



Jacobio Pharmaceuticals 2021 Annual Results





以**患者受益**为导向
利用最新的科研成果研发**全球首创新药**

Focus on a **patient-centric** approach
Develop **transformative medicine** through scientific breakthroughs
and innovative technology

Our Strategy



In-house R&D

Focus on in-house R&D leveraging our **allosteric inhibitor tech platform** rather than in-licensing



Global FIC

Aim for **“first-in-class”** drugs over me-too drugs



Global Market

Explore MNC partnership to capture global market



Full Function Pharma

Expand areas of manufacturing and commercialization in China



Our Progression

Clinical Milestones

JAB-3312 (SHP2i)

- Mono – Dose expansion initiated
- Globally initiated 4 combos
 - combo w/PD-1 (Pembrolizumab)
 - combo w/MEKi (Binimetinib)
 - combo w/KRAS G12Ci
 - *Sotorasib, Global*
 - *JAB-21822, China*
 - combo w/EGFRi (Osimertinib)

JAB-21822 (KRAS G12Ci)

- RP2D determined
- Pivotal trial to be initiated in 2022 2H (NSCLC)
- NSCLC
 - mono 2/3L,
 - mono STK-11 co-mutant 1L,
 - combo w/PD-1 CRC (combo w/EGFR mAb)

JAB-3068 (SHP2i)

- China PhIIa mono trial enrollment completed
- combo w/ PD-1 (JS001) in China

JAB-8263 (BETi)

- Dose escalation ongoing
- Solid tumor and hematological malignancy in US ad China

JAB-21822 NDA

Commercial Manufacturing
First Product Launch

New INDs

- JAB-21822 (KRAS G12Ci)
- JAB-2485 (Aurora Ai)
- JAB-BX102 (CD73 antibody)
- JAB-6343 (FGFR4i)

AbbVie Partnership
regarding SHP2

HKEX listing

2021-2022 March

2023-2024

2020

2015

Founded in 2015
Set the goal of global
innovation



Our Team



Yinxiang Wang Ph.D.
Chief Executive Officer
Chairman of our Board



Xiaojie Wang, EMBA
President of Administration



Yunyan Hu, M.S.
Senior Vice President



**Andrea Wang-Gillam
M.D, Ph.D.**
CMO and Senior Vice President



Haijun Wang, Ph.D.
Senior Vice President of Data
and Information Management

- Chairman and CEO
- Former Board Director and CEO of Beta Pharma
- Chairman on the New Drug R&D Committee of China for PhIRDA (2017-2019)
- Vice Chairman on the Anti-tumor Drug Professional Committee of Chinese Pharmaceutical Association
- Over 20 years of experience in the industry



Wayne Long, Ph.D.
Vice President of
Chemistry



Yuli Ding, M.S.
Vice President of
Clinical Development



Bin Fan, Ph.D.
Vice President of
Clinical Pharmacology



Qiao Li, Ph.D.
Vice President of Biostatistics
and Data Science



Yanping Wang, M.S.
Vice President of
Pharmacology



Hong Cao, M.S.
Vice President of CMC

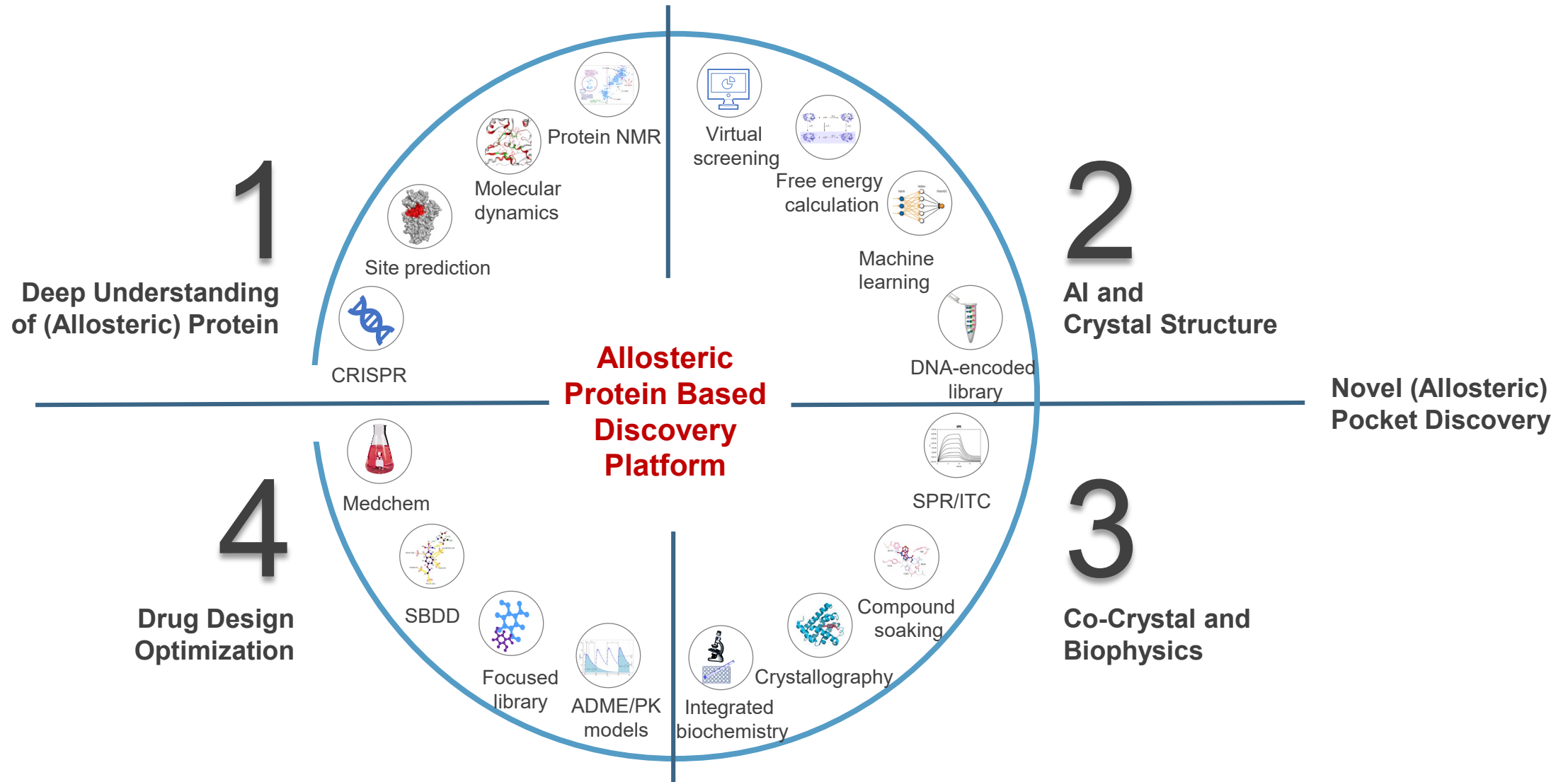


Jian Kang, M.D.
Vice President of
Intellectual Property



Tiffany Yang, M.S.
Vice President of
Human Resources

Our Allosteric Inhibitor Technology Platform



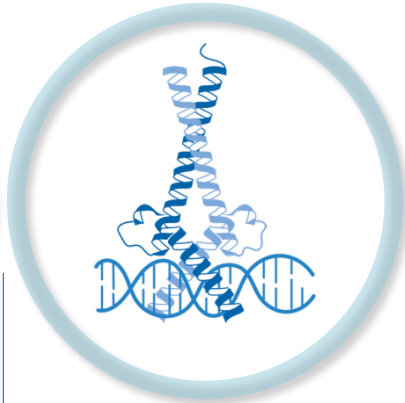
We have validated our discovery and validation platform and gained competitive advantages.

Advance Novel Drug Development in Key Oncogenic Pathways



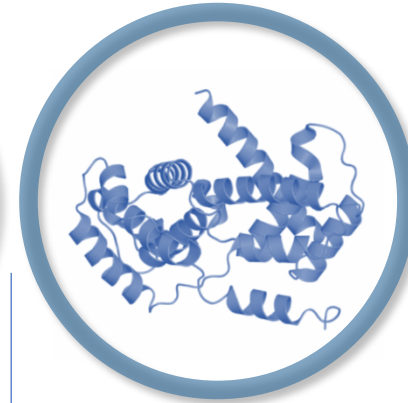
RAS

The beating heart of cancer



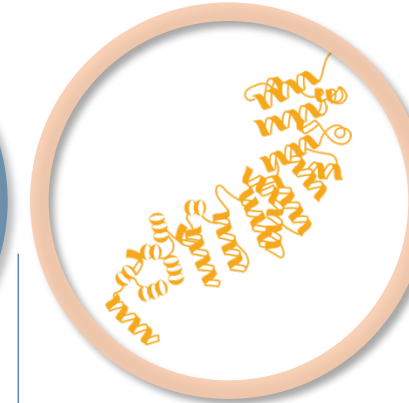
MYC

The master regulator



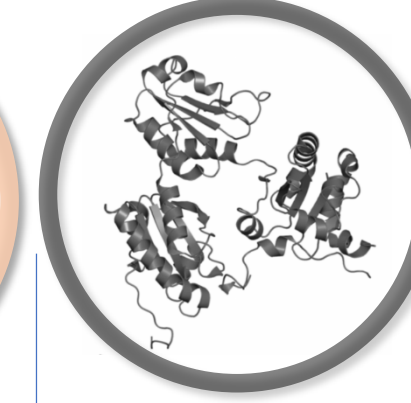
I/O pathways

The Defensive barrier



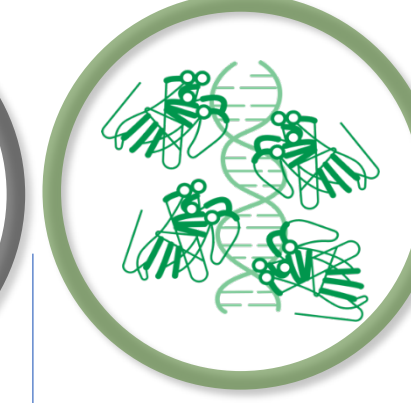
RB pathway

The brake pads of cancer



Tumor Metabolic Pathways

The suppliers of cancer



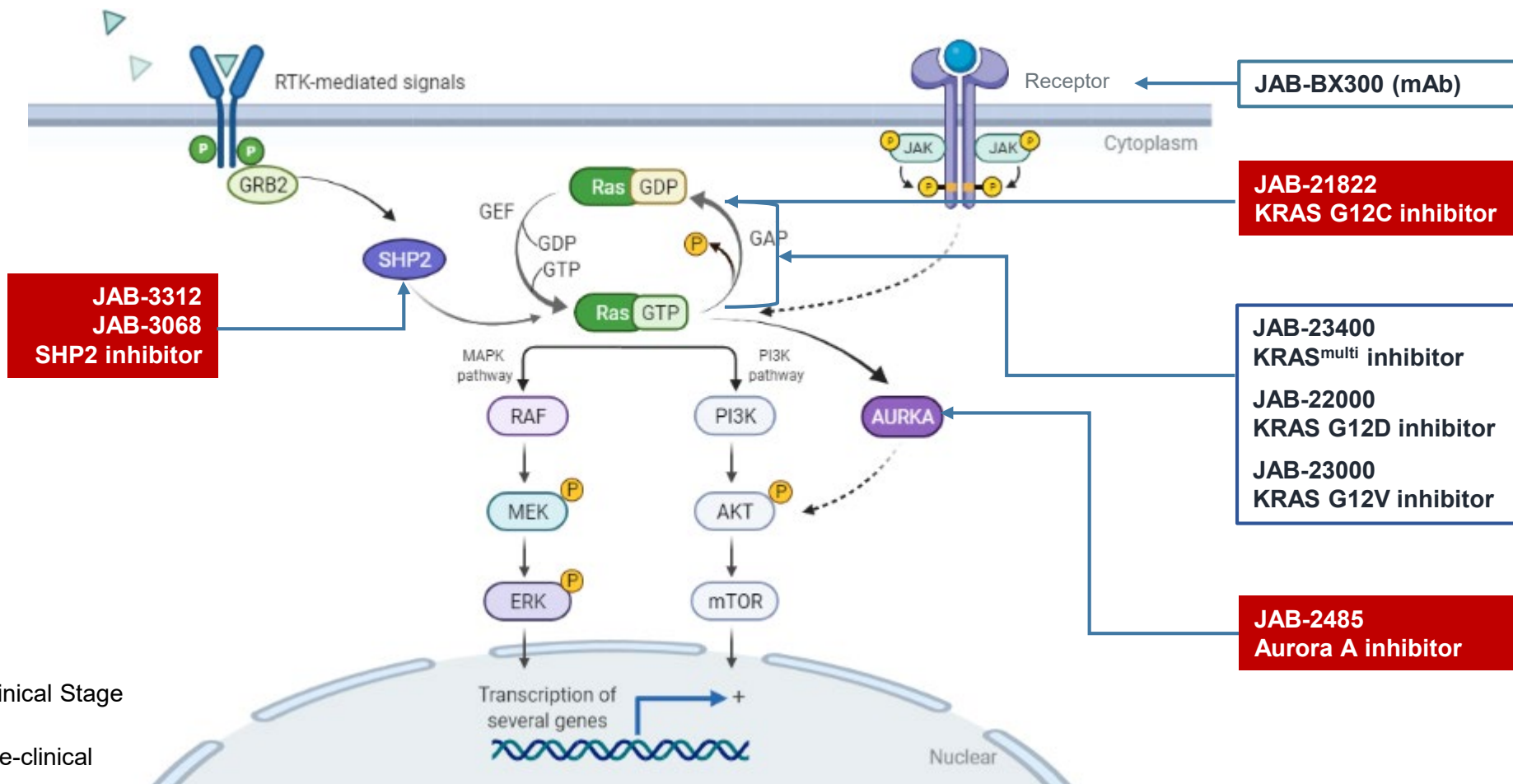
p53

The guardian of the genome

The “BIG 6” pathways cover 70%-80% cancer.

1. Nusse R, Clevers H. *Cell* 2017;169(6):985–99;
2. Lane DP. *Nature* 1992;358(6381):15–6;
3. Waters AM, Der CJ. *Cold Spring Harb Perspect Med* 2018;8(9):a031435;
4. Dang CV, et al. *Cell* 2012;149(1):22–35.

Deep Dive in RAS Pathway to Address Unmet Need



● Clinical Stage
○ Pre-clinical

Our SHP2 Inhibitor JAB-3312 as a Potential Best-in-Class Drug

TOP3 SHP2 inhibitor
(in order of US FDA IND filing date)

Company Name	Company	US FDA IND
TNO-155	Novartis	May 2017 (US)
JAB-3068 JAB-3312	Jacobio/ AbbVie	Dec 2017 (US)
RMC-4630	Revolution Medicines/Sanofi	Sep 2018 (US)

SHP2 Functions in the Downstream of anti-PD-1 and Upstream of KRAS with Tremendous Market

New cases estimated worldwide with RAS/MAPK pathway alterations

5.5 million (worldwide)	BRAF class III/ NF1 LOF mutant solid tumors	KRASi / EGFRi MEKi combo
New cases estimated worldwide per annum	90+% with limited/no treatment options	SHP2 mono strategies
		SHP2 combo strategies

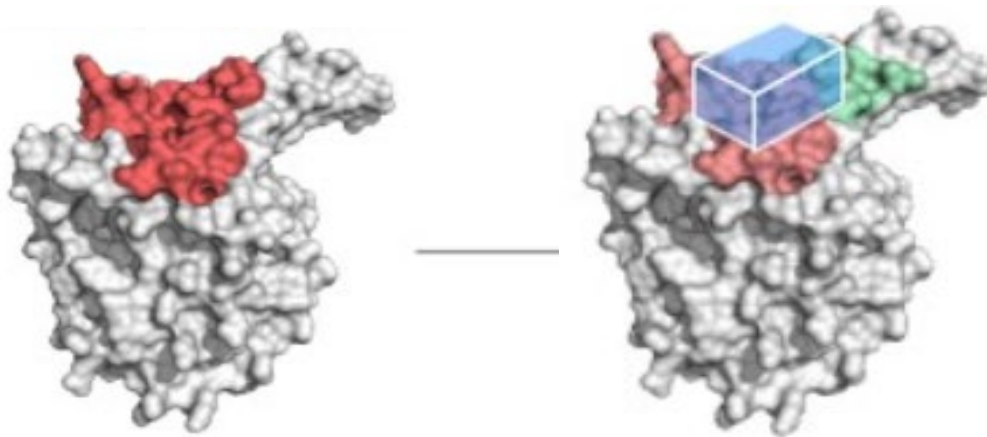
Market potential and clinical efficacy of SHP2 in the PD-(L)1 pathway

<30%	3.37 million pts (worldwide in 2019)	PD-(L)1 combo
ORR of PD-(L)1 monotherapy in most cancer types	Number of patients with the 5 largest cancer types who did not respond to PD-(L)1	SHP2 combo strategies

Our SHP2 Inhibitor JAB-3312 as a Potential Best-in-Class Drug

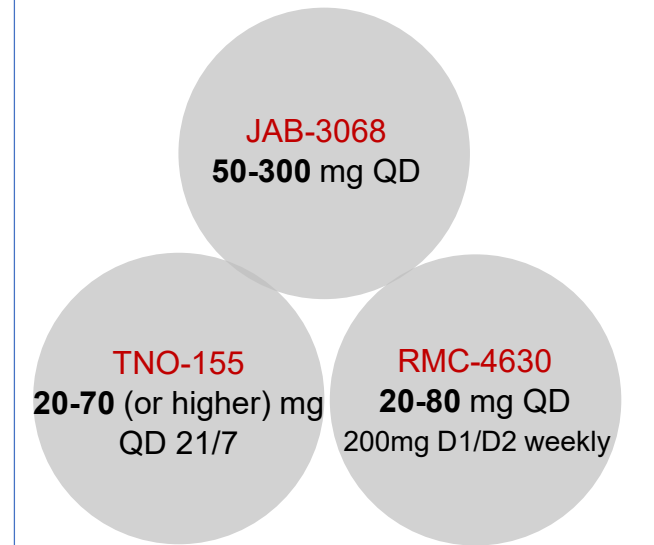
Preclinical comparison

	JAB-3312	RMC-4550 In-house or ref
SHP2 biochemical IC ₅₀ (nM)	1.5	10.4
Binding kinetics KD (nM)	0.206	13.6
Cellular p-ERK IC ₅₀ in NCI-H358 (nM)	3.64	28 (ref)
Cellular p-ERK IC ₅₀ in KYSE-520 (nM)	0.32	9.1 (ref)
Cellular proliferation KYSE-520 IC ₅₀ (nM)	3.5	127



Clinical dose

JAB-3312
1-8mg QD



SHP2 Inhibitor-Global Development Plan

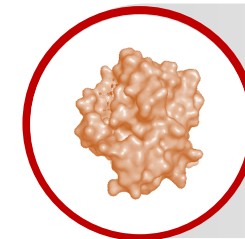
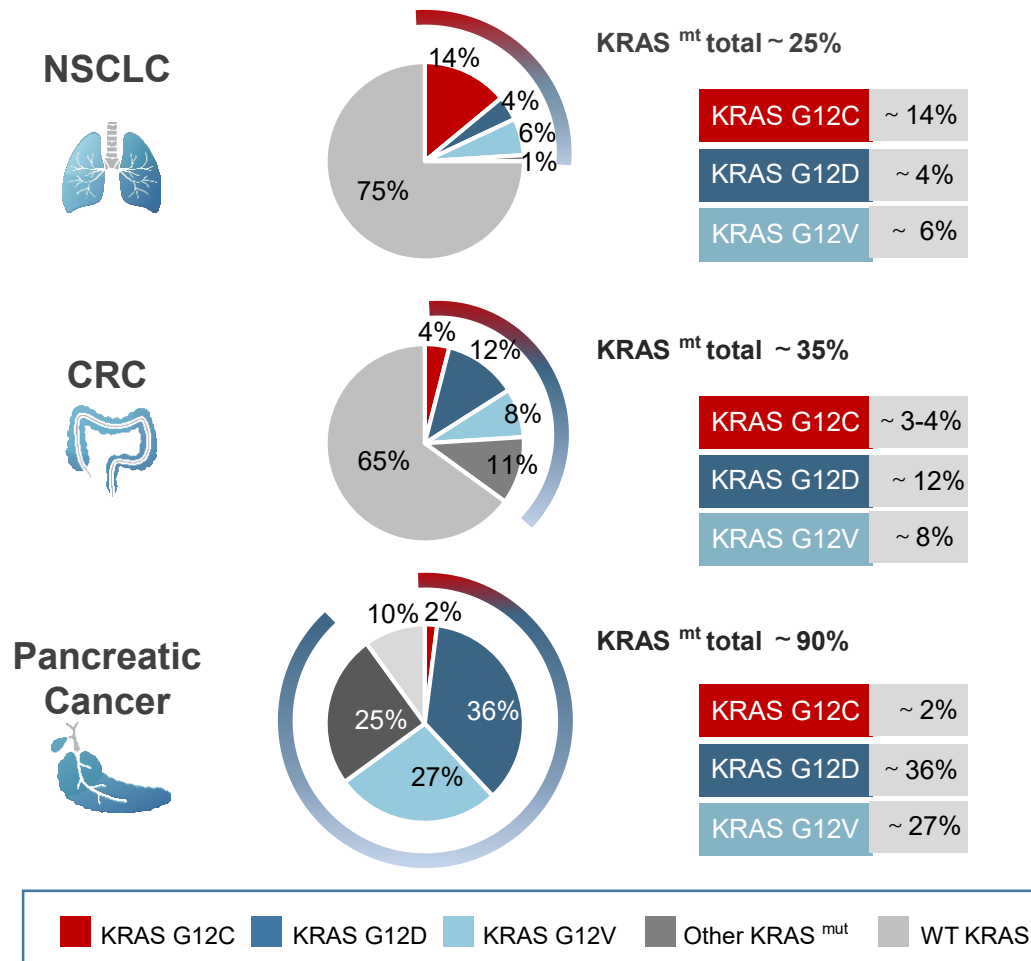
Given the unique dual-blockade mechanism of SHP2 against RAS pathway and PD-(L)1 pathway, we plan to develop our SHP2 inhibitors as monotherapy and as a backbone for various combination therapies

Asset	Regimen	Indications	Phase I	Phase IIa	Recent development
JAB-3312 SHP2 abbvie	Mono	Solid tumors	US trial		
	Mono	Solid tumors	China trial		Phase IIa initiated with FPI in Jan 2022
	Mono	BRAF class 3/ NF1 LOF mutant solid tumors	US trial		Phase IIa initiated with FPI in Dec 2021
	Combo w/PD-1 mAb	NSCLC, HNSCC, ESCC	Global trial		Phase IIa Initiated in Feb 2022
	Combo w/EGFRi	Osimertinib resistant NSCLC	Global trial		FPI in Jan 2022
	Combo w/KRAS G12Ci	KRAS G12C mut NSCLC	Global trial		FPI in Jan 2022
	Combo w/MEKi	KRAS mut CRC/ Pancreatic cancer	Global trial		
JAB-3068 SHP2 abbvie	Mono	Solid tumors	US trial		
	Mono	ESCC, HNSCC, NSCLC	China trial		
	Combo w/PD-1 mAb	ESCC, HNSCC, NSCLC	China trial		

CRC = colorectal cancer; ESCC = esophageal squamous cell carcinoma; HNSCC = head and neck squamous cell carcinoma; NSCLC = non-small cell lung cancer
¹ We assume the data of Phase IIb registrational trial will be used for NDA submissions of JAB-3068 and JAB-3312 in China and the U.S.

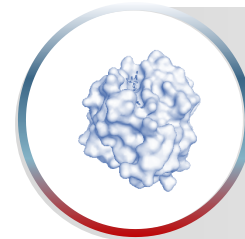
Our Diversified KRAS Inhibitor Portfolio

KRAS Prevalence In Tumors With High Unmet Need ¹⁻³



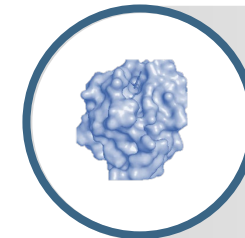
JAB-21822 KRAS G12Ci

Currently in Phase IIa
Pivotal trial to be initiated in China in 2022 H2



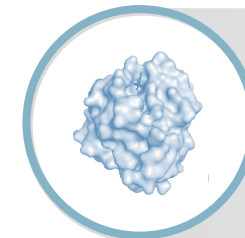
JAB-23400 KRAS^{multi} inhibitor

Candidate nominated in Feb 2022
Target to submit IND in 2023



JAB-22000 KRAS G12Di

Lead optimization stage
Target to submit IND in 2023



JAB-23000 KRAS G12Vi

Hit-to-lead stage
Target to submit IND during 2023 to 2024

1. ZehirA, et al. *Nat Med.* 2017;23(6)703-713.
2. KrakstadC, et al. *PLoSOne.* 2012;7(12):e52795.
3. NIH TCGA: *The Cancer Genome Atlas.* February 11, 2021. <https://www.cbiportal.org>.

KRAS G12Ci JAB-21822-Global Development Plan

Accelerate the clinical development of JAB-21822 utilizing strong internal capacities and extensive external resources

- Achieved FPI in China within 2 months after IND approval
- Monotherapy and combination therapies are being rapidly advanced in clinical setting, sustaining competitive advantage.

Asset	Regimen	Indications	IND	Phase I	Phase IIa	Recent development	Upcoming Milestone (expected)
JAB-21822 KRAS G12C	Mono	NSCLC, CRC	Global trial			FPI in Sep 2021	
	Mono	NSCLC, CRC	China trial			Dose expansion initiated with FPI in Mar 2022	Pivot trial to be initiated in 2022 2H
	Mono	NSCLC with STK-11 co-mutation	Global trial			IND approved in Oct 2021	FPI (2022 2H)
	Combo w/PD-1 mAb	NSCLC	China trial			IND approved in Oct 2021	
	Combo w/SHP2i	NSCLC	China trial			IND approved in Feb 2022	FPI (2022 Q2)
	Combo w/EGFR mAb	CRC	China trial			FPI in Feb 2022	

Preliminary Clinical Results

- RP2D determined
- Impressive preliminary efficacy
- Safe and well tolerated

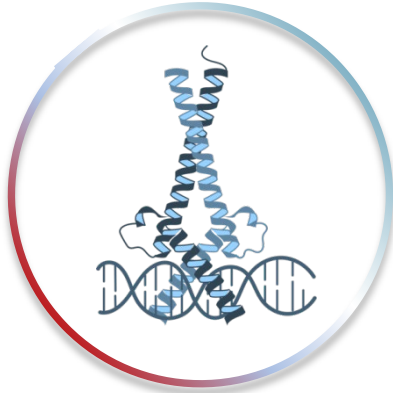
Exploring potential global partnering deal while Jacobio retains the greater China right

Synergistic Combinatorial Strategy among In-house Assets



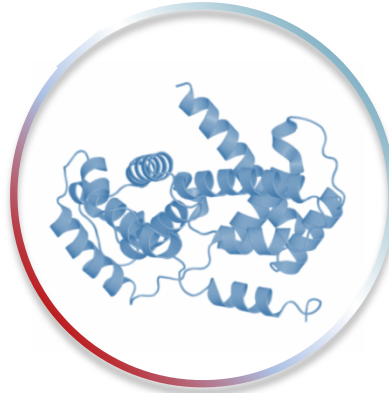
RAS

JAB-3312/3068
JAB-21822
JAB-23400
JAB-22000
JAB-23000
JAB-BX300



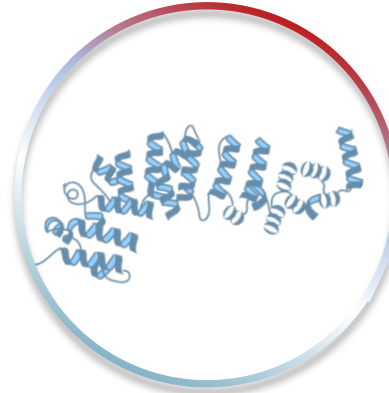
MYC

JAB-8263
JAB-2485



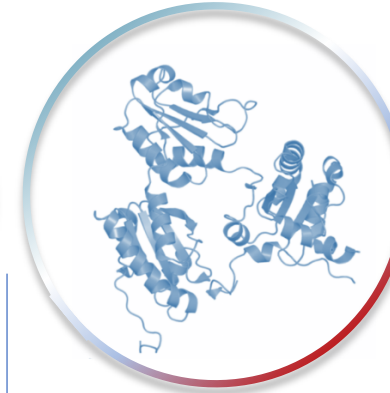
I/O pathways

JAB-3312/3068
JAB-BX102
JAB-BX300
JAB-26000



RB pathway

JAB-2485



Tumor Metabolic Pathways

JAB-24000



P53

JAB-30000

1. Nusse R, Clevers H. *Cell* 2017;169(6):985–99;
2. Lane DP. *Nature* 1992;358(6381):15–6;
3. Waters AM, Der CJ. *Cold Spring Harb Perspect Med* 2018;8(9):a031435;
4. Dang CV, et al. *Cell* 2012;149(1):22–35.

Our Diverse Pipeline Targeting Critical Pathways

Asset	Target & pathway	Indications	Combo Strategy	Pre-clinical	Phase I	Phase IIa
JAB-3312/ JAB-3068	SHP2 (RAS pathway, I/O)	Solid tumors, including NSCLC, HNSCC, ESCC	KRASI , PD-1, EGFRi	US, China		
JAB-21822	KRAS G12C (RAS pathway)	NSCLC, CRC	SHP2i , PD-1, EGFR mAb	US, China, Europe		
JAB-8263	BET (MYC pathway)	Solid tumors Blood tumors	Aurora Ai , JAKi, PD-1	US, China		
JAB-2485	Aurora A (RB pathway)	Solid tumors	BETi , SHP2i , KRASI	US		
JAB-BX102	CD73 mAb (I/O)	Solid tumors	PD-1	US, China		
JAB-24114	Undisclosed (Tumor metabolic pathway)	NSCLC, HNSCC	KRASI , P53i , BETi , PD-1			
JAB-BX300	Undisclosed (RAS pathway)	PDAC, CRC	SHP2i , KRASI			
JAB-26766	Undisclosed (I/O)	SCLC, HNSCC, ESCC	SHP2i , PD-1			
JAB-23400	KRAS ^{MULTI} (RAS pathway)	PDAC, CRC, NSCLC	SHP2i , SOS1i, Aurora Ai			
JAB-22000	KRAS G12D (RAS pathway)	PDAC, CRC, NSCLC	SHP2i , SOS1i, Aurora Ai			
JAB-23000	KRAS G12V (RAS pathway)	PDAC, CRC, NSCLC	SHP2i , SOS1i, Aurora Ai			
JAB-30000	P53 (P53 pathway)	Solid tumors	KRASI , JAB-24114 , PD-1			

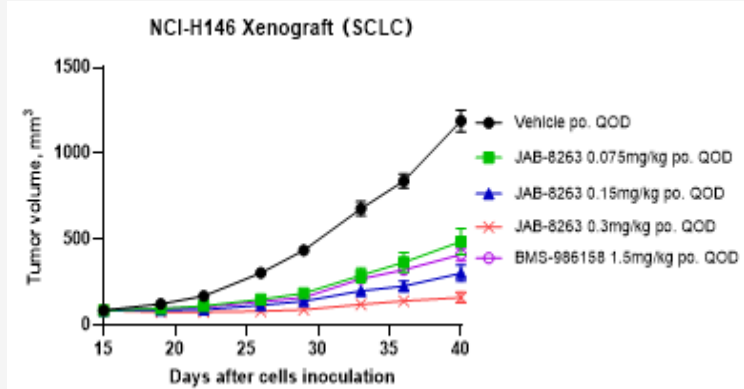
FIC/FIC potential

BIC potential

Potent BET Inhibitor JAB-8263

Asset	Regimen	Indications	IND	Phase I	Recent development	Upcoming Milestone (expected)
JAB-8263 BET	Mono	Solid tumors	US trial			RP2D to be determined in 2022 2H
	Mono	Solid tumors	China trial		FPI in Feb 2022	
	Mono Combo w/JAKi	MF and AML	China trial		FPI in Apr 2021	

Strong Antitumor Effect



- Highly effective in solid and hematologic cancer models
- Greater anti-tumor response at lower dose

JAB-8263(0.15 mg/kg) TGI=80% BMS-986158(1.5 mg/kg) TGI=71%

Preliminary Clinical Results

- Early signals of clinical benefit
- Wide safety window
- Favorable PK profile

Market Potential

To date, there has been no approved and marketed BET inhibitors globally.

Indications:

Solid Tumor

NMC, NSCLC, SCLC, CRPC, ESCC, Ovarian Cancer (4.7 million by 2030)

Hematologic Tumors

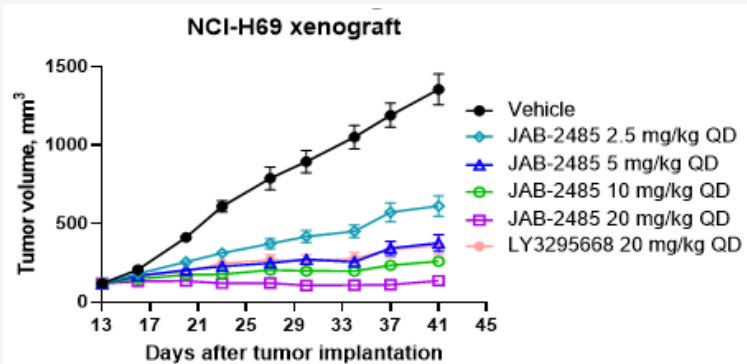
MF (127,500 by 2030) AML (187,600 by 2030)

Highly Selective Aurora A Inhibitor JAB-2485

Asset	Regimen	Indications	IND	Phase I	Recent development	Upcoming Milestone (expected)
JAB-2485 Aurora A	Mono	Solid tumors	<i>US trial</i>		IND approved in Jan 2022	FPI (2022 2H)

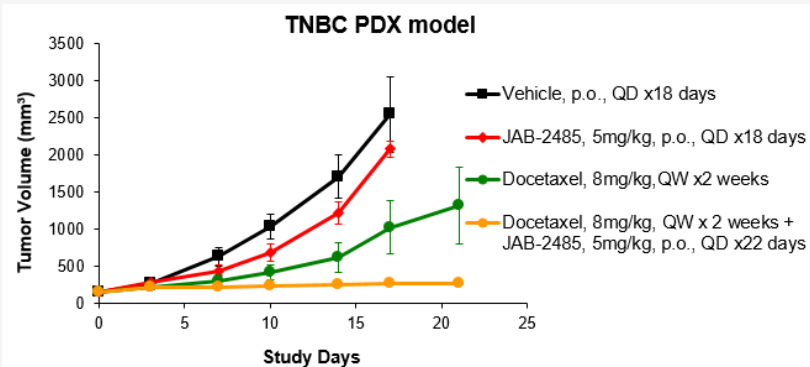
- **No** Aurora A kinase inhibitor had been **approved and marketed** globally.
- **JAB-2485** is one of top 3 highly selective Aurora A inhibitors in clinical stage

Strong Anti-tumor Effect in Xenografts



- Greater anti-tumor response at lower dose
 JAB-2485 (5 mg/kg) TGI=79% LY3295668 (20 mg/kg) TGI=79%

Combinational Anti-tumor Effect



- Significant synergistic effect when combining with chemotherapy.

- **More specificity:** ~2000-fold selectivity over Aurora B in biochemical and cell-based assay

- **Indications**
 Small cell lung cancer (330,000 new cases in 2020)

Triple negative breast cancer (339,000 new cases in 2020)

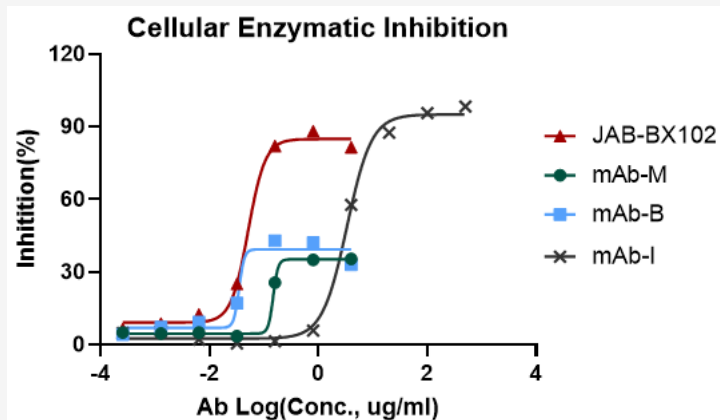
Highly Differentiated Anti-CD73 Antibody JAB-BX102

Clinical Plan

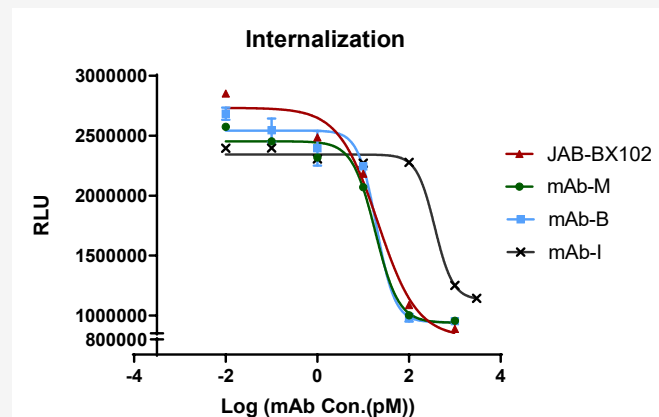
Asset	Regimen	Indications	IND	Phase I	Recent development	Upcoming Milestone (expected)
JAB-BX102 CD73 mAb (I/O)	Mono Combo w/PD-1 mAb	Solid tumors	<i>US trial</i>		IND approved in Oct 2021	FPI (2022 1H)
	Mono	Solid tumors	<i>China trial</i>		IND submitted in Jan 2022	

Excellent Efficacy & Safety

- Direct inhibition of CD73 enzyme activity – better efficacy



- Efficiently eliminate CD73 from the cell surface by inducing internalization of surface CD73






- ✓ Recognize the N-terminal epitope of CD73
- ✓ In vivo PK/PD model, complete CD73 inhibition **without the “hook effect”**

Key Pipeline Assets with First-in-Class Potential




Global TOP3 Potential

Clinical Stage

JAB-3068/3312




-  SHP2 inhibitor
-  2nd FDA IND
-  Clinical - Phase IIa

JAB-2485




-  Aurora A inhibitor
-  2nd FDA IND
-  Clinical - Phase I

2022 IND




JAB-24114

-  Tumor metabolic Pathway
-  One Program in Ph I
-  2022 IND

JAB-26766


-  I/O
-  One Program in Ph I
-  2022 IND

JAB-BX300 (mAb)




-  RAS pathway
-  One Program in Ph I
-  2022 IND

2023-2024 IND




JAB-23400

-  KRAS^{multi} inhibitor
-  No IND Globally
-  2023 IND




JAB-22000


-  KRAS G12D inhibitor
-  No IND Globally
-  2023 IND

JAB-30000

-  P53 inhibitor
-  No IND Globally
-  2023-2024 IND

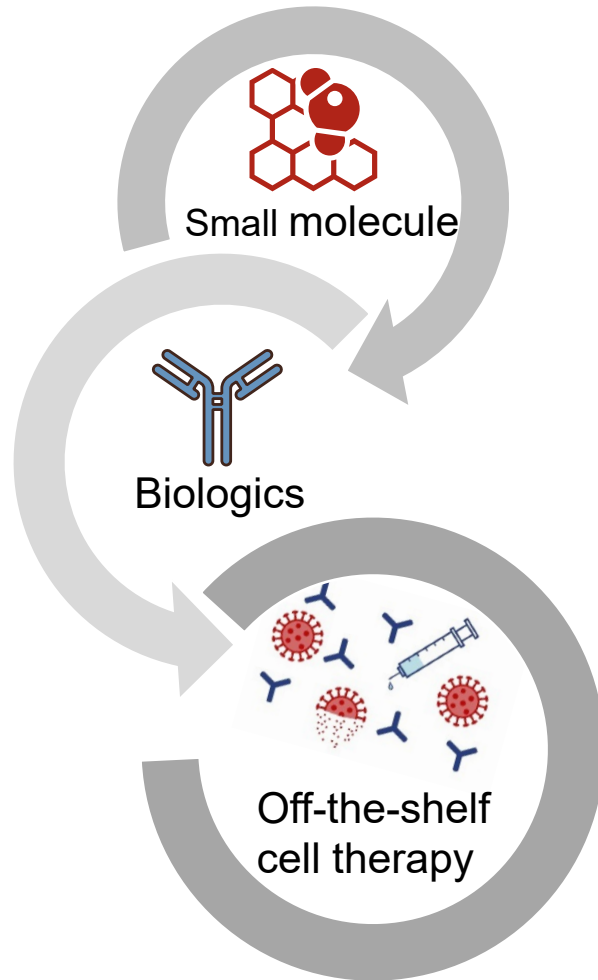
JAB-23000

-  KRAS G12V inhibitor
-  No IND Globally
-  2023-2024 IND

The background features a warm sunset gradient from light blue at the top to orange and yellow at the bottom. A central glowing sphere is composed of a network of white nodes and connecting lines, resembling a molecular or data structure. Two hands, one in the foreground and one slightly behind, are shown in silhouette, holding the sphere. The scene is decorated with numerous white starburst light effects scattered across the sky.

Expanding Our Pipeline to Off-the-shelf Cell Therapies

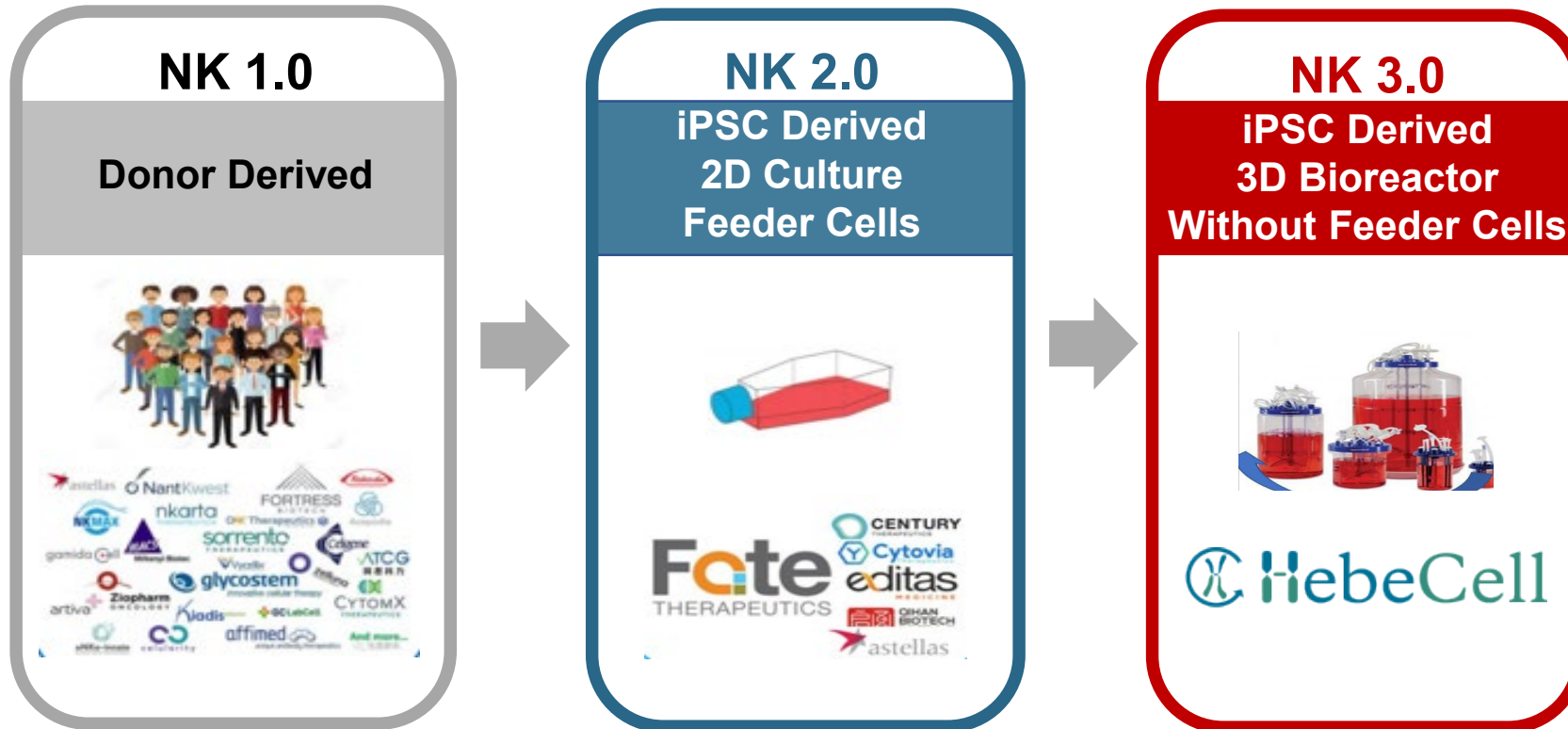
Strategic Layout in Cell Therapy



- Collaborate with Hebecell to develop the **next-generation** iPSC-NK cell therapy
- Key transaction terms:
 - \$25M of consideration in total
 - 19.74% of the share capital (fully-diluted)
 - Dr. Wang Yinxiang was appointed as Chairman of Hebecell
- **Boston R&D Center** - around 30 employees
- **Beijing R&D Center** - established in Q4 2021
- **Expected IND – 2023**
- Enhance our capability to **explore clinical value** of **combination** therapies between our current programs and off-the-shelf cell therapies

iPSC Derived-NK -- Hebecell Platform

3D Culture Without Feeder Cells



✓
3D Bioreactor
Cost-effective
industrial scalability

iPSC-NK Is Changing the Game in Cell Therapy

Universal, Off-the-Shelf Cell Products Derived from Renewable Master Cell Lines

	Autologous CAR -T	iPSC Derived NK
Safety		
Graft Versus Host Disease Risk(GvHD)	Low	Low
Cytokine Release Syndrome (CRS) or NeurotoxicityRisk	High	Low
Manufacturing		
Off-the-shelf Product	-	++
Cost of Manufacturing	+++++	+
Ease of Gene Editing	++	+++++
Master Cell Bank	-	+++
Homogeneous Product	+	+++
Batch to Batch Variation	Yes	No
Multiple Dosing	No	Yes
Efficacy		
Persistence	+++++	++
CAR-Independent Tumor Cytotoxicity	-	+

Advantages of iPSC-NK

- Off-the-shelf Availability
- Uniform Product
- Patient Accessibility
- Multiplexed Engineering
- Lower GvHD/CRS risk

A person in a white lab coat is pointing at a futuristic digital interface. The interface features various data visualization elements: a pie chart, a line graph with a y-axis from 0 to 100, a bar chart, a world map, and a grid of icons representing different medical and scientific fields. The overall aesthetic is clean and professional, with a light blue and white color palette.

On Target to Capture the Global Market

AbbVie Partnership Expedited Our Global Development



Transformative Collaboration

- Leverage a partner's global clinical, regulatory, medical, patient advocacy and commercial footprint
- **Rights of Parties**
 AbbVie – Worldwide
 (except for PRC, Hong Kong and Macau)
 Jacobio - PRC, Hong Kong and Macau

Financial Arrangement

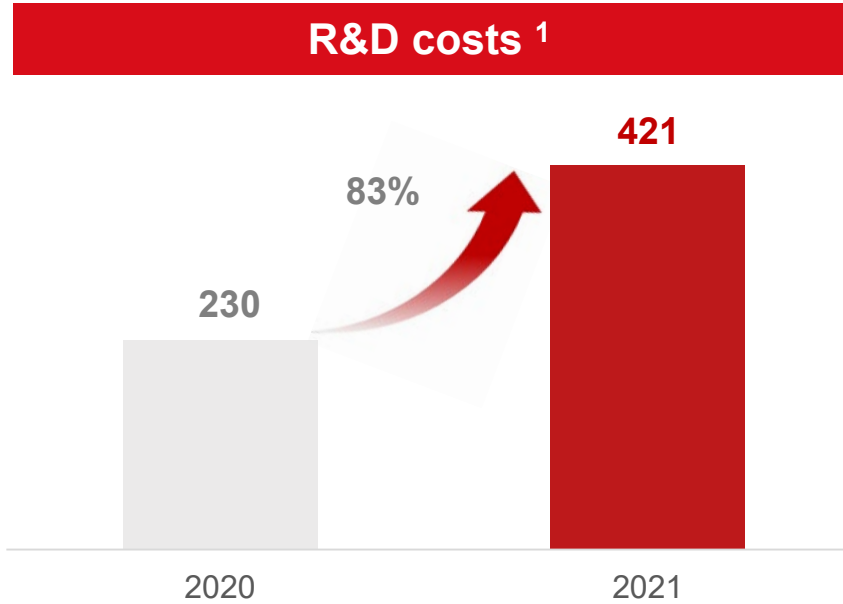
- **Upfront Payment (Received)**
\$45mm
- **Milestone Payments**
up to \$810mm - \$20mm received
- **Royalties**
Low-to-mid Double-digit percentages
- AbbVie will **reimburse** costs of global clinical development (incl. China) pre registrational trials



Financial Information

SELECT COMPANY FINANCIALS

(RMB mm)



Financial Position ²

Cash and Bank Balances @ 12/31/2021	1,538
Cash Runway	18-24 months

¹ R&D costs = Cost of revenue + Research and development expenses.
All R&D costs in relation to AbbVie Collaboration were recorded in “Cost of revenue” account.

² As of June 30, 2021, the Group did not have any interest-bearing bank and other borrowing.

Our Expansion

Global Headquarter
Beijing, China

2021 HC: 262 employees

In-house R&D

Allosteric Inhibitor
Tech Platform

Global FIC

6 Clinical Stage Assets
(2 Global TOP 3)
15+ Pipeline Assets
(9 Global TOP 3 Potential)

Global Market

Global Partnership
License-out Deals

Full Function Pharma

First NDA in 2023-2024



Beijing Headquarter
(under construction)



Beijing R&D Center



Beijing Clinical Office
(Dazu)



Beijing Clinical office
(Shoudong)



Shanghai Clinical Office



U.S. R&D Center
in MA



Hebecell R&D Center
in MA