

InnoCare Pharma 2019 Annual Report Presentation

April 2020

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To Become a **Global Biopharmaceutical Leader** that Develops and Delivers **Innovative Therapies** for Patients **Worldwide**

Oncology



Autoimmune

Our Therapeutic Focus



I N N O C A

Fully-integrated Biopharma Company



Drug Discovery

All Products Developed In-house

- 90+ research scientists
- Beijing R&D center 8,300 m²
 Chemistry, biology and CMC labs



- 800 m² AAALAC-like animal facility
- Nanjing R&D center 3,350 m²
 - A state-of-the-art solid-state research lab
- Diagnostic and biology platform

Target Identification

Protein Structure Aided Drug Design Prof. Yigong Shi

- Expertise in structure biology
- Deep understanding of cancer biology Novel Target Identification
- Prof. Zemin Zhang
- Single cell sequencing platform
- Big Data analysis



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3 Clinical stage assets

- Potential best-in-class BTK inhibitor targeting 2020
 - market launch
- 1 IND Submitted and Accepted
- 5 at IND enabling stage



Commercialization

- Developing Sales & Marketing strategy
- Building Sales & Marketing force
 - Sales and marketing head in-place
- Team of ~120 by product launch in 2020
- Unrivalled medical collaboration



Clinical Development

Unparalleled Clinical Execution

- ~70 Clinical development personnel
- All China trials managed in-house
- 180+ Patient Enrollment within a year
- 100+ Clinical sites initiated
- 10+ trials ongoing

Manufacturing



50,000 m² manufacturing facility in Guangzhou

- Designed to comply with both Chinese and international drug manufacturing standards
- Consisted of 46 employees as of Apr, 2020
- Est Completion: 2020

A Robust Product Pipeline



Balanced Drug Portfolio Targeting Both Proven and Novel Pathways



Registrational trials

Abbreviations: CLL = Chronic Lymphocytic Leukemia; SLL = Small Lymphocytic Lymphoma; MCL = Mantle Cell Lymphoma; MZL = Marginal Zone Lymphoma; CNSL = Central Nervous System Lymphoma; GCB = Germinal Center B-cell; DLBCL = Diffuse Large B-Cell Lymphoma; WM = Waldenstrom's Macroglobulinemia; FL = Follicular Lymphoma; SLE = Systemic Lupus Erythematosus; HCC = Hepatocellular Carcinoma.

Notes:

- 3. Initiation of Phase II trials for cholangiocarcinoma have begun, patient screening is expected to begin in second quarter of 2020
- 4. Expect to complete the Phase I trial for HCC in the fourth quarter of 2020
- 5. IND application for NTRK fusion-positive cancers submitted to the NMPA in the first quarter of 2020
- 6. Expect to submit an IND application for autoimmune diseases to the NMPA in the second half of 2020
- 7. The Company also has four undisclosed IND-enabling stage candidates currently under development

^{1.} Denotes the Company's Core Product Candidate, orelabrutinib (ICP-022)

For indications of *rtr* CLU/SLL and *rtr* MCL, the registrational trial for NDA submission is the Phase II clinical trial based on the communications with the NMPA. Confirmatory Phase III clinical trials will be required after the Company receives conditional approvals from the NMPA based on the results of these two registrational Phase I and Phase I clinical trials





Recent Development and Upcoming Milestones





Key Financials Updates





¹ Cash balance = investments measured at fair value through profit or loss + investments measured at amortised cost (both are wealth management products) + cash and bank balance. Net cash = cash balance – convertible loan (0, RMB957mm, and RMB1,117mm as of 31-Dec-2017, 31-Dec-2018, and 31-Dec-2019, respectively) – loans and borrowings – loans from a related party



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Section 1

Business Highlights





BTK Inhibitor: Large Market Potential

- Bruton's Tyrosine Kinase ("BTK") is a key component of the B-cell receptor signaling pathway, which is an important regulator of cell proliferation and cell survival in various lymphomas (mainly NHL). BTK inhibitors block B-cell receptor ("BCR") induced BTK activation and its downstream signaling. Successful blockage of BTK activation would result in growth inhibition and cell death of B-cells
- BTK is a proven target for the treatment of malignant B lymphomas with significant market potential
 - Only 3 BTK inhibitors approved globally and only 1 approved in China
 - BTK inhibitor global sales reached US\$4.5 billion in 2018
 - Currently approved BTK inhibitors, however, have demonstrated common toxicities, some of which are believed to be attributable to the off-target effects of these drugs, such as diarrhea, bleeding and atrial fibrillation
- Potential to treat autoimmune diseases such as systemic lupus erythematosus, rheumatoid arthritis, multiple sclerosis, pemphigus and lupus nephritis





China BTK Inhibitors Market Size



Advantages and Highlights



Orelabrutinib is a potential best-in-class late-stage BTK inhibitor **Improved Target Selectivity** Our "Point-of-Differentiation" Significant inhibition of only BTK by >90% Orelabrutinib and NO significant inhibition of other kinases Johnson-Johnson **Ibrutinib** M INNOCARE abbvie BeiGene Significant inhibition of kinases other than AstraZeneca Acalabrutinib BTK Zanubrutinib Ibrutinib Orelabrutinib Acalabrutinib Zanubrutinib Õ Late-stage / Approved Favorable PK/PD Profile and Better Target Occupancy The better bioavailability of orelabrutinib tablet enables **Target Selectivity** Once-daily administration at low dosage • Near 100% 24-hr BTK occupancy in blood • Safety Improved Safety and Robust Efficacy Profile Once-daily

BUSINESS HIGHLIGHTS 11





Improved Target Selectivity

KINOMEscan dendrogram



Source: "Potency and Selectivity of BTK Inhibitors in Clinical Development for B-Cell Malignancies" by Kaptein, A., et. al, Blood, 2018, 132 (Suppl 1) 1871; DOI: 10.1182/blood-2018-99-109973





Favorable PK/PD Profile

Post-dosing plasma exposure profile



Sources: Bruton tyrosine kinase inhibitor ibrutinib (PCI-32765) has significant activity in patients with relapsed/refractory B-cell malignancies, Advani R.H., et al. Journal of Clinical Oncology, 2013; 31(1):88-94. doi: 10.1200/JCO.2012.42.7906. Acalabrutinib (ACP-196) in Relapsed Chronic Lymphocytic Leukemia, Byrd J.C., et al, The New England Journal of Medicine, 2016; 374(4):323-32. doi: 10.1056/NEJMoa1509981 BeiGene corporate presentation dated June 5, 2019, http://hkexir.beigene.com/media/1238/bgne-investordeck-20190605.pdf





Better Target Occupancy

BTK occupancy



Abbreviations: SAD = single ascending dose; MAD = multiple ascending dose

Sources: "Acalabrutinib (ACP-196) in Relapsed Chronic Lymphocytic Leukemia" by Byrd J.C., et al. The New England Journal of Medicine, 2016; 374(4):323-32. doi: 10.1056/NEJMoa1509981; Company filings



Improved Safety and Robust Efficacy Profile



Abbreviations: CR=complete response, PR=partial response, PR-L= partial response with lymphocytosis, SD=stable disease, PD=progressive disease, ORR=objective response rate, DRC=disease control rate, DOR=duration of response

Sources: Imbruvica Prescribing Information, Jan 2019

Pooled Analysis of Safety Data from Clinical Trials Evaluating Acalabrutinib Monotherapy in Hematologic Malignancies, John C. Byrd, et al., Blood, 2017; 130:4326

NDA/BLA Multi-disciplinary Review and Evaluation, 210259Orig1s000, Center for Drug Evaluation and Research

Pooled Analysis of Safety Data from Monotherapy Studies of the Bruton Tyrosine Kinase (BTK) Inhibitor, Zanubrutinib (BGB-3111), in B-Cell Malignancies, S. Tam C., et al., European Hematology Association, Jun 15, 2019; 266776, PS1159

"Safety Analysis of Four Randomized Controlled Studies of Ibrutinib in Patients with Chronic Lymphocytic or Mantle Cell Lymphoma" by Susan O'Brien, et al., Original Study, 2018; 18(10), 648-657. e15

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Rapid Clinical Development for Treatment of B-cell Malignancies





Clinical Development Plan

Indication(s)	Type of Therapy	IND	Phase I	Phase II ⁽¹⁾	Phase III ⁽²⁾	NDA Filing
China						
r/r CLL/SLL	Mono					Accepted and given priority review status 1/2020
r/r MCL	Mono					Accepted and given priority review status 3/2020
r/r MZL	Mono					
r/r CNSL	Mono					
r/r WM	Mono					
1L: CLL/SLL ⁽³⁾	Mono				•	
r/r non-GCB DLBCL (double mutation)	Mono					
FL	Combo with CD20					
U.S.						
B-cell malignancies (basket)	Mono	US Development	t Status			

Registrational trials

Abbreviations: CLL = Chronic Lymphocytic Leukemia; SLL = Small Lymphocytic Lymphoma; MCL = Mantle Cell Lymphoma; MZL = Marginal Zone Lymphoma; CNSL = Central Nervous System Lymphoma; GCB = Germinal Center B-cell; DLBCL = Diffuse Large B-Cell Lymphoma; MXH = Waldenstrom's Macroglobulinemia; FL = Follicular Lymphoma.

Notes:

Notes: 1. Some indication(s) may not require a non-registrational Phase II clinical trial prior to the beginning of registration Phase II or III clinical trials 2. Some trials may require the completion of a Phase III clinical trial to submit an NDA application 3. Received approval from the NMPA to initiate a Phase III trial of Orelabrutinb as a first-line treatment for CLL/SLL

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Prevalence of SLE and other major autoimmune diseases (RA, MS, Psoriasis and LN) expected to grow rapidly



Abbreviations: LN = lupus nephritis, MS = multiple sclerosis, RA = rheumatoid arthritis

Rapidly Growing SLE Therapeutic Market Size



Other major autoimmune diseases Global Prevalence (MM) 129.4 129.4 12.9 13.7 2018A 2030E

SLE Competitive Landscape: Orelabrutinib vs. Other BTKi at Clinical Stage

Generic Name/Drug Code	Company	Global Filing Status
Orelabrutinib	👬 INNOCARE	Phase I (China)
Fenebrutinib	Roche	Phase II
Evobrutinib	Merck KGaA	Phase II
ABBV-105	AbbVie	Phase II
BIIB068	Biogen	Phase I
AC0058	ACEA Pharma	Phase I
SN1011	SinoMab	Phase I

1 NO BTKi approved for the treatment of SLE in the global market

Huge unmet medical needs

Orelabrutinib (ICP-022) : Potential First-in-class BTK Inhibitor for Autoimmune Diseases (Cont'd)



Robust Pre-clinical Efficacy Profile in Both SLE and RA

- Initiating a Phase Ib/IIa trial in combination with standard of care treatment for SLE in China •
- Explore orelabrutinib in other autoimmune diseases, such as LN, MS and pemphigus



Significant reduction of SLE-associated biomarkers · Improvement of survival in MRL/Ipr mice



Abbreviations: Anti-dsDNA = Anti-double-standard DNA: mpk = mg/kg.



- Dose-dependent reduction of proinflammatory cytokines, ameliorated arthritis histopathology scores
- Prevention of joint destruction

Representative micro-computed tomography images of rat ankle joints













Normal Vehicle Dex

Orelabrutinib Orelabrutinib Orelabrutinib 0.5mg/kg QD 1mg/kg QD 3mg/kg QD 10mg/kg QD

Orelabrutinib reduced erosive bone changes and prevented bone loss

Vehicle-treated group showed severe and widespread bone loss

Orelabrutinib's pre-clinical efficacy in arthritis rat model







Favorable Pre-clinical Profile with Improved Target Selectivity and High FGFR Inhibitory Potency



Source: Perera T. et al, Molecular Cancer Therapeutics 2017, 16(6), 1010-20. Doi: 10.1158/1535-7163.MCT-16-0589

ICP-192: Potential Best-in-class Pan-FGFR Inhibitor (cont'd)



Completed Phase I clinical trials and commenced Phase II clinical trials

Advantages and Highlights



Clinical program



³ ICP-105: Potential First-in-class FGFR4 Inhibitor







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Section 2

Growth Strategies

Growth Strategies



Commercialization Strategy



In a Staggered Approach Corresponding with the Launch Timeline of Orelabrutinib



World-class Manufacturing Facility



To Meet Commercial Scale Production and Comply with GMP Requirements





Guangzhou Subsidiary





Appendix

Other Information

Top-notch Executives & Advisors





- 20+ years of experience in research and development and company management in the pharmaceutical industry
- Former CEO and CSO of BioDuro, a PPD Company
- · Former Head of Early Development Team, Cardiovascular Diseases at Merck US
- · Former Fellow at The Howard Hughes Medical Institute
- The 17th President of the Sino-American Pharmaceutical Professional Association (SAPA)

Dr. Jisong Cui Co-founder and CEO





Co-founder, President of Scientific Advisory Board W!AS 浙江西湖高等研究院

ЕМВО

the U.S. and European Molecular Biology Organization

Professor of Tsinghua University and Princeton University

OF ARTS & SCIENCES **PRINCETON** K

AMERICAN ACADEMY

NIVERSITY



- 28 years of experience in clinical development
- Roche, Former Senior Medical Director
- Pfizer, Former Senior Associate Director
- University of Missouri-Kansas City, Former Fellow



Dr. Rick Xu Roche



CFO

UBS AG, Former Healthcare Equity **Research Analyst** Merrill Lynch Asia, Former Equity Research

Research Analyst

Analyst Mehta Partners LLC. Former Equity

Shaojing Tong

🛣 UBS bank of america 🧇 🕅



中国科学院

Elite Structural Biologist

- 20+ years of drug discovery experience
- · BioDuro, Former Executive Director of Medicinal Chemistry
- Pfizer, Former Principal Scientist
- Albert Einstein College of Medicine, Former Postdoctoral Researcher

Dr. Xiangyang Chen



BIODURO Pizer EINSTEIN STEIN



Prof. Zemin Zhang Scientific Advisory Board Member



Professor at Peking University

Former head of the bioinformatics division at Genentech Inc., USA





Director of the Clinical Immunology Center / Rheumatism Immunology



Scientific Advisory Board Member

- Professor emeritus at Institute of Advanced Study, Princeton
- US National Academy of Sciences member
- GM of Becton Dickinson's ٠ Greater China business
- Former CEO and president of Novartis Pharmaceuticals China





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Prof. Yigong Shi

Experienced Core Management Team



Research









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BetaPharma

Retter Medicine Retter Life

Bristol-Myers Squibb

Dr. Jean Wang Vice President of Formulation



Dr. Norman Kong

Executive Director of Chemistry

Roche





Zuopeng Wang Senior Director of PM

Clinical Development



Dr. Renbin Zhao

Executive Director of Regulatory Affairs

Dr. Richard Liu

Vice President of Biology



Reinna Zhang Director of Clinical Operation



Alan Zhu Director of Medical Research



Dr. Jason Zhang Director of Pharmacology



Dr. Rock Lv Associate Director of Medical Research



Corporate History and Milestones





Six Pre-clinical Drug Candidates







	Year Ended 31 December				
RMB'000	2017	2018	2019		
Revenue	102	1,617	1,247		
Gross Profit	102	1,617	1,247		
Other Income and Gains	11,424	31,395	104,449		
Selling and Distribution Expenses	-	(558)	(3,458)		
Research and Development Costs	(62,882)	(149,726)	(213,123)		
Administrative Expenses	(14,644)	(17,523)	(63,623)		
Other Expenses	(542)	(27,979)	(159,909)		
Fair Value Changes of Convertible Redeemable Preferred Shares	(272,686)	(387,804)	(1,814,018)		
Finance Costs	(2,537)	(3,441)	(1,916)		
Share of Profits and Losses of Joint Ventures	31	(4)	_		
Loss Before Tax	(341,734)	(554,023)	(2,150,351)		
Loss for the Year / Period	(341,734)	(554,023)	(2,150,351)		
Loss for the Year / Period Excluding Fair Value Changes	(69,048)	(166,219)	(336,333)		

Revenue was mainly generated from providing research and development services to biopharmaceutical companies; no product sales have been generated to date

Other Income and Gains

- Includes RMB10.4mm, RMB 17.5mm and RMB 28.3mm of government grants in FY2017, FY2018 and FY2019 respectively;
- Mainly comprised of government grants received from the PRC local government authorities to support our R&D activities. All conditions related to these government grants have been fulfilled



- Depreciation and Amortisation
 al Expenses
 Third Party Contracting Cost
- Direct Clinical Trial ExpensesShare-Based Compensation

Fair Value Changes of Convertible Redeemable Preferred Shares represents fair value increase of preferred shares issued by us from prior financing rounds



	As at 31 December		
RMB'000	2017	2018	2019
Non-Current Assets			
Property, Plant and Equipment	2,362	4,908	48,479
Goodwill	3,125	3,125	3,125
Other Intangible Assets	36,580	36,947	37,011
Right-of-use Assets	9,716	13,053	86,311
Investments in Joint Ventures	1,163	1,159	1,159
Other Non-current Assets	880	78,463	30,861
Total Non-current Assets	53,826	137,655	206,946

Current Assets

Total Current Assets	53,575	2,063,504	2,408,747
Cash and Bank Balances	36,874	1,876,618	2,291,773
Investments Measured at Amortised Cost	10,023	-	-
Investments Measured at Fair Value through Profit or Loss	-	169,054	80,347 -
Deposits, Prepayments and Other Receivables	6,678	17,788	36,590
Trade Receivables	-	44	37

Cash and cash equivalents as of 31 December 2019 amounted to RMB2,372mm, which includes:

 Investments Measured at Fair Value through Profit or Loss and Investments Measured at Amortised Cost (wealth management products denominated in RMB)

Cash and Bank Balance

-



	As at 31 December		
RMB'000	2017	2018	2019
Current Liabilities			
Trade Payables	2,958	2,193	8,197
Loans and Borrowings	25,000	50,395	-
Other Payables and Accruals	21,086	5,397	41,528
Deferred Income	2,234	90	645
Lease Liabilities	2,801	5,332	6,204
Loans from a Related Party	51,331	8,882	9,098
Total Current Liabilities	105,410	72,289	65,672
Net Current (Liabilities) / Assets	(51,835)	1,991,215	2,343,075
Total Assets Less Current Liabilities	1,991	2,128,870	2,550,021

Non-current Liabilities

Non-current Liabilities			
Convertible Redeemable Preferred Shares	330,316	1,934,750	4,213,772
Convertible Loan	-	957,269	1,117,176
Loans and Borrowings	50,220	-	-
Lease Liabilities	7,063	7,791	3,394
Deferred Income	420	61,398	157,389
Deferred Tax Liabilities	6,036	6,036	6,036
Total Non-current Liabilities	394,055	2,967,244	5,497,767
Equity			
Share Capital	3	3	4
Reserves	(392,067)	(904,304)	(3,004,714)
Non-controlling Interests	_	65,927	56,964
Total Equity	(392,064)	(838,374)	(2,947,746)

Convertible Redeemable Preferred Shares Represents fair value of preferred shares issued by us from prior financing rounds

Convertible Loan

 In August 2018, Guangzhou InnoCare Pharma Tech Co., Ltd. ("Guangzhou InnoCare") was jointly established by Guangzhou Kaide Technology Development Limited ("Guangzhou Kaide") and a subsidiary of the Company

 Guangzhou Kaide provided Guangzhou InnoCare with a RMB930 million convertible Ioan, which bears interest at 6.5% per annum and is due on 31 December 2024. Guangzhou Kaide has an option to convert the Ioan into ordinary shares of Guangzhou InnoCare under certain conditions

• The amount represents the fair value of the convertible loan



	Year Ended 31 December		
RMB'000	2017	2018	2019
Cash Flows from Operating			
Loss before Tax	(341,734)	(554,023)	(2,150,351)
Adjustments for			
Finance Costs	2,537	3,441	1,916
Share of Profits and Losses of Joint Ventures	(31)	4	-
Interest Income	(213)	(8,416)	(72,047)
Fair Value Changes of Convertible Loan	-	27,269	159,907
Fair Value Changes of Convertible Redeemable Preferred Shares	272,686	387,804	1,814,018
Depreciation of Property, Plant and Equipment	552	1,078	1,462
Depreciation of Right-of-use Assets	3,149	4,219	7,204
Amortisation of Other Intangible Assets	17	91	400
Share Based Payment Expenses	10,395	65,215	65,804
	(52,642)	(73,318)	(171,687)
Decrease/(Increase) in Trade Receivables	2	(44)	7
Decrease/(Increase) in Deposits, Prepayments and Other Receivables	12,497	(11,111)	(17,455)
Increase/(Decrease) in Trade Payables	2,795	(765)	6,004
Decrease/(Increase) in Other Payables and Accruals	(4,945)	311	36,132
Decrease/(Increase) in Deferred Income	(7,276)	58,834	(3,454)
Cash Used in Operations	(49,569)	(26,093)	(150,453)
Interest Received	213	8,416	70,700
Net Cash Flows Used in Operating Activities	(49,356)	(17,677)	(79,753)

	Year Ended 31 December		
RMB ² 000	2017	2018	2019
Cash Flow from Investing Activities			
Investment income in wealth management products	809	1,337	3,772
Receipt of government grant for property, plant and equipment	-	-	100,000
Purchase of Investments	(143,430)	(483,500)	(1,087,000)
Purchase of Items of Property, Plant and Equipment	(1,417)	(3,624)	(45,033)
Purchase of Other Intangible Assets	-	(16,458)	(464)
Increase in Other Non-current Assets	(880)	(77,583)	(29,536)
Proceeds Upon Maturity of Investments in Wealth Management Products	170,224	323,133	1,171,935
(Increase) / Decrease in Time Deposits	-	(631,414)	(66,206)
Investment in a Joint Venture	(132)	-	-
Net Cash Flows from / (Used in) Investing Activities	25,173	(888,109)	47,468
Cash Flows from Financing Activities			
Proceeds from issue of shares	-	-	9,342
Proceeds from Issue of Convertible Redeemable Preferred Shares	31,029	1,165,184	412,672
Proceeds form Convertible Loan	-	930,000	-
Loans from a Related Party	43,794	-	-
Repayment of Loans from a Related Party	-	(31,508)	-
Repayment of Loans from Third Parties	(20,000)	(25,000)	(50,000)
Finance Expense Paid	(1,823)	(3,080)	(2,222)
Capital Injection from a Non-controlling Shareholder of a subsidiary	-	70,000	-
Acquisition of Non-controlling Interests	(22,955)	-	_
Principal Portion of Lease Payments	(3,235)	(4,296)	(6,851)
Net Cash flows from Financing Activities	26,810	2,101,300	362,941
Net Increase in Cash and Cash Equivalents	2,627	1,195,514	330,656
Cash and Cash Equivalents at the Beginning of the Year/Period	32,228	36,874	1,245,204
Effect of Foreign Exchange Rate Change, Net	2,019	12,816	18,293
Cash and Cash Equivalents at the End of the Year/Period	36,874	1,245,204	1,594,153