ASCENTAGE PHARMA GROUP INTERNATIONAL

亞盛醫藥集團

(Incorporated in the Cayman Islands with limited liability)

(Stock Code: 6855)

Voluntary Announcement

Ascentage Pharma's Novel Potent EED Inhibitor APG-5918 Obtained IND Clearance by the US FDA and will launch First-in-Human Study

Ascentage Pharma Group International (the "**Company**" or "**Ascentage Pharma**") is pleased to announce that the novel inhibitor of the embryonic ectoderm development (EED) protein under its development, APG-5918, has been cleared by the US Food and Drug Administration (FDA) to launch a first-in-human (FIH) study that will assess the safety, pharmacokinetics, and preliminary efficacy of APG-5918 in patients with late stage solid tumors or hematologic malignancies. Professor Joseph Paul Eder, Clinical Director of the Early Drug Development Program at Yale Cancer Center will be the Principal Investigator of this multicentric clinical trial.

This multicenter, open-label Phase I clinical study is designed to assess the safety and tolerability, and determine the dose-limiting toxicity, maximum tolerated dose (MTD), and recommended Phase II dose (RP2D), of orally administered APG-5918 in patients with late stage solid tumors or hematologic malignancies.

EZH2, which is highly expressed in multiple tumors in humans, was found to promote the development and progression of tumors, making the targeted inhibition of EZH2's methyltransferase activity an effective mechanistic approach for cancer treatment. However, the secondary mutation of EZH2 may lead to acquired drug resistances, while the homologous EZH1 also has methyltransferase activity that could limit the effects of EZH2 inhibitors. Furthermore, EED protein can stimulate the methyltransferase activity of EZH2, thus making the allosteric targeting of EED an effective approach for PRC2 inhibition. EED inhibitors have shown good therapeutic potential in many kinds of solid tumors and hematologic malignancies.

Discovered and developed by Ascentage Pharma, APG-5918 is an orally active, novel, potent, selective, small-molecule EED inhibitor with high binding affinity. As an allosteric inhibitor, APG-5918 selectively binds to the EED protein. By regulating tumor epigenetics and microenvironment, APG-5918 can potentially overcome tumor resistance and deliver complete and durable tumor regression.

About Ascentage Pharma

Ascentage Pharma is a China-based, globally focused, clinical-stage biotechnology company engaged in developing novel therapies for cancers, CHB (Chronic hepatitis B), and age-related diseases. On October 28, 2019, Ascentage Pharma became listed on the Main Board of The Stock Exchange of Hong Kong Limited with the stock code: 6855.HK.

Ascentage Pharma has its own platform for developing therapeutics that inhibit protein-protein interactions to restore apoptosis or programmed cell death. The Company has built a pipeline of eight type I small molecule clinical drug candidates which have entered the clinical development stage, including novel, highly potent Bcl-2, and dual Bcl-2/Bcl-xL inhibitors, as well as candidates aimed at IAP and MDM2-p53 pathways, and next-generation tyrosine kinase inhibitors (TKIs). Ascentage Pharma is also the only company in the world with active clinical programs targeting all three known classes of key apoptosis regulators. The Company is conducting more than 50 Phase I/II clinical trials in China, the US, Australia and Europe. Olverembatinib, the Company's core drug candidate developed for the treatment of drug-resistant chronic myeloid leukemia (CML), was granted Priority Review status and a Breakthrough Therapy Designation (BTD) by the Center for Drug Evaluation (CDE) of China National Medical Products Administration (NMPA), and is already approved for the indication. In addition, Olverembatinib has also been granted an Orphan Drug Designation (ODD) and a Fast Track Designation (FTD) by the US FDA, and an Orphan Designation by the EU. As at the date of this announcement, Ascentage Pharma has obtained a total of 15 ODDs from the US FDA and 1 ODD from the EU for four of the Company's investigational drug candidates. The Company has been designated for multiple major national R&D projects in China, including five Major New Drug Development Projects, one Enterprise Innovative Drug Incubator Base status, four Innovative Drug Research and Development Programs, and one Major Project for the Prevention and Treatment of Infectious Diseases.

Leveraging its robust research and development capabilities, Ascentage Pharma has built a portfolio of global intellectual property rights, and entered into global partnerships with numerous leading biotechnology and pharmaceutical companies and research institutes such as UNITY Biotechnology, MD Anderson Cancer Center, Mayo Clinic, Dana-Farber Cancer Institute, MSD, AstraZeneca and Pfizer. The Company has built a global and talented team with experience in the research and development of innovative drugs and clinical development, and is setting up its commercial manufacturing and sales and marketing teams with high standards. Ascentage Pharma aims to continuously strengthen its research and development capabilities and accelerate the clinical development progress of its product pipeline to fulfil its mission of 'addressing unmet clinical needs of patients in China and around the world' for the benefit of more patients.

Cautionary Statement required by Rule 18A.05 of the Rules Governing the Listing of Securities on The Stock Exchange of Hong Kong Limited: We cannot guarantee that we will be able to obtain further approval for, or ultimately market APG-5918 successfully.

By order of the Board
Ascentage Pharma Group International
Dr. Yang Dajun
Chairman and Executive Director

Suzhou, People's Republic of China, June 29, 2022

As at the date of this announcement, the Board of Directors of the Company comprises Dr. Yang Dajun as Chairman and executive Director, Dr. Wang Shaomeng and Dr. Lu Simon Dazhong as non-executive Directors, and Mr. Ye Changqing, Dr. Yin Zheng, Mr. Ren Wei and Dr. David Sidransky as independent non-executive Directors.