

InnoCare Pharma (9969.HK, 688428.SH) 2023 Interim Results NDR

August 2023

Disclaimer



These materials are for information purposes only and do not constitute or form part of an offer or invitation to sell or issue or the solicitation of an offer or invitation to buy or subscribe for securities of InnoCare Pharma Limited (the "Company") or any of its holding company or subsidiaries in any jurisdiction. No part of these materials shall form the basis of or be relied upon in connection with any contract or commitment whatsoever.

The information or opinions contained in these materials has not been independently verified. No representation or warranty, whether expressed or implied, is made as to, and no reliance should be placed on, the fairness, accuracy, completeness or correctness of such information or opinions contained herein. The information and opinions contained in these materials are provided as of the date of the presentation, are subject to change without notice and will not be updated or otherwise revised to reflect any developments, which may occur after the date of the presentation. The Company, any of its affiliates, directors, supervisors, senior managers, officers, employees, advisers and their respective representatives shall not have any liability whatsoever (in negligence or otherwise) for any loss howsoever arising from or in reliance upon any information contained or presented in or derived from these materials or otherwise arising in connection with these materials.

These materials contain statements that reflect the Company's current beliefs and expectations about the future as of the respective dates indicated herein. These forward-looking statements are based on a number of assumptions about the Company's operations and businesses and on factors beyond the Company's control, and are subject to significant risks and uncertainties, and, accordingly, the actual results may differ materially from these forward-looking statements. You should not place undue reliance on any of such forward-looking information. The Company assumes no obligation to update or otherwise revise these forward-looking statements for new information, events or circumstances that emerge subsequent to such dates.





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

Oncology



Autoimmune

Our Therapeutic Focus

Strategy Execution Delivered Strong Growth & Development in 2023H1



Commercialization

- Total revenue reached RMB 378mn, +53.5% yoy growth
- Orelabrutinib sales +47.8% yoy growth
- Orelabrutinib rapid market penetration and hospital coverage after NRDL inclusion
- Highly experienced commercial team in hematology
- Tafasitamab
- Approved for Urgent Clinical Use in the Hainan Province
- Approved in Hong Kong
- Access for Urgent Clinical Use in Big Bay Area

Progress of Internal R&D Pipeline

Orelabrutinib

- r/r MZL NDA approved, first and only BTKi approved in China; r/r MCL approved in SG
- r/r MCL US registrational trial finished patients enrollment, NDA submission in mid-2024
- IL CLL/SLL registrational Phase III finished patients enrollment, NDA submission 2Q2024
- 1L DLBCL-MCD registrational Phase III ongoing
- ITP PoC in PII, PIII registrational trial initiated
- SLE Plla positive, Pllb enrollment ongoing, interim results expected by end of 2024
- MS PII: 24-week results: 92.3% relative new Gd+T1 lesion reduction at 80mg QD compared to placebo arm
- ICP-248 achieved FPI with excellent efficacy
- ICP-332 PII for AD will finish patients enrollment in Sept., result readout by end of 2023
- ICP-488 PI in healthy finished; early PoC in psoriasis cohorts started, PII initiated
- ICP-723 registrational trial ongoing, IND approved for pediatric arm
- ICP-192 registration trial for cholangiocarcinoma

License-in/Collaboration

- ICP-B04, Tafasitamab+LEN
 Finished enrollment in registrational trial, NDA submission 2Q2024
- ICP-B02 (CD3*CD20) Good efficacy observed in IV and SC cohorts
- ICP-B05 (CCR8) PI dose escalation ongoing

Platform

- Guangzhou manufacture facility is producing majority of commercial Orelabrutinib & all other clinical drug products
- Beijing biologics CMC facility started to operation
- Removed "B" in HKEx

Focus concerted efforts towards Company's 2.0 objectives Continue corporate culture of cost sensitive, strong execution & innovation

Research & Development *Product Pipeline – Liquid Cancer*



	Drug	Target	Indication(c)	Piabte	hts IND Enabling	Pichts IND Enabling		xpansion Pivotal Trial		al Trial	Expected	Market
	Didg	raiget	maleation(s)	rights		PHIa	PHIb	PH2*	PH2**	PH3	Filing	Market
			r/r CLL/SLL	$\langle $	NDA approved: 2	25 Dec 2020				,		
			r/r MCL	3	NDA approved: 2	25 Dec 2020						
			r/r MZL	3	NDA approved: 2	21 Apr 2023						
	ICP-022/ Orelabrutinib	ВТК	1L: CLL/SLL	3							2024	
			1L: MCL	$\langle $							Ŷ	
Liquid			1L: MCD DLBCL	3							X	
Cancer			r/r MCL		U.S. Developme	ent Status					2024	
	ICP-B04/ Tafasitamab	CD19	Tafa + LEN, r/r DLBC	al 🐝							2024	★ ^{HK}
	ICP-B02	CD3 x CD20	Hemato-oncology	$\langle $	Dose escalating	in IV&SC	•					
	ICP-248	BCL2	NHL/ALL/ Combo	3	Dose escalating							
	ICP-490	E3 Ligase	MM / DLBCL / Hemato-oncology	3	Dose escalating							
	ICP-B05	CCR8	Hemato-oncology		Dose escalating							

Comprehensive Coverage for Hemato-oncology



INNOCARE

Commercialization Review Increasing Sales Momentum in Orelabrutinib



Significant Growth of Net Sales



Successful Commercialization Strategy

- Net sales achieved RMB 321mn in 2023H1
- Swift implementation of NRDL¹ at local level
- Experienced and effective in-house commercial team
- Rapid coverage of hemato-oncology market in China
- First and only BTKi for MZL in China



¹ Indications included in NRDL: r/r Mantle Cell Lymphoma ("MCL") and r/r Chronic Lymphocytic Leukemia/Small Cell leukemia ("CLL/SLL") FPI to NDA took 1.5 years while FPI to launch to the market took 2.5 years

Strategies to Cover DLBCL





	Drug	Torgot	Indication	Diabte	IND Enchling	Dose Escalation	Dose E	xpansion	Pivota	al Trial	
	Drug	Target	Indication	Rights		PHIa	PHIb	PH2*	PH2**	PH3	
	ICP-022/		1L: DLBCL - MCD	3							_
	Orelabrutini b		Combo w/ CD20 r/r DLBCL	3	Combo w/ MIL-62 ((basket)					
	ICP-B04/ Tafasitamab	CD19	Tafa+LEN, r/r DLBCL	**							нк
DLBCL	ICP-B02	CD3 x CD20	DLBCL/Hemato- oncology	3							
			DLBCL/Hemato- oncology	$\langle $							
	ICP-490	E3 ligase	Combo w/ CD19 DLBCL/Hemato- oncology	3							
	ICP-248	BCL2	Combo w/ Orela r/r DLBCL	3							
Registrationa	al trials	linical Stage	Pre-clinical Stage	+ Listed drug							



Improved Safety and Robust Efficacy Profile, No severe AF case observed after 850+ patient dosed.

Efficacy Profile

r/r CLL/SLL

Safety Profile

	Orelabrutinib (ICP-CL-00103,	Ibrutinib Resonate	Acalabrutinib ASCEND	Zanubrutinib (BGB-3111-	Adverse events of special interest	Orelabrutinib N=550* (%)	Ibrutinib N= 1,476 ¹ (%)	Acalabrutinib N= 1,029 ² (%)	Zanubrutinib N= 629 ^{3,4} (%)
Median	N=80)'	(n=195) ²	(n=155) ³	205, N=91) *	Any grade diarrhea	6.0%	43.8%	31%	20%
Follow-up Time	47 months	44 months	36 months	34 months	onths >= Grade 3 Atrial fibrillation	0	4.0%	1.1%	0.6%
ORR	93.8%	91%	93%	87.9%	Second primary malignancies	0.4%	10%	12%	9%
CR / CRi	30%	9%	5%	6.6 %	Major	1 19/	40/ #	0.70/	20/
PR / nPR	52.5%	78%	78%	69.2%	hemorrhage	1.1%	4%"	2.1%	3%
PR-L	11.3%	4%	10%	12.1%	≥ Grade 3 Infection	9.6%	21%	19%	23%

r/r MCL (N=106, median follow time of 39.4 months)

- 83% patients achieved ORR and 87.7% patients achieved disease control.
- CR rate, by conventional CT method, increased to 36.8% and it was expected a higher rate of in depth response may occur with prolonged treatment.
- The median PES was 27.4 month and the median OS was not reached.

r/r MZL (N=90, median follow time of 24.3 months)

- First BTKi for MZL in China.
- ORR was 58.9% assessed by independent review committee ("IRC").
- The median duration of response ("DOR") was 34.3 months (95% CI).
- The estimated 12-month PFS and OS were 82.8% and 91%.

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, 210259Orig15000, Center for Drug Evaluation and Research Pooled Analysis of Safety Data from Monotherapy Studies of the Bruton Tyrosine Kinase (BTK) Inhibitor, Zanubrufnib (BGB-3111), in B-Cell Malignancies, S. Tarr C., et al., European I Jun 15, 2019; 202776, PS1159; Vu V, et al. J Hematol Oncol. 2020 Way 11.3(11):45; Huang X, et al. Cancer Med. 2018 Apr;7(4):1043-55; Byrd JC, et al. 2017 ASCO poster 272. et al., European Hematology Association, Gria P, et al. J Clin Oncol. 2020 May 27, JCC1903355 Stafety Analysis of Four Randomized Controlled Studies of Itrutinibi In Patients with Chronic Lymphocytic or Manife Cell Lymphocytic by Susan O'Brien, et al., Original Study, 2018; 18(10), 648-657, e15

Efficacy data cut off data: 2022.12.30 (MCL & CLL/SLL); 2022.10.9 (WM)

Safety profile Note: Data cut off date 2022.12.24

^{*2} Grade 3, serious, or any grade central nervous system bleeding events, two cases, one with intracranial hemorrhage (65-year old patient with >10 hypertension) and the other with vitreous hemorrhage which was assessed as unlikely related to the treatment of orelabrutinib.** Data cutoff date Octobe 31, 2020. § one AML and one bladder cancer (based on TEAEs irrespective of causality assessment).¥ ≥ Grade 3, serious, or any grade central nervous system bleeding events. # From 2.838 pts who received ibrutinib in 27 clinical trials ## Bruising and petechiae excluded.1 Imbruvica US prescribing in Calquence US prescribing information3 Brukinza US prescribing information 4 Brukinza NDA Multi-Discipline Review





Current Status and Further Development

- Registrational trial for r/rDLBCL finished enrollment in mainland China, NDA submission 2Q2024, NDA approval in 1-2Q2025
- Approved for Urgent Clinical Use in the Hainan Province
- BLA was approved in Hong Kong and approved for pilot use in GBA

Competitive	Landscape: Sel	ected Novel T	Therapy in r/r DLBCL
-------------	----------------	---------------	----------------------

Company	Target	Therapy	Phase	ORR (%)	CR (%)	mDOR (m)	mPFS (m)	mOS (m)
Incyte/InnoCare	CD19	Tafasitamab + Lenalidomide	Approved ex-China	57.5	40	43.9	11.6	33.5
ADC Therapeutics	CD19 ADC	Loncastuximab tesirine	Approved ex-China	48.3	24.1	10.25	4.93	9.92
Roche	CD79b ADC	Polatuzumab vedotin + BR vs BR	Approved	42 vs 18	23 vs 3	12.6 vs 7.7	9.5 vs 3.7	12.4 vs 4.7
Roche	CD20/CD3	Glofitamab	BLA	52	39	10.4	3.8	11.5
Amgen/ Beigene	CD19/CD3	Blinatumomab	II	43	19	11.6	3.7	5.0
Regeneron/ Zai Lab	CD20/CD3	Mosunetuzumab	II	33	21	N/A	N/A	N/A
AbbVie	BCL2	Venetoclax+R+Pola	II	65	31	5.8	4.4	11
				L/				Leeeee

3

Major Program Update

ICP-248 (BCL2 Inhibitor): Best Combination Partner for Orelabrutinib









- Dose escalation of IV cohorts completed, 1st SC cohort completed
- Good efficacy observed in both IV and SC cohorts in FL and DLBCL patients
- Well tolerated with no DLT observed, low grade and manageable CRS
- SC formulation improves safety and convenience
- Significant potential across a broad range of indications in NHL as mono or combo therapies.



Superior anti-tumor activity





Major Program Update ICP-490: Highly Potent Next Generation CRBN Modulator





500·

DMSO ICP-490

+

Source a: Reference: a: Jan, M., Sperling, A. S., and Ebert, B. L. (2021). Cancer therapies based on targeted protein degradation - lessons learned with lenalidomide. Nature Reviews Clinical Oncology 18, 401-417.

Synergistic

Combinations

DLBCL.

etc.)

Autoimmune Disease Strategy





Orelabrutinib (ICP-022): Phase III Registrational Trial for ITP Initiated





- Phase II: 40% patients met the primary endpoint at 50mg QD
- Phase III: registrational trial being initiated in China
- Frontline BTK inhibitor gets approved for AID
- Considering global markets

Orelabrutinib (ICP-022): SLE Phase IIa Positive Results Lead to Further Development B & CARE



SLE Prevalence



Global (million number of patients)

SLE Phase II Study Results¹



placebo = orelabrutinib 50 mg = orelabrutinib 80 mg = orelabrutinib 100 mg

- SLE Responder Index ("SRI")-4 response rates increased in a dose dependent manner
- Trends of reduction in proteinuria level and improvement of immunologic bio-markers²
- The only BTK inhibitor ever shown efficacy in Phase II SLE trials
- PIIb enrollment ongoing, interim results expected by end of 2024

¹ The Phase IIa trial evaluated the safety and efficacy of Orelabrutinib plus standard of care verse placebo plus standard of care ("**SoC**") in patients with mild to moderate SLE ² Reduced immunoglobulin G and increased complements C3 and C4 were observed

Major Program Update: MS Phase II Results Orelabrutinib (ICP-022): Potential Best-in-class BTKi for Multiple Sclerosis



Key Findings

- All orelabrutinib groups achieved T1 new lesion control after 4 weeks of treatment, and the effect is sustained up to 24 weeks
- 92.3% relative reduction achieved in cumulative number of new Gd + T1 lesions 24 weeks at 80mg QD compared to placebo arm
- Best-in-class profile



Cumulative number of New Gd+ T1 Lesion from Week 4 to Week 24	Placebo / Orela 50mg QD (N=27)	Orela 50mg QD (N=30)	Orela 50mg BID (N=29)	Orela 80mg QD (N=29)
Adjusted mean cumulative number (95% CI) of lesions from W4 to W24	6.45 (3.62, 11.52)	2.10 (0.62, 7.11)	1.08 (0.30, 3.81)	0.50 (0.09, 2.74)
Percent reduction		67.4 (-22.0, 91.3)	83.3 (33.2, 95.8)	<mark>92.3</mark> (56.5, 98.6)
P-value		0.0958	0.0114	0.0037

Notes: The adjusted mean cumulative number, percent reduction (orelabrutinib vs placebo) associated with the 95%CI and p-value are estimated from a poisson regression model with a pearson scale parameter with a log link function and offset by log number of scans as of that visit. Baseline number of Gd+ T1 brain lesions is included in the model as a continuous covariate.

The above analyses are based on PHS population that includes all randomized subjects, but excludes the subjects who missed any one of the three MRI data points within first 12 weeks due to Covid-19 or unexpected events including Ukraine war and early termination per US FDA partial clinical hold.

nn

ICP-332:TYK2 JH1 Inhibitor for Oral Treatment of AD and Other Indications



Atopic Dermatitis



SelectivityDrugTYK2 vs. JAK1
(fold)TYK2 vs. JAK2
(fold)JAK1 vs. JAK2
(fold)ICP-332~4010

Evaluate JAK1/TYK2 inhibitor for AD and other indications

ICP-332 (TYK-2, JH1)

- Phase I study: SAD, MAD, food effect completed
 - Demonstrated a dose proportional and favorable PK profile, no significant food effect observed
 - Safe and well-tolerated, no significant decrease of platelet and hemoglobin (JAK2-related AE) observed and no DLT observed
- Phase II trial for atopic dermatitis (80 and 120 mg QD doses) will finish patients enrollment in Sept. 2023, study readout by end of 2023

Strategies: Targeting Type 2 Inflammation by JAK-Inhibitor



ICP-488:TYK2 JH2 Inhibitor for Oral Treatment for Psoriasis and Other Indications



ICP-488 (TYK-2, JH2)

- An oral, potent and allosteric TYK2 inhibitor that selectively binds to the JH2 pseudokinase domain with no activities on JAK1-3
- Phase I study
 - Completed SAD (maximum dosage to 36mg),
 MAD and food effects arms, no DLT
 observed
 - 2 cohorts of psoriasis patients for early PoC
- Phase II being initiated
 - Potential to show significant advantages in safety profiles verse other JAK family inhibitors.

Solid Tumor Strategy





ICP-723: Favorable Clinical Results with Potential Best-in-Class Profile



ICP-723 (Zurletrectinib,TRK)

- 2nd generation TRKi overcomes acquired resistance to 1st generation TRKi
- Phase I study demonstrated favorable PK profile and excellent anti-tumor activity
- No DLTs observed in Phase I dose escalation study (1-20 mg)
- Phase II registration trial for NTRK gene abnormalities ongoing, 80-90% ORR, NDA submission expected by end of 2024
- 1 PR in larotrectinib-resistant patient
- IND for pediatric patients approved
- Exploring in patients with ROS1 mutations

NTRK Gene Fusion Mutation is an Oncogenic Driver for a Variety of Cancer Types





Major Program Update ICP-192: Promising Safety and Efficacy in Phase II Trials



ICP-192 (Gunagratinib, FGFR)

- Finished phase I dose-escalation 2 mg to 26 mg, no DLT observed
- Safe and well-tolerated in patients with advanced solid tumors
- Registrational trial is ongoing at 20 mg in cholangiocarcinoma
- Exploring multiple other indications in solid tumor

A Glance at FGFR Mutation by Solid Tumor Types Worldwide



Frequency of All Currently Known FGFR 1, 2, 3 and 4 Aberrations



Source: Frost & Sullivan Analysis

Source: Helsten et al., 2015, Clinical Cancer Research



ICP-189: Potentially Best-in-Class and First-in-Class SHP2 Inhibitor with Large Potential in Combinational Treatments





Research & Development *Product Pipeline – Autoimmune & Solid Tumor*





Anticipated Milestones & Catalysts in Next 12 Months



Liquid	 Orelabrutinib 1L CLL/SLL NDA submission r/r MCL NDA submission in the U.S. 			
Cancer	 ICP-248 Preliminary data readout; U.S. IND filing Combo with Orelabrutinib for CLL/SLL i Phase II pivotal study in r/r CLL/SLL 	n the U.S. and CN		
Auto-immune Diseases	 Orelabrutinib Complete SLE PIIb patient enrollment, interim readout by end of 2024 Complete ITP PIII patient enrollment MS path-forward 	 ICP-332 Phase II AD data readout Phase III study initiation ICP-488 PoC in psoriasis; PII psoriasis initiation 		
Solid Tumors	 ICP-189 Phase I data readout Start combo study with EGFR in NSCLC 	 ICP-723: Complete patient enrollment of registrational trial; NDA submission ICP-192: Strive to complete patient enrollment 		

Company 2.0 Objective: Provide More Innovative Drugs to Patients

≥ 6 commercial products

- Marketed: Orela-Hema^①, Tafa^{*} (Hainan, HK, GBA)
- 2025-6: Tafa² (China mainland), ICP-723³, ICP-192⁴

2027-8: Orela-AID⁽⁵⁾ (*ITP, SLE, MS*); ICP-248⁽⁶⁾, ICP-332⁽⁷⁾, ICP-488⁽⁸⁾, ICP-490, ICP-189, ICP-B02, ICP-B05.....

- A recognized leader in hematology
- A strong competitor in **autoimmune diseases** and solid tumor
- Additional 5-10 well-positioned assets in R&D, unique research platforms
- Powerful engine in R&D, BD, manufacturing and commercialization platforms, operational excellence
- 3-4 products globalization (out-license, partnership, etc.)
- Annual revenue reaches significant numbers

2028

Now

Financial Review Key Financials for 2023H1





R&D Costs



(RMB mn)





¹ Non-HKFRS: excluding foreign exchange and share-based compensation impact

² Cash and cash equivalents = investments measured at fair value investments, cash and bank balance, interest receivable

* Successful STAR Board listing on Sept. 21, 2022



科学驱动创新 患者所需为本

Science Drives Innovation for the Benefit of Patients