亞盛醫藥
Ascentage Pharma

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 Announces that the China NMPA Accepted and Granted Priority Review Designation to a New Drug Application for Olverembatinib for the Treatment of Drug-Resistant CML

Ascentage Pharma Group International (the "**Company**" or "**Ascentage Pharma**") is pleased to announce that the Center for Drug Evaluation (CDE) of China National Medical Products Administration (NMPA) has accepted and granted Priority Review designation (the public notice period of which ended on July 18, 2022) to a New Drug Application (NDA) that will support the full approval of olverembatinib in patients with chronic-phase chronic myeloid leukemia (CML-CP) who are resistant and/or intolerant of first-and second-generation tyrosine kinase inhibitors (TKIs). Following the conditional NDA approval in November 2021, the acceptance for the latest application marks another milestone and will potentially bring olverembatinib to benefit a broader population of patients with chronic myeloid leukemia (CML). Ascentage Pharma and Innovent Biologics, Inc. are mutually committed to the commercialization of olverembatinib in China market.

In November 2021, the NMPA granted a conditional approval to olverembatinib for the treatment of adult patients with TKI-resistant CML-CP or CML-AP harboring the T315I mutation as confirmed by a validated diagnostic test, thus filling an urgent treatment gap that have long hindered the clinical outcome in Chinese patients with TKI-resistant CML harboring the T315I mutation.

The acceptance and Priority Review designation for this application are based on the results from an open-label, randomized, controlled, confirmatory Phase II pivotal study (HQP1351CC203) which previously served as the basis for a Breakthrough Therapy designation to olverembatinib by the CDE in March 2021. The study is designed to evaluate the efficacy and safety of olverembatinib in patients with CML-CP who are resistant and/or intolerant to first-and second-generation TKIs, with the event-free survival (EFS) as its primary endpoint. A total of 144 patients were enrolled and randomized to either receive olverembatinib or the control group to receive the current best available treatment (BAT). Results show that olverembatinib significantly improved the EFS compared to the control group and has met the pre-specified superiority criteria. Detailed results from this study will be released at an upcoming relevant academic conference.

CML is a hematologic malignancy of the white blood cells. The introduction of BCR-ABL TKIs have significantly improved the clinical practice of CML. However, acquired resistance to TKIs remains a major challenge in the treatment of CML. BCR-ABL tyrosine kinase mutations represent a key mechanism of acquired drug resistance and there is an urgent unmet medical need for a safe and effective treatment option. Olverembatinib is a potentially global best-in-class drug that has been developed by Ascentage Pharma and supported by the National Major New Drug Discovery and Manufacturing Program in China. As the first and only third-generation BCR-ABL inhibitor approved for the treatment of drug-resistant CML in China, olverembatinib is able to effectively target a spectrum of BCR-ABL mutants, including T315I.

About Olverembatinib

Developed by Ascentage Pharma with support from the National Major New Drug Discovery and Manufacturing Program in China, the orally active, third-generation BCR-ABL inhibitor olverembatinib is the first China-approved third-generation BCR-ABL inhibitor targeting drug-resistant CML. Olverembatinib can effectively target a spectrum of BCR-ABL mutants, including the T315I mutation.

In November 2021, olverembatinib was approved by the NMPA of China for the treatment of adult patients with TKI-resistant CML-CP or CML-AP harboring the T315I mutation as confirmed by a validated diagnostic test. In March 2021, it was granted the Breakthrough Therapy designation by the CDE for the treatment of patients with CML-CP who are resistant and/or intolerant of first-and second-generation TKIs.

In overseas, olverembatinib was cleared by the US FDA in July 2019 to directly enter a Phase Ib study. Since 2018, the clinical results of olverembatinib have been selected for oral presentations at the American Society of Hematology (ASH) Annual Meetings for four consecutive years, and was nominated for "Best of ASH" in 2019. As at the date of this announcement, olverembatinib has been granted one Fast Track designation and three Orphan Drug designations from the US FDA for the treatment of CML, acute lymphoblastic leukemia (ALL), and acute myeloid leukemia (AML); and one Orphan Designation from the European Medicines Agency (EMA) of the European Union for the treatment of CML.

In July 2021, Ascentage Pharma (6855.HK) and Innovent Biologics, Inc. (1801.HK) reached the agreement regarding the joint development and commercialization of olverembatinib in China.

* *Olverembatinib has not been approved for any indication in the U.S.*

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.

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

Suzhou, People's Republic of China, July 19, 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.