Shanghai Henlius Biotech, Inc.
上海復宏漢霖生物技術股份有限公司
(A joint stock company incorporated in the People's Republic of China with limited liability)
(Stock code: 2696)

VOLUNTARY ANNOUNCEMENT

THE FIRST SUBJECT HAS BEEN DOSED IN A PHASE 1 CLINICAL STUDY IN HEALTHY CHINESE MALE SUBJECTS OF IPILIMUMAB BIOSIMILAR HLX13 (RECOMBINANT ANTI-CTLA-4 FULLY HUMAN MONOCLONAL ANTIBODY INJECTION)

A. INTRODUCTION

This announcement is made by Shanghai Henlius Biotech, Inc. (the "**Company**") on a voluntary basis to inform the shareholders and potential investors of the Company about the latest business development of the Company.

The board of directors of the Company (the "**Board**") is pleased to announce that, recently, the first subject has been dosed in a phase 1 clinical study in healthy Chinese male subjects of Ipilimumab biosimilar HLX13 (recombinant anti-CTLA-4 fully human monoclonal antibody injection) ("**HLX13**") independently developed by the Company.

B. CLINICAL TRIAL DESIGN AND PURPOSE

This randomised, single intravenous infusion, phase 1 parallel study aims to compare the pharmacokinetics, safety, and immunogenicity between HLX13 and ipilimumab (YERVOY®, US-, EU-, and China-approved) in healthy Chinese adult male subjects. This study consists of two parts. Part 1 is an open-label, randomised, single-dose, two-arm, parallel study aiming to evaluate the pharmacokinetic parameters, safety, and immunogenicity after single intravenous infusion of HLX13 or EU-approved YERVOY®, and to optimise the study design of Part 2. Part 2 is a double-blind, randomised, single-dose, four-arm, parallel study aiming to evaluate the similarity in pharmacokinetics among HLX13 and US-, EU-, and China-approved YERVOY®, as well as their safety and immunogenicity. Subjects in the respective groups will receive 0.3 mg/kg of HLX13, or 0.3 mg/kg of US-, EU-, or China-approved YERVOY® via intravenous infusion. The primary endpoints of this study are the area under the serum concentration-time curve from time 0 to infinity ($AUC_{0\text{-}inf}$) and the maximum serum concentration (C_{max}). Secondary endpoints include other pharmacokinetic parameters, safety, and immunogenicity.

C. ABOUT HLX13

HLX13 is an Ipilimumab biosimilar independently developed by the Company, and is intended to be used for the treatment of Melanoma, Renal Cell Carcinoma, Colorectal Cancer, Hepatocellular Carcinoma, Non-Small Cell Lung Cancer, Malignant Pleural Mesothelioma and Esophageal Squamous Cell Cancer. Ipilimumab is a fully human, anti-CTLA-4 (cytotoxic T-lymphocytes associated antigen 4, also known as CD152), IgG1 monoclonal antibody with κ light chain. CTLA-4 mainly expresses in regulatory T cells (Treg) and activated T cells and is designed to inhibit the growth of T cells and the creation of the cytokine (IL-2 and IFN-γ) by competing with CD28 for the B7 ligands (B7-1 and B7-2) attached to antigen-presenting cells. By blocking the binding between CTLA-4 and the ligands, Ipilimumab can elevate the immune response and in turn achieve the goal of killing tumors. According to the Technical Guidelines for R&D and Evaluation of Biosimilars (Trial), the Company conducted comprehensive comparability studies between HLX13 and reference product Ipilimumab in terms of the chemical manufacture and control (CMC), pre-clinical pharmacology, toxicology and pharmacokinetics. Results from these studies showed that there is a high similarity or no significant difference between HLX13 and reference product Ipilimumab.

D. MARKET CONDITION

As at the date of this announcement, the Ipilimumab commercially available in mainland China (excluding Hong Kong, Macau and Taiwan regions, same as below) includes Yervoy® of Bristol-Myers Squibb only (approved for commercialization in June 2021). According to the information of IQVIA CHPA and IQVIA MIDAS™ (IQVIA is a global provider of professional information and strategic consulting services in the pharmaceutical and healthcare industry), the sales of Ipilimumab in mainland China and worldwide in 2022 are approximately RMB10.98 million and US$2.346 billion, respectively.

WARNING STATEMENT WITH REFERENCE TO THE REQUIREMENTS UNDER RULE 18A.05 OF THE RULES GOVERNING THE LISTING OF SECURITIES ON THE STOCK EXCHANGE OF HONG KONG LIMITED: The Company cannot guarantee the successful development and commercialization of HLX13. Shareholders and potential investors of the Company are advised to exercise caution when dealing in the shares of the Company.

On behalf of the Board
Shanghai Henlius Biotech, Inc.
Wenjie Zhang
Chairman

Hong Kong, 13 December 2023

As at the date of this announcement, the board of directors of the Company comprises Mr. Wenjie Zhang as the chairman and executive director, Mr. Jun Zhu as the executive director, Mr. Qiyu Chen, Mr. Yifang Wu, Ms. Xiaohui Guan, Mr. Deyong Wen and Dr. Xingli Wang as the non-executive directors, and Mr. Tak Young So, Dr. Lik Yuen Chan, Dr. Guoping Zhao and Dr. Ruilin Song as the independent non-executive directors.