies.

According to the Trial Administrative Measures of Overseas Securities Offering and Listing by Domestic Companies (《境內企業境外發行證券和上市管理試行辦法》) and five relevant guidelines, issued by the CSRC on February 17, 2023 and effective from March 31, 2023 (hereinafter referred to as the “Overseas Listing Trial Measures”), where a domestic company seeks overseas securities issuance and listing, the issuer shall file with the CSRC in accordance with the Overseas Listing Trial Measures. If an issuer procures an overseas initial public offering or listing, it shall file with the CSRC within three (3) business days after submitting application documents for overseas securities issuance and listing.

According to the Provisions on Strengthening Confidentiality and Archives Administration of Overseas Securities Offering and Listing by Domestic Companies (《關於加強境內企業境外發行證券和上市相關保密和檔案管理工作的規定》) jointly issued by the CSRC and other departments on February 24, 2023 and effective on March 31, 2023, in the overseas offering and listing activities of domestic enterprises, domestic enterprises, and securities companies and securities service institutions that provide corresponding services shall strictly comply with the applicable laws and regulations of the PRC and satisfy the requirements of these Provisions, enhance the legal awareness of safeguarding state secrets and strengthening archives administration, establish and improve the confidentiality and archives work system, and take necessary measures to fulfill the confidentiality and archives administration obligations, and shall not divulge state secrets or work secrets of state organs, or harm the interests of the state or the public. A domestic enterprise that, either directly or through its overseas listed entity, publicly discloses or provides to relevant securities companies, securities service institutions, overseas regulators, and other entities and individuals, any documents and materials that involve state secrets or work secrets of state organs, shall obtain approval from the competent department with the power of examination and approval according to the law, and report to the administrative department of confidentiality at the same level

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for filing. A domestic enterprise that, either directly or through its overseas listed entity, publicly discloses or provides to relevant securities companies, securities service institutions, overseas regulators, and other entities and individuals, other documents and materials whose divulgence will have adverse impact on national security or public interest, shall strictly undergo the relevant procedures in accordance with the relevant regulations of the state.

Regulations in Relation to the “Full Circulation” of H Share

In accordance with the Guidelines for the “Full Circulation” Program for Domestic Unlisted Shares of H-share Listed Companies (《H股公司境內未上市股份申請“全流通”業務指引》) which was promulgated by the CSRC on November 14, 2019 and came into effect on the same date, which was partly revised on August 10, 2023 according to the Decision on Revising and Abolishing Part of Securities and Futures Policy Documents by CSRC (《中國證券監督管理委員會關於修改、廢止部分證券期貨制度文件的決定》), the term “full circulation” means the circulation of domestically unlisted shares (including domestically unlisted shares held by domestic shareholders prior to the listing abroad, additional domestically unlisted shares issued domestically after listing abroad and unlisted shares held by foreign shareholders) of H-share companies on the Stock Exchange. On the premise of complying with relevant laws and regulations as well as policies governing state-owned asset management, foreign investment and industrial supervision, the shareholders of domestically unlisted shares may determine the number and proportion of shares under application for circulation through negotiation at their discretion and entrust a H-share company to file an application for “full circulation”. After the domestically unlisted shares are listed for circulation on the Stock Exchange, they shall not be transferred back to the mainland China. A shareholder of domestically unlisted shares may reduce or increase its holding of the shares involved that are circulating on the Stock Exchange according to relevant business rules. H share companies shall submit a report on the relevant information to the CSRC within 15 days from completion of re-registration of the shares involved in the application to the CSDC.

In accordance with the Notice on Promulgation of the Implementing Rules for “Full Circulation” of H-shares (《關於發佈<H股“全流通”業務實施細則>的通知》) which was promulgated by the CSDC and Shenzhen Stock Exchange on December 31, 2019 and came into effect on the same date, it shall apply to the relevant businesses involved in “full circulation” of H-shares, such as cross-border re-registration, custodian and maintenance of holding details, entrustment of transactions and order routing, settlement, management of clearing participants, services of nominee holders etc. Upon completion of information disclosure by a H-share listed company approved by the CSRC to participate in “full circulation” of H-shares, such company shall register anew its fully tradable H-shares free from pledge, freezing, restriction of transfer and other restrictive status with the Hong Kong share registration authorities to have them become shares that can be listed and circulated on the Stock Exchange. The relevant securities shall be deposited centrally with the CSDC in China. The CSDC, as the nominee of the aforesaid securities, shall handle the business such as the depository and maintenance of holding details as well as cross-border clearing and settlement involved in the “full circulation” of H-shares, and provide services of nominee for investors. H-share listed companies shall obtain the authorization from investors and select a domestic securities company to participate in the “full circulation” of H-shares. Investors submit the trading orders for the “full circulation” of H-shares through a domestic securities company. The domestic securities company shall select a Hong Kong securities company through which investors’ trading instructions shall be reported to the Stock Exchange for trading. After transactions are concluded, the CSDC and China Securities Depository and Clearing (Hong Kong) Company Limited shall handle cross-border clearing and settlement of relevant shares and funds. The settlement currency of H-share “full circulation” transaction business is Hong Kong dollars. Where an H-share listed company entrusts the CSDC to distribute cash dividends, it shall file an application with the CSDC. The H-share listed company, when distributing cash dividends, may claim the details of the shares held by relevant investors on the equity registration date for cash dividends from the CSDC. If an investor obtains “fully tradable” non-H-shares listed on the Stock Exchange due to the equity distribution or conversion of H-shares under full circulation, the

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investor may sell but cannot purchase such securities; if the investor obtains the right to subscribe for shares listed on the Stock Exchange and such right is listed on the Stock Exchange, the investor may sell but shall not exercise such right.

In accordance with the Notice of China Securities Depository and Clearing Co., Ltd Shenzhen Branch on the Release of China Securities Depository and Clearing Co., Ltd Shenzhen Branch H-shares “Full Circulation” Business Guidelines (《中國結算深圳分公司關於發佈〈中國證券登記結算有限責任公司深圳分公司H股“全流通”業務指南〉的通知》) which was promulgated by CSDC Shenzhen Branch on September 20, 2024 and came into effect on September 23, 2024, newly amended in June 2025 and came into effect on June 30, 2025, it specified the business preparation, account arrangement, cross-border share transfer registration and overseas centralized custody, etc. And China Securities Depository and Clearing (Hong Kong) Company Limited also promulgated the Guide to the Program for Full Circulation of H-shares (《中國證券登記結算(香港)有限公司H股“全流通”業務指南》) to specify the relevant escrow, custody, agent service of China Securities Depository and Clearing (Hong Kong) Company Limited, arrangement for settlement and delivery and other relevant matters.

OVERVIEW OF LAWS AND REGULATIONS IN THE UNITED STATES

This section summarizes the principal laws and regulations in the United States that are relevant to our business.

Laws and Regulations in Relation to New Drug

U.S. Government Regulation of Drug and Biological Products

In the United States, the FDA regulates drugs under the Federal Food, Drug and Cosmetic Act and its implementing regulations, and biologics under the Federal Food, Drug and Cosmetic Act and the Public Health Service Act and their implementing regulations. Both drugs and biologics are also subject to other federal, state and local statutes and regulations, such as those related to competition. The process of obtaining regulatory approvals and the subsequent compliance with appropriate federal, state, and local statutes and regulations requires the expenditure of substantial time and financial resources. Failure to comply with the applicable U.S. requirements at any time during the product development process, approval process or following approval may subject an applicant to administrative actions or judicial sanctions. These actions and sanctions could include, among other actions, the FDA’s refusal to approve pending applications, withdrawal of an approval, license revocation, a clinical hold, untitled or warning letters, voluntary or mandatory product recalls or market withdrawals, product seizures, total or partial suspension of production or distribution, injunctions, fines, refusals of government contracts, restitution, disgorgement and civil or criminal fines or penalties. Any agency or judicial enforcement action could have a material adverse effect on our business, the market acceptance of our products and our reputation.

Once a product candidate is identified for development, it enters preclinical studies, which includes laboratory evaluations of product chemistry, toxicity, formulation and stability, as well as animal studies. IND-enabling nonclinical safety and toxicology studies are generally required to be conducted in accordance with FDA’s Good Laboratory Practice regulations. A sponsor of an IND must submit the results of the preclinical tests, manufacturing information, analytical data, the clinical trial protocol, and any available clinical data or literature to the FDA. The IND automatically becomes effective 30 days after receipt by the FDA, unless the FDA raises concerns or questions and places the trial on a clinical hold within that 30-day period. The FDA may also impose clinical holds or partial clinical holds at any time during clinical trials due to safety concerns or non-compliance.

All clinical trials, which involve the administration of the investigational product to humans, must be conducted under the supervision of one or more qualified investigators in accordance with Good Clinical Practice regulations, including the requirement that all research subjects provide

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informed consent in writing before their participation in any clinical trial. Further, an Institutional Review Board (the “IRB”), must review and approve the plan for any clinical trial before it commences at any institution, and the IRB must conduct continuing review and reapprove the study at least annually. Each new clinical protocol and any amendments to the protocol must be submitted for FDA review, and to the IRBs for approval.

An IRB can suspend or terminate approval of a clinical trial at its institution if the trial is not being conducted in accordance with the IRB’s requirements or if the product has been associated with unexpected serious harm to subjects.

Clinical trials generally are conducted in three sequential phases, known as Phase I, Phase II and Phase III, and may overlap. The FDA does not require a rigid, formal sequence of Phase I, then Phase II, then Phase III in every case. Rather, once an IND is in effect and FDA has not imposed a clinical hold, the sponsor may proceed with the studies set out in the IND, subject to the applicable regulatory requirements and the adequacy of the supporting data for the proposed investigation.

- Phase I clinical trials generally involve a small number of healthy volunteers or disease-affected patients who are initially exposed to a single dose and then multiple doses of the product candidate. The primary purpose of these clinical trials is to assess the metabolism, pharmacologic action, side effect tolerability and safety of the product candidate.
- Phase II clinical trials involve studies in disease-affected patients to evaluate proof of concept and/or determine the dose required to produce the desired benefits. At the same time, safety and further PK and PD information is collected, possible adverse effects and safety risks are identified and a preliminary evaluation of efficacy is conducted.
- Phase III clinical trials generally involve a large number of patients at multiple sites and are designed to provide the data necessary to demonstrate the effectiveness of the product for its intended use, its safety in use and to establish the overall benefit/risk relationship of the product and provide an adequate basis for product labeling.

Progress reports detailing the results of the clinical trials must be submitted at least annually to the FDA. Safety reports must be submitted to the FDA and the investigators 15 calendar days after the trial sponsor determines that the information qualifies for reporting. The sponsor also must notify FDA of any unexpected fatal or life-threatening suspected adverse reaction as soon as possible but in no case later than 7 calendar days after the sponsor’s initial receipt of the information. Sponsors of applicable clinical trials of FDA-regulated products, including drugs, are required to register and disclose certain clinical trial information, which is publicly available at www.clinicaltrials.gov.

Concurrent with clinical trials, companies usually complete additional animal studies and must also finalize a process for manufacturing the product in commercial quantities in accordance with cGMP requirements. The process of obtaining regulatory approvals and compliance with appropriate federal, state, local and foreign statutes and regulations requires the expenditure of substantial time and financial resources. Failure to comply with the applicable U.S. requirements may subject an applicant to administrative or judicial sanctions.

U.S. Review and Approval Processes

The results of product development, preclinical studies and clinical trials, along with descriptions of the manufacturing process, analytical tests conducted on the product, proposed labeling and other relevant information, are submitted to the FDA as part of an NDA or a BLA. Unless deferred or waived, NDAs or BLAs, or supplements must contain data adequate to assess the safety and effectiveness of the product for the claimed indications in all relevant pediatric

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subpopulations and to support dosing and administration for each pediatric subpopulation for which the product is safe and effective. The submission of an NDA or a BLA is subject to the payment of a substantial user fee and an annual prescription drug product program fee.

Within 60 days of its receipt, the FDA reviews the NDA/BLA to ensure that it is sufficiently complete for substantive review before it accepts the NDA/BLA for filing. After accepting the NDA/BLA filing, the FDA begins an in-depth substantive review to determine, among other things, whether a product is safe and effective for its intended use. The FDA also evaluates whether the product's manufacturing is cGMP-compliant to assure the product's identity, strength, quality and purity. Before approving the NDA/BLA, the FDA typically will inspect whether the manufacturing processes and facilities are in compliance with cGMP requirements and adequate to assure consistent production of the product within required specifications. The FDA may refer the NDA/BLA to an advisory committee, a panel of experts, for review whether the application should be approved and under what conditions and considers such recommendations when making decisions.

The FDA may refuse to approve the NDA/BLA if the applicable regulatory criteria are not satisfied or may require additional clinical data or other data and information. The FDA will issue a complete response letter describing all of the specific deficiencies that the FDA identified in the NDA/BLA that must be satisfactorily addressed before it can be approved. The deficiencies identified may be minor, for example, requiring labeling changes, or major, for example, requiring additional clinical trials. Additionally, the complete response letter may include recommended actions that the applicant might take to place the application in a condition for approval. The applicant may either resubmit the NDA/BLA, addressing all of the deficiencies identified in the letter, or withdraw the application or request an opportunity for a hearing.

The regulatory approval may be limited to specific diseases and dosages or the indications for use may otherwise be limited, which could restrict the commercial value of the product. Further, the FDA may require that certain contraindications, warnings or precautions be included in the product labeling. In addition, the FDA may require post-approval studies, including Phase IV clinical trials, to further assess a product's safety and effectiveness after NDA/BLA approval and may require testing and surveillance programs to monitor the safety of approved products that have been commercialized.

Expedited Development and Review Programs

The FDA has various programs that are intended to expedite or streamline the process for the development and FDA review of drugs that are intended for the treatment of serious or life-threatening diseases or conditions and demonstrate the potential to address unmet medical needs. The purpose of these programs is to provide important new drugs to patients earlier than under standard FDA review procedures.

Fast Track Designation

To be eligible for a fast-track designation, the FDA must determine, based on the request of a sponsor, that a drug is intended to treat a serious or life-threatening disease or condition for which there is no effective treatment and demonstrates the potential to address an unmet medical need for the disease or condition. Under the fast-track program, the sponsor of a drug candidate may request FDA to designate the product for a specific indication as a fast-track product concurrent with or after the filing of the IND for the drug candidate. The FDA must make a fast-track designation determination within 60 days after receipt of the sponsor's request.

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In addition to other benefits, such as the ability to use surrogate endpoints and have more interactions with FDA, FDA may initiate review of sections of a fast-track product’s NDA before the application is complete. This rolling review is available if the applicant provides, and FDA approves, a schedule for the submission of the remaining information and the applicant pays applicable user fees. However, FDA’s time period goal for reviewing a fast-track application does not begin until the last section of the NDA is submitted. In addition, the fast-track designation may be withdrawn by FDA if FDA believes that the designation is no longer supported by data emerging in the clinical trial process.

Orphan Drug Designation

Under The Orphan Drug Act of 1983, the FDA may grant orphan drug designation to drugs or biologic candidates intended to treat a rare disease or condition generally affecting fewer than 200,000 individuals in the U.S. or for which a manufacturer has no reasonable expectation of recovering drug treatment research and development costs. The first applicant to receive FDA approval for the disease or indication for which it has orphan drug designation is entitled to a seven-year exclusive marketing period. During the exclusivity period, the FDA may not approve any other applications to market the same product for the same disease or condition except in limited circumstances.

Accelerated Approval

Under FDA’s accelerated approval regulations, the FDA may approve a drug or biologic candidate for a serious or life-threatening illness that provides meaningful therapeutic benefit to patients over existing treatments and demonstrates an effect on either 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 (“IMM”), that is reasonably likely to predict an effect on IMM or other clinical benefit, taking into account the severity, rarity, or prevalence of the disease or condition and the availability or lack of alternative treatments. A product candidate approved on this basis is subject to rigorous post-marketing compliance requirements, including the completion of post-approval clinical trial to confirm the effect on the clinical endpoint. Failure to conduct required post-approval studies, or to confirm a clinical benefit during post-marketing studies, will allow the FDA to withdraw the product from the market on an expedited basis. All promotional materials for product candidates approved under accelerated regulations are subject to prior review by the FDA.

Breakthrough Therapy Designation

A drug or biologic may be eligible for designation as a breakthrough therapy if the product is intended, alone or in combination with one or more other drugs or biologics, to treat a serious or life-threatening condition and preliminary clinical evidence indicates that the product may demonstrate substantial improvement over currently approved therapies on one or more clinically significant endpoints, such as substantial treatment effects observed early in clinical development. A sponsor may request that a product be designated as a breakthrough therapy concurrently with, or at any time after, the submission of IND, and the FDA must determine if the candidate qualifies for such designation within 60 days of receipt of the request. If so designated, the FDA shall act to expedite the development and review of the product’s marketing application, including by meeting with the sponsor throughout the product’s development, providing timely advice to the sponsor to ensure that the development program to gather pre-clinical and clinical data is as efficient as practicable.

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Priority Review

The FDA may give a priority review designation to drugs that offer major advances in treatment or provide a treatment where no adequate therapy exists. A priority review means that the goal for the FDA to review an application is six months, rather than the standard review of ten months under the Prescription Drug User Fee Act guidelines. These six- and ten-month review periods are measured from the “filing” date rather than the receipt date for NDAs for new molecular entities, which typically adds approximately two months to the timeline for review and decision from the date of submission. Most products that are eligible for fast-track designation are also likely to be considered appropriate to receive a priority review.

Post-Marketing Requirements

Following the approval of a new product, the manufacturer and the approved product are subject to continuing regulation by the FDA, including, among other things, monitoring and record-keeping activities, reporting of adverse experiences, complying with promotion and advertising requirements, which include restrictions on promoting products for unapproved uses or patient populations (known as “off-label use”) and limitations on industry-sponsored scientific and educational activities. Although physicians may prescribe legally available products for off-label uses, manufacturers may not market or promote such uses. The FDA and other agencies 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, including investigation by federal and state authorities. Prescription drug promotional materials must be submitted to the FDA in conjunction with their first use or first publication. Further, if there are any modifications to the drug or biologic, including changes in indications, labeling or manufacturing processes or facilities, the applicant may be required to submit and obtain FDA approval of a new NDA/BLA or NDA/BLA supplement, which may require the development of additional data or preclinical studies and clinical trials. The FDA may also place other conditions on approvals including the requirement for a risk evaluation and mitigation strategy (“REMS”), to assure the safe use of the product. If the FDA concludes a REMS is needed, the sponsor of the NDA/BLA must submit a proposed REMS. The FDA will not approve the NDA/BLA without an approved REMS, if required. A REMS 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. Any of these limitations on approval or marketing could restrict the commercial promotion, distribution, prescription or dispensing of products. Product approvals may be withdrawn for non-compliance with regulatory standards or if problems occur following initial marketing.

FDA regulations require that products be manufactured in specific approved facilities and in accordance with cGMP regulations. These manufacturers must comply with cGMP regulations that require, among other things, quality control and quality assurance, the maintenance of records and documentation, and the obligation to investigate and correct any deviations from cGMP.

Manufacturers and other entities involved in the manufacture and distribution of approved drugs or biologics 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 requirements and other laws. Accordingly, manufacturers must continue to expend time, money and effort in the area of production and quality control to maintain cGMP compliance. The discovery of violative conditions, including failure to conform to cGMP regulations, could result in enforcement actions, and the discovery of problems with a product after approval may result in restrictions on a product, manufacturer or holder of an approved NDA/BLA, including recall.

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Once an approval is granted, the FDA may issue enforcement letters or withdraw the approval of the product if compliance with regulatory requirements and standards is not maintained or if problems occur after the drug or biologic reaches the market. Corrective action could delay drug or biologic distribution and require significant time and financial expenditures. Later discovery of previously unknown problems with a drug or biologic, including AEs 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 or other restrictions under a REMS program. Other potential consequences include, among other things:

- restrictions on the marketing or manufacturing of the drug or biologic, suspension of the approval, complete withdrawal of the drug from the market or product recalls;
- fines, warning letters or holds on post-approval clinical trials;
- refusal of the FDA to approve applications or supplements to approved applications, or suspension or revocation of drug or biologic approvals; drug or biologic seizure or detention, or refusal to permit the import or export of drugs; or
- injunctions or the imposition of civil or criminal penalties.

Patent Term Extension

After approval, owners of relevant drug or biological product patents may apply for up to a five-year patent extension to restore a portion of patent term lost during product clinical development and FDA review of an NDA or a BLA if approval of the application is the first permitted commercial marketing or use of a biologic containing the active ingredient under the Drug Price Competition and Patent Term Restoration Act of 1984, commonly referred to as the Hatch-Waxman Act. The allowable patent term extension is calculated as one-half of the product’s testing phase, which is the time between IND and NDA/BLA submission, and all of the review phase, which is the time between NDA/BLA submission and approval, minus any applicant delays, up to a maximum of five years. The time can be shortened if the FDA determines that the applicant did not pursue approval with due diligence. The total patent term after the extension may not exceed more than 14 years from the date of FDA approval of the product. Only one patent related to claiming each approved product is eligible for term extension, only those claims covering the approved product, a method for using it, or a method for manufacturing it may be extended, and the patent holder must apply for extension within 60 days of approval. The United States Patent and Trademark Office, in consultation with the FDA, reviews and approves the application for patent term restoration. For patents that might expire during the application phase, the patent owner may request an interim patent term extension. An interim patent term extension increases the patent term by one year and may be renewed up to four times. For each interim patent term extension granted, the post-approval patent term extension is reduced by one year. The director of the United States Patent and Trademark Office must determine that approval of the drug candidate covered by the patent for which a patent term extension is being sought is likely. Interim patent term extensions are not available for a drug candidate for which an NDA or a BLA has not been submitted.