

# ANTENGENE INVESTOR PRESENTATION

TREATING PATIENTS BEYOND BORDERS

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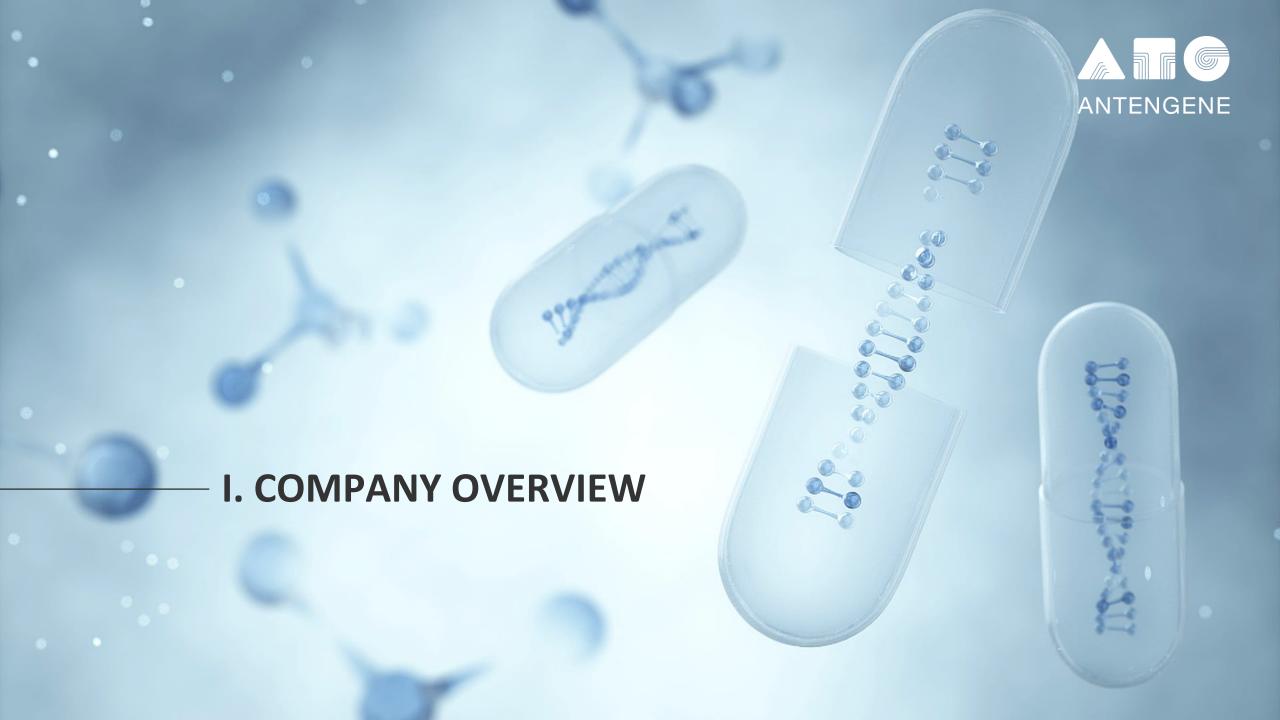
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# **Realizing Our Vision of Treating Patients Beyond Borders**



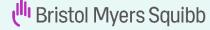
# **Commercialization in Multiple Markets**

- XPOVIO® approved in mainland China, Australia, Korea and Singapore
- Expecting approvals in Hong Kong and Taiwan markets in 2022
- ~188 person commercial team in Greater China and APAC

# **Clinical and Regulatory Operations**

- Multi-regional clinical trials with 24 INDs obtained across regions including mainland China, Australia, and US
- Studies ongoing in China, Australia and US including programs with wholly owned global rights

# **Global Partnerships**















**Globally** 





**Regions Expecting Selinexor Approval in 2022** 

Regions with Ongoing Clinical Trials

# **Global Team of Industry Veterans**



### Global Management Team with Deep Local Experience and Insights Working Seamlessly Across Regions











Eitan Liu

**Chief Operating Officer** 











Yiqiang Zhao, M.D., Ph.D.

**Executive Director, Clinical Development** 











Jasmine Sun, M.D., MPH

Corporate Vice President, Head of Clinical Operations













#### Zhinuan Yu, Ph.D.

Corporate Vice President, Biometrics & Regulatory **Enabling Functions** 











Jay Mei, M.D., Ph.D. Founder / Chairman / Chief Executive Officer











Bo Shan, Ph.D.

Chief Scientific Officer







Donald Lung, JD, MBA

**Chief Financial Officer** 



Yijun Yang, Ph.D., Sc.D

Corporate Vice President, Head of Clinical Enabling Functions & Operational Excellence











Godfrey Guo, M.D.

**Executive Director, Medical** 







**Thomas Karalis** 

Corporate Vice President, Head of Asia Pacific Markets













Lixin Yu

Head of Hematology Business Unit, China

















**Track Record of Antengene Management Team** 











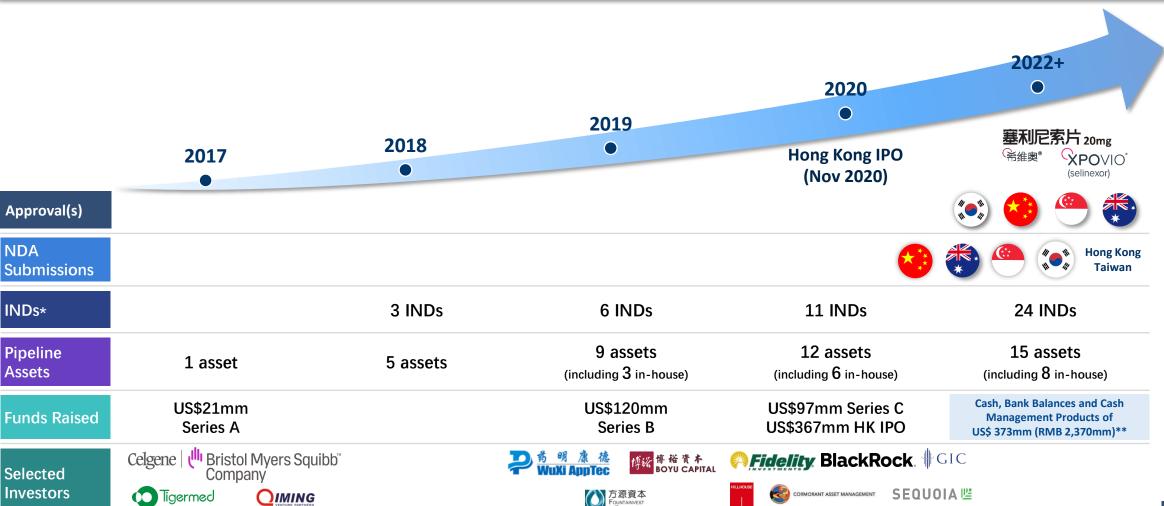




# **Dual Engine of Growth Through Value-adding Partnerships and In-house Discovery**



## Rich Pipeline with Commercialization Across Multiple Markets, Driven by Strong Execution Capabilities

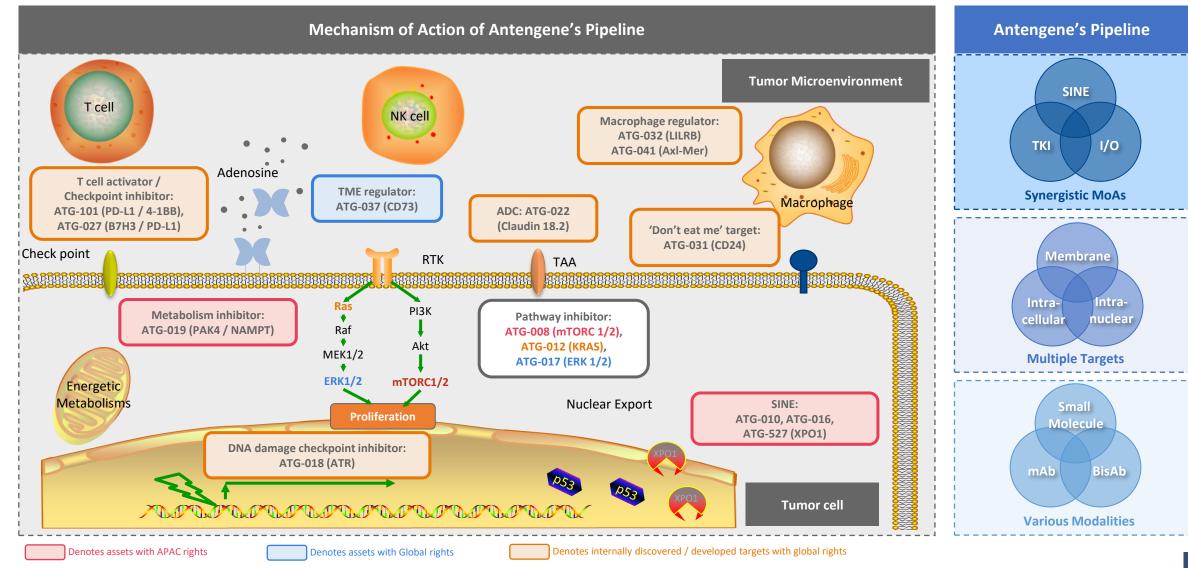


<sup>\*</sup> Total # of IND/CTA approvals obtained

<sup>\*\*</sup> As of 31st December 2021; USD/RMB exchange rate of 1/6.35 as of 17th March 2022

# The Antengene Pipeline Includes Multiple Novel Immuno-oncology and Targeted Assets – Allowing Broad Proprietary Combinations





# **2021** Achievements and Recent Corporate Updates



### Cash, Bank Balances and Cash Management Products – US\$ 373mm (RMB 2,370mm)

# Commercial Launch of XPOVIO® across Asia



- ✓ 2021 Revenue of **RMB28.8 mm**, primarily contributed by NPP Program
- ✓ NDA Approval by **China's NMPA** in Dec 2021
- ✓ NDA Approval by **Australia's TGA** and **Singapore's HSA** in Mar 2022
- ✓ NDA Approval by **South Korea's MFDS** in Jul 2021
- ✓ Expansion of commercial team to 188 members across APAC
- ✓ CSCO Diagnosis and Treatment Guidelines 2021 inclusion for multiple selinexor regimens in multiple myeloma and lymphoma

# **Late Stage Clinical Programs**



- √ ATG-010 Selinexor
  - 5 registrational studies on-going in mainland China for ATG-010 (selinexor), including 3 global trials in collaboration with Karyopharm
  - Data presentation in ASCO 2021 and ASH 2021

# **Global Rights Assets**



- √ 4 Global rights assets in clinical development
  - ATG-017 (ERK1/2 small molecule inhibitor)
  - ATG-101 (PD-L1/4-1BB bispecific antibody)
  - ATG-037 (CD73 small molecule inhibitor)
  - ATG-018 (ATR small molecule inhibitor)
- ✓ Research data presentation in multiple medical conferences
  - SITC Annual Meeting Nov 2021
    - o ATG-017 (ERK1/2 small molecule inhibitor)
    - ATG-101 (PD-L1/4-1BB bispecific antibody)
  - AACR Annual Meeting Apr 2022
    - o ATG-037 (CD73 small molecule inhibitor)
    - o ATG-018 (ATR small molecule inhibitor)
    - o ATG-022 (Claudin 18.2 ADC)
    - o ATG-012 (KRAS G12C small molecule inhibitor)
- √ 3 in-house discovered molecules with novel targets in the macrophage space disclosed
  - ATG-031 (CD24 monoclonal antibody)
  - ATG-032 (LILRB monoclonal antibody)
  - ATG-041 (Axl-Mer small molecule inhibitor)

## **Business Development**



- Announced global clinical collaboration with Bristol Myers Squibb to evaluate ATG-017 (ERK1/2 inhibitor) in combination with nivolumab
- In-licensed worldwide rights of a CD73 small molecule inhibitor from Calithera Biosciences
- Announced research collaboration with Legochem Biosciences

## **Corporate Operations**



- Inauguration of our manufacturing center at the Binhai Life Science and Healthcare Industrial Zone in Shaoxing
- Entered into a framework agreement for the construction of a drug discovery and manufacturing center for antibody biologics in Hangzhou Qiantang area

<sup>\*</sup> USD/RMB exchange rate of 1/6.35 as of 17th March 2022

<sup>\*\*</sup> Cash and bank balances of US\$ 358mm (RMB 2,275mm) and Cash Management Products of US\$ 15.1mm (RMB 95.7mm) as of 31st December 2021



# Pipeline of Near-to-midterm Drug Candidates with First-in-class / Best-in-class Potential



# of APAC Rights Assets:		5		of Antengene-Spor		16	# of Re	gistrational Trials:		5
Assets	Target (Modality)	Indication	Pre-clinical	Phase I	Phase II	Phase III	NDA	Commercialization	Antengene Rights	Partner
		R/R Multiple Myeloma	Combo with bortezomib and		··	*	E	STORM (US, SK, SG & AU NDA approved) BOSTON (US , SG & AU sNDA approved)		
	XPO1 (Small molecule)	R/R Diffuse Large B-cell Lymphoma	Monotherapy (SEARCH)  Combo with R-GDP (DLBCL-03	Ab and dexamethasone (STOM	*	*		SADAL (US, SK & SG sNDA approved)		
ATG-010 (Selinexor) <sup>1</sup>		R/R NHL R/R T-cell & NK-cell Lymphoma	Combo with lenalidomide + ri							Wan onbarm'
		Myelofibrosis  Maintenance Endometrial  Cancer	Monotherapy (MF 035)  Monotherapy (SIENDO)		*				APAC <sup>2</sup>	Karyopharm Cherapoutica
		Advanced Liposarcoma	Monotherapy (SEAL)							
		Recurrent Glioblastoma ND& Recurrent Glioblastoma	Monotherapy (KING)  Monotherapy (GBM-029)							
ATG-016 Eltanexor)	XPO1 (Small molecule)	R/R MDS	Monotherapy (HATCH)  Monotherapy (KCP-8602-801)							
Entanexory	(Smail molecule)	Advanced Solid Tumors	Monotherapy (REACH)		CRC PrC					
ATG-008 (Onatasertib)	mTORC1/2 (Small molecule)	2L+ HBV+ Hepatocellular Carcinoma Advanced Solid Tumors and Hepatocellular Carcinoma	Monotherapy (TORCH)  Combo with anti-PD-1 mAb (1	TORCH-2)*		<b>&gt;</b>			APAC <sup>3</sup>	Colgene ( Bristol Myers Squit Company
		R/R Diffuse Large B-cell Lymphoma	Combo with ATG-010 (MATCH	н)	•					ANTENGENE
ATG-527 (Verdinexor)	XPO1 (Small molecule)	Lupus, Anti-viral	Monotherapy (EVOLVE)						APAC <sup>2</sup>	Karyopharm Therapeutics
ATG-019 (KPT-9274)	PAK4/NAMPT (Small molecule)	Advanced Solid Tumors & NHL	Monotherapy <u>+</u> niacin (TEACH	H)					APAC	ANTENGENE

<sup>1 (</sup>s)NDA approved by US FDA, China NMPA, Australia TGA, South Korea MFDS and Singapore HSA; China Hong Kong and China Taiwan NDA submissions are completed;

\* Investigator-initiated trials; R/R = relapsed/refractory; ND = newly diagnosed; MDS = myelodysplastic syndrome; CRC = colorectal cancer; PrC = prostate cancer; CAEBV = chronic active Epstein-Barr virus; NHL = non-Hodgkin lymphoma; Hem/Onc = hematological malignancies and solid tumors; SK= South Korea; R-GDP: Rituximab, Gemcitabine, Dexamethasone &Cisplatin; in GBM-029 trial, the combination regimen is with standard of care (SoC) therapy for newly diagnosed glioblastoma or recurrent glioblastoma, including radiation therapy, temozolomide, lomustine, bevacizumab, tumor treating fields, or carmustine

<sup>&</sup>lt;sup>2</sup> Antengene has rights for Greater China (mainland China, Hong Kong, Taiwan, Macau), Australia, New Zealand, South Korea, and the ASEAN Countries;

<sup>&</sup>lt;sup>3</sup> Antengene has rights for Greater China, South Korea, Singapore, Malaysia, Indonesia, Vietnam, Laos, Cambodia, the Philippines, Thailand and Mongolia;

<sup>&</sup>lt;sup>4</sup> Most advanced trial status in Antengene territories and the trials are responsible by Antengene;

<sup>&</sup>lt;sup>5</sup> Most advanced trial status in partner territories in the rest of the world and the trials are conducted by our licensing partners

# An Early-stage In-house Pipeline with Transformational Potential





<sup>&</sup>lt;sup>1</sup>Licensed from AstraZeneca and Antengene has obtained exclusive global rights to develop, commercialize and manufacture ATG-017;

<sup>&</sup>lt;sup>2</sup> Licensed from Origincell and Antengene has obtained exclusive global rights to develop, commercialize and manufacture ATG-101;

<sup>&</sup>lt;sup>3</sup> Licensed from Calithera Biosciences and Antengene has obtained exclusive global rights to develop, commercialize and manufacture ATG-037

<sup>\*</sup> ATG-037 IND equivalent in Australia = institutional scientific and ethics review before governmental notification Hem/Onc = hematological malignancies and solid tumors

# These Exciting Assets are in Clinical Stage Now .... with Worldwide Rights





## **ATG-017**



# **ATG-101**



### **ATG-037**



**Target** 

ERK1/2 (Small molecule)



CD73 (Small molecule)



- √ Higher potency and dual IoC and PoA activity with slow off-rate kinetics
- ✓ Lower efficacious dose with a higher max absorbable dose/dose ratio
- √ Broad therapeutic potential (targeting RAS/MAPK pathway)
- √ Multiple combination opportunities

- ✓ Shown potent PD-L1 crosslinking-dependent 4-1BB agonist activity, with the potential for delivery of enhanced therapeutic efficacy, whilst mitigating risk of hepatoxicity
- ✓ Demonstrated significant anti-tumor activity in animal models of resistant tumors as well as those that progressed on anti-PD-1/L1 treatment.
- √ Displayed an excellent safety profile in GLP toxicology studies

- ✓ Orally bioavailable small molecule that completely overcomes 'hook effect' common in other anti-CD73 antibodies
  - √ Tissue penetrance not achievable with mAbs
  - Promising preclinical efficacy as a monotherapy and strong potential synergy with Antengene pipeline candidates



**Status** 

Completed 5 cohorts in solid tumors of ERASER trial, expanded cohort at 20 mg bid with potential escalation to 30 mg bid, first combination module agreed (nivolumab), hematology cohorts to commence post MTD

Phase 1 clinical trial application for solid tumors and B-NHL approved by Bellberry HREC in Australia, US IND completed, cohort 2 commenced, China IND approved

Phase 1 STAMINA-001 Trial in-progress in Australia



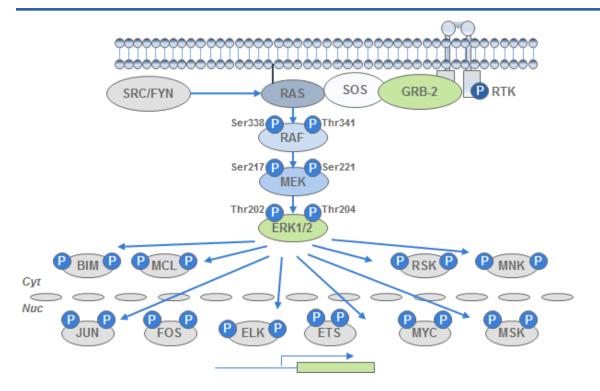
- 1. RASm NSCLC, Pancreatic, CRC, and Melanoma
- 2. I/O combinations

- 1. Resensitize prior CPI responders (NSCLC, SCLC, GI, H/N SCC, melanoma)
- 2. Efficacy in disease with previously limited CPI activity
- 3. Multiple combination opportunities

- 1. Monotherapy opportunity where immune suppressed TME is critical
- 2. Extremely broad opportunities both as monotherapy and combination with existing and future I/O

# ATG-017: A Small Molecule ERK1 / 2 Inhibitor with Best-In-Class Potential





# **Key Highlights**

- RAS / MAPK signaling pathway drives cell survival and proliferation
- Dysfunction in the signaling pathway is a major trigger for the development of most cancer types



# **Synergy with Antengene Pipeline Assets**

■ ERKi + KRASi

ATG-017 + ATG-012 in solid tumors

■ ERKi + I/O:

ATG-017 + ATG-101 in solid tumors

# ATG-017: A Small Molecule ERK1 / 2 Inhibitor with Best-In-Class Potential



## **Clinical Trial Overview**

Trial	Indication	Details
ERASER	Advanced solid tumors and hematologic malignancies with RAS / MAPK alternations	<ul> <li>Phase I, open-label, multicenter dose finding study to investigate the safety, PK and preliminary efficacy of ATG-017 monotherapy</li> <li>Completed the first 5 cohorts in solid tumors</li> </ul>

## **Competitive Advantages**

### **Best-in-class potential**

 ATG-017 is a potent and selective small molecule extracellular signal-regulated kinases 1 and 2 (ERK1/2) inhibitor with bestin-class potential

### **Leading in Clinical Development**

• First-in-human Phase I trial investigating safety and preliminary efficacy among patients with solid tumors and hematological malignancies; proceeding smoothly through dose escalation

### **Broad Therapeutic Potential**

 ATG-017 has great clinical potential as a monotherapy or combination in multiple solid tumors and hematological malignancies that harbor activating alterations in the RAS-MAPK pathway

# ATG-017: A Small Molecule ERK1 / 2 Inhibitor with Best-In-Class Potential



		ATG-017	GDC0994	BVD523	LY3214996	Differentiation
Potent ERK inhibitor with activity in relevant MAPK	<ul> <li>ERK potency and kinetics:</li> <li>A375 Cell pRSK / pERK IC<sub>50</sub> (uM)</li> <li>Mechanism of Action</li> <li>Cell proliferation Calu 6 / A375 GI<sub>50</sub> (uM)</li> <li>T<sup>1/2</sup> (non-phosphorylated/phosphorylated ERK)</li> </ul>	0.006 / 0.002  IoC and PoA 0.2 / 0.06  194 / 277 mins	0.09 / 0.03 IoC and PoA 2.3 / 0.15 1.2 / 0.8 mins	0.16 / 3  IoC 0.5 / 0.19  2.8 / 26 mins	0.32 / NT loC + PoA (tbc) 1.1 / NT 2.44 / 10.2 mins	ATG-017 more potent in vitro and has dual IoC and PoA activity with slow off rate kinetics
models	Efficacy Calu6 @ 50 mg/Kg >100%	>100% TGI (regression)	>100% TGI (100mg/kg QD)	93% TGI	~15 hrs cover at >1 x pRSK IC50 @ 50 mg/Kg; planning PD/efficacy	ATG-017 shows regression at 50 mg/Kg
Flexibility to allow optimal pathway inhibition	Predicted Dose to Man <100 mg  Max absorbable dose/Dose ratio >10  Human half life	20 mg BID 233 8 hrs (predicted)	200-400 mg BID*/** 0.5 23 hrs*	600 mg BID* 0.2 15 hrs (predicted)	ND	ATG-017 is a lower dose compound with a higher MAD:Dose ratio

<sup>\*</sup>clinical data from publications

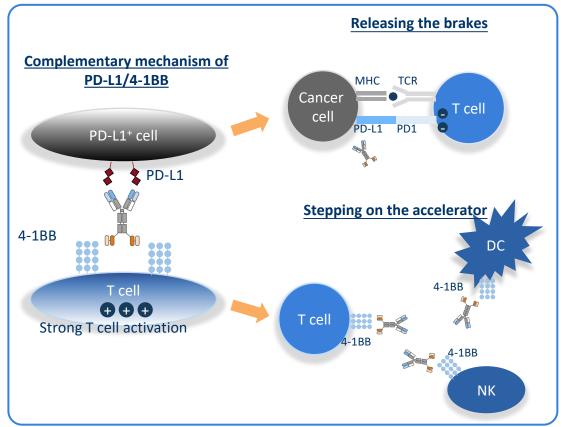
<sup>\*</sup>denendent on dosing regimen

# ATG-101: A Unique Bispecific Antibody Targeting PD-L1 and 4-1BB with Best-In-Class **Potential**



ATG-101: Complementary Mechanism of PD-L1 and 4-1BB Binding mAbs of PD-L1 and 4-1BB both active, clinical potential for 1+1>2

- Efficacy of PD-1/PD-L1 targeting well-demonstrated over the past decade
- 4-1BB is a T cell co-stimulatory receptor, the benefits of which have yet to be realized in the clinic
- The concept of PD-L1×4-1BB bispecific antibody like ATG-101 has demonstrated promising activity in early clinical trials with an acceptable safety profile (GEN1046, NCT03917381)
- Novel design of ATG-101: high affinity of PD-L1 and conditional activation of 4-1BB, results in 4-1BB agonist activity only when crosslinked by PD-L1-positive tumor cells → reduced risk of 4-1BB related liver toxicity





Synergy with Antengene Pipeline Assets

■ SINE + I/O:

Selinexor + ATG-101 in solid tumors and lymphoma

■ ERKi + I/O:

ATG-017 + ATG-101 in solid tumors

■ CD73 + I/O:

ATG-037 + ATG-101 in solid tumors

■ mTORi + I/O:

ATG-008 + ATG-101 in solid tumors

# ATG-101: A Unique Bispecific Antibody Targeting PD-L1 and 4-1BB with Best-In-Class Potential



## **Clinical Trial Overview**

Trial	Indication	Details
PROBE	Metastatic / advanced solid tumors and non-Hodgkin lymphoma	<ul> <li>Phase I, first in human trial</li> <li>Belberry Human Research Ethics Committee in Australia has approved our clinical trial application in July</li> <li>First patient dosed in January 2022</li> </ul>

### **Dose Escalation**

- Adv. Solid tumors (regardless of PD-L1 expression, not HCC) OR R/R B-NHL
- Exhausted available standard therapies

### **Dose Expansion**

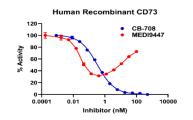
- Adv. Solid tumors of any histology (except HCC) and *CPI-exposed*:
  - a. DP following prior response/SD for ≥6mths to anti-PD-1/PD-L1
  - b. Best response of SD <6mths or DP after anti-PD-1/PD-L1
- Adv. Solid tumors / hematological malignancies with specific histologies below who have failed prior therapies but are *CPI naïve*:
  - a. TNBC
  - b. GBM
  - c. Gastric cancer, GEJ, oesophageal cancer
  - d. HPV+ HNSCC
  - e. Cervical cancer
  - f. B-NHL

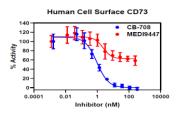
# ATG-037: An Orally Available, Small Molecule CD73 Inhibitor with Best-In-Class Potential

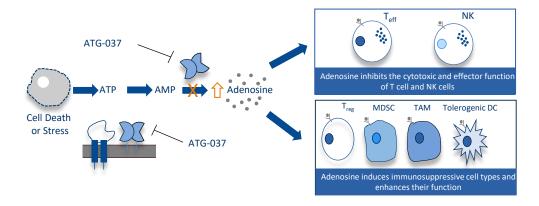


### **Differentiated Small Molecule Inhibitor of CD73**

- CD73 is the ecto-5'-nucleotidase, catalyzing the conversion of AMP to adenosine, a key immune suppressive molecule in the tumor microenvironment
- An orally available small molecule CD73 inhibitor in development
- Completely blocks CD73 activity, and overcome the "hook effect" commonly seen in anti-CD73 antibodies

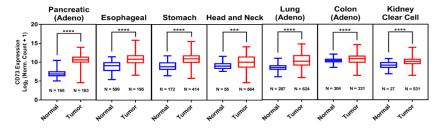






### **Broad Indication Potential**

 Pancreatic, esophageal, gastric, NSCLC, CRC, ovarian, prostate, head and neck, etc.



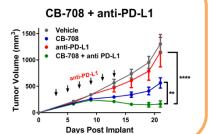
### **Advanced Drug Development**

- GLP toxicology studies completed: well tolerated in rodent and dog
- Potential large therapeutic window observed
- IND by end of year/ early next year



■ CD73 + I/O:

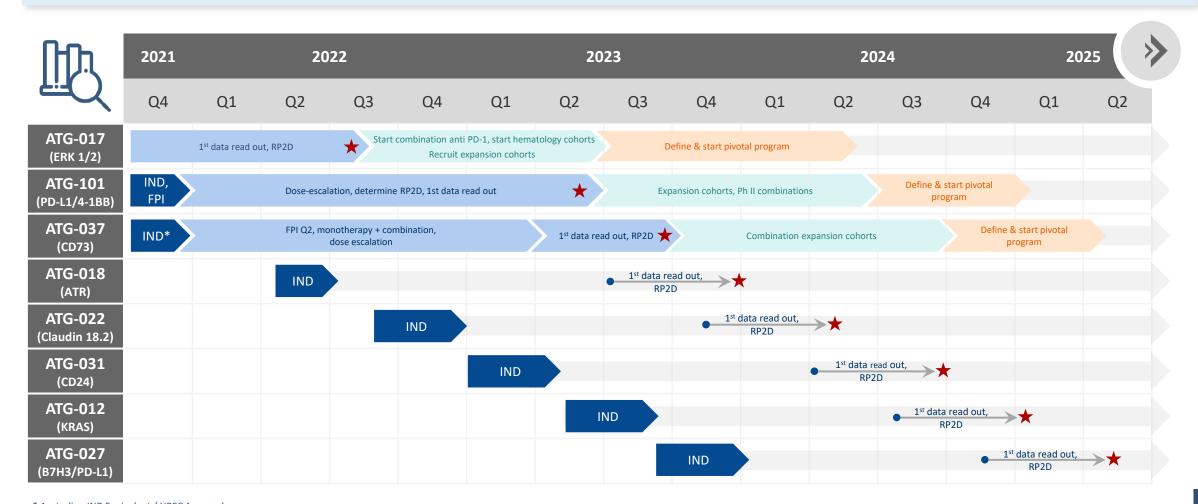
ATG-037 + ATG-101 in Solid Tumors



# A Series of INDs, FIH studies and Data Readouts Spanning 2022 to 2025



### Early data on safety, PK, PD through 2022 with ATG-017, ATG-101 and ATG-037



# **Strong Progress with Clinical Development Objectives for 2021/22**







- 1. Execute China selinexor bridging studies in MM
- 2. NDA submissions in Mainland China, Hong Kong, Taiwan, Singapore, South Korea, Australia
- 3. Define priority indication expansion for selinexor (endometrial cancer, myelofibrosis)
- 4. Critical data readouts with selinexor MM
- 5. Progress ATG-008 monotherapy and combination studies, define RP2D for ATG-008 and define potential registration path in combination with anti-PD-1
- 6. ATG-017 dose-escalation well-advanced and next steps defined (monotherapy expansion, combination with nivolumab, commence hematology cohorts)
- 7. Commence Phase II program ATG-016
- 8. Define Phase Ib signal seeking approach with ATG-527
- 9. Commence Clinical Activities in the USA
- 10. Complete IND and HREC submission for Phase I ATG-101, first patients enrolled
- 11. ATG-037 define FIH study design, secure HREC approval Q1, first patients enrolled Q2
- 12. ATG-018 define FIH study design, HREC submissions Q2

- Execute key Selinexor trial programs (SEARCH, BENCH, DLBCL 030, MATCH, SWATCH, SIENDO)
- 2. Complete extension of TORCH-2 program with ATG-008 combination with ICI and design pivotal regulatory path program
- 3. Define RP2D with ATG-017 monotherapy, RP2D combination with nivolumab, assess hematology program, progress through expansion cohorts, commence program in US and China
- 4. Progress through dose cohorts with ATG-101, early clinical, PK and PD data, commence program in US and China
- 5. Execute first dose cohorts with ATG-037, early monotherapy safety/PK/PD data, complete China IND
- 6. Secure Australian HREC approval for ATG-018 and commence FIH study
- 7. Complete 1-2 additional INDs within 2022



# And Next a Series of In-house Discovered, Best or First-in class Assets IND-ready within 6-18 Months





**AACR** 

American Association

for Cancer Research



**AACR** 

American Association

for Cancer Research



ATG-012







Target	ATR	Claudin 18.2	KRAS G12C	CD24	B7H3/PD-L1
Indication	Hematology/solid tumors	Solid tumors	Solid tumors	Hematology/solid tumors	Hematology/solid tumors
Modality	Small molecule	ADC	Small molecule	Monoclonal antibody	Bispecific antibody
Differentiation	<ul> <li>✓ Orally available</li> <li>✓ Promising therapeutic agent for patients with recombination deficiencies/genetic alterations</li> <li>✓ Better in vivo efficacy compared with benchmark</li> </ul>	<ul> <li>✓ High affinity antibody (sub-nanomolar grade) to enable treatment of patients with a broad range of Claudin 18.2 expression levels</li> <li>✓ Better in vivo efficacy compared with benchmark ADC</li> </ul>	<ul> <li>✓ Good cell potency</li> <li>✓ Strong in vivo synergism when used with in-house combination regimens</li> <li>✓ Better TDI and hERG profile compared with benchmark</li> <li>✓ Better in vivo efficacy compared with benchmark</li> </ul>	chemotherapy, rituximab and CPI	<ul><li>✓ ADCC/CDC effect to kill tumor</li><li>✓ In vivo POC completed</li></ul>
Status	IND Approved in Australia  Data presented in April 2022 in:	IND planned for 2022  Data presented in April 2022 in:	IND planned for 2023  Data presented in April 2022 in:	IND planned for 2023	IND planned for 2023

AACR

American Association

for Cancer Research



# 塞利尼索片 20mg









Approved in South Korea July 30<sup>th</sup>, 2021

**Commercial Launch** 

**Dec 2021** 

#### **Indications:**

- rrMM XPOVIO® in combination with dexamethasone (Xd)
- rrDLBCL XPOVIO® as monotherapy (X)

**Target National Reimbursement** 

2023



Approved in Mainland China December 14<sup>th</sup>, 2021

**Commercial Launch** 

**May 2022** 

Indications:

• rrMM - XPOVIO® in combination with dexamethasone (Xd)

**Target National Reimbursement** 

2024



Approved in Singapore March 1st, 2022

**Commercial Launch** 

May 2022

#### **Indications:**

- rrMM XPOVIO® in combination with bortezomib and dexamethasone (XVd)
- rrMM XPOVIO® in combination with dexamethasone (Xd)
- rrDLBCL XPOVIO® as monotherapy (X)

**Self-pay Market** 



Approved in Australia March 9<sup>th</sup>, 2022

Commercial Launch
May 2022

#### Indications:

- rrMM XPOVIO® in combination with bortezomib and dexamethasone (XVd)
- rrMM XPOVIO® in combination with dexamethasone (Xd)

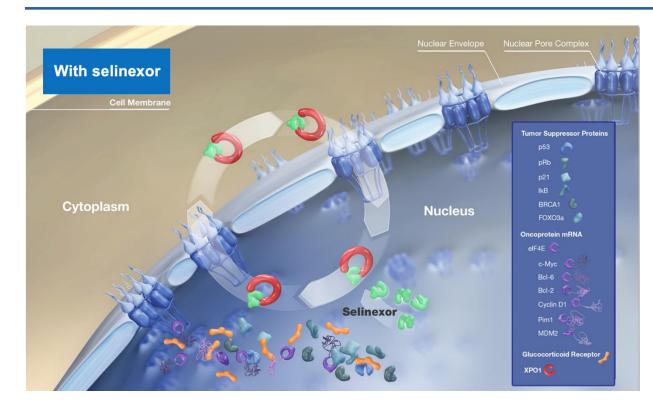
Target National Reimbursement 2022 — Early 2023





# ATG-010 (Selinexor) - First-in-class and Only-in-class SINE Compound with Unique MoA and Differentiated Profile





# **Key Highlights**

- 1<sup>st</sup> and only XPO1 inhibitor approved by FDA for treating R/R MM and R/R DLBCL
- 1<sup>st</sup> and only FDA-approved drug for the treatment of both R/R MM and R/R DLBCL
- Only single-agent, oral therapy approved by the FDA to treat R/R DLBCL
- Recommended by NCCN and CSCO guidelines for R/R MM and R/R DLBCL treatment



# **Synergy with Antengene Pipeline Assets**

■ SINE + mTORi

Selinexor + ATG-008 in diffuse large B-cell lymphoma (MATCH)

■ SINE + I/O:

Selinexor + ATG-101 in solid tumors and lymphoma

# Clinical Benefits Validated by Selinexor's Completed and Ongoing Studies in Multiple Myeloma and DLBCL







# **BOSTON**

(SVd)

### Selinexor Dosage: 100mg QW

- 1-3 prior therapies
- ORR: 76% (SVd) vs. 62% (Vd)
- CR rate: 17% (SVd) vs. 10% (Vd)
- mPFS: 13.93 mos (SVd) vs. 9.46 mos (Vd)
- mDOR: 20.3 mos (SVd) vs. 12.9 mos (Vd)
- Improved efficacy achieved when receiving 40% less bortezomib and 25% less dexamethasone

# **STOMP**

(SVd/SPd/SRd/SKd/etc.)

- 11 combinations
- ORR (study arm vs. benchmark data):
  - SKd: 78% vs. 23% (Kd)
  - SDd: **73%** vs. 29% (D)
  - SPd: 65% (pts dosed at RP2D) vs.
    - 29% (Pd)
  - SRd: 92% vs. 67% (Rd)

# **STORM**

(Sd)

### Selinexor Dosage: 80mg BIW

**FDA Approved** 

- mOS (≥MR): 15.6 mos
- Penta refractory (median # of prior therapies: 8)
  - ORR: 25%
- mPFS: 3.7 mos
- mOS: 8.6 mos

# SADAL

(S)

### Selinexor Dosage: 60mg BIW

- 2-5 prior lines
- ORR: 29%
- CR rate: 13%
- mDOR: 9.3 mos
- mOS: 9.0 mos
- mOS (≥MR): Not reached
- mOS (SD): 18.3 mos

### **FDA Approved**

**Multiple Myeloma** 

### **FDA Approved**

**Diffuse Large** 

**B-cell Lymphoma** 

Source: Dimopoulos M, et al. ASCO 2020. Abstract 8501; Gasparetto C, et al. ASH 2020. Abstract 1363.; Gasparetto C, et al. ASH 2020. Abstract 1393.; Kyprolis Package Insert; Study PX-171-003 A1; Lonial et al. Lancet 2016.; Pomalyst Package Insert.; Stewart et al. NEJM 2015.; Revlimid® (lenalidomide), Pomalyst® (pomalidomide), Velcade® (bortezomib), Kyprolis® (carfilzomib) or Darzalex® (daratumumab).; Chari A, Voqi DT, Dimopoulos M, et al. Results of the Pivotal STORM Study (Part 2) in Penta-Refractory Multiple Myeloma (MM): Deep and Durable Responses with Oral Selinexor Plus Low Dose Dexamethasorne in Patients with PentaMM. Blood 2018; FDA label for XPOVIO® (selinexor); Kalakonda N, et al. is currently in press and publication expected in the near term (Lancet Haematology 2020).

<sup>\*</sup>Some of the information in this presentation is from third-party medical professionals and for academic purposes only. Antengene is not responsible for the contents published by such external sources. \*\*Data shown for SDd and SPd in STOMP are from patients not previously exposed to D and patients dosed at RP2D respectively.

# Multiple Selinexor Containing Regimens Recommended by NCCN, ESMO, CSCO, and CMDA Guidelines







European Society for Medical Oncology



### **Multiple Myeloma**

### 1-3 Prior Therapies

- SVd QW
- SDd
- SPd
- SKd
- > 3 Prior Therapies (whose disease is refractory to at Least Two PIs, IMIDs, and an anti-CD38 mAb)
- Sd

### **Diffuse Large B-cell Lymphoma**

# 3L+ (including patients with disease progression after transplant or CAR T-cell therapy)

S monotherapy

### **Multiple Myeloma**

### **2L Option After VRD**

- R sensitive (SVd)
- R refractory (SVd)
- V sensitive (SVd)

### **2L Option After DaraRD**

- R sensitive (SVd)
- R refractory (SVd)

### **2L Option After DaraVMP or DaraVTD**

V sensitive (SVd)

### **Second or Subsequent Relapse**

- R refractory and PI sensitive (SVd)
- Triple-class refractory (Sd)

### **Multiple Myeloma**

### Relapsed/Refractory

- SVd
- SPd
- SDd
- SKd

### **Diffuse Large B-cell Lymphoma**

### Relapsed/Refractory

S monotherapy



### **Multiple Myeloma**

### Relapsed/Refractory

- SVd
- SPd
- SDd
- SKd

<sup>\*</sup> Some of the information in this presentation is from third-party studies and only provided to medical research professionals for academic purpose. Antengene is not responsible for the contents published by such external sources.

<sup>\*\*</sup> Approved for RRMM by the US FDA, European EMA, Chinese NMPA, Korean MFDS, Singaporean HSA and Australian TGA. Approved for RRDLBCL by the US FDA, Korean MFDS and Singaporean HSA. As of Mar 9, 2022.

<sup>\*\*\*</sup> Indications for Selinexor undergoing clinical trial / planned study include: PTCL, NKTL, AML, MDS, MF, Endometrial Cancer

# Dose Reduction can be Used to Manage Patients, while Optimizing Outcomes with PFS of 16.6 Months



### The median dosage of XPOVIO in the BOSTON trial was 80 mg (range: 30-137 mg) taken once weekly<sup>1</sup>

### Key efficacy endpoints for patients (N=195) with and without XPOVIO dose restrictions in the BOSTON trial<sup>2</sup>

	ITT Patient Population	Patients with Dose Reduction
Patient population	N = 195	n=126
% of ITT arm	100	65
mPFS, mo	13.9 (95% CI: 11.7, NE)	16.6 (95% CI: 12.9, NE)
ORR, %	76.4	81.7
≥VGPR, %	44.6	51.6
mDOR, mo	20.3 months (95% CI: 12.6, NE)	Not evaluable (95% CI: 13.8, NE)

### **Limitation of Subgroup Analyses:**

- This post hoc analysis was exploratory in nature, not included in the study objectives, and does not control for type 1 error
- This post hoc analysis was not powered or adjusted for multiplicity to assess PFS across other prespecified subgroups
- This post hoc analysis was underpowered to detect clinically meaningful differences in treatment effect

65% of patients in the XVd arm had an XPOVIO dose reduction (126/195 patients) vs 35% of those without a dose reduction (65/195 patients)<sup>2</sup>

Source: Karyopharm Investor Presentation dated December 8th, 2021

1. XPOVIO. Prescribing information. Karyopharm Therapeutics Inc; 2021. 2. Jagganath, et al. ASH 2021

<sup>\*</sup> Response was evaluable in 125 of 126 patients receiving a dose modification for selinexor and 66 of the 69 patients who did not receive a dose modification for selinexor. Overall response rate is the proportion of patients who achieve a partial response or better, before IRC-confirmed progressive disease or initiating a new MM treatment or crossover. CI=confidence interval, IRC=independent review committee; ITT=intent to treat; mDOR=median duration of response; mo=month; mPFS=median progression-free survival; NE=not evaluable; ORR=overall response rate; VGPR=very good partial response.

<sup>\*\*</sup> Some of the information in this presentation is from third-party studies and only provided to medical research professionals for academic purpose. Antengene is not responsible for the contents published by such external sources.

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<sup>\*\*\*\*</sup> Indications for Selinexor undergoing clinical trial / planned study include: PTCL, NKTL, AML, MDS, MF, Endometrial Cancer

# **XPOVIO Evolving into a Standard of Care with Dose and Schedule Redefined Over Time to Improve Efficacy and Patient Experience**



From the STORM trial to the BOSTON trial to the STOMP trial, XPOVIO dosing has been continually refined to help optimize the patient experience

**US FDA Approval Date: July 2019** 

**US FDA Approval Date: Dec 2020** 

Ongoing/Completed

1<sup>st</sup> approval in MM Dose: 160mg (80 mg, twice weekly)

> Xd **STORM**

Phase 2b, single-arm, open-label, multi-center study

Patients with penta-refractory RRMM

2<sup>nd</sup> approval in MM Dose: 100mg, once weekly

> XVd **BOSTON**

Phase 3, 2-arm, active comparatorcontrolled, open-label, multi-center study

After at least 1 prior therapy in MM

Phase 1/2 study in MM Dose Range: 60-100mg, once weekly

SPd, SKd, SDd **STOMP** 

Phase 1/2, open-label, multi-center study

Patients with RRMM (dose escalation/expansion)

**Once Weekly** 

(previously twice weekly)

**Lower Dose** 

(previously a higher dose)

**XPOVIO-based Triplets** 

(previously a doublet)

**Earlier Lines** 

(previously only in later lines)

**Supportive Care** 

(active symptom management)

Source: Karyopharm Investor Presentation dated February 8th, 2022

<sup>\*</sup> STOMP was designed to study selinexor in combination with other MOAs across multiple triplet and quadruplet regimens, including XVd. MM=multiple myeloma; MOA=mechanism of action; RRMM=relapsed or refractory multiple myeloma.

<sup>\*\*</sup> Combinations other than XVd and Xd are not promoted by Karyopharm, but may be considered for future indication updates.

<sup>\*\*\*</sup> Combinations other than Xd are not promoted by Antengene, but may be considered for future indication updates

<sup>\*\*\*\*</sup> Some of the information in this presentation is from third-party studies and only provided to medical research professionals for academic purpose. Antengene is not responsible for the contents published by such external sources.

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<sup>\*\*\*\*\*\*</sup> Indications for Selinexor undergoing clinical trial / planned study include: PTCL, NKTL, AML, MDS, MF, Endometrial Cancer

# **Broad and Deep Potential for Selinexor / SINE Beyond Multiple Myeloma**



Incidence / Prevalence China (APAC) 19,600 (1,900) / 68,600 (8,740)	MF	Global Pivotal Study Ongoing
49,000 / 57,937 (3,100) / (9,300)	MDS	Signal Detection Studies/IITs in Preparation in China
84,000 / 116,280 (3,200) / (3,520) (AML)	Leukemia	Signal Detection Studies/IITs in Preparation in China
86,000 / 204,910 (9,100) / (53,000)	Endometrial Cancer	<ol> <li>Global Study</li> <li>Partner in the US announced top-line results in Phase III Study</li> <li>Potentially first solid tumor indication for Selinexor</li> </ol>
50,585 84,463 (9,199) / (34,658) (DLBCL + TCL)	Lymphoma (i.e., DLBCL, TCL)	1. Approved in the US for 3L DLBCL; pivotal study ongoing in China 2. Recommended by NCCN and CSCO guidelines 3. Multiple studies (SADAL, SEARCH, XPORT-030, TOUCH, RWD)
21,000 / 54,800 (6,000) / (23,500)	Multiple Myeloma	1. Approved in the US for 2L+ MM and approved in China for rrMM 2. Recommended by NCCN, CSCO, ESMO, CPA guidelines as 2L+ therapy 3. Multiple studies (BOSTON, BENCH, STORM, STOMP, MARCH, RWD)
Total: Total: 310,185 / 586,990		

Source: Antengene research

(32,499)

(132,718)

<sup>\*</sup> Investigator Initiated Trials (IIT)

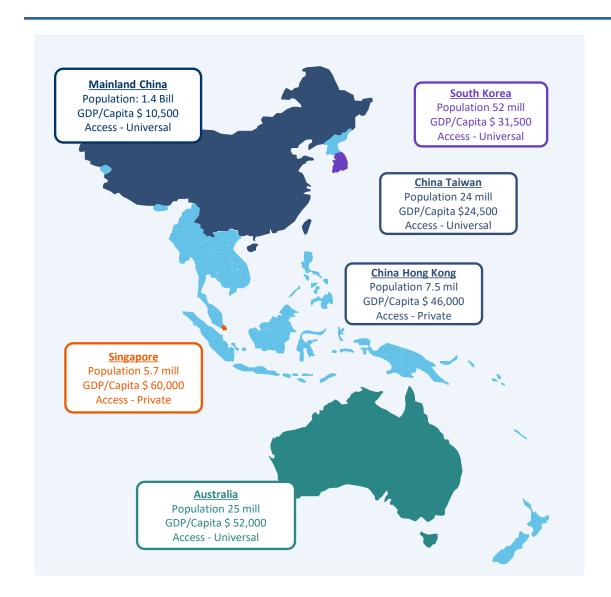
<sup>\*\*</sup> Some of the information in this presentation is from third-party studies and only provided to medical research professionals for academic purpose. Antengene is not responsible for the contents published by such external sources.

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<sup>\*\*\*\*</sup> Indications for Selinexor undergoing clinical trial / planned study include: PTCL, NKTL, AML, MDS, MF, Endometrial Cancer

# **Antengene is Focused on Markets with Greatest Commercialization Potential**







Commercialization strategy focuses on selection of 6 core stage 1 markets defined by 3 parameters for success:

- High unmet need
- Reimbursed markets
- High GDP/capita



Initial focus is to build Antengene presence in the core markets



Ensure successful commercial launch of Xpovio®



Expand portfolio in core markets and expand Antengene presence to other stage 2 ASEAN markets

## Commercial Team with a Proven Track Record of Success



### **Commercialization Strategy**



John F. Chin

■ 30+ years of experience in the pharmaceutical industry, instrumental involvement in the commercial launch and lifecycle management of REVLIMID®, one of the industry's most successful oncology products globally

#### **Commercialization in China**



Lixin Yu

- 30+ years of experience in commercialization for Oncological **Products in China**
- Successful launch cases in hematological, global-local products in Multiple Myeloma, Lymphoma and Leukemia

#### **Commercialization in APAC**



**Thomas Karalis** 

- 30+ years of experience in the pharmaceutical industry, achieved multiple regulatory and reimbursement milestones in APAC
- Lead to successful launches of REVLIMID®, POMALYST® and ABRAXANE® in APAC markets

Track record of Antengene commercial team in hematology:













#### **China Marketing**



Frank Sun Director, Marketing and Commercial Channels. Hematology BU China Deep industry experience in hematology product launch in mainland China, market development and team management

#### **China Sales**



Chen Wei National Director, Hematology BU China Deep industry experience in commercializing hematology products in mainland China

#### **China Medical Affairs**



Godfrev Guo Executive Director, Medical, China Seasoned experience in Hematology & Skin Cancer, proven track record for the launch of a series of novel medicines, including Zelboraf, Hemlibra, Gazyva and Polivy, as well as the expansion of new indications



**Austin Wang** Associate Director, MSL, China Extensive experience in working with key KOLs, deep medical insights in CN Hema. market and landscape

#### **AU/NZ Commercialization**



AU, US and EU Commercial, Govt Affairs and Market Access leadership roles in Hematology, Oncology and Specialty Therapeutics

Michele Robbins



**GM of South Korea** 

market development and team management Minyoung Kim



Wendy Lau

#### **HK/SG Commercialization**



Asia Marketing Lead in Hematology Oncology and Specialty Care **Business Unit** Management

ASEAN, Central East

#### **APAC Medical Affairs**



AU. US and Global Medical Affairs leadership roles. Extensive clinical/translational research background in Hematology and Oncology

Tamara Etto

### **APAC Commercialization**



Sathya Walisinghe

Extensive ANZ, US and APAC commercial experience including Global Marketing CAR T Launch and strong background in Hematology & Oncology

# **Antengene's Commercial Ready Infrastructure Across Asia Pacific**



### Pan APAC Team with Proven Track Record



Commercial team of ~200 by the end of 2021 with experience in successfully launching multiple products across the APAC region



Track Record of
Antengene
Commercial Team in
Hematology











### **Pre-approval KOL Engagement**



- Selinexor patient experience gained in APAC markets through Early Access Programs
- **Clinical Trial Participation** 
  - Australian participation in BOSTON and SADAL registrational studies
- **Investigator Initiated Trials**
- Advisory Boards
- Market Research
- Disease Symposia

# Limited Availability of Reimbursed Triplet Regimens in Early RRMM in APAC Reimbursed Markets



1 Regimen

DVd



2 Regimens

KRd and NRd

v.s.

8 Triplet Regimens
Commonly Used in
the US 2L & 3L
Therapies

Taiwan Markets

3 Regimens

DVd, DRd and IRd

# Fewer Myeloma Medicines Approved in China Compared to the US

Launching with less competition in China



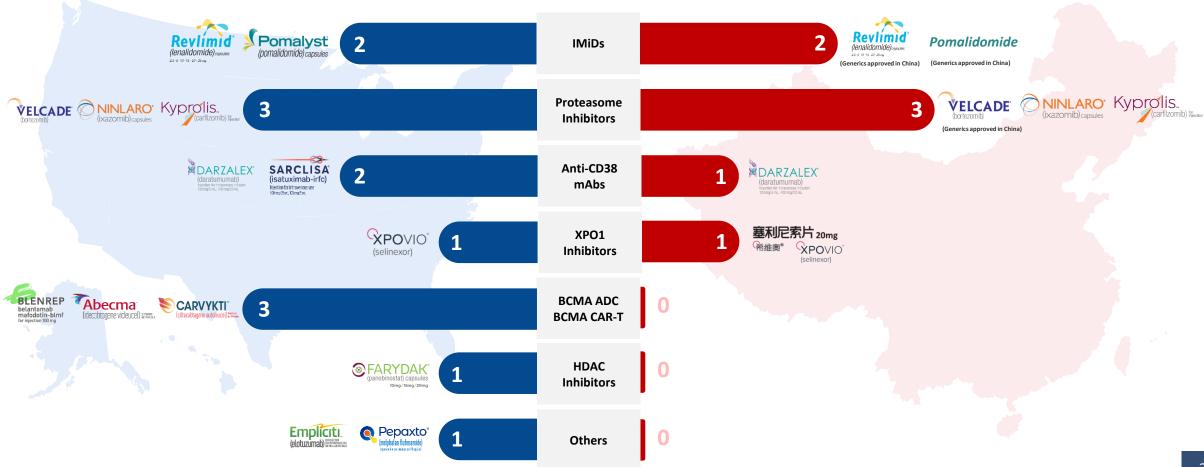
### **United States**



V.S.



## **Mainland China**



# Strong KOL Support and Engagement Paving the Way for a Successful Launch in China 🧥 🔟







# 塞利尼索片 20mg



2022 Target Hospitals Coverage: 600

#### **China Launch Readiness**

- Advancing Healthcare Professional (HCP) / Key Opinion Leader (KOL) advocacy
  - 600+ HCP experience
  - 1300+ Patient Experience
- Named Patient Program (NPP) patient access in mainland China / Hong Kong markets
- A number of **Investigator Initiated Trials** in planning / ongoing in China in a broad range of indications, expanding the breadth of selinexor in China
- Publication plan in academic journals on NPP real world study data
- Encouraging signs of **KOL support and usage** in China:
  - Multiple combinations (triplets and quadruplets) being used by physicians in China
  - Large proportion of lymphoma usage
  - Gathered real world data and feedback

2Q2022

China Launch

卡非佐米+泊马度胺+

泊马度胺+地塞米松(2A类)

地塞米松 (2A类)

地塞米松(2B类)

硼替佐米(2类)

泊马度胺(2类)

Selinexor+ 地塞米松 +

Selinexor+ 地塞米松 +

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# **Asia Pacific Markets – Accelerating and Executing on XPOVIO® Launch Plans**





### **Launch Readiness**

- Commercial presence in Australia, South Korea, Singapore, Hong Kong & Taiwan
- NDA regulatory approvals in Australia, South Korea & Singapore
- Advancing National Reimbursement in Australia and South Korea
- Building **KOL advocacy and experience**:
- >250 patients treated with XPOVIO® via access program
- Ongoing advisory boards and medical education programs
- 2 IITs advancing



## South Korea – 2022

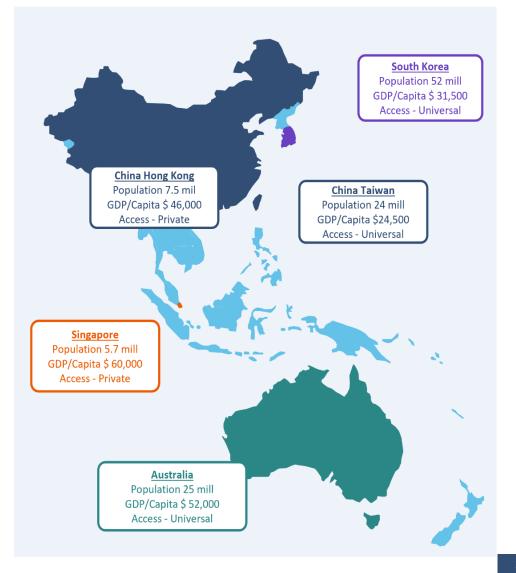
- Pre reimbursement launch Dec 21
- Patient Access Programs Initiated
- Hospital Formulary submissions
- XPOVIO® patient case study sharing
- Nurse led patient support programs





### Australia – 2022

- Local & International KOL led medical education
- MSAG Guideline inclusion
- Broaden physician experience with **Patient familiarization Program**
- Myeloma Australia/ALLG integrated patient support



# **Strong Foundation for Successful Launches Across China and Asia Pacific Markets**





**Progressing through regulatory and reimbursement** approvals

**NDAs Submissions** 





Physician experience with XPOVIO®

Patients on

Access programs

Best in class team established

**Commercial and** Medical Personnel

Rapid engagement with KOLs across the region





# **2022** Will Be a Transformational Year for Antengene



## Commercialization across China and APAC, with multiple data read outs of FIH studies of global rights assets



### Selinexor Commercial Launch across APAC



- Commercial launch: South Korea (MM Sd; DLBCL)
- Commercial launch: Mainland China
- Commercial launch: Australia (MM SVd & Sd)
- Commercial launch : Singapore (MM SVd & Sd; DLBCL)
- Commercial launch : Hong Kong (MM Sd)
- Commercial launch : **Taiwan** (MM SVd & Sd; DLBCL)
- Reimbursement submission: **South Korea** (MM Sd; DLBCL)
- Reimbursement approval: Australia (MM SVd & Sd)





- Interim data read out on pivotal trial: ATG-016 (XPO1) (MDS)
- Interim data read out: ATG-008 (mTORC1/2) (TORCH-2 in combination with PD-1)
- Preliminary data read out in First-inhuman (FIH) studies of global rights assets:
  - ATG-017 (ERK1/2) (ERASER)
  - ATG-101 (PDL1/41BB bispecific) (PROBE)
- IND submission: ATG-018 (ATR) and ATG-022 (Claudin 18.2 ADC)
  - Progressing through dosing cohorts in First-in-human (FIH) studies of ATG-037 (CD73) and ATG-018 (ATR)

# Multiple Regulatory Filings



- Selinexor (ATG-010) sNDA filing in **South Korea** (MM SVd)
- Selinexor (ATG-010) sNDA filing in **Australia** (DLBCL)
- Selinexor (ATG-010) sNDA filing in mainland China (DLBCL)
  - Supplementary NDA filing of selinexor (ATG-010) packaging at Shaoxing manufacturing site

# Advancing Pipeline & Discovery Programs



- Attain **IND-readines**s in-house developed program: **ATG-031** (CD24)
- Multiple novel ADC PCC nominations
  - Initiate 2-3 new discovery projects
  - Continuous BD efforts will bring in innovative assets





# **Steady Stream of Catalysts Continue to Drive Value for Investors**



### 2021 Revenue: RMB28.8 mm; 2022 Revenue Target: RMB180 to 200 mm



### **De-risked Biotech with Revenue Generation Capabilities**

• Starting with 2 disease areas with multiple indication expansion / blue ocean opportunities with an only-in-class asset in unique markets



### **Broad and Deep Pipeline of Differentiated Global Rights Assets**

 10 global rights assets with highly-differentiated, combinational and synergistic mechanism of action with series of upcoming data read outs



### **Following the Science and Pursuit of Synergistic Partnerships**

 Continued BD discussions on new assets, clinical collaborations and technology platforms to power internal discovery efforts



### **Efficiently Utilizing Cash Provided by Our Strong Base of Global Shareholders**

 US\$373 mm (RMB2,370mm) of cash, bank balances and cash management products as of 31st December 2021



ANTENGENE CORPORATION LIMITED (SEHK: 6996.HK)

JULY 2022

# THANK YOU

TREATING PATIENTS BEYOND BORDERS