

*Hong Kong Exchanges and Clearing Limited and The Stock Exchange of Hong Kong Limited take no responsibility for the contents of this announcement, make no representation as to its accuracy or completeness and expressly disclaim any liability whatsoever for any loss howsoever arising from or in reliance upon the whole or any part of the contents of this announcement.*



**Antengene Corporation Limited**

**德琪醫藥有限公司**

*(Incorporated in the Cayman Islands with limited liability)*

**(Stock Code: 6996)**

**VOLUNTARY ANNOUNCEMENT**

**FDA APPROVAL OF XPOVIO® (SELINEXOR) AS A TREATMENT  
FOR PATIENTS WITH MULTIPLE MYELOMA  
AFTER AT LEAST ONE PRIOR THERAPY**

Antengene Corporation Limited (the “**Company**”, together with its subsidiaries, the “**Group**”) hereby informs the shareholders and potential investors of the Company of the attached press release that Karyopharm Therapeutics Inc., the Company’s partner in the United States (the “**U.S.**”), has received the approval of XPOVIO® (selinexor) by the U.S. Food and Drug Administration (the “**FDA**”) as a treatment for patients with multiple myeloma after at least one prior therapy.

This is a voluntary announcement made by the Company. The Group cannot guarantee that XPOVIO® (selinexor) will ultimately be successfully developed and marketed. Shareholders and potential investors of the Company are advised to exercise caution when dealing in the shares of the Company.

By order of the Board  
**Antengene Corporation Limited**  
**Dr. Jay Mei**  
*Chairman*

Hong Kong, December 20, 2020

*As at the date of this announcement, the board of directors of the Company comprises Dr. Jay Mei, Mr. John F. Chin and Mr. Yiteng Liu as executive directors; Mr. Xubo Hu, Mr. Zhen Li and Mr. Yanling Cao as non-executive directors; and Mr. Mark J. Alles, Ms. Jing Qian and Mr. Sheng Tang as independent non-executive directors.*

## **Antengene Announces its U.S. Partner, Karyopharm Therapeutics Inc., has received FDA Approval of XPOVIO® (Selinexor) as a Treatment for Patients with Multiple Myeloma After At Least One Prior Therapy**

- Oral XPOVIO® Approval as Combination Therapy in Patients with Multiple Myeloma After At Least One Prior Therapy Significantly Expands the XPOVIO® Addressable Patient Population –
- Oral XPOVIO® is Now the Only Approved Multiple Myeloma Drug Indicated as Part of a Once-Weekly Bortezomib Combination Regimen –
- First Multiple Myeloma Drug with a New Mechanism of Action Approved by the FDA in the Second-Line Setting Since 2016 –
- FDA Approval Comes Approximately Three Months Ahead of Target PDUFA Date –

Shanghai and Hong Kong, PRC, December 20, 2020 – Antengene Corporation Limited (“**Antengene**”, SEHK: 6996.HK) today announced that the U.S. Food and Drug Administration (FDA) has approved the supplemental New Drug Application (sNDA) submitted by its partner Karyopharm Therapeutics Inc. (Nasdaq: KPTI) for oral XPOVIO® (selinexor, ATG-010), a first-in-class, oral Selective Inhibitor of Nuclear Export (SINE) medicine, in combination with bortezomib and dexamethasone for the treatment of adult patients with multiple myeloma who have received at least one prior therapy. This indication was approved three months ahead of its March Prescription Drug User Fee Act (PDUFA) date based on the result of a confirmatory Phase 3 BOSTON study.

XPOVIO® was previously approved under the FDA’s Accelerated Approval Program for the treatment of adult patients with relapsed or refractory multiple myeloma (rrMM) who have received at least four prior therapies and whose disease is refractory to at least two proteasome inhibitors, at least two immunomodulatory agents, and an anti-CD38 monoclonal antibody.

XPOVIO®, a first-in-class and only-in-class oral SINE compound, and now the first and only drug approved by FDA for use in both multiple myeloma and diffuse large B-cell lymphoma, discovered and developed by Karyopharm, is currently being developed by Antengene, who has the exclusive development and commercial rights in certain Asia-Pacific markets, including China. In December 2020, National Comprehensive Cancer Network (NCCN®) added three different XPOVIO® combination regimens to its Clinical Practice Guidelines in Oncology (NCCN® Guidelines) for previously treated multiple myeloma.

Antengene has conducted two Phase 2 registrational clinical trials of XPOVIO® in China for relapsed or refractory multiple myeloma (MARCH) and for relapsed or refractory diffuse large B-cell lymphoma (SEARCH). A Phase 3 randomized, controlled, open-label and multicenter BENCH trial has also received the investigational new drug approval from the National Medical Products Administration (NMPA) in China. Antengene is working on making XPOVIO® commercially available in Asia Pacific regions and has submitted the New Drug Applications (NDAs) for XPOVIO® to the Health Sciences Authority (HSA) of Singapore and the Australian Therapeutic Goods Administration (TGA) for three indications, including the treatment of patients with multiple myeloma who have received at least one prior therapy.

“This is the third approved indication for XPOVIO<sup>®</sup>, highlighting its potential of clinical application and broad market prospects. XPOVIO<sup>®</sup> offers patients with rrMM and their physicians the first oral SINE compound and makes more treatment options available to cancer patients with critical unmet medical needs.” Dr. Jay Mei, M.D., Ph.D., Founder, Chairman and CEO of Antengene said, “We plan to launch XPOVIO<sup>®</sup> as soon as possible for the treatment of patients with rrMM and rrDLBCL in APAC following regulatory approvals, by leveraging our established commercial infrastructure and seasoned international team.”

### **About the Phase 3 BOSTON Study**

The approval of XPOVIO<sup>®</sup> in combination with once-weekly bortezomib plus low-dose dexamethasone (SVd) is supported by the results of the multi-center, Phase 3, randomized study, which evaluated 402 adult patients with relapsed or refractory multiple myeloma who had received one to three prior lines of therapy. The study was designed to compare the efficacy, safety and certain health-related quality of life parameters of once-weekly XPOVIO<sup>®</sup> (selinexor) in combination with once-weekly bortezomib plus low-dose dexamethasone (SVd) versus twice-weekly bortezomib plus dexamethasone (Vd). The primary endpoint of the study was progression-free survival (PFS) and key secondary endpoints included overall response rate (ORR), rate of peripheral neuropathy, and others. Additionally, the BOSTON study allowed for patients on the Vd control arm to crossover to the SVd arm following objective (quantitative) progression of disease verified by an Independent Review Committee (IRC). The BOSTON study was conducted at over 150 clinical sites internationally.

Although the study had one of the highest proportions of patients with high-risk cytogenetics (~50%) as compared with other bortezomib studies in previously treated myeloma, the median PFS in the SVd arm was 13.9 months compared to 9.5 months in the Vd arm, representing a 4.4 month increase in median PFS (hazard ratio [HR] of 0.70; p=0.0075). The SVd group also demonstrated a significantly greater ORR compared to the Vd group (76.4% vs. 62.3%, p=0.0012). Importantly, SVd therapy compared to Vd therapy showed consistent PFS benefit and higher ORR across several important subgroups.

In addition, the following results favored SVd therapy as compared to Vd therapy:

- SVd therapy demonstrated a significantly higher rate of deep responses, defined as  $\geq$  Very Good Partial Response compared to Vd therapy (44.6% vs. 32.4%) as well as a longer median duration of response (20.3 months vs. 12.9 months). Additionally, 17% of patients on the SVd arm achieved a Complete Response or a Stringent Complete Response as compared to 10% of patients receiving Vd therapy. All responses were confirmed by an IRC.
- Peripheral neuropathy (PN) rates were significantly lower on SVd compared to Vd (32% vs. 47%). In addition, PN rates  $\geq$  Grade 2 were also significantly lower in the SVd arm compared to Vd (21% vs. 34%).

The most common adverse reactions were cytopenias, along with gastrointestinal and constitutional symptoms and were consistent with those previously reported from other selinexor studies. Most adverse reactions were manageable with dose modifications and/or standard supportive care. The most common non-hematologic adverse reactions were nausea (50%), fatigue (42%), decreased appetite (35%), and diarrhea (32%) and were mostly Grade 1 and 2 events. The most common Grade 3 and 4 adverse reactions were thrombocytopenia (43%), lymphopenia (38%), anemia (17%), and fatigue (13%).

## **About XPOVIO® (selinexor, ATG-010)**

XPOVIO® (selinexor, ATG-010), a first-in-class and only-in-class oral selective inhibitor of nuclear export compound discovered and developed by Karyopharm, is currently being developed by Antengene, who has the exclusive development and commercial rights in certain Asia-Pacific markets, including China. In July 2019, the US Food and Drug Administration (FDA) approved selinexor (XPOVIO®) in combination with low-dose dexamethasone for the treatment of relapsed/refractory multiple myeloma (rrMM) and in June 2020 approved selinexor (XPOVIO®) as a single-agent for the treatment of relapsed/refractory diffuse large B-cell lymphoma (rrDLBCL). A Marketing Authorization Application (MAA) has also been submitted to the European Medicines Agency (EMA) with a request for conditional approval of selinexor (XPOVIO®) in this same rrMM indication. On December 18, 2020, the supplemental New Drug Application (sNDA) requesting an expansion of its indication to include the treatment for patients with multiple myeloma after at least one prior therapy was approved by the FDA. Selinexor (XPOVIO®) is so far the first and only oral SINE compound approved by the FDA. Selinexor (XPOVIO®) is also being evaluated in several other mid-and later-phase clinical trials across multiple solid tumor indications, including liposarcoma and endometrial cancer. In November 2020, at the Connective Tissue Oncology Society 2020 Annual Meeting (CTOS 2020), Antengene's partner, Karyopharm, presented positive results from the Phase 3 randomized, double blind, placebo controlled, cross-over SEAL study evaluating single agent, oral selinexor (XPOVIO®) versus matching placebo in patients with liposarcoma. Karyopharm also recently announced that the ongoing Phase 3 SIENDO study of selinexor (XPOVIO®) in patients with endometrial cancer passed planned interim futility analysis and that Data and Safety Monitoring Board (DSMB) recommended the study should proceed as planned without any modifications. Top-line SIENDO study results are expected in the second half of 2021.

Antengene is conducting two registrational Phase 2 clinical trials of selinexor (XPOVIO®) in China for relapsed/refractory multiple myeloma (MARCH) and for relapsed/refractory diffuse large B-cell lymphoma (SEARCH), and has initiated clinical trials for high prevalence cancer types in the Asia Pacific region including peripheral T-cell lymphoma and NK/T-cell lymphoma (TOUCH) and KRAS-mutant non-small cell lung cancer (TRUMP).

## **About Antengene**

Antengene is a leading clinical-stage Asia-Pacific biopharmaceutical company focused on innovative oncology medicines. Antengene aims to provide the most advanced anti-cancer drugs to patients in China, the Asia Pacific Region and around the world. Since its establishment, Antengene has built a pipeline of 12 clinical and pre-clinical stage assets and obtained 11 investigational new drug approvals in Asia Pacific. The vision of Antengene is to "Treat Patients Beyond Borders". Antengene aims to address significant unmet medical needs by discovering, developing and commercializing first-in-class/best-in-class therapeutics.

## **Forward-looking Statements**

The forward-looking statements made in this article relate only to the events or information as of the date on which the statements are made in this article. Except as required by law, we undertake no obligation to update or revise publicly any forward-looking statements, whether as a result of new information, future events or otherwise, after the date on which the statements are made or to reflect the occurrence of unanticipated events. You should read this article completely and with the understanding that our actual future results or performance may be materially different from what we expect. In this article, statements of, or references to, our intentions or those of any of our Directors or our Company are made as of the date of this article. Any of these intentions may alter in light of future development.

- \* XPOVIO® is a registered trademark of Karyopharm Therapeutics Inc.
- \* NCCN® is a registered trademark of National Comprehensive Cancer Network.