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### Sihuan Pharmaceutical Holdings Group Ltd.

四環醫藥控股集團有限公司 (incorporated in Bermuda with limited liability) (Stock Code: 0460)

# VOLUNTARY ANNOUNCEMENT PHASE III CLINICAL TRIALS OF XUANZHU BIOPHARM'S CLASS 1 INNOVATIVE DRUG BIROCICLIB REACHED PRIMARY ENDPOINT

The board of directors (the "**Board**") of Sihuan Pharmaceutical Holdings Group Ltd. (the "**Company**" or "**Sihuan Pharmaceutical**", together with its subsidiaries, the "**Group**") is pleased to announce that the Phase III clinical trial of Birociclib (XZP-3287, CDK4/6 inhibitor) ("**Birociclib**"), a class 1 innovative drug under development by the Group's non-wholly owned subsidiary Xuanzhu Biopharmaceutical Co., Ltd. ("**Xuanzhu Biopharm**"), used in combination with Fulvestrant in advanced breast cancer patients with hormone receptor-positive (HR+)/human epidermal growth factor receptor 2 negative(HER2-) who progressed from endocrine therapy, reached its primary endpoint. In addition, the Phase II clinical trial application of Birociclib used in combination with abitherone acetate and prednisone in the treatment of advanced or metastatic prostate cancer has been accepted by the Center for Drug Evaluation of the National Medical Products Administration of the People's Republic of China ("**China**").

Xuanzhu Biopharm recently held the Independent Data Monitoring Committee (IDMC) meeting to review the interim analysis data of the Phase III clinical trial (Birociclib combined with Fulvestrant). IDMC experts unanimously agreed that the interim analysis results showed that the trial reached the preset boundary of interim analysis, which showed significant statistical differences and clinical significance compared to the control group. It is recommended to submit marketing application earlier. In addition, Birociclib is also the first domestically developed CDK4/6 inhibitor to be planned for clinical research in the treatment of advanced prostate cancer.

#### **About Birociclib**

The phase I clinical data demonstrated that Birociclib, a novel selective CDK4/6 inhibitor, has the potential to overcome endocrine therapy resistance in hormone receptor-positive (HR+) breast cancer patients. Birociclib monotherapy also exhibited efficacy in patients with advanced breast cancer who have failed in multiple lines of treatment. Pre-clinical data showed that Birociclib has a unique pharmacokinetic profile, which allows it to go through the blood-brain barrier effectively, and thus it is expected to have potential efficacy in breast cancer patients with brain metastases and brain tumor patients. Moreover, due to the novel targeting mechanism of CDK4/6, Birociclib is able to demonstrate significant clinical benefits and extensive market prospect through multiple novel Birociclib-based combination therapies.

#### About Xuanzhu Biopharm

Xuanzhu Biopharm is the innovative drug subsidiary of Sihuan Pharmaceutical. It is an innovative pharmaceutical company with roots in China and a global perspective, focusing on major diseases such as digestion, oncology and non-alcoholic steatohepatitis, and is committed to the continuous research and development, production and commercialization of class 1 drugs with core proprietary intellectual property rights to address unmet clinical medical needs. The company has a team with extensive experience in the development and industrialisation of innovative drugs. Having been involved in the research of digestion, oncology and non-alcoholic steatohepatitis for many years, the company has a deep understanding and international vision of the development of new drugs in related fields and their future development direction. The company has two R&D platforms: small molecule chemistry and large molecule biologics. The dual engines drive the company's innovation and development, forming a product pipeline that covers small molecule, monoclonal antibodies, bispecific antibodies, antibody-drug conjugate (ADC) and other types of products. The company adheres to the strategic concept of "innovation driven, promoting the development of new drugs in China and serving human health", and the values of "open innovation, courageous commitment, overcoming difficulties and scientific rigor". With a focus on unmet clinical needs, the company continues to develop new class 1 drugs with international competitiveness, and is committed to developing into a first-class innovative pharmaceutical company with independent research and development, production, and sales capabilities.

## **About Sihuan Pharmaceutical**

Founded in 2001 and listed on the Main Board of The Stock Exchange of Hong Kong Limited in 2010, Sihuan Pharmaceutical is an international medical aesthetic and biopharmaceutical company led and driven by innovation, with an independent and leading research and development technology platform, a rich global product pipeline, strong product registration capability, a full dosage form production platform with high efficiency and low cost and a mature and excellent sales system. Adhering to the overall strategic objective of "full promotion of a two-wheel drive strategy of its medical aesthetics and biopharmaceutical businesses", Sihuan Pharmaceutical endeavours to build itself into a leading medical aesthetics and biopharmaceutical company in China.

This announcement is being made by the Group on a voluntary basis to update the investing public on the Group's latest business development, and does not constitute, and is not intended to be, an advertisement regarding the use of any medicine, surgical appliance, treatment or orally consumed product.

> By order of the Board Sihuan Pharmaceutical Holdings Group Ltd. Dr. Che Fengsheng Chairman and Executive Director

Hong Kong, 26 April 2023

As at the date of this announcement, the executive directors of the Company are Dr. Che Fengsheng (Chairman), Dr. Guo Weicheng (Deputy Chairman and Chief Executive Officer), Dr. Zhang Jionglong, Ms. Chen Yanling and Ms. Miao Guili; and the independent non-executive directors of the Company are Mr. Tsang Wah Kwong, Dr. Zhu Xun and Mr. Wang Guan.