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 Releases Encouraging Clinical Results of Bcl-2 Inhibitor Lisaftoclax (APG-2575) in Chinese Patients with Relapsed/Refractory Non-Hodgkin Lymphomas

Ascentage Pharma Group International (the "**Company**" or "**Ascentage Pharma**") is pleased to announce that it has released results from a Phase I clinical study of the Company's novel Bcl-2 inhibitor lisaftoclax (APG-2575) in Chinese patients with relapsed/refractory non-Hodgkin lymphomas (r/r NHLs) at the 2022 European Hematology Association Hybrid Congress (EHA 2022).

The EHA Congress is the largest international gathering of the hematology field in Europe. It attracts over 10,000 clinical experts and researchers from over 100 countries every year to share and explore innovative ideas and latest scientific and clinical research results in relation to hematology.

The data presented at this year's EHA Congress show that lisaftoclax was well tolerated at doses of up to 800 mg/day, without evidence of tumor lysis syndrome (TLS). In addition, lisaftoclax demonstrated preliminary efficacy in a range of relapsed/refractory hematologic malignancies, including chronic lymphocytic leukemia/small lymphocytic lymphoma (CLL/SLL), marginal zone lymphoma (MZL), mantle cell lymphoma (MCL), and T-cell NHL, having achieved 4 complete responses (CRs) and 8 partial responses (PRs) in 32 efficacy evaluable patients. In the 11 efficacy evaluable patients with CLL (all of whom were heavily pretreated and had failed prior therapies such as chemoimmunotherapies and Bruton's tyrosine kinase BTK inhibitors, and the majority had at least 1 type of adverse prognostic

factors such as 17p deletion/TP53 mutation), there were 8 efficacy evaluable patients in cohorts received 200 mg or higher doses, including 3 CRs and 4 PRs, thus demonstrating an ORR of 87.5%. In the 6 efficacy evaluable patients with MCL, the ORR was 33.3% (1 CR). In the 4 efficacy evaluable patients with MZL, the ORR was 50%. In the 3 efficacy evaluable patients with T-cell NHL, the ORR was 33.3%.

Lisaftoclax (APG-2575) is a novel, orally administered small-molecule Bcl-2 selective inhibitor being developed by Ascentage Pharma to treat hematologic malignancies and solid tumors by selectively blocking the antiapoptotic protein Bcl-2 and hence restoring the normal apoptosis process in cancer cells. Lisaftoclax (APG-2575) is the first China-developed Bcl-2 inhibitor entering clinical development in China, and the second entering pivotal studies globally. Lisaftoclax (APG-2575) is being studied in multiple clinical studies in a range of solid tumors and hematologic malignancies, and has shown enormous clinical potential.

Highlights of this abstract on lisaftoclax (APG-2575) are as follows:

Preliminary Results of a Phase 1 Study of Novel Bcl-2 Inhibitor Lisaftoclax (APG-2575) in Chinese Patients (pts) with Relapsed or Refractory (r/r) Non-Hodgkin Lymphomas (NHLs)

Abstract number: P1106

Highlights:

This multicenter, single-agent, Phase I trial consists of a dose escalation and a dose expansion and is the first-in-human study of lisaftoclax in adults with R/R NHL in China. The objective of this study was to evaluate the safety, tolerability, pharmacokinetics (PK), pharmacodynamics (PD), and preliminary efficacy of lisaftoclax in Chinese patients with R/R NHL.

Patients were dosed with lisaftoclax orally once daily in 28-day cycles. A daily ramp-up schedule was used in patients with CLL or NHL with medium/high-risk TLS.

As of January 1, 2022, 40 patients had been treated with lisaftoclax (dose range 20–800 mg), including 20 who remain in the trial. NHL subtypes included: 12 CLL, 9 follicular lymphoma (FL), 7 MCL, 4 MZL, 4 T-cell lymphoma, 2 diffuse large B-cell lymphoma, and 2 Waldenström macroglobulinemia.

With a median treatment duration of 4 cycles, 4 of 32 evaluable patients achieved CR and 8 achieved PR, resulting in an ORR of 37.5%. In the 11 evaluable patients with CLL, the ORR was 63.6%, the CR rate was 27.3%, and the PR rate was 36.4%. In patients with CLL treated at doses ≥ 200 mg, the ORR was 87.5%. In the 6 evaluable patients with MCL, 1 patient achieved CR and 1 achieved PR. In the 4 evaluable patients with MZL, 2 achieved PR. In the 3 evaluable patients with T-cell NHL, 1 achieved PR.

Lisaftoclax was generally well tolerated. Most treatment-emergent adverse events (TEAEs) were grade 1–2 (67.5%). Dose-limiting toxicity (DLT) and laboratory/clinical TLS were not observed in any of the dose cohorts, and there were no dose-reduction or discontinuation due to intolerance.

Conclusions:

Lisaftoclax showed promising antitumor activity and favorable responses in subtypes of NHL including CLL/SLL, MZL, MCL, T-cell NHL. In addition, lisaftoclax demonstrated a favorable safety profile, with no TLS or DLT observed during the dose ramp-up from 20 mg–800 mg/day.

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-2575 successfully.

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

Suzhou, People's Republic of China, June 13, 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.