

HUTCHISON CHINA MEDITECH

**Company Overview** 

(AIM/Nasdaq: HCM)

**June 2017** 

# CHI-MED

# Safe harbor statement & disclaimer

This presentation contains forward-looking statements within the meaning of the "safe harbor" provisions of the U.S. Private Securities Litigation Reform Act of 1995. These forwardlooking statements can be identified by words like "will," "expects," "anticipates," "future," "intends," "plans," "believes," "estimates," "pipeline," "could," "potential," "believe," "first-in-class," "best-in-class," "designed to," "objective," "quidance," "pursue," or similar terms, or by express or implied discussions regarding potential drug candidates, potential indications for drug candidates or by discussions of strategy, plans, expectations or intentions. You should not place undue reliance on these statements. Such forward-looking statements are based on the current beliefs and expectations of management regarding future events, and are subject to significant known and unknown risks and uncertainties. Should one or more of these risks or uncertainties materialize, or should underlying assumptions prove incorrect, actual results may vary materially from those set forth in the forward-looking statements. There can be no quarantee that any of our drug candidates will be approved for sale in any market, or that any approvals which are obtained will be obtained at any particular time, or that any such drug candidates will achieve any particular revenue or net income levels. In particular, management's expectations could be affected by, among other things: unexpected regulatory actions or delays or government regulation generally; the uncertainties inherent in research and development, including the inability to meet our key study assumptions regarding enrollment rates, timing and availability of subjects meeting a study's inclusion and exclusion criteria and funding requirements, changes to clinical protocols, unexpected adverse events or safety, quality or manufacturing issues; the inability of a drug candidate to meet the primary or secondary endpoint of a study; the inability of a drug candidate to obtain regulatory approval in different jurisdictions or gain commercial acceptance after obtaining regulatory approval; global trends toward health care cost containment, including ongoing pricing pressures; uncertainties regarding actual or potential legal proceedings, including, among others, actual or potential product liability litigation, litigation and investigations regarding sales and marketing practices, intellectual property disputes, and government investigations generally; and general economic and industry conditions, including uncertainties regarding the effects of the persistently weak economic and financial environment in many countries and uncertainties regarding future global exchange rates. For further discussion of these and other risks, see Chi-Med's filings with the U.S. Securities and Exchange Commission and on AIM. Chi-Med is providing the information in this presentation as of this date and does not undertake any obligation to update any forward-looking statements as a result of new information, future events or otherwise.

In addition, this presentation contains statistical data and estimates that we obtained from industry publications and reports generated by third-party market research firms, including Frost & Sullivan, an independent market research firm, and publicly available data. All patient population, market size and market share estimates are based on Frost & Sullivan research, unless otherwise noted. Although we believe that the publications, reports and surveys are reliable, we have not independently verified the data. Such data involves risks and uncertainties and are subject to change based on various factors, including those discussed above.

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All references to "Chi-Med" as used throughout this presentation refer to Hutchison China MediTech Limited and its subsidiaries. This presentation should be read in conjunction with Chi-Med's final results for the year ended December 31, 2016, copies of which are available on Chi-Med's website (<a href="https://www.chi-med.com">www.chi-med.com</a>).



# A risk-balanced global-focused BioPharma

## **Innovation Platform**

Broad late-stage pipeline

- $\checkmark$  (8 oncology) drug candidates in 30 studies worldwide.
- ✓ 1<sup>st</sup> positive Ph.III result fruquintinib Launch 2018.
- ✓ 7 further Phase III trials; 3 underway & 4 in-planning.
- √ ~330-person Scientific Team.

### **Commercial Platform**

Solid cash flow from operations

- √ ≥3,300-person China Sales Team (~2,200 med. reps).
- ✓ To commercialise Innovation Platform drugs in China.
- ✓ 2016 sales<sup>[1]</sup> up 21% to \$627.4 million.
- ✓ 2016 net income<sup>[2]</sup> up 180% to \$70.3 million.<sup>[3]</sup>



# Exceptional scale for pre-approval biotech

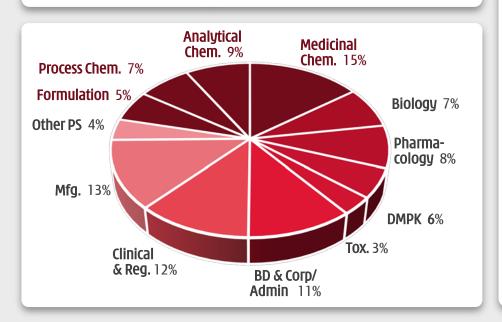




One of the leading China-based innovators in oncology & immunology

# ~330 SCIENTISTS & STAFF[1]

- ✓ 208 with advanced technical degrees
- **√** 26 M.D.S
- √ 54 doctorate degrees



#### **OUR ADVANTAGES**

- ✓ Large-scale fully integrated in house platform chemistry, biology, pharmacology, DMPK, toxicology, CMC, clinical & regulatory, and translational organizations working together seamlessly and continuously.
- ✓ China clinical speed

  major unmet medical needs (3.4 million new cancer patients / year<sup>[2]</sup>),
  rapid development and regulatory support. Allows for study of
  multiple indications and proof-of-concept in China.
- ✓ **Competitive costs** overall clinical costs, particularly pre-PoC, a fraction of US or Europe.
- ✓ **Constancy of purpose**Over 15 years with continuous financial support.

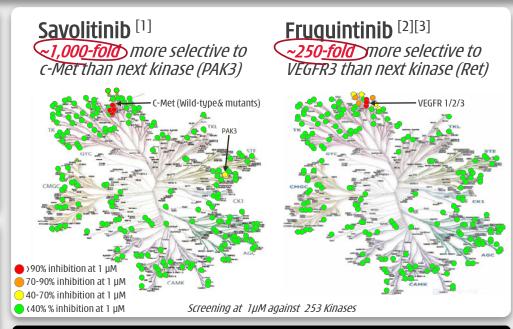
<sup>[1]</sup> Headcount as of December 31, 2016; Chem. = Chemistry; DMPK = Drug, Metabolism, & Pharmacokinetics; Tox. = Drug Safety Evaluation; PS = Pharmaceutical Science (CMC); Mfg = Manufacturing; Reg. = Regulatory; C&R = Clinical & Regulatory; BD = Business Development; [2] Frost & Sullivan.

# Chemistry is our edge

### Seriously selective small molecules

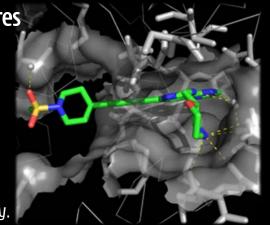


- 1. Fragment-based design of Novel Chemical Entities.
- Internally designed (all 8) clinical drug candidates.
- Use of co-crystal structures.
- Focus on small molecule interactions with tyrosine kinases - proteins/enzymes involved in cell signaling.
- 2. Total focus/discipline in designing and progressing drug candidates with superior kinase selectivity.
- Optimize binding to on target protein, minimize offtarget protein binding.
- No off-target kinase inhibition gives compound the chance to be more potent, attaining better target coverage with less toxicity.
- Combinability clean compounds allow for combinations with other tyrosine kinase inhibitors ("TKIs"), immunotherapy & chemotherapy agents.



# Use of co-crystal structures Focus on small molecule interactions with kinases ✓ Optimize binding to ontarget protein, for potency.

 Minimize binding to offtarget proteins for selectivity



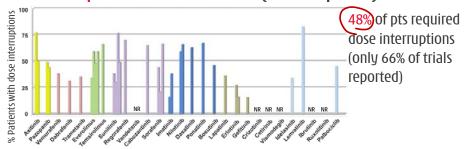
# Superior selectivity = Better tolerability



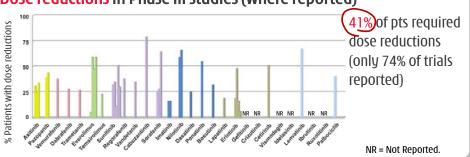
### More patient use = prolonged/total target coverage = better efficacy

- 3. Better tolerability important for sustained usage... Review of 28 FDA approved small molecule oncology targeted therapies revealed high incidence of toxicity<sup>[1]</sup>
- Pronounced in drugs with narrow therapeutic index (i.e. efficacious dose at or near MTD).
- Combination trials even harder 64% with grade 3-4 toxicities vs. 37% in monotherapy trials.

#### **Dose interruptions in Phase III studies (where reported)**



#### Dose reductions in Phase III studies (where reported)



# 4. ...whereas 1<sup>st</sup> gen. multi-kinase inhibitors require substantial dose modifications (interruptions/reductions).

Drug - targets	<b>2016 Sales</b>	Phase III Study	Dose Interruptions	Dose <u>Reductions</u>
Sunitinib (Sutent®) -VEGFR1,2,3, PDGFRβ, FLT3, CSF-1R, c-Kit, Ret	\$1.10b	1L RCC – Sunitinib vs. placebo	54% vs 39%	52% vs 27% (Gr 3/4 AE: 77% vs 55%)
Sorafenib (Nexavar®) – RAF, VEGFR2, PDGFRβ, Flt3, c-Klt, FGFR1	\$0.87b	1L RCC – Sorafenib Vs. placebo		(Gr 3/4 AE: 38% vs 28%)
Axitinib (Inlyta®) – VEGFR1,2,3, PDGFRα, c-kit	\$0.40b	2L RCC - Axitinib Vs. Sorafenib	Dose Mods: 55% vs 62%	34% vs 54%
Pazopanib (Votrient®) - VEGFR1,2,3, c- KIT, ITK, LCK, PDGFRα,β, FGFR1,3, c-Fms	\$0.73b	1L/2L RCC – Pazopanib vs. placebo	42%	36%
Regorafenib (Stivarga®) - VEGFR1,2,3, Raf, Ret, PDGFR, c-Kit	\$0.31b	2L CRC - Regorafenib vs. placebo	61%	38%
Lenvatinib (Lenvima®) – VEGFR1,2,3, Ret, PDGFR, c-Kit, FGFR1,2,3,4	\$0.20b	DTC - Lenvatinib vs. placebo	82% vs 18%	68% vs 5%
Cabozantinib (Cometriq®) – AXL, c-Kit, FLT-3, MET, RET, TIE-2, TrkB, VEGFR1,2,3	\$0.14b	2L RCC – Cabozantinib vs. everolimus		62% vs 25%
Savolitinib – c-Met (Ph I/Ib/II)		Several open-label studies	28%	8%
Fruquintinib – VEGFR1,2,3 (Ph II)		≥3L CRC - Fruquintinib vs. placebo	<b>34%</b> vs. 13%	28% vs. 13%
Fruquintinib - VEGFR1,2,3 (Ph II)		3L NSCLC – Fruquintinib vs. placebo	13% vs. 0%	13% vs. 0%
Sulfatinib - VEGFR 1,2,3, FGFR1		Several open-label studies	34%	17%
Epitinib – EGFR (Ph I/II)		NSCLC w/brain mets - Epitinib (Ph I/Ib)	13%	6%

<sup>[1]</sup> FDA approved btw Jan '02 to Feb '15. Roda D et al. "Are Doses and Schedules of Small-Molecule Targeted Anticancer Drugs Recommended by Phase I Studies Realistic?" Clinical Cancer Research 2016 May 1;22(9):2127-32.

# 30 active clinical trials on 8 drug candidates



1<sup>st</sup> positive pivotal readout - 4 lead candidates all in pivotal Ph.III in 2017

Prog	ıram	Target	Partner	Study number/Indication	Latest Status	Line	Target patient	Combo therapy	Site	Preclin.	Ph.I	Proof-of-conc	ept Pivota	al/Ph.III
1				1. Papillary renal cell carcinoma	Report Ph.II Feb. 2017; Ph.III start H12017	1st	c-Met-driven		Global					*
			AstraZenec	2. Papillary renal cell carcinoma	NCI Ph.II – savo vs. sunitinib vs. cabozan. vs. crizot.	All	c-Met-driven		US		į	<b>)</b>		
				3. Papillary renal cell carcinoma	Ph.Ib enrolling (dose finding)	-	All	durvalumab (PD-L1)	UK		i,		*	
				4. Clear cell renal cell carcinoma	Start when Study 2/4 begin Ph.Ib expansion stage	2nd	VEGF TKI refractory		UK				*	
			<u>ئۇ</u> .	5. Clear cell renal cell carcinoma	Ph.Ib enrolling (dose finding)	2nd	VEGF TKI refractory	durvalumab (PD-L1)	UK		į,		*	
Savol	itinih		Ze	6. Non-small cell lung cancer	Ph.IIb expans'n enrolling; Pivotal decision 2017	2nd	EGFR TKI refractory	Tagrisso® (T790M)	Global		i i	i	*	
(AZD6		c-Met	Ž	7. Non-small cell lung cancer	Ph.II enrolling	3rd	EGFR/T790M TKI	Tagrisso® (T790M)	Global		i i	•	*	
(AZDO	JU 7 <del>4</del> j		C	8. Non-small cell lung cancer	Ph.II enrolling	2nd	EGFR TKI refractory	Iressa® (EGFR)	China			<u> </u>	*	
			نة	9. Non-small cell lung cancer	Ph.II enrolling	1st	c-Met+/Ex.14skip		China		i)		*	
			A	10. Pulmonary sarcomatoid ca.	Ph.II enrolling	1st	c-Met+/Ex.14skip		China		•		*	
			2	11. Gastric cancer	Ph.Ib enrolling	3rd/All	c-Met+		SK/PRC		!		*	
				12. Gastric cancer	Ph.Ib enrolling	2nd	c-Met+	docetaxel (chemo)	SK		i	<b>)</b>	*	
				13. Gastric cancer	Ph.Ib enrolling	2nd	c-Met O/E	docetaxel (chemo)	SK				*	
_				14. Colorectal cancer	Ph.III met all endpoints; NDA mid 2017	3rd	All		China		1	I		
			- 0	15. Non-small cell lung cancer	Ph.III enrolling	3rd	All		China			n/a i	-	*
Frugui	intinib	VEGFR	Lilly (in China	16. Non-small cell lung cancer	Ph.Ib enrolling (dose finding)	1st		Iressa® (EGFR)	China		•			*
		1/2/3		17. Caucasian bridging	Ph.I dose escalation start 2017	-	All comers	,	US		i i			
			only)	18. Gastric cancer	Ph.III (w/ interim analysis) start 2017	2nd	All	paclitaxel (chemo)	China			ı		*
_								. , ,				_;		
				19. Pancreatic NET	Ph.III enrolling	1st			China		į	1		*
		VEGFR/		20. Non-pancreatic NET	Ph.III enrolling	1st			China					*
Sulfa	tinib	CSF1R/		21. Caucasian bridging	Ph.I dose escalation enrolling	-	All comers		US		- 7			
		FGFR1		22. Medullary thyroid ca.	Ph.II enrolling	2nd	Radiotherapy ref.		China					*
				23. Differentiated thyroid ca.	Ph.II enrolling	2nd	Radiotherapy ref.		China		<u> </u>	_ ; •		*
				24. Biliary tract cancer	Ph.II enrolling	2nd	Gemcitabine ref.		China					*
				25. Non-small cell lung cancer	Ph.III start 2017	1st	EGFRm+ brain mets		China					*
Epiti	inib	EGFRm+		26. Glioblastoma	Ph.II start 2017	-	ear air. Sidiii ilicis		China					*
				201 dilopidatorila	That state 2017				Cillia					

### 4 pivotal Phase III studies active & 4 more to start in 2017

Oncology Immunology

# Next wave of innovation now in proof-of-concept



4 novel 2<sup>nd</sup> wave drug candidates in Phase Ib/II studies or about to start

Program	Target	Partner	Study number/Indication	Latest Status	Line	Target patient	Combo therapy	Site Pre	eclin. Ph.I	Proof-of-conce	pt Pivotal/P	h.III
Theliatinib	EGFR WT		27. Solid tumors	Ph.I dose escalation enrolling (continuing)	-	All comers		China	•			*
Hieliauliib	EGFK WI		28. Esophageal cancer	Ph.Ib expansion enrolling	1st	EGFR WT		China				*
			29. Rheumatoid arthritis	Ph. I complete; preparing for Ph.II in 2017	-	Methotrexate ref.		Aus	•			*
HMPL-523	Syk		30. Immunology	Ph.I dose escalation start 2017	-	Healthy volunteers		China				*
HIMPL-323	Jyk		31. Hematological cancers	Ph.I enrolling; target complete Ph.I 2017	2nd/3rd	All comers		Aus	•		*	
			32. Lymphoma	Ph.I dose escalation enrolling	-	All comers		China			*	
HMPL-689	ΡΙ3Κδ		33. Hematological cancers	Ph.I dose escalation (PK analysis)	-	Healthy volunteers		Aus	•			*
HI-IPL-007	PIDIO		34. Lymphoma	Ph.I dose escalation start 2017	2nd/3rd	All comers		China				*
HMPL-453	FGFR		35. Solid tumors	Ph.I dose escalation	-	All comers		Aus			*	
HMPL-433	1/2/3		36. Solid tumors	Ph.I dose escalation start 2017	-	All comers		China			*	
HM004-659	NF- <sub>K</sub> B	Nestle	Ulcerative colitis (Induction)	HMPL-004 reformulation; Re-submit IND 2017	2nd	5ASA refractory		China				*
HM004-037	$(TNF-\alpha)$	Science	Ulcerative colitis (Maintenance)	Await positive Ph.II in Ulcerative Colitis (Induction)	2nd	5ASA refractory		China				*
		Nestlē										
NSP DC2	TBD	Health	Immunology	Preclinical complete end 2017				China				*
		Science										
Multiple	TBD		Oncology	Four small molecule/antibody programs in preclin.				TBD			*	

Oncology Immunology

~2,900 patients/subjects treated in studies to date on our drug candidates, with about 711 dosed in 2016 (2015: 705).

Notes: \* = when an NDA submission is possible based on the receipt of favorable clinical data; Proof-of-concept = Phase lb/II study (the dashed lines delineate the start and end of Phase Ib); combo = in combination with; brain mets = brain metastasis; VEGFR = vascular endothelial growth factor receptor; TKI = tyrosine kinase inhibitor; EGFR = epidermal growth factor receptor; NET = neuroendocrine tumors; ref = refractory, which means resistant to prior treatment; T790M= EGFR resistance mutation; EGFRm+ = epidermal growth factor receptor wild-type; 5ASA = 5-aminosalicyclic acids; chemo = chemotherapy; c-Met+ = c-Met gene amplification; c-Met 0/E = c-Met over-expression; MS = Multiple Sclerosis; RA = Rheumatoid Arthritis; Aus = Australia; SK = South Korea; PRC = People's Republic of China; UK = United Kingdom; US = United States; EU = Europe; Global = >1 country; MTC = Medullary Thyroid Cancer; DTC = Differentiated Thyroid Cancer.

# 8 shots at pivotal success



First positive pivotal Ph.III readout – fruquintinib in colorectal cancer

					Breakthrough Therapy ("BTT") potential	Est. Pivotal Read-out (if not BTT)
SAVO	Papillary renal cell carcinoma (c-Met-driven)	Pivotal Phase III	U.S., EU5, Japan	Initiating In H1 2017	Depends on est. c-Met as -ve prognostic 2017	H1 2019
SAVU	NSCLC -2L Tagrisso combo (T790M+/- & c-Met+)	Pivotal Phase II/III	U.S., EU5, Japan	Decision based on Ph.IIb data (2017)	Depends on strength of Ph.IIb data set (H1 2017)	H2 2019
	3L (or above) Colorectal cancer ("CRC")	Pivotal Phase III	China	Complete <b>Met All Endpoints</b>		March 3 <sup>rd</sup> 2017
FRUQ	3L Non-small cell lung cancer ("NSCLC")	Pivotal Phase III	China	Enrolling		H1 2018
	2L Gastric cancer combo w/ Taxol	Pivotal Phase III	China	Initiating in 2017		H2 2019
SULF	Pancreatic neuroendocrine tumors	Pivotal Phase III	China	Enrolling		H2 2018
JOLI	Non-pancreatic neuroendocrine tumors	Pivotal Phase III	China	Enrolling		H2 2018
EPIT	1L EGFR-mutant NSCLC with brain metastasis	Pivotal Phase III	China	Initiating in 2017		H1 2019

# Major market potential



### CRC peak net income of ~\$20-35m in China is only the start for fruq.

		Pot. launch Year / Territory	Incidence (New pts./yr.) <sup>[1]</sup>	Approx. WAC [2] of various reference TKIs (US\$/month)	Median PFS (months) <sup>[3]</sup>	Potential Peak (US\$) Sales Net income
CAVO	Papillary renal cell carcinoma (c-Met-driven)	<b>2020</b> Global	~25,000	\$11,600 (Sutent®) \$10,500 (Afinitor®)	6.2 Ph.II (actual)	
SAVO	NSCLC -2L Tagrisso combo (T790M+/- & c-Met+)	<b>2019</b> Global	~35,000 - 40,000	\$15,100 (Tagrisso®)	TBD	
	3L (or above) Colorectal cancer ("CRC")	2018 Chi <u>na</u>	~50,000 - 60,000	\$14,000 (Regorafenib - global) \$2,900 (Apatinib - China off label)	3.7 Ph.II (actual)	<b>~\$110-160m</b>
FRUQ	3L Non-small cell lung cancer ("NSCLC")	<b>2019</b> China	~60,000 - 70,000	No approved TKIs \$2,900 (Apatinib - China off label)	3.8 Ph.II (actual)	
	2L Gastric cancer combo w/ Taxol	<b>2020</b> China	~250,000 - 300,000	\$2,900 (Apatinib)	3.7 Ph.II (actual)	
SULF	Pancreatic neuroendocrine tumors	<b>2019</b> China	~5,000 - 6,000	\$11,000 (Sutent®/Afinitor® – global) \$5,500 (Somatuline® – global)	19.4 Ph.II (actual)	
JULF	Non-pancreatic neuroendocrine tumors	<b>2019</b> China	~50,000 - 60,000	\$11,000 (Sutent®/Afinitor® - global) \$5,500 (Somatuline® - global)	13.4 Ph.II (actual)	
EPIT	1L EGFR-mutant NSCLC with brain metastasis	<b>2020</b> China	~30,000 - 40,000	\$15,100 (Tagrisso®) - <i>Brain pen. <sup>[5]</sup></i> \$1,100 (Iressa®) - <i>min. brain pen.</i> \$850 (Conmana®) - <i>min. brain pen</i>	TBD	

# Apatinib/icotinib - Local company TKIs in China [1]



### Major un-met medical need in China - fruquintinib's opportunity

#### Manufacturer Listing Location/Ticker Market Capitalisation (\$US -- Feb 22, 2017) Founded Company 2015 Revenue (US\$ million / 2013-15 CAGR) 2015 R&D Spending (US\$ million / % of Revenues) 2015 Net Profit (US\$ million / 2013-15 CAGR) Commercial Team (# Medical Reps @ end 2015)

#### Molecular Target / Innovation source Formulation Therapy Total Daily Dose (regime)

	Monthly Cost (28 day cycle) Current (US\$)
Patient	Reimbursement (Note: Likely only for est. 40-50% of people enrolled in Medical Insurance Scheme for Urban Employees
costs	Population in mkts. w/ reimbursement (million / % China Po
COSCS	Patient Assistance Program ("PAP") Partner

PAP Starting Date

potential

Monthly Cost (28 day cycle) -- at Launch (US\$)

	PAP Details
Anrikot	Approved Indication (Appr. Indic.)
4arket	Median Progression Free Survival (months / vs. comparator
otontial	Incidence (Overall indication) (Est. New patients/year)

Diagnosed (Overall indication) (Est. New patients/year)

Addressable Patients (Appr. indication) (Est. New ptnts /uear)

	Addiessable Fatients (Appl. Indication) (Est. New pines, year)
Sales	China FDA Approval (competitive approvals?)
כאופכ	China NDA Review Time (months)
History	Launch Date
ווואנטוץ	Year 1 (Revenues US\$ million/ Est. Penetration in Appr. Indic.)
since	Year 2 (Revenues US\$ million/ Est. Penetration in Appr. Indic.)
JIIICC	Year 3 (Revenues US\$ million/ Est. Penetration in Appr. Indic.)
launch	Year 4 (Revenues US\$ million/ Est. Penetration in Appr. Indic.)
iddiicii	Voor E (Dovonuos LIC\$ million/ Est Donotration in Appr Indic)

ATAN®						
Apatinib						
Jiangsu Hengrui Medic	ine					
Shanghai:	600276.SS					
\$15.9 billion						
1970						
1,479	23%					
142 (10% of Rev	1.)					
345	32%					
5,491						

VEGFR2 (licensed in from U	J.S. Co. <sup>[3]</sup> )			
Oral tablet				
850mg				
(425mg twice dail	y) )			
~2,900				
~2,900				
None				
None	0%			
PhIRDA [2]				
June 2015				
Free drug after 3 paid cycles				
(i.e. 3 months)				

Gastric cancer ("GC"), third-line				
2.6	1.8 (pbo)			
~660,000 (GC)				
~395,000				
~40,000-50,00	00			

<b>6</b>	a.
Conmana	9
Icotinib	
Betta Pharma	3
Shenzhen:	300558.SZ
\$3.8 billion	
2003	
145	38%
19 (13% of Re	V.)
55	39%
296	

375mg (125mg
three times a day)
~1,900
~850
5 Provinces (Zhejiang; Hunan; Guangxi;
Gansu; Inner Mongolia); 2 Cities (Qingdao;
Shenzhen)
240 17%
Phirda
July 2011
Free drug after 6 paid cycles

EGFR (licensed in from U.S.)

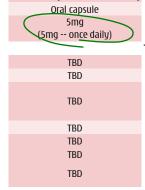
Oral tablet

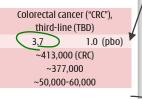
Non-small cell lung cancer ("NSCLC"),							
> second-line / first-line EGFRm positive							
4.6 / 9.5 3.4 / 9.5 (Iressa®)							
~625,000 (NSCLC)							
~600,000 / ~220,000							
~150.000-170.000 / ~220.000							

(i.e. 6 months)

June 2011 (	multiple appr. EGF	R TKIs)	TBI	)
	(10)			
	August 2011			
2011	9	1%		
2012	48	2%		
2013	78	3%		
2014	116	5%		
2015	( 145	( (0)		

Fruquintin	ib	
Chi-Med <sup>[4]</sup>		
LSE/Nasdaq:	HCM	
\$1.6 billior	1	
2000		
178	na	
56 (31% of	Rev.)	
8	na	
~2,200	)	K





(only appr. 3L CRC drug) **TBD** 

2018 (Estimated)

**TBD TBD** 

TBD

**TBD** 

Αp	atinib	per	etr	ation
	in <5	yeaı	S	
V	<b>7</b> \$10	U IIII	IIIUI	i saies

✓ Apatinib used in 3<sup>rd</sup> line NSCLC, CRC, etc.

Icotinib penetr. low -

#### Chi-Med investing all resources into R&D

#### **Chi-Med Commercial** Platform is important

#### Fruquintinib highly potent vs. other TKIs

- √ 5mq/day vs. 850mq & 375mq
- ✓ Once daily optimal vs. twice/thrice daily

#### Frug. robust clinical efficacy vs. other TKIs

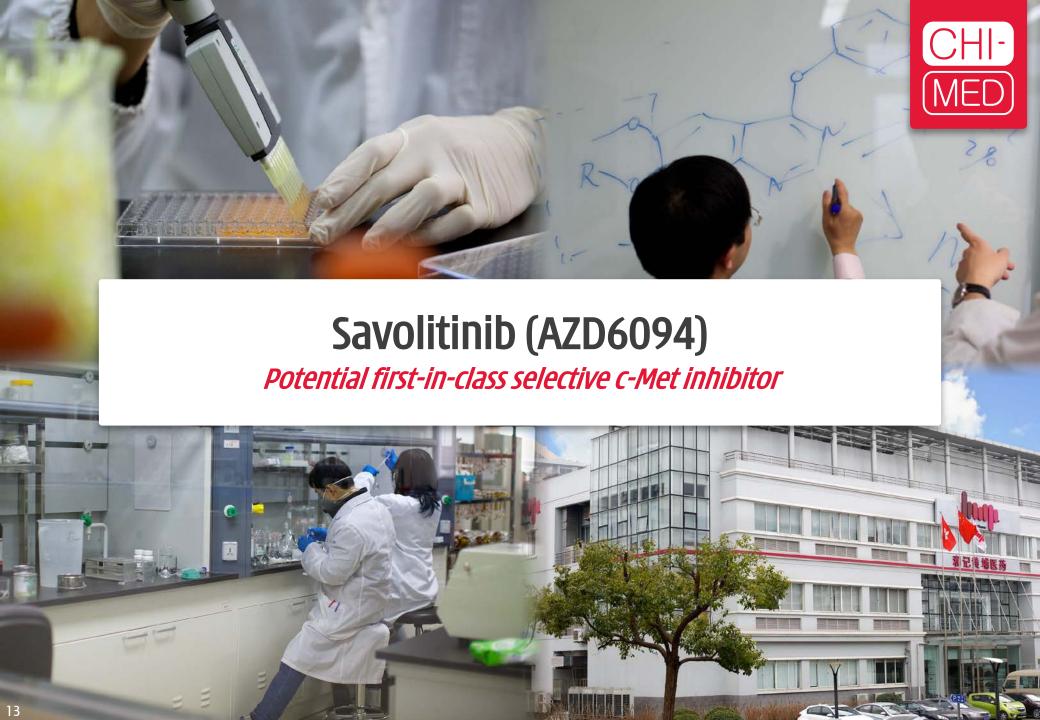
#### China major TKI market potential due to unmet medical need

√ \\$100 million cales

# high - off-label use

b/c Iressa®/Tarceva®

[1] China Cancer Registry; Betta Pharma IPO prospectus; China 2010/2015 census; Goldman Sachs; [2] PhIRDA = China Pharmaceutical Innovation & Research Development Association; [3] Advenchen Labs, California; [4] HMP = Hutchison MediPharma



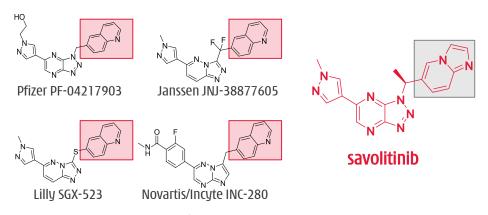
# Savolitinib (AZD6094)





### Potential global first-in-class selective c-Met inhibitor

- 1. In strong position to become first selective c-MET inhibitor approved globally.
  - ✓ Clear clinical efficacy observed in non-small cell lung ("NSCLC"), kidney, gastric and colorectal cancers.
  - ✓ Partnered with AstraZeneca key comp. advantages in NSCLC (Tagrisso® combo.) & molecular selection.
- 3. Savolitinib design eliminates renal toxicity first generation of selective c-MET inhibitors encountered >460 patients treated to-date with no renal toxicity.



2-quinolinone metabolite in humans in 1st gen c-Met compounds has dramatically reduced solubility and appeared to crystallize in the kidney resulting in obstructive toxicity.

#### 2. c-Met is aberrant in many tumor settings.<sup>[3]</sup>

	(C-MET			New Case	es (2015)
Indication	Amplifi- cation	Mutation	Over- Expression	Global	China
Gastric	10%	1%	41%	1,034,000	679,000
Lung (Non-small cell)	8-10%[1]	8%	67%	1,690,000	575,000
Head & Neck		11%	46%	740,000	135,000
Colorectal	10%		65%	1,477,000	376,000
Renal cell Carcinoma (Papillary)	40-70%	100%[2]		50,000	7,000
Renal cell Carcinoma (Clear cell)			79%	270,000	60,000
Esophagus	8%		92%	496,000	251,000

#### 4. AstraZeneca collaboration & 2016 amendment.

- 2011 global licensing agreement: \$20m up front; \$120m in development/approvals milestones (\$20m paid by Jun'16); significant commercial milestones; ex-China tiered royalty 9-13%, AZ pay 100% development cost; China 30% royalty, AZ pay 75% development cost (Chi-Med 25%).
- 2016 amendment: Chi-Med pay \$50m towards joint development costs, over 3 years; in return for ex-China royalty +5% points (to 14% to 18%).

# c-MET +ve PRCC - unmet medical need



#### 1. No treatment choices for non-ccRCC patients.

### Approved therapies in RCC [3]

Good efficacy in ccRCC; Multiple treatment options

FIRST LINE – clear-cell RCC [4]	ORR	mPFS	mos
Placebo (avg. multiple studies)	~2%	~3.5	~15.0
Interferon-α	6%	5.0	21.8
<b>Nexavar®</b> (VEGFR, multi-kinase SM) (avg. multiple studies)	~12%	~6.0	~21.0
<b>Sutent®</b> (VEGFR, multi-kinase SM) (avg. multiple studies)	~28%	~10.5	~27.0
<b>Votrient®</b> (VEGFR, multi-kinase SM)	31%	10.5	28.4
SECOND LINE – clear-cell RCC			
Placebo (avg. multiple studies)	~0%	~2.0	~14.0
<b>Afinitor®</b> (mtor). <i>(Meteor)</i>	3%	3.9	16.5
<b>Afinitor®</b> (mTOR). <i>(CheckMate025)</i>	5%	4.4	19.6
<b>Inlyta®</b> (VEGFR, multi-kinase SM)	23%	8.3	20.1
<b>Opdivo®</b> (PD-1 mAb). <i>(CheckMate025)</i>	25%	4.6	25.0
Cabometyx® (VEGFR/MET, multi-kinase SM). (METEOR)	17%	7.4	21.4

### Nothing approved in non-ccRCC

NCCN guideline - "Patients should enter clinical trials"

FIRST LINE - non clear-cell RCC	Oitit	mPFS	
<b>Sutent®</b> (VEGFR, multi-kinase SM) <sup>[4]</sup>	9%	6.1	16.2
<b>Afinitor®</b> (mTOR) <sup>[4]</sup>	3%	4.1	14.9
SECOND LINE – non-clear-cell RCC			
<b>Sutent®</b> (VEGFR, multi-kinase SM) <sup>[4]</sup>	10%	1.8 2.8	na
<b>Afinitor®</b> (mTOR) <sup>[4]</sup>	9%	2.8	na

2. RCC est. ~\$3.3 bln. market by 2020 [1]

Clear-cell RCC (~\$2.7b)

~80% of RCC ~ 270k new patients/yr.<sup>[2]</sup>

Non-Clear-cell RCC (~\$0.6b)

~20% of RCC ~ 70k new patients/yr.<sup>[2]</sup> 3. Two crucial questions:

c-MET +ve
Papillary RCC
(~\$0.2-0.3b)

~7% of RCC
~ 25k new patients/yr.[2]

C-MET -ve
Papillary RCC
(~\$0.2-0.3b)

~7% of RCC
~ 25k new patients/yr.[2]

Other non-ccRCC (~\$0.1-0.2b) ~5% of RCC

~5% of RCC ~ 20k new patients/yr.<sup>[2]</sup> Question 1: Does savolitinib provide meaningful benefit to patients w/ c-MET+ve PRCC?

Answer: Phase II data (next page)

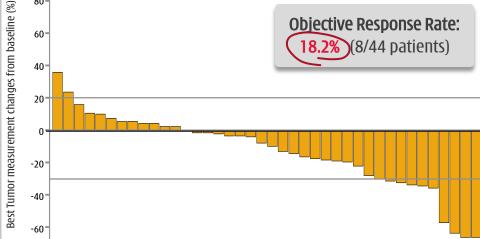
Question 2: Is c-MET +ve status predictive of worse outcome (PFS/OS) in PRCC patients?

Answer: >300 pt.
PRCC Molecular
Epidemiology Study
(late 2017 readout).

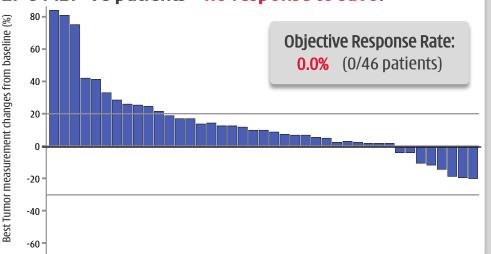
## Savolitinib – PRCC Phase II

### Clear efficacy & durable response in c-MET +ve PRCC patients





### c-MET -ve patients - no response to savo.



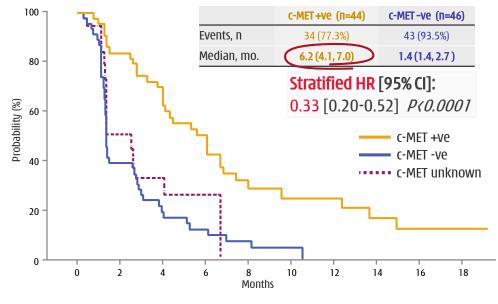
### 3. Disease Control Rate ("DCR") - big advantage in c-MET +ve with DCR 73.2% vs. c-MET -ve 28.2%.^

Tumor responses in the overall treatment population and by MET status

RECIST response, n (%)	c-MET +ve (n=44)	c-MET -ve (n=46)	c-MET unknown (n=19)	Total (n=109)
Partial Response <sup>†</sup>	8 (18.2%)*	0 (0.0%)	0 (0.0%)	8 (7.3%)
Stable Disease	22 (50.0%)	11 (23.9%)	5 (26.3%)	38 (34.9%)
Progressive Disease	11 (25.0%)	28 (60.9%)	9 (47.3%)	48 (44.0%)
Not Evaluable	3 (6.8%)	7 (15.2%)	5 (26.3%)	15 (13.8%)

<sup>\*</sup>P=0.002 versus MET-independent subgroup (Fisher exact test). Responses assessed according to RECIST version 1.1. †Unconfirmed responses excluded. ^ evaluable patients

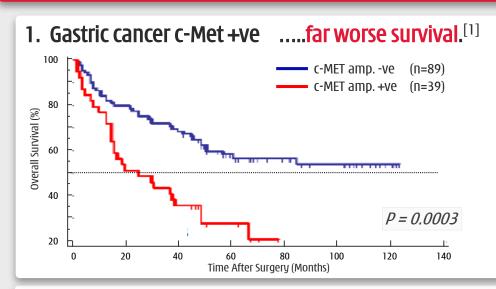
#### 4. Median PFS - big advantage in c-MET +ve patients.

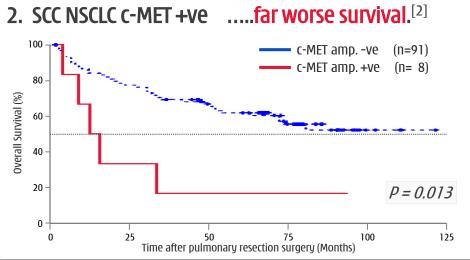


# **c-MET** positive status



### Shown to be a predictor of very poor patient outcome in many cancers





#### 3. PRCC Molecular Epidemiology Study ("MES") Plan:

- → A pooled analysis of historical data Correlation of c-METpositive PRCC with treatment outcomes.
- → 3 collaborations GETUG<sup>[3]</sup> (France); IMDC<sup>[4]</sup> (N. America, EU, Asia, New Zealand); & Asan GU (Korea). Total >300 patient data.
- $\rightarrow$  Timing MES to be conducted Q1-Q3 2017- Results (end 2017.)







#### PRCC Patient Data

(n >300)

- → Tissue samples for c-MET testing
- → Medical records treatment history/outcomes

#### 4. How we will use the MES data set?

- → Possible breakthrough therapy submission with clear evidence that c-MET +ve PRCC has far worse treatment outcome/survival than c-MET -ve.
- → Clarity on PFS/OS treatment outcome of c-MET +ve patients how do c-MET +ve PRCC patients (vs. c-MET -ve) respond to sunitinib and other approved RCC therapies.

[1] c-MET amplification: gene copy number of ≥4. J Shi et al. Frequent Gene Amplification Predicts Poor Prognosis in Gastric Cancer. Int. J. Mol. Sci. 2012, 13, 4714-4726; [2] SCC NSCLC = squamous cell carcinoma non-small cell lung cancer. (~20-30% of NSCLC) -- c-MET gene amplification: >15 copies in >10% of tumor cells with 4-10 copies in a gene cluster. H Go et al. High MET Gene Copy Number Leads to Shorter Survival in Patients with Non-Small cell Lung Cancer. J. Thorac. Oncol. 2010, 5, 303-313.; [3] GETUG = Groupe Français d'Etude des Tumeurs Uro-Genitales; [4] IMDC = International Metastatic Renal Cell Carcinoma Database Consortium.

## Savolitinib – PRCC Phase II



### Safe & very well tolerated – apparent advantage over other RCC TKIs<sup>[7]</sup>

		PRCC PHASE II Savolitinib	COMPARZ F	PHASE III [1] Pazopanib	METEOR P	HASE III <sup>[2]</sup> Everolimus	SINGLE-ARM PHASE III [3] Sunitinib	
		1L/2L (n=109)	1L (n=548)	1L (n=554)	2L (n=331)	2L (n=322)	2L (n=106)	
MSKCC Risk Group	Favorable Intermediate Poor Missing	14% 45% 9% 32%	27% 59% 9% 4%	27% 58% 12% 3%	45% 42% 12% 0%	46% 4T% 13% 0%	58% 42% <sup>[6]</sup> 0%	Better safety data of risk patient popular  ✓ Only 14% "favoral
Number of prior systemic therapies	0 1 ≥2	55% 23% 22%	100% 0% 0%	100% 0% 0%	0% 71% 29%	0% 70% 30%	0% 100% 0%	
Grade ≥3 AEs:	Any AE Any treatment-related AE [4]	19%	77% <sup>[5]</sup>	76%[5]	68%	58%		
All Grade≥3 AEs with ≥5% incidence (AND selected savolitinib AEs for comparison)	Hypertension Fatigue Hand-foot-syndrome Diarrhea	TR AES 0% 2% 0% 0%	TR AES 15% 17% 12% 8%	TR AES 15% 11% 6% 9%	All AEs 15% 9% 8% 11%	All AES 3% 7% <1% 2%	6% 11% 7%	Superior safety pro TKIs - Most ≥3 G3 A ✓ Hypertension: 0%
Hematologic Abnormalities Grade≥3 AEs with ≥5% incidence:	Neutropenia Thrombocytopenia Lymphocytopenia Leukopenia Anemia	0% 0% 0% 0% (1%	20% 24% 14% 6% 7%	5% 4% 5% 1% 2%	0% 0% 0% 0% 5%	0% 0% 0% 0% 16%	16% 6%	✓ Fatigue: 2% vs. 6~ ✓ Diarrhea: 0% vs. ~ ✓ Anemia: <1% vs. 7
Lab Abnormalities Grade≥3 AEs with ≥5% incidence:	Increased ALT Increased AST Hypophosphatemia Hyponatremia Hypokalemia	5% 3% 0% 3% 0%	4% 3% 9% 7% 1%	17% 12% 4% 7% 3%	2% 2% 4% 0% 5%	(1% (1% 2% 0% 2%		≈ ALT/AST Increase: ✓ Other Lab Abnorn
Tolerability	Hyperglycemia  Treatment discontinuation due to any AE:  Dose reduction due to AE:	0% 8% 13%	20%	24%	12%	5% 11% 25%	11%	Highly tolerable vs.  ✓ Discontinued: 8% ✓ Dose reduction: 1

#### despite higher ation:

able" vs. 27-58%.

#### ofile vs. other AES $\approx$ 0-2%:

- % vs. 6~17%.
- ~12%.
- ~10%.
- 7~16%.
- e: 3-5% vs. 0~17%.
- rm: 0% vs. ≤9%.

#### s. other TKIs:

- % vs. 10~24%.
- 13% vs. 44-62%.

<sup>[1]</sup> RJ Motzer et al, *Pazopanib versus Sunitinib in Metastatic Renal-Cell Carcinoma*, N Engl J Med 369;8, Aug 22, 2013; [2] TK Choueiri et al, *Cabozantinib versus everolimus in advanced renal cell carcinoma (METEOR)*, Lancet Oncol.17;7, Jun 5, 2016; [3] RJ Motzer et al, *Sunitinib in Patients with Metastatic Renal Cell Carcinoma*, JAMA 295;21 Jun 7, 2006; [4] As assessed by investigator. [5] Includes Grade 5AEs; [6] includes Intermediate & Poor. TR AEs = Treatment-Related Adverse Events; [7] RCC = Renal Cell Carcinoma, TKIs = Tyrosine Kinase Inhibitors.

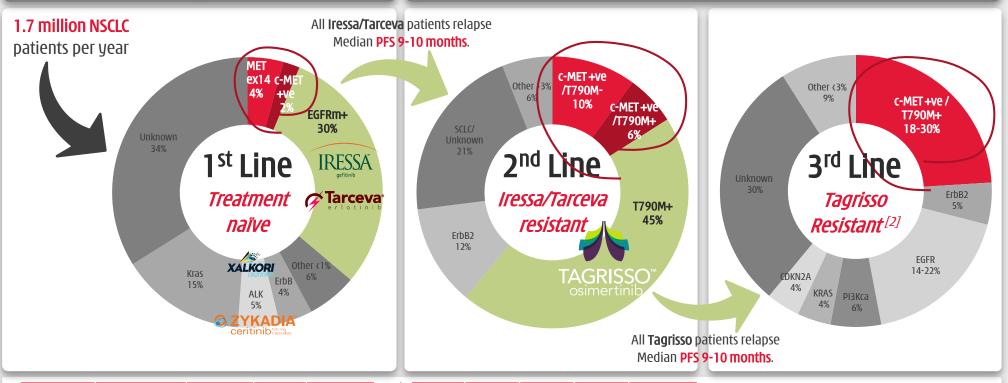
# Savolitinib



Biggest opportunity is c-MET +ve non-small cell lung cancer ("NSCLC")

### **Primary NSCLC**

### **Resistance-driven NSCLC**



	Target	Launch	2016 (\$m)	Est. <sup>[1]</sup> Ptnt. Treat	Launch	Q4 2015 (\$m) <sup>[3]</sup>	H1 2016 (\$m) <sup>[3]</sup>	H2 2016 (\$m) <sup>[3]</sup>	Est. <sup>[3]</sup> Ptnt. Treat
Iressa	EGFRM+	2003	513	~20,000					
Tarceva	EGFRM+	2004	1,137	~50,000					
Tagrisso	EGFRm+/T790M	2018/19?			Dec-15	~20	143	280	~5,000
Xalkori	ALK/ROS1/MET	2011	561	Est. peak					
Zykadia	ALK	2015	91						
Total Sales			2,302	~\$3.0b		~20	143	280	

<sup>[1]</sup> general estimate based on mPFS ~9 mo. average cost/cycle ~\$2,500-3,000; [2] based on aggregate rocelitinib/Tagrisso data published at 2016/2017 ASCO; [3] AstraZeneca 2016 results...

# Savolitinib – 1<sup>st</sup> Line NSCLC



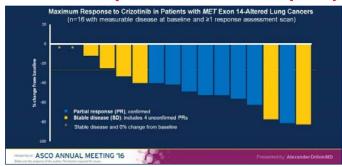
### Xalkori® (crizotinib) proof-of-concept in Exon 14 skip 1L NSCLC

1. Xalkori® is a multi-kinase inhibitor with ALK, ROS1, & MET inhibition – savolitinib is uniquely selective and (10x) more potent against c-Met.

IC <sub>50</sub> (nM)	Savolitinib	Xalkori® (crizotinib)	Savolitinib vs. Xalkori®
EBC1 Viability	2	19	10x
EBC1 pMET	1	39	40x
293T MET (wild type)	7	79	11x
293T MET (Ex14del)	9	140	16X

2. 1<sup>st</sup> line NSCLC - Xalkori® MET Exon14 skipping - 2016 ASCO - strong response (~50% ORR) but > 1/3<sup>rd</sup> of responses not durable (4/12)<sup>[1]</sup>.

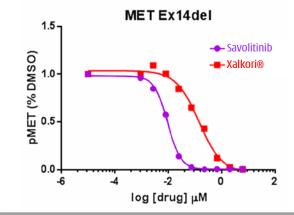




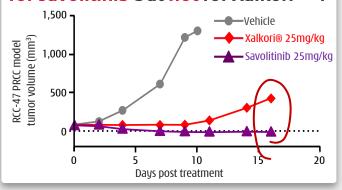
3. Multi-center retrospective analysis of 148 pts. w/ NSCLC MET Exon14 [2]

34 pts. with $\underline{\text{metastatic NSCLC}}\text{w/}\text{MET Exon14}\text{skipping:}$	No exposure to c-MET TKI:	median OS = 8.1 months			
	(c-MET+ve pts. mOS = 5.2 mo.; c-MET-ve pts. mOS = 10.5 mo.; p=0.06)				
27 pts. with $\underline{\text{metastatic NSCLC}}\text{w/MET Exon14}\text{skipping:}$	Exposure to c-MET TKI:	median OS = 24.6 months			
	(22pts. treated with Xalkori® with median PFS = 7.4 mo.)				

4. Savolitinib versus Xalkori® in MET Ex14del mutant cells<sup>[3]</sup> - better target coverage.



5. Durable tumour cell suppression for savolitinib but not for Xalkori<sup>®[4]</sup>.

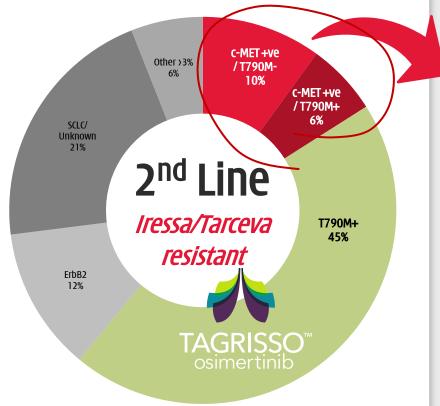


# Savolitinib - 2<sup>nd</sup> Line NSCLC Phase Ib/II



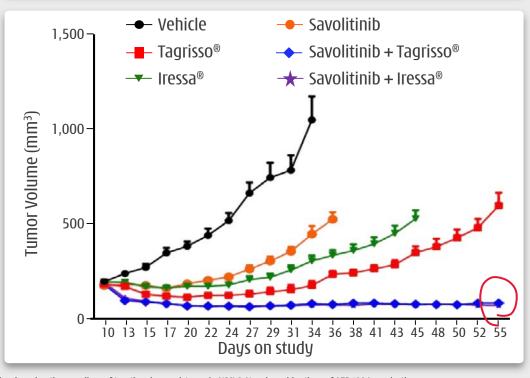
### Very strong early signal emerging – Clear competitive edge for savolitinib

1. 2<sup>nd</sup> Line NSCLC is the fastest & most attractive indication for savolitinib to go after. Also important unmet medical need and potential Breakthrough Therapy area.



#### 2. Potential in EGFR TKI resistant NSCLC:

- Must shut down both EGFR & c-Met signaling pathways;
- ✓ Prolonged tumor growth suppression by combining savolitinib with Tagrisso® (osimetinib - EGFR/T790M) or Iressa® (gefitinib/EGFR) in T790M-, c-MET +ve patients.

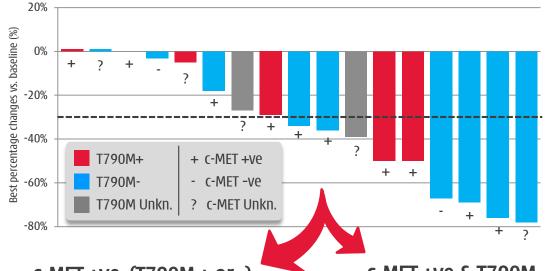


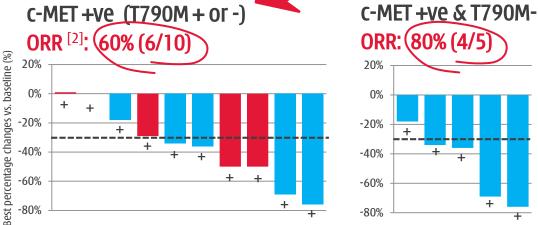
# Savolitinib – 2<sup>nd</sup> Line NSCLC



### Clear anti-tumor effect in NSCLC patients - Phase IIb to complete 2017

1. TATTON efficacy...Phase I/IIa exploring combo treatment of 600/800mg savolitinib & 80mg Tagrisso® daily.[1]





- 2. Particularly encouraging efficacy in 32 yr. old NSCLC patient w/ c-Met +ve & T790M-.
- ✓ Rapidly progressing bone & lung mets. Major solid tumor.
- ✓ Primary progression on prior EGFR TKI (i.e. Tarceva resist.).
- Brief response to platinum doublet.



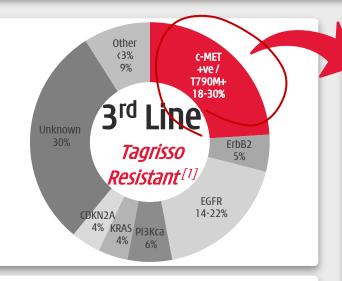
3. TATTON safety – savolitinib & Tagrisso® combo treatment at full doses. No major toxicity.

Number of events, n	600 (n =		800 (n =	
Adverse Event occurring in over three instances at any dose	Any Gr.	Gr.≥ 3	Any Gr.	Gr.≥3
Vomiting	7	0	3	0
Nausea	3	0	6	1
Rash	4	0	3	0
Pyrexia	3	0	3	0
White blood cell count decrease	4	0	1	1
Decreased appetite	1	0	3	0

# Savolitinib – 3<sup>rd</sup> Line NSCLC – Tagrisso® resistant



### T790M+ & c-Met+ unmet medical need starting to emerge



3/3 patients with T790M+/c-MET+ responded to savo/Tagrisso® combo.









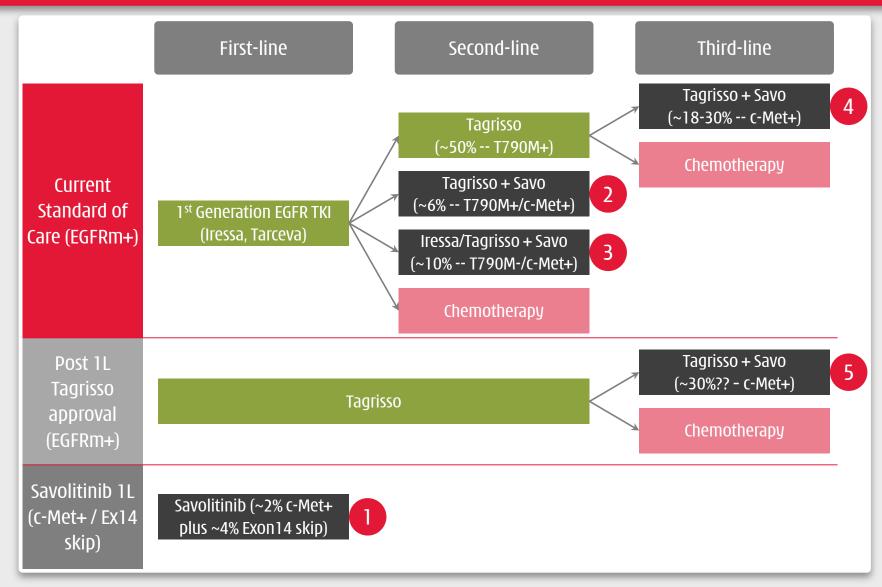
Pt	EGFR mutation	# Prior Therapies	Prior 3 <sup>rd</sup> gen TKI	TISSUE (NGS, FISH)	PLASMA CTDNA (NGS)
1	L858R	1		<i>MET</i> amp, T790 WT	<i>MET</i> amp, T790M ND
2	Del19	1		-	T790M ND
3	Del19	2	Υ	-	T790M ND
4	L858R (de novo T790M)	2	Υ	<i>MET</i> amp, <i>EGFR</i> amp T790M (germline)	-
5	L858R	3	Υ	T790wt, <i>EGFR</i> amp	T790M ND
6	L858R	4	Υ	T790 WT	T790M ND
7	Del19	3	Υ	-	T790M ND
8*	Del19	3		T790M/C797S	T790M/C797S
9	L858R	4	Υ	T790 WT	-
10	Del19	3	Υ	-	<i>PIK3CA</i> E545K, <i>PIK3CA</i> amp, T790M ND
11	Del19	2	Υ	<i>MET</i> amp, <i>EGFR</i> amp, T790 WT	T790M ND
12	Del19	2	Υ	-	T790M/C797S
13	Del19	9		T790 WT	-
7	Del19	2	Υ	T790 WT	T790M ND
1	Del19	1		T790 WT	<i>FGFR1</i> D60N, <i>FGFR1</i> amp, T790M ND
16	L858R	2		<i>MET</i> amp, T790 WT	<i>MET, EGFR</i> amp, T790M ND
17	L858R	3	Υ	T790 WT	T790M ND
18	Del19 (de novo T790M)	3		SCLC, T790 WT	T790M ND, <i>EGFR</i> amp
19	Del19	3	Υ	T790 WT	T790M/C797S, <i>MET</i> amp, <i>EGFR</i> amp
20	L858R	2		<i>MET</i> amp, <i>EGFR</i> amp, T790 WT	-
21	L858R	3		-	T790M/C797S, <i>EGFR</i> amp
22*	L858R	1		MET amp, T790 WT	-
23	Del19	4	Υ	-	T790M/C797S

(-) testing not performed; EGFR - Epidermal Growth Factor Receptor; TKI- Tyrosine Kinase Inhibitor; amp- amplification; WT- wild type; ND- not detected

### Savolitinib - NSCLC



### Five clear opportunities for savolitinib in the NSCLC treatment algorithm



# Savolitinib - Gastric cancer



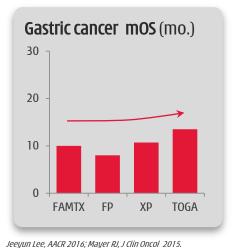
### A major problem in east Asian countries – Japan, South Korea and China

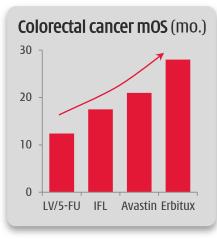
1. Gastric (stomach) cancer is the 5<sup>th</sup> most common cancer globally - 723,000 deaths/year.

	Est. Age Standardised Rates (cases/100,000)	New cases ('000)	Deaths ('000)	5-year Prevalence ('000)
World	17.0	952	723	1,538
South Korea	41.8	22	17	32
Japan	29.9	38	29	56
China	22.7	405	325	594
EU-28	9.0	82	58	119
USA	6.8	21	12	32

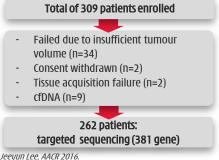
Jeeyun Lee, AACCR 2016; IARC, WHO 2012; Jung KW, Cancer Research Treatment 2013; World Cancer Research Fund International.

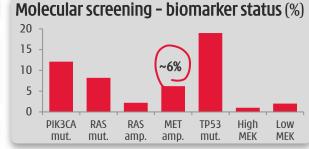
# 2. Little progress in gastric cancer in improving overall survival ("OS") in first-line palliative setting.

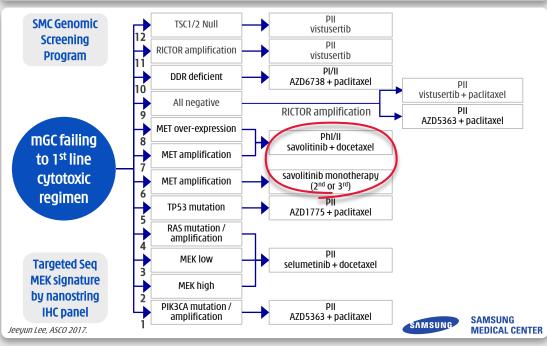




#### 3. VIKTORY – umbrella trial in gastric cancer *(South Korea).*



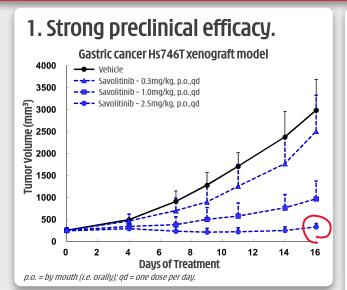


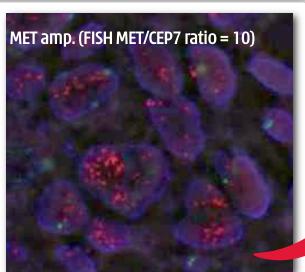


# Savolitinib - Gastric cancer



### VIKTORY trial - very promising early clinical results in c-MET +ve patients



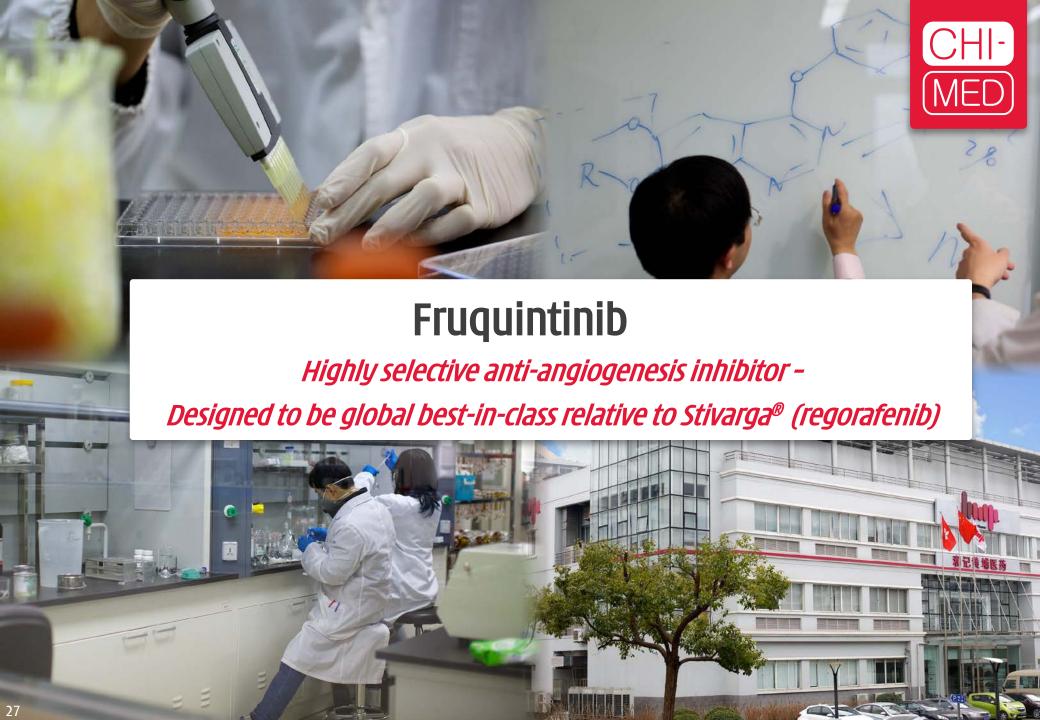


Jeeyun Lee, AACR 2016

#### 2. VIKTORY trial - 34-year old male; surgery ruled-out; failed 4-cycles XELOX.







# Fruquintinib – 24hr full target coverage

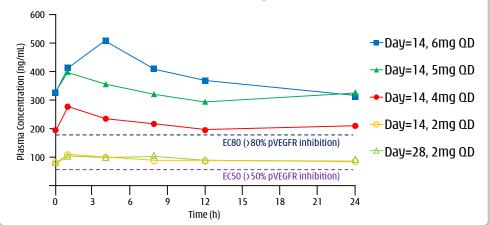


### The most selective VEGFR inhibitor in clinical trials globally<sup>[1]</sup>

# 1. Substantial progress made in 2016 - fruquintinib China NDA submission mid-2017.

- ✓ Validation of R&D approach designed to only inhibit VEGFR1,2,3, facilitating **full target coverage & combinations**.
- ✓ Pivotal Phase III in 3L CRC met all endpoints NDA submit mid-2017.
- ✓ **Pivotal Ph. III** trial in **3L NSCLC well underway** since Q4 2015 initiation.
- ✓ Ph.Ib Taxol® combo in 2L gastric cancer dose finding complete. Phase III pivotal study starting 2017.
- ✓ Ph.II Iressa® combo trial in 1L EGFRm+ NSCLC started early 2017.
- ✓ China GMP **production facility operational** to support launch.

# 2. Only inhibits VEGFR - limits off-target toxicity & allows for full & sustained target inhibition.



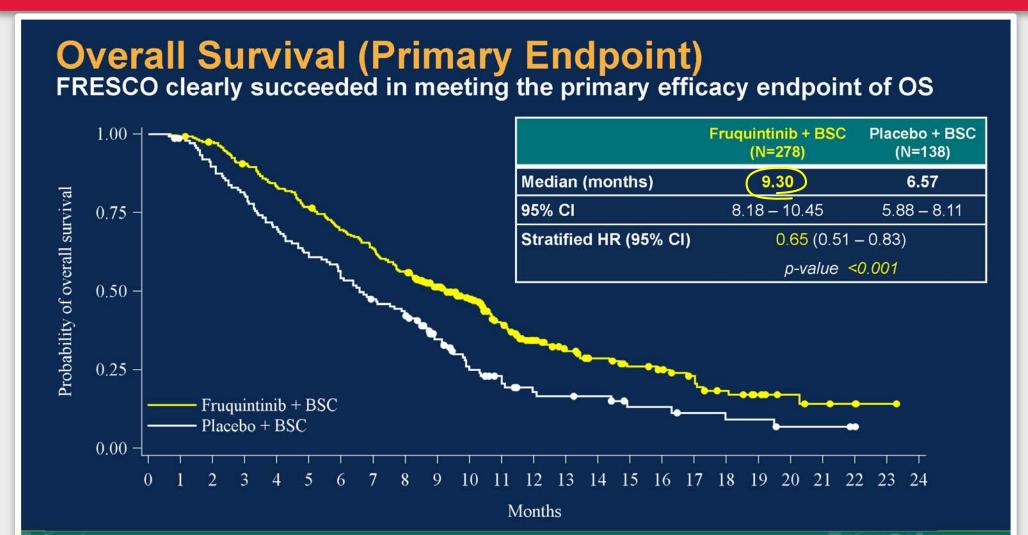
#### 3. Selectivity and potency superior to competitor drugs.

	Sutent® (sunitinib)	Nexavar® (sorafenib)	Stivarga® (regorafenib)	Tivozanib	Fruquintinib
Kinase profile	VEGFR1,2,3, PDGFRβ, FLT3, CSF-1R, c-Kit, Ret	RAF, VEGFR2, PDGFRB, Flt3, c-Kit, FGFR1	VEGFR1,2,3, Raf, Ret, PDGFR, c- <u>Kit</u>	VEGFR1,2,3, BRK, PDGFRα, PDGFRβ, c-Kit, Tie2, EphB2	VEGFR1,2,3
AUC at ED50/ED60 in mouse (ng/mL*hr)	2,058	25,473	na	1,640	898
MTD in human (mg/day)	50, qd	400, bid	160, qd	1.5, qd	4, qd; 6, 3wk/1wk
AUC, 0~24h at Steady state MTD (ng/mL*hr)	592	47,780 x2 (D28)	58,270 (D21)	1,180 (D28)	5,000~6,000 (D28)
Efficacy in Phase I	22 patients PR: 4 (18%), DCR: 27%	45 patients <sup>[2]</sup> PR: 1 (2%), DCR: 58%	53 patients PR: 3 (6%), DCR: 66%	37 evaluable patients PR: 1 (3%) DCR: 51%	34 evaluable patients PR: 13 (38%), DCR: 82%

# Fruquintinib - Third-line colorectal cancer

Lilley CHI
MED

Best-in-class efficacy/safety - Phase III FRESCO data at ASCO 2017<sup>[1]</sup>



PRESENTED AT: ASCO ANNUAL MEETING '17

#ASCO1

Presented by: Jin Li, MD PhD

June 5, 2017



# Fruquintinib - FRESCO efficacy in 3L CRC

	Fruquii	Fruquintinib		<sup>T</sup> enib	Regora	afenib	Regorafenib		
Patients (n)  Complete Response, n (%)  Partial Response, n (%)  Stable Disease, n (%) <b>Disease Control Rate, n (%)</b>	FRES	6CO	CONC	UR	CON	CUR	CORRECT		
	Mainlan	d China	Chinese Patien China, Hong Kor	The second secon	Mainland China Taiwan, Vietnar		CORRECT  Global  Regorafenib Pla 505 2  0.0% 0 1.0% 0 42.8% 14 41.0% +26.1 14  1.9 +0.2 1 (0.000001) 0.49  6.4 +1.4 5 0.0052	al	
Treatment arms	Fruquintinib	Placebo	Regorafenib	Placebo	Regorafenib	Placebo	Regorafenib	Placebo	
Patients (n)	278	138	112	60	136	68	505	255	
Complete Response, n (%)	0.4%	0.0%	0.0%	0.0%	0.0%	0.0%	0.0%	0.0%	
Partial Response, n (%)	4.3%	0.0%	3.6%	0.0%	4.4%	0.0%	1.0%	0.4%	
Stable Disease, n (%)	57.6%	12.3%	40.2%	6.7%	45.6%	7.4%	42.8%	14.5%	
Disease Control Rate, n (%)	62.2% (+49	9 12.3%	45.5% (+38	.8 6.7%	51.5% (+44	7.4%	41.0% (+26.	14.9%	
Median Progression Free Survival (mPFS) (mo.)	3.7 (+1.9	1.8	2.0 (+0,	3 1.7	3.2 (+1)	.5 1.7	1.9 (+0.2	1.7	
mPFS p-value	⟨0.0⟩	01	not publ	ished	(0.0)	001	⟨0.000	001	
mPFS Hazard Ratio	0.2	6	0.32	2	0.3	31	0.49	)	
Median Overall Survival (mOS) (mo.)	9.3 +2.	6.6	8.4 (+2.	2 6.2	8.8 (+2.	.5 6.3	6.4 +1.4	5.0	
mOS p-value	(0.0)	01	not publ	ished	0.00	002	0.00	52	
mOS Hazard Ratio	0.6	55	0.56	5	0.5	55	0.7	7	

- Good fruquintinib efficacy over regorafenib in Chinese patients specifically in terms of Disease Control Rate; median Progression Free Survival and median Overall Survival.
- FRESCO is a fully-powered Phase III registration study (n=416) whereas CONCUR was an under-powered Asia region study (n=204, including only 129 mainland Chinese patients<sup>[2]</sup>).
- CONCUR results should be regarded as directional only China approval resulted from CORRECT study (n=760).

# Fruquintinib - FRESCO safety in 3L CRC



## Fruquintinib high VEGFR selectivity - no off-target AEs & more tolerable

atients (n)  G3 AE (Safety population)  AES:  All (Safety population)  AES:  All (Safety population)  AES:  APPOPULATION  AES:  ALT increased, ≥G3  BOOD bilirubin increased, ≥G3  BOOD bilirubin increased, ≥G3	FRE	intinib :SCO nd China	Regorafenib CONCUR Chinese Patients (Mainland China, Hong Kong, Taiwan) <sup>[1]</sup>			
Treatment arms	Fruquintinib	Placebo	Regorafenib	Placebo		
Patients (n)	278	138	112	60		
≥G3 AE (Safety population)	61.1%	19.7%	69.6%	46.7%		
SAE (Safety population)	15.5%	5.8%	31.3%	26.7%		
VEGFR on-target related AEs:						
Hypertension ≥G3	21.2%	2.2%	12.5%	8.3%		
Hand Foot Syndrome (Palmar-plantar), ≥G3	10.8%	0.0%	17.0%	0.0%		
Off-target (i.e. non-VEGFR) related AEs:						
Hypophosphatemia, ≥G3	0.0%	0.0%	8.0%	0.0%		
Hypokalemia, ≥G3	0.7%	0.7%	6.3%	0.0%		
Rash/desquamation, ≥G3	0.0%	0.0%	4.4%	0.0%		
Lipase increase, ≥G3	0.0%	0.0%	6.3%	1.7%		
Hepatic function (Liver function) AEs:						
ALT increased, ≥G3	0.7%	1.5%	7.1%	3.3%		
AST increased, ≥G3	0.4%	0.7%	8.9%	0.0%		
Blood bilirubin increased, ≥G3	1.4%	1.5%	8.9%	8.3%		
NOTE: Baseline Characteristics Liver metastasis	66.5%	73.9%	na	na		
Tolerability:						
AE Leading to dose interuption	35.3%	10.2%	68.8%	25.0%		
AE Leading to dose reduction	47.1%	13.1%	23.2%	0.0%		
AE Leading to treatment discontinuation	15.1%	5.8%	14.3%	6.7%		

#### Fruquintinib far more selective than regorafenib

BIOCHEMICAL ACTIVITY	Fruquintinib IC <sub>so</sub> (nmol/L)	Regorafenib IC <sub>so</sub> (nmol/L)
On-Target Kinases:	-111 (	-111
VEGFR1	33	13
VEGFR2	35	4.2
VEGFR3	0.5	46
Off-Target Kinases:		
Ret	128	1.5
FGFR1	181	202
c-kit	458	$\overline{7}$
PDGFRβ	>10,000	22
RAF-1	>10,000	2.5
B-RAF	>10,000	28
B-RAF <sup>V600E</sup>	>10,000	$\bigcirc 19$

#### Regorafenib liver toxicity Black-box warning:

- → Increased liver function test monitoring (weekly if elevated) & remedial dose interruption.
- → 3L CRC China 65-75% liver metastasis weaker pts.

STIVARGA (regorafenib) tablets, oral Initial U.S. Approval: 2012

#### WARNING: HEPATOTOXICITY

See full prescribing information for complete boxed warning. Severe and sometimes fatal hepatotoxicity has been observed in clinical trials. Monitor hepatic function prior to and during treatment. Interrupt and then reduce or discontinue Stivarga for hepatotoxicity as manifested by elevated liver function tests or hepatocellular necrosis, depending upon severity and persistence. (2.2, 5.1)

🚺 [1] Efficacy & safety of regorafenib monotherapy in Chinese patients with previously treated metastatic colorectal cancer: subgroup analysis of the CONCUR trial; R.

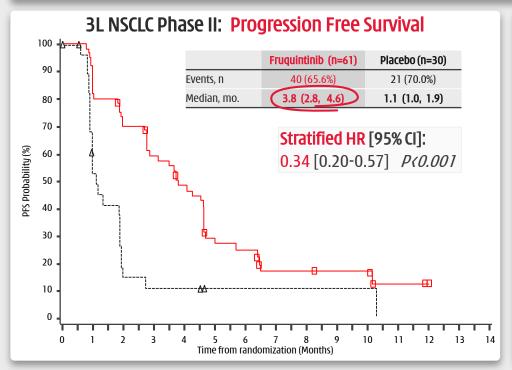
# Fruquintinib - Third-line NSCLC

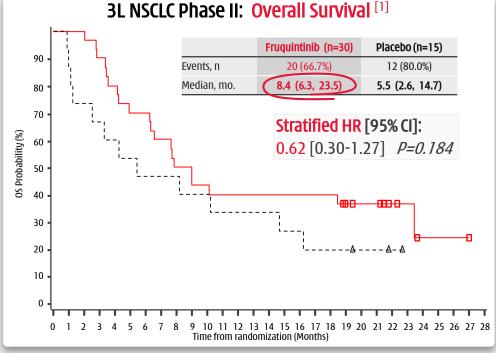




- Non-small cell lung cancer ("NSCLC") Phase II PoC.
  - ✓ 91 3<sup>rd</sup> line only pts. enrolled in ~9 months (Jun'14-Mar '15).
  - Clearly met primary endpoint of reduction in risk of progression.
    \$10 million success milestone from Lilly in Q4 2015.
  - ✓ **AEs consistent** with the known safety profile and generally superior versus 3L colorectal cancer Phase II with lower >Gr.3 AEs (32.8% vs. 66.0%) and dose reductions (13.1% vs. 27.7%).

Patients, %	Fruquintinib (n=61)	Placebo (n=30)
All AEs, any grade	61 (100%)	27 (90.0%)
All AEs, grade ≥3	20 (32.8%)	6 (20.0%)
Hypertension, grade ≥3	5 (8.2%)	1 (3.3%)
Hand-foot syndrome ("HFS"), grade ≥3	3 (4.9%)	0
All other AEs, grade ≥3 (each)	≤2 (≤3.3%)	0
Leading to dose interruption	9 (14.8%)	0
Leading to dose reduction	8 (13.1%)	0
Leading to treatment discontinuation	6 (9.8%)	1 (3.3%)





# Fruquintinib - Third-line NSCLC is competitive Liley



### ...but we believe fruquintinib is well positioned

(MED)

Anlotinib (Sinobiopharm) is about 12-18 months ahead of fruquintinib in 3L NSCLC - their Phase III will reported at ASCO 2017. However, anlotinib Phase II seems to have been in abnormally healthy 3L NSCLC patients (32% placebo DCR<sup>[1]</sup>; 0% brain mets; & only 20% EGFRm<sup>[2]</sup>) so close analysis of their Phase III results will be critical.

Third-line NSCLC	(Blinded Independent   Fruquintinib Phase III   (Local Physician   ISINORIODHAPMI   (FGEP mut + IAT)   WT (		Apatinib Phase II (EGFR WT only) [JIANGSU HENGRUI]		(FGEP WT only)		Lenvatinib Phase II [EISAI]									
Timing				FPI Q4-2015			LPI Q2-2016; Topline Q2-2017		Failed on mPFS Primary endpoint		_		FPI Q1-2015			
		Fruquin.	Placebo	Fruquin. Placebo	Anlotinib	Placebo	Anlotinib	Placebo	Apatinib	Placebo	Apatinib	Placebo	Apatinib	Placebo	Lenvatinib	Placebo
patients (n)		61	31	520 (enrolling)	60	57	294	143	48	80	90	45	417 (enr	rolling)	89	46
Complete Response ("CR")		0 (0%)	0 (0%)		0 (0%)	0 (0%)	0 (0%)	0 (0%)			0 (0%)	0 (0%)			0 (0%)	0 (0%)
Partial Response ("PR")		10 (16%)	0 (0%)		6 (10%)	0 (0%)	27 (9%)	1 (1%)			18 (20%)	1 (2%)			9 (10%)	1 (2%)
Stable Disease ("SD")		33 (54%)	5 (16%)		44 (73%)	18 (32%)	211 (72%)	52 (36%)			44 (49%)	10 (22%)			58 (65%)	12 (26%)
Disease Control Rate ("DCR")		43 (71%)	5 (16%)		50 (83%)	18 (32%)	238 (81%)	53 (37%)			62 (69%)	11 (24%)			67 (65%)	13 (28%)
median Progression Free Survival ("PFS") (m)		3.8	1.2		4.8	1.2	5.4	1.4	Failed mPF	S endpoint	4.7	1.9			4.8	1.8
value		(0.	.001		(0.0	001	<0.0	001			(0.0	001			(0.0	001
Hazard Ratio ("HR")		0.2	275		0.3	20	0.3	25			0.2	!78			0.4	100
median Overall Survival ("OS") (m)		7.7	9.7	mOS Primary endpoint	10.3	6.3	9.6	6.3					mOS Primar	y endpoint	8.7	5.5
P value		0.2	264		0.0	75	0.0	)18								
HR		0.7	743		0.6	56	0.	68								
G3 Adverse Events ("AE")		22 (36%)	8 (27%)		13 (22%)	3 (5%)									61 (69%)	23 (51%)
SAE		6 (10%)	4 (13%)		7 (12%)	8 (14%)									46 (52%)	21 (47%)
HFS >G3, n (%)		3 (5%)	0 (0%)		2 (3%)											
Fatigue >G3, n (%)		2 (3%)	0 (0%)													
Hypertension >G3, n (%)		5 (8%)	1 (3%)		5 (8%)											
Diarrhea >G3, n (%)		1 (2%)	0 (0%)	Phase III			Study	Ph.III					Secon	nd trv	Globa	al price 2k/mo
Proteinuria >G3, n (%)		1 (2%)	0 (0%)				_							_	1.5	2/ /
Triglicerides >G3, n (%)				Top-line	3 (5%)		data:						at 3 <sup>rd</sup>	IIne	~\$13.	.2K/M0
				results			(1) Why	hiah					NSCLO			
AE leading to dose interruption		8 (13%)	0 (0%)										NOCLU	• -		
AE leading to dose reduction		8 (13%)	0 (0%)	2018	6 (10%)	0 (0%)	placebo	DCR?					only	wild-		
AE leading to treatment discontinue		4 (7%)	1 (3%)				(2) EGF	Rm					_		22 (25%)	8 (18%)
								.,,,					type i	EGFR		
	0	4 (7%)	1 (3%)		7 (12%)	3 (5%)	ratio?				20 (22%)	12 (27%)	patie	ntc	17 (19%)	11 (24%)
ECOG PS, n (%)	1	57 (93%)	29 (97%)		47 (78%)	49 (86%)	(3) Brai	'n			70 (78%)	33 (73%)	patie	טווו	63 (71%)	29 (63%)
	2				6 (10%)	5 (9%)							(~40-	60%)	8 (9%)	6 (13%)
5tage, n (%)	IIIB				6 (10%)	2 (4%)	mets?						( 70	33/0)		
nuge, 11 (/0)	IV				54 (90%)	55 (96%)	(4) Cen.	soring? 🥒								
							(5) AEs.	_								
Brain metastases			12%			0%)	(J) MES.									
	+ve	30 (49%)	15 (48%)		12 (20%)	9 (15%)										
EGFR Mutation, n (%)	-ve (WT)	27 (44%)	13 (42%)		48 (80%)	48 (85%)					90 (100%)	45 (100%)				
	unkn.	4 (7%)	3 (10%)		0 (0%)	0 (0%)										

<sup>[1]</sup> DCR = Disease Control Rate; [2] EGFR Mutation positive - In China 40-60% of NSCLC patients harbor EGFR mutation (compared to 10-15% of Caucasian patients in the Wes

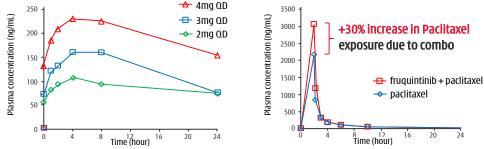
# Fruguintinib - Gastric combo with paclitaxel Liley



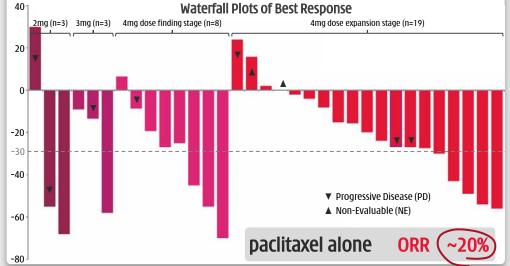


Clear efficacy, safety as expected & +30% incr. in paclitaxel exposure

1. Dose proportional increase of fruquintinib AUC at steady state. Over (30%) increase in paclitaxel drug exposure (mean  $AUC_{0-8}$ ) following multiple dose fruquintinib.



ORR of (36%) (10/32) & DCR of 68% in efficacy evaluable pts. Fruguintinib 4mq,  $\geq$ 16 wk. PFS of 50% &  $\geq$ 7 mo. OS of 50%.

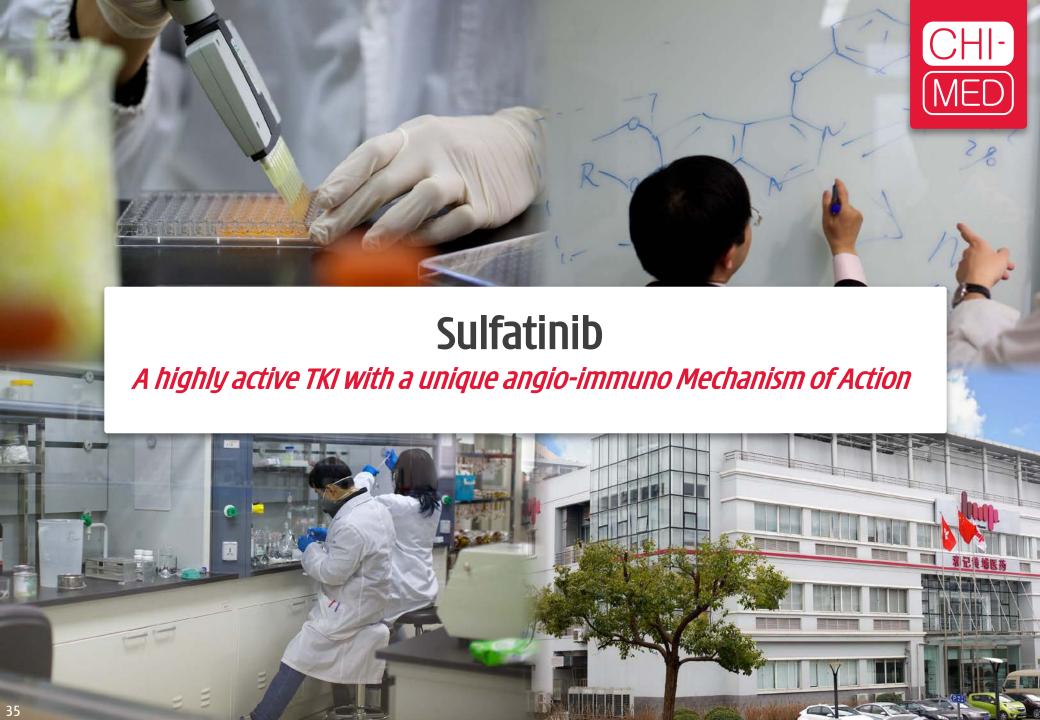


3. Encouragingly low level of dose reduction/interruption. Actual mean administered dose in the first cycle was 3.32mg/day for fruquintinib (83.0% planned dose) & 78.6 mg/m2/week for paclitaxel (98.3% planned dose).

Characteristics (Unit)	Drug Expansion Stage (N=19) Fruquintinib 4 mg + paclitaxel 80 mg/m²	
	Drug interruption	Drug reduction
Dose modification with Fruquintinib N (%)	2 (10.5%)	2 (10.5%)
Dose modification with Paclitaxel N (%)	5 (26.3%)	1 (5.3%)

4. AE profile in-line with expectations. Neutropenia - a paclitaxel driven AE - with 57.9% Grade >3 AEs. Similar to 60% level seen in RAINBOW study of ramcirumab (VEGF mAb) combo with paclitaxel in second-line Gastric cancer.

Drug related grade 3 or 4 AEs (NCI-CTCAE v 4.0) term	Dose Expansion Stage(N=19) Fruquintinib 4 mg + paclitaxel 80 mg/m²	
Neutropenia	11 (57.9%)	
Leukopenia	4 (21.0%)	
Hypertension	2 (10.6%)	
PLT decreased	1 (5.3%)	
Anemia	1 (5.3%)	
HFSR	1 (5.3%)	
Mucositis oral	1 (5.3%)	
Hepatic disorder	1 (5.3%)	
Upper gastrointestinal hemorrhage	1 (5.3%)	

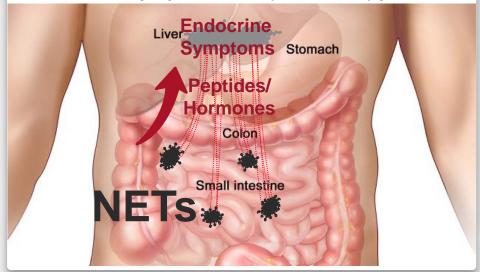


# **Neuroendocrine tumors ("NET")**

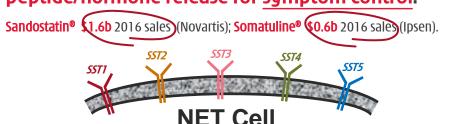
### Sulfatinib potential advantages



1. NETs release peptides & hormones that cause endocrine symptoms such as hot flushes, diarrhea, nausea, heart palpitations & (abdominal) pain.



2. Somatostatin analogues ("SSTA"): Inhibit peptide/hormone release for <u>symptom control</u>.<sup>[3]</sup>



# 3. Available NET therapies – control symptoms/tumor growth but provide minimal tumour shrinkage:

- Sandostatin® & Somatuline® (SSTAs) are used primarily for symptom control in early stage NET (Ki67 <10%) SSTAs do provide some tumor growth control (DCR/mPFS) but almost no tumor shrinkage (ORR);
- Lutathera® radio nucleotide SSTA delivers radiation to NET via SST receptors very effective <40 mo. mPFS & ~18% ORR in midgut NET (~21% of NETs) with MoA potential in other NETs. Primary issues around logistics half-life 3 days requiring efficient product supply systems not very practical for broad scale usage in developing world;
- Sutent® & Afinitor® in pancreatic NET & certain lung/GI NETs provide tumor growth control (DCR/mPFS) but low tumor shrinkage (<10% ORR)

#### 4. Emerging advantages of sulfatinib:

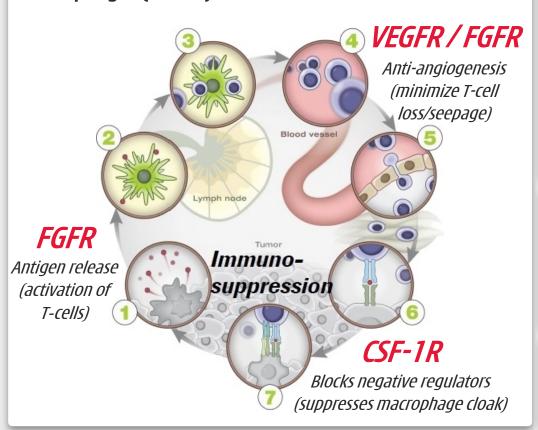
- Broad spectrum NET efficacy:
  - (1) Tumor control & shrinkage across all NET sub-types;
  - (2) Unique angio-immuno MoA 2L usage (post failure on 1L therapy);
  - (3) Efficacy in ~20% of NET patients without overexpressed SST receptors.
- Convenience/cost:
  - (1) **Oral formulation** vs. very short half-life (3 days) injection (Lutathera®);
  - (2) **Cost/pricing vs. Lutathera**<sup>®</sup> est. >\$200k/yr.; Sutent® \$140k/yr.

# Sulfatinib's unique angio-immuno kinase profile



Multi-dimensional global development program, initially for NETs<sup>[1]</sup>

Sulfatinib's unique angio-immuno kinase profile & MoA<sup>[2]</sup> activates & enhances the body's immune system, namely T-cells, via VEGFR/FGFR while inhibiting the prod- uction of macrophages (CSF-1R) which cloak cancer cells.



# Activity 1: Fast/first approval in China for all NET [1] patients - 2x pivotal Phase III trials in progress

	Pancreatic NET Phase III	Non-Pancreatic NET Phase III		
Primary site	Pancreas	GI, lung, other or unknown		
Population	Unresectable or metastatic disease; well differentiated (G1/G2); ≤2 prior systemic drugs.			
# of Sites	20-30	O (China)		
# of Patients	~195 ~270			
Study design	Double-blind. Randomized 2:1 to sulfatinib or placebo, until PD. Predefined interim analysis.			
Dosage	Sulfatinib 300mg QD, 28 days per cycle (vs. placebo)			
Primary Endpoint	Progression-Free Survival (PFS) by BICR evaluation			
Secondary Endpoints	Overall Survival (OS), ORR, safety, etc.			
First Patient In / Readout	March 2016 / 20 <u>18</u> December 2015 / 20 <u>18</u>			

### **Activity 2: Global development**

- U.S. Phase I bridging in Caucasian patients almost complete -RP2D<sup>[3]</sup> expected to be same as China - 300mg QD.
- U.S. Phase II in planning, expect to start in 2017 focusing on areas of NET unmet medical need/BTT<sup>[4]</sup> opportunity.

### Activity 3: Exploratory PoC<sup>[5]</sup> in other indications

China Ph.II studies underway in: (a) Medullary thyroid cancer; (b) Differentiated thyroid cancer; and (c) Biliary tract cancer.

# Activity 1: China NET - Phase II (ENETS 2017 [7])



PDs or

**Deaths** 

(% pts)

51.9%

(42/81)

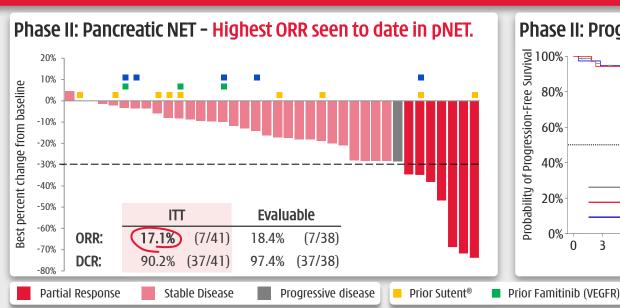
39.0%

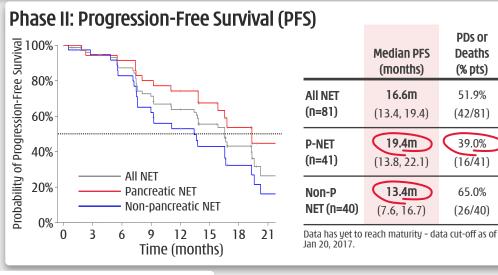
(16/41)

65.0%

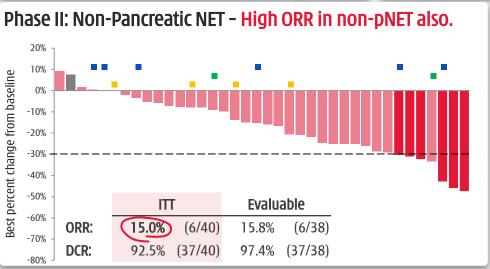
(26/40)

Efficacy in pNET & non-pNET; & patients who failed on Sutent®/Afinitor®





Prior Afinitor®



### Phase II: Safety - Well tolerated - Adverse Events manageable.

	Grade ≥3 (≥4pts) n (%)	Adverse Events ("AEs") - Regardless of causality	N=81 n (%)
Hypertension	25 (30.9)	Any AE	81 (100.0)
Proteinuria	11 (13.6)	Grade ≥3 AE	63 (77.8)
Hyperuricemia	8 (9.9)	Any SAE	21 (25.9)
Hypertriglyceridemia	7 (8.6)	Any drug-related AE	81 (100)
Diarrhea	6 (7.4)	Any drug-related grade ≥3 AE	58 (71.6)
ALT increased	5 (6.2)	Any drug related SAE	10 (12.3)
Anemia	4 (4.9)	Drug related AE leading to:	
Hypokalemia	4 (4.9)	dose interruption	40 (49.4)
Hepatic function	4 (4.9)	dose reduction	20 (24.7)
abnormal	4 (4.7)	drug withdrawal	7 (8.6)

# Activity 1: China NET - Phase II (ENETS 2017 [1])

Tumor devascularization & central necrosis



Patient 1

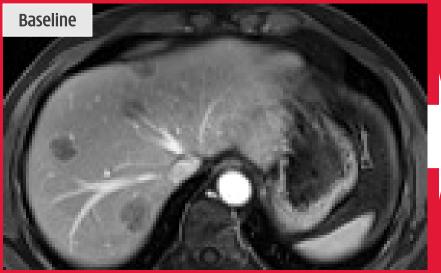
Duodenum NET G2

w/ multiple liver & retroperitoneal lymph node metastases





Patient 2
Rectum NET G2



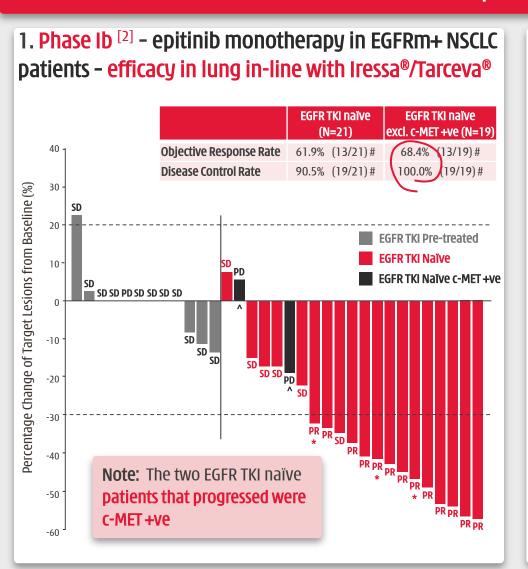


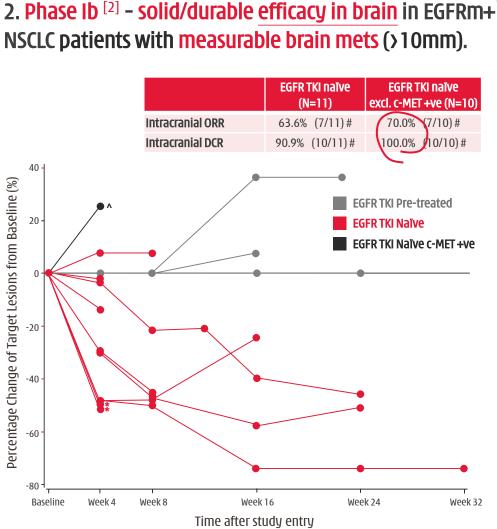


# Epitinib – Blood-brain-barrier penetrating TKI



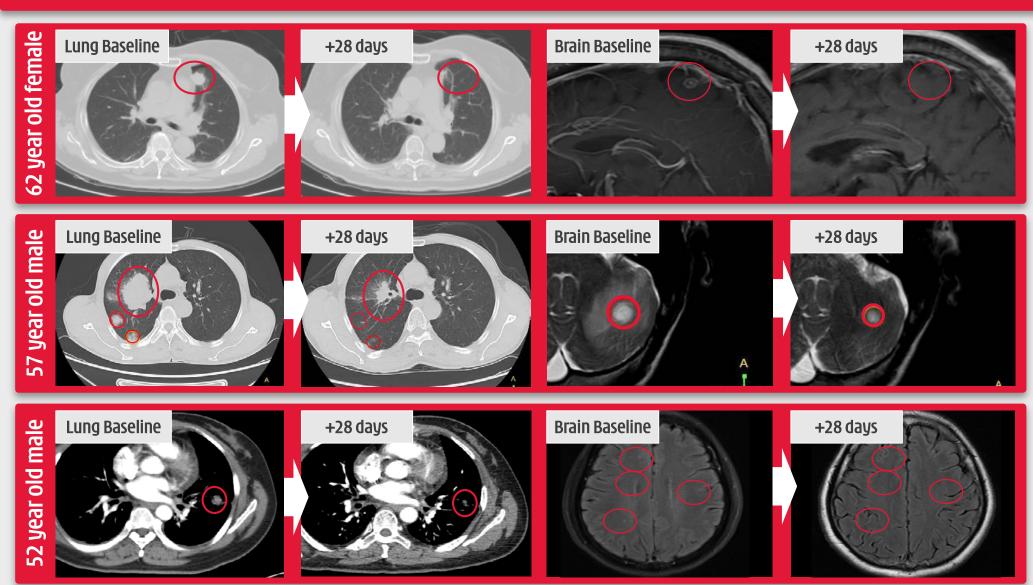
Unmet medical need for ~50% NSCLC patients that develop brain mets<sup>[1]</sup>





# CHI-

# Epitinib - Powerful Phase Ib efficacy



## Epitinib - Safe & well tolerated

### Pivotal Phase III study to initiate in 2017



3. Epitinib well tolerated by patients<sup>[1]</sup> w/advanced solid tumours. Safety profile is consistent with that of approved EGFR-TKIs (e.g. Iressa®/ Tarceva®).

Dose Escalation Stage (n=35*)				
(Drug related AEs reported > 10%)				
All Grades Grade 3/4				
Adverse Event ("AE") n (%) n (%)				
Skin rash	21 (60.0%)	1 (2.9%)		

12 (34.3%)

11 (31.4%)

10 (28.6%)

5 (14.3%)

5 (14.3%)

5 (14.3%)

4 (11.4%)

4 (11.4%)

4 (11.4%)

Diarrhea

AST increase

**ALT increase** 

**Stomatitis** 

**Pruritus** 

Total bilirubin increase

**Exfoliative dermatitis** 

**Hyper-pigmentation** 

Gamma-GGT increase

Conjugated bilirubin

(Drug related AEs reported >10%)					
All Grades Grade 3 Adverse Event ("AE") n (%) n (%)					
Skin rash	31 (83.8%)	2 (5.4%			
Hyper-pigmentation	18 (48.6%)	1 (2.7%			
ALT increase	15 (40.5%)	7 (18.9%			
AST increase	15 (40.5%)	4 (10.8%			
ASP increase	11 (29.7%)	1 (2.7%			
Diarrhea	10 (27.0%)	-			
Proteinuria	10 (27.0%)	-			
Total bilirubin increase	9 (24.3%)	1 (2.7%			
Hyperuricemia	9 (24.3%)	2 (5.4%			
Gamma-GGT increase	7 (18.9%)	4 (10.8%			
Stomatitic	4 (14 20/)	_			

Dose Expansion Stage (n=37)

### 4. Now moving into Phase III pivotal study in China.

- Phase III in first-line NSCLC with brain metastasis to start:
- Published positive Phase Ib expansion results at World Conference on Lung Cancer Dec 2016, Vienna.

1 (2.9%)

1 (2.9%)

2 (5.7%)

2 (5.7%)

1 (2.9%)

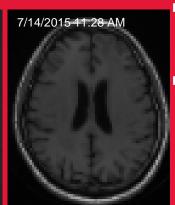
- → China FDA Phase III clinical trial cleared in July 2016 initiating Phase III in 2017.
- Glioblastoma (primary brain tumors):
- **Phase II proof-of-concept planning underway**, initiating 2017.

### CASE STUDY - EGFR-TKI pretreated patient

- Man, 58 y.o., diagnosed with NSCLC adenocarcinoma (Exon21 L858R) on Dec 12, 2014.
- Tumor lesions located at left lung upper lobe, bone & brain cT1bN3M1.
- 3 days prior brain radiotherapy, followed by Iressa® for 5.5 months with most recent progression in the brain.

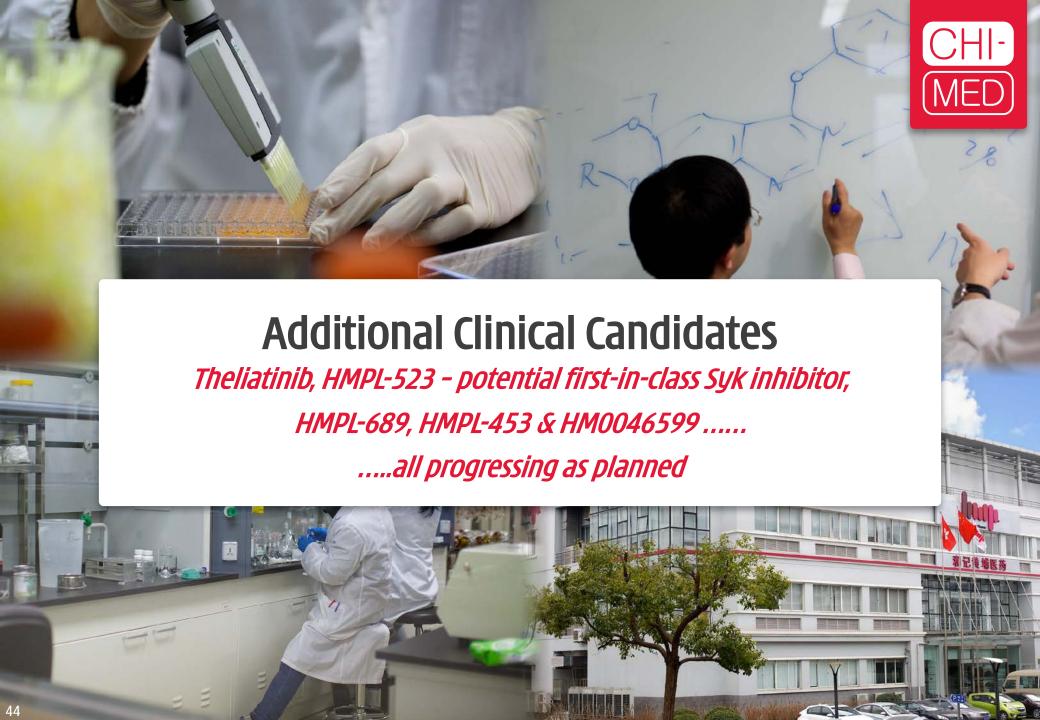


6/1/2015 1:48 PI





- Patient presented walking with crutch assistance.
  - Epitinib 160 mg q.d. began on Jun 17, 2015. Achieved stable disease in both intracranial & extracranial lesions from week 8, & could walk without assistance.
- Remained on stable disease for 43 weeks until disease progression (pleural effusion).



# Theliatinib – encouraging activity observed



### Potent & highly selective TKI – strong affinity to wild-type EGFR kinase

# 1. Major unmet medical need for wild-type EGFR activation tumors.

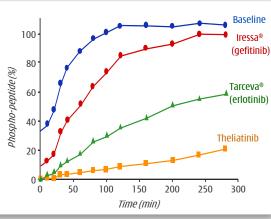
■ EGFR activation affects multiple tumor types. Current EGFR TKIs are less effective in treating solid tumors with wild-type EGFR activation (gene amplification & protein over expression).

Phase Ib expansion study on theliatinib in esophageal cancer is currently underway in China.
TKIS approved:

Tumor Types	Wild-type: Gene \ Amplification	wild-type: Over Expression	Mutations Mutations
NSCLC	29%	62%	10-30%
Esophagus	8-30%	30-90%	12% (esophageal adenocarcinoma)
Stomach	29%	44-52%	₹5%
Glioblastoma	36-51%	54-66%	27-54% (EGFR variant III)
Colorectal	4.5%	53%	8%
Head and neck	10-30%	66-84%	42% (EGFR variant III)  MAbs approved: Frbitux® Vectibix®

# 2. Superior anti-tumor activity of theliatinib in pre-clinical studies with wild-type EGFR.

- 5-10-fold more potent than Tarceva<sup>®</sup>.
- Sustained target occupancy.



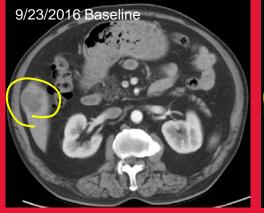
# 3. Esophageal cancer (EC): No effective treatment options.

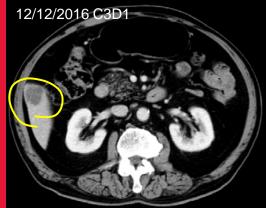
Major issue in Asia with poor prognosis: 5 year survival 10-20%

	new cases/year	deaths/year
U.S.	16,940	15,690
China	477,900	375,000

### CASE STUDY - EGFR protein over expression

- May 4, 2016: Man, 62, stage IV esophageal squamous cell cancer cT3N0M1with liver metastasis. High protein overexpression - EGFR IHC local test: >75% of tumor cells 3+.
- May 4 to Sep 23, 2016: nimotuzumab/placebo + paclitaxel + cisplatin 6 cycles with best tumor response: PD.
- Oct 11, 2016: began theliatinib 400mg daily.
- Dec 12, 2016: Cycle 3 Day 1 (C3D1) tumor assessment: Target lesion (liver metastasis) shrank -33% (36mm to 23mm diameter) - unconfirmed PR.
- Jan 23, 2017: Withdrew from study due to AEs Gr 1 (diarrhea/pruritus/dental ulcer), Gr 2 (epifolliculitis/dermatitis).





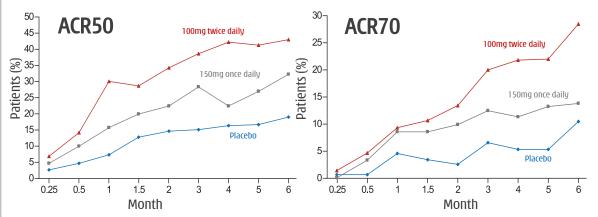
### HMPL-523 - superiority vs. fostamatinib



### Superior selectivity, better target coverage & efficacy



...but GI toxicity, infection & 23% put on antihypertensives.

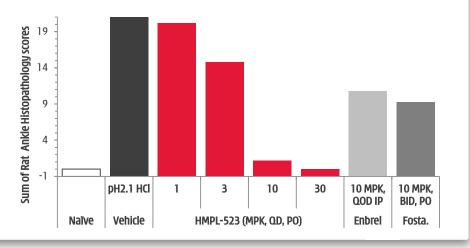


Percent of patients	Placebo (n = 153)	150mg QD (n = 152)	100mg BID (n = 152)
Diarrhea	3.0%	11.8%†	19.1%†
Upper respiratory infection	7.1	7.2	14.5 †
Urinary tract infection	4.6	3.3	5.9
Nausea	4.6	5.9	4.6
Neutropenia	0.7	6.6 †	5.9 †
Headache	5.2	6.6	5.9
Abdominal pain	2.6	6.6 †	5.9 †
ALT >3x ULN	2.0	3.9	3.9
Dizziness	2.0	2.6	4.6
Hypothyroidism	2.6	2.6	3.3
Cough	2.6	2.0	3.3
$\dagger P \in 0.05$ for comparison with placebo group; ALT = alanine aminotransferase.			

#### 2. HMPL-523 - far superior selectivity to fostamatinib......

Selectivity	HMPL-523 IC <sub>50</sub> (nM)	fostamatinib IC <sub>50</sub> (nM)
Syk enzyme	25 ± 5 (n=10)*	54 ± 16 (n=10)*
JAK 1,2,3 enzyme	>300, >300, >300*	120, 30, 480*
FGFR 1,2,3	>3,000, >3,000, >3,000	89, 22, 32*
FLT3 enzyme	63*	9*
LYN enzyme	921*	160*
Ret enzyme	>3,000*	5**
KDR enzyme	390 ± 38 (n=3)*	61 ± 2 (n=3)*
KDR cell	5,501 ± 1,607 (n=3)*	422 ± 126 (n=3)*

#### .and very strong efficacy in preclinical RA models.

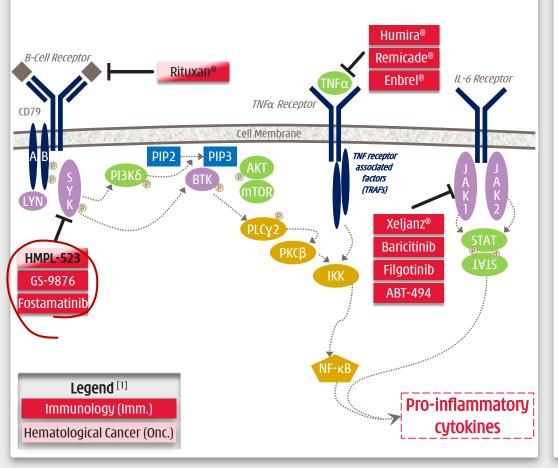


## HMPL-523 - immunology potential



### Potential first-in-class Syk inhibitor in immunology - Phase II in planning

1. Syk, the most upstream B-cell pathway kinase target is clinically validated in rheumatoid arthritis ("RA"), but currently Chi-Med & Gilead are the only companies pursuing.



2. RA expected to be a \$45 billion market in 2020 with B-cell pathway; anti-TNF; & JAK the main focus.

(Methotrexate-IR: placebo adjusted)	ACR20	ACR50	ACR70	2016 Sales (\$billion) <sup>[2]</sup>
B-Cell receptor mAbs				(, , , , , , , , , , , , , , , , , , ,
Rituxan® (24-Week)	33%	21%	11%	1.6
Anti-TNFα/NF-κB mAbs				
Humira® (24-Week)	33%	29%	18%	16.1
Remicade® (24-Week)	30%	22%	8%	7.0
Enbrel® (24-Week)	44%	36%	15%	8.3
JAK Inhibitors Small molecules				
Xeljanz® (24-Week)	25%	23%	13%	0.9
Xeljanz® (12-Week)	28%	21%	8%	0.9
baricitinib 4mg QD (12-Week)	30%	28%	14%	n/a
filgotinib 100mg BID (12-Week)	35%	40%	23%	n/a
ABT-494 24mg QD (12-Week)	32%	24%	18%	n/a
Syk Inhibitor Small molecule				
fostamatinib 100mg BID (24-Week)	32%	24%	18%	n/a

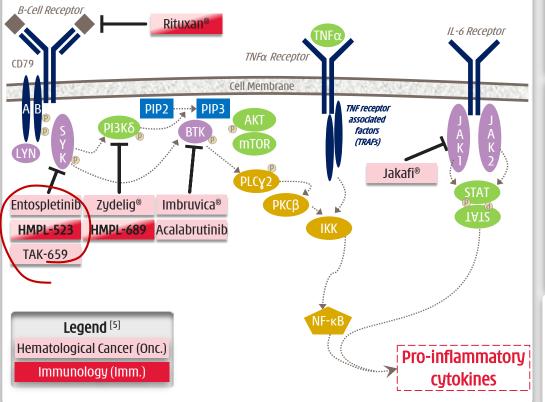
- 3. Substantial market potential remains in RA.
- mAbs intravenous administration and shut down immune system for 4-6 weeks - high infection / lymphoma risks.
- First-in-class JAKs in RA limited by compound-related tox.
- Syk inhibition shown to benefit patients but fostamatinib failed due to major off-target toxicity.

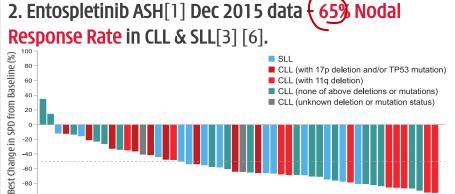
### HMPL-523 - hematological malignancies



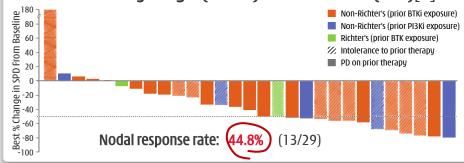
### Syk exciting target emerging in oncology - Lymphoma Phase I ongoing

- 1. The B-cell signaling is critical in hematological cancer with three breakthrough therapies recently approved.
- Sales in 2016 of Imbruvica® were \$1.8 billion; Zydelig® \$0.2 billion; Jakafi® \$0.6 billion; & Rituxan® \$6.5 billion[2].





3. Entospletinib potential for overcoming resistance/ intolerance to Zydelig® (PI3Kδ) & Imbruvica® (BTK)[6].



- 4. Entospletinib not a perfect compound[6].
- Poor solubility/oral absorption & high variation in drug exposure.
- Some CYP[4] inhibition & increased risk of drug-drug interaction.
- 66% Grade ≥3 AEs, 49% SAEs, 46% drug interruption & 20% disco.

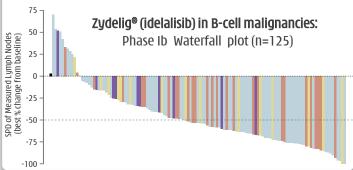
### **HMPL-689**



### Designed to be a best-in-class inhibitor of PI3Kδ - Phase I started

#### 1. PI3Kδ now a proven target.

- PI3Kδ activation associated with allergy, inflammation & oncology.
- Evidence that PI3Kδ inhibitors effective in ibrutinib-resistant mutant population.



### 3. HMPL-689 -- Important asset.

Designed to improve on existing PI3Kδ inhibitors:

- Improved isoform selectivity (sparing PI3Kγ).
- Improved potency at whole blood level (>5x more potent than idelalisib) to cut compound related toxicity.
- Improved PK properties particularly efflux and drug/drug interaction due to CYP inhibition / induction, critical for combo therapy.

### 2. PI3Kδ inhibitors being developed in a very broad range of indications.

Compound		Indication	Status	Issue
Zydelig®	Gilead	Chronic lymphocytic leukaemia, non-Hodgkin's lymphoma	Registered	High incidence of liver
(idelalisib)	Sciences	Hodgkin's lymphoma	Phase II Trial	toxicity seen with
РІЗКδ	54.4	Waldenstrom's hypergammaglobulinaemia	Preclinical	idelalisib (150mg bid)
AMG-319 PI3Kδ	Amgen	B-cell lymphoma, non-Hodgkin's lymphoma, T-cell lymphoma, chronic lymphocytic leukaemia	Phase I Trial	
duvelisib <sup>[1]</sup>		B-cell lymphoma, non-Hodgkin's lymphoma, chronic lymphocytic leukaemia	Phase III Trial	Need to spare PI3Ky serious infection seen
(IPI-145) ADDVIE /	Asthma, rheumatoid arthritis	Phase II Trial	with duvelisib due to	
ΡΙ3Κγ/δ	Infinity	COPD, SLE, psoriasis, MS transplant rejection, allergy, acute lymphocytic leukaemia, T-cell lymphoma	Phase I Trial	strong immune suppression

### 4. HMPL-689 more potent and more selective than idelalisib & duvelisib.

Enzyme IC <sub>50</sub> (nM)	HMPL-689	Zydelig <sup>®</sup>	duvelisib
ΡΙ3Κδ	0.8 (n = 3)	2	1
PI3K <sub>γ</sub> (fold vs. PI3Kδ)	114 (142x)	104 <b>(52x)</b>	2 (2X)
PI3Kα (fold vs. PI3Kδ)	>1,000 (>1,250x)	866 <b>(433x)</b>	143 <b>(143x)</b>
PI3Kδ human <u>whole blood</u> CD63+	3	14	15
PI3Kβ (fold vs. PI3Kδ)	87 <b>(109x)</b>	293 <b>(147x)</b>	8 (8X)

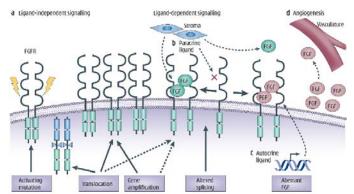
### **HMPL-453**



### Designed to be a first-in-class FGFR1/2/3 inhibitor - Phase I started

#### FGFR genetic alterations are oncogenic drivers

- FGF/FGFR signaling normally involved in embryonic development, tissue repair, angiogenesis, neuroendocrine and metabolism homeostasis.
- Multiple oncogenic driver genetic alterations in FGFR pathway: gene amplification, mutation, translocation, fusion, splicing, etc.

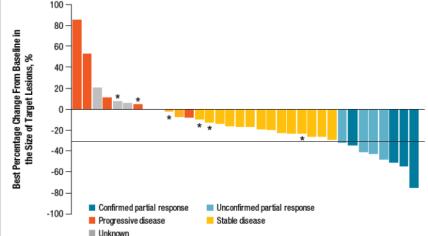


 Diverse and complicated genetic changes and multiple tumor types with low incidence

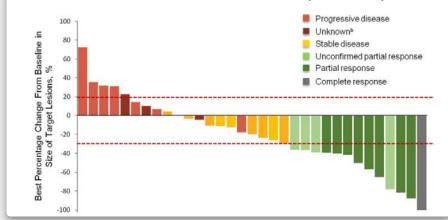
	Gene amplification	Gene translocation	Gene mutation
FGFR1	Lung squamous (7~15%) H&N squamous (10~17%) Esophageal squamous (9%) Breast (10~15%)	Lung squamous (n/a) Glioblastoma (n/a) Myeloproliferative syndrome (n/a) Breast (n/a)	Gastric (4%) Pilocytic astrocytoma (5~8%)
FGFR2	Gastric (5~10%) Breast (4%)	Intra-hepatic cholangiocarcinoma (14%) Breast (n/a)	Endometrial (12~14%) Lung squamous (5%)
FGFR3	Bladder (n/a) Salivary adenoid cystic (n/a)	Bladder (3~6%); Lung squamous (3%); Glioblastoma (3%) Myeloma (15~20%)	Bladder (60~80% NMIBC; 15~20 MIBC) Cervical (5%)

# Cholangiocarcinoma (CCA) and bladder cancer have made much progress in clinic to date

BGJ398 Phase II PoC in cholangiocarcinoma (2016 ASCO GI).



BGJ398 Phase II PoC in bladder cancer (2016 ASCO).





### Chi-Med's Commercial Platform in China



### Long track record of commercial success - important source of cash

### 2 National household name brands



# Focus on largest disease categories

Most common disease diagnosed/treated in rural hospitals[1]:

Cold/Flu: 86%
Cardiovascular: 78%
Diabetes: 46%
Gl: 45%

# Major commercial & production scale

~2,200 Rx & ~1,200 OTC sales people in about 300<sup>[2]</sup> cities & towns in China.

Drugs in ~18,700 hospitals detailing ~87,000 doctors.

Sold ~4.5 billion doses of medicine in 2016.

### Leadership market shares

Market leader in the subcategories/markets in which we compete<sup>[3]</sup>:

SXBX pill: [4][5] ~12% Rx Cardiovascular TCM Banlangen: [6] ~51%

Banlangen:<sup>[6]</sup>
OTC Anti-viral /flu TCM

FFDS tablet:<sup>[7]</sup> ~32% OTC Angina TCM

### JVs with 3 leading China Pharmas







### Commercial Platform Performance - 2003-2016<sup>[8][9]</sup>

					IF	RS						US G	AAP		15-16
(US\$ millions)	03	04	05	06	07	08	09	10	11	12	13	14	15	16	Growth
Sales	21.9	27.9	65.1	101.4	119.0	155.8	197.0	236.4	278.6	360.7	402.3	465.4	518.9	627.4	21%
Prescription Drugs	17.2	21.8	23.3	23.2	28.1	39.5	54.4	71.2	92.4	116.5	138.2	204.9	286.6	372.3	30%
Consumer Health	4.7	6.1	41.8	78.2	90.9	116.3	142.6	165.2	186.2	244.2	264.1	260.5	232.3	255.1	10%
Total Sales Growth	n/a	27%	133%	56%	17%	31%	26%	20%	18%	29%	n/a	16%	11%	21%	
Net (loss)/Income after tax	(10.7)	(3.6)	2.2	6.7	11.2	14.7	21.5	27.9	30.1	33.1	39.7	48.8	54.1	144.1[11]	167%
Prescription Drugs	(0.4)	1.3	1.9	1.3	1.9	2.8	6.0	11.9	14.2	17.7	22.4	26.5	31.9	122.2	284%
Consumer Health	(10.3)	(4.9)	0.3	5.4	9.3	11.9	15.5	16.0	15.9	15.4	17.2	22.3	22.2	21.9	-1%
% Margin	-48.9%	-12.9%	3.4%	6.6%	9.4%	9.4%	10.9%	11.8%	10.8%	9.2%	9.9%	10.5%	10.4%	23.0%	
Net (loss)/income attrib. to Chi-Med	(5.7)	(3.7)	(0.5)	1.2	4.5[10]	5.9[10]	9.3[10]	12.6[10]	13.6[10]	14.6[10]	18.2[10]	22.8[10]	25.2[10]	70.3[11]	180%
Prescription Drugs	(0.2)	0.6	1.0	0.7	0.9	1.4	3.0	5.9	7.1	8.8	11.2	13.2	15.9	61.1	284%
Consumer Health	(5.5)	(4.3)	(1.5)	0.5	3.6	4.5	6.3	6.7	6.5	5.8	7.0	9.6	9.3	9.2	0%
Net (loss)/income attrib. to Chi-Med growth	n/a	-35%	-86%	340%	275%	31%	58%	35%	8%	7%	n/a	26%	10%	180%	

# A powerful Rx Commercial Platform in China



### Chi-Med management run all day-to-day operations

- National Coverage:
  - ~300 cities & towns.
  - ~18,700 hospitals.
  - ~87,000 doctors.
- New team of 143 CNS reps built since 2015.



490 (23%)

**NORTH** 

Pop'n: 320m (23%)

CV Medical Reps: CNS Medical Reps: HSP Sales staff:

32 (22%)

### **WEST**

Pop'n: 100m (7%)

CV Medical Reps: 76 (4%)
CNS Medical Reps: 5 (3%)
HSP Sales staff: 0 (0%)

**81** (4%)



**568** (26%)

(6%)

**EAST** 

Pop'n: 393m (28%)

CV Medical Reps: 808 (40%) CNS Medical Reps: 61 (43%) HSP Sales staff: 31 (100%)

### **SOUTHWEST**

Pop'n: 190m (14%)

CV Medical Reps: 112 (6%) CNS Medical Reps: 12 (9%) HSP Sales staff: 0 (0%)

### **CENTRAL-SOUTH**

Pop'n: 383m (28%)

CV Medical Reps: 535 (27%)
CNS Medical Reps: 33 (23%)
HSP Sales staff: 0 (0%)

## Deep portfolio of household name drugs



>200 products - Top 7 represent 63% of sales<sup>[1]</sup> and 92% of gross profit<sup>[1]</sup>

Main Prod	ducts SALES <sup>[2]</sup>	2011	2012	2013	2014	2015	2016
- 小型	SXBX pill Coronary artery disease (Rx) 12% National market share Patent expiry 2029	<b>79,438</b> +32%	102,215 +29%	<b>123,587</b> +21%	138,848 +12%	1 <b>59,326</b> +15%	195,371 +23%
TOTAL STATE OF THE PARTY OF THE	FFDS tablet Angina (OTC) 32% National market share	<b>57,001</b> -3%	60,181 +6%	<b>69,996</b> +16%	76,297 +9%	60,154 -21%	<b>59,906</b> <i>0%</i>
	Banlangen granules Anti-viral/flu (OTC) 51% National market share	57,278 +8%	65,381 +14%	<b>72,300</b> +11%	<b>55,573</b> -23%	<b>54,793</b> -1%	<b>56,664</b> +3%
Seroquel XR	Seroquel tablets Bi-polar/Schizophrenia (Rx) 5% National market share	n/a	n/a	n/a	n/a	21,131	34,380 +63%
◎ 藤川清片 毎川清片 ※	<b>NXQ tablet</b> Cerebrovascular disease (Rx) Proprietary formulation	<b>3,741</b> +55%	<b>6,933</b> +85%	10,142 +46%	<b>14,681</b> +45%	17,581 +20%	<b>21,000</b> +19%
日本中国社 一	KYQ granules Periodontitis (OTC) >90% National market share	15,412 +22%	16,351 +6%	16,318 <i>0%</i>	18,370 +13%	1 <b>7,051</b> -7%	17,210 +1%
胆宁片	Danning tablet Gallbladder/stone (RX) Patent expiry 2027	9,914 +22%	11,648 +17%	12,364 +6%	13,822 +12%	13,526 -2%	9,041 -33%

[1] Based on aggregate sales and gross profit of consolidated subsidiaries and non-consolidated joint ventures; [2] Rx = prescription drug; OTC = over-the-counter drug; SXBX pill = She Xiang Bao Xin pill; FFDS tablet = Fu Fang Dan Shen tablet; NXQ table = Nao Xin Qing tablet; KYQ granules = Kou Yan Qing granules; Market shares according to Frost & Sullivan.



### 2017 Guidance



### Over performance in 2016 - Strong Commercial Platform & property gain

(US\$ millions)	2016 Actual	2017 Guidance
Revenues	216.1	225.0 - 240.0
Innovation Platform		
Revenue	35.2	35.0 - 40.0
Innovation platform operating expenses	(76.1)	(85.0) - <u>(90.0)</u>
Commercial Platform		
Sales (consolidated)	180.9	190.0 - 200.0
Sales of non-consolidated joint ventures	446.5	480.0 - 500.0
Net income attributable to Chi-Med – Total	70.3	46.0 - 50.0
- Core business	29.9	32.0 - 34.0
- One-time property compensation gain	<b>40.4</b> [1]	<b>14.0 - 16.0</b> <sup>[2]</sup>
Chi-Med Group Costs		
General & admin. expenses (incl. interest/tax)	(17.9)	(18.0) - (19.0)
Net (Loss)/Income Attributable to Chi-Med	11.7	(13.0) - (28.0)

### **Balance Sheet**

- \$173.7m available cash resources (Dec 31, 2016) at Chi-Med Group level.
  - $\checkmark$  \$103.7m cash & cash equiv. & ST invest.[3]
  - ✓ \$70m unutilized banking facilities.
- ~\$40m dividend from JV to Group level in mid-2017.
  - ✓ JVs no bank borrowings.
  - ✓ JV cash \$91.0m before dividend payout (Dec. 31, 2016).
- \$46.8m bank borrowings as at December 31, 2016.



## Expected 2017 catalysts

### ■ Target to publish data on 4 drug candidates in 5 Phase II-III studies:

- ✓ Savolitinib: 1. Phase II median overall survival data in PRCC patients;
  - 2. Phase IIb data in second-line NSCLC combinations with Tagrisso® & Iressa®;
  - 3. Phase II dose finding data in ccRCC combination with durvalumab (PD-L1).
- ✓ Fruquintinib: 4. Phase III FRESCO study full data set publication in CRC patients.
- Sulfatinib:
  5. Preliminary Phase II proof-of-concept data in medullary and differentiated thyroid cancer patients.
- ✓ HMPL-523 (Syk):
   6. Preliminary Phase Ib proof-of-concept data in hematological cancer patients.

### Target to achieve multiple late-stage/global clinical & regulatory milestones by end of 2017:

- ✓ Savolitinib: 1. Initiate global Phase III study in PRCC patients;
  - 2. Initiate **global Phase III study in second-line NSCLC** in combination with Tagrisso®;
- ✓ Fruquintinib:
  3. Submit New Drug Application ("NDA") in China in third-line CRC;
  - 4. Initiate China Phase III study in second-line gastric cancer patients;
  - Complete enrollment of Phase III FALUCA study in third-line NSCLC;
  - 6. Initiate **U.S. Phase I bridging study** in Caucasian patients.
- ✓ **Epitinib**: 7. Initiate **China Phase III study in first-line EGFR-mutant NSCLC** patients with brain metastasis;
  - 8. Initiate China Phase II study in glioblastoma (primary brain cancer).
- ✓ **Sulfatinib:** 9. Initiate **U.S. Phase II study in NET** patients.
- ✓ HMPL-523: 10. Initiate Australian Phase Ib/II expansion study in hematological cancer patients.
- ✓ HMPL-689 (PI3Kδ): 11. Initiate Phase I studies in China in hematological cancer patients.
- ✓ HMPL-453 (FGFR-1/2/3): 12. Initiate Phase I studies in Australia/China in solid tumor patients.



# **Appendices**

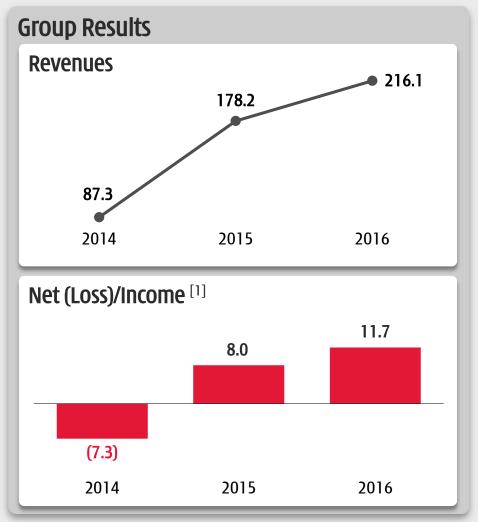
### **2016 Financial Results**



### Record net income - despite <\$76 million innovation platform investment

### **Financial Summary**

				Cha	nge
	2014	2015	2016	14-15	15-16
Revenues	87.3	178.2	216.1	104%	21%
Unconsolidated JV Revenues	398.4	392.7	446.5		
Net (Loss)/Income [1]					
Innovation Platform	(22.2)	(3.8)	(40.7)	83%	~10x
Base HMP Operations	(13.8)	(0.0)	(36.5)		
50% share of Nestlé JV (NSP) [2]	(8.4)	(3.8)	(4.2)		
Commercial Platform (Con't. Operations)	22.8	25.2	70.3	10%	180%
Prescription Drugs Business	13.2	15.9	61.1		
- Base business	13.2	15.9	20.7	20%	30%
- Land compensation (SHPL) <sup>[3]</sup>	-	-	40.4		
Consumer Health Business	9.6	9.3	9.2	-4%	0%
Chi-Med Group Costs	(9.0)	(13.4)	(17.9)	-49%	-34%
General & administrative Expenses	(6.4)	(10.9)	(12.6)		
Interest/Tax	(2.6)	(2.5)	(5.3)		
Discontinued Operations	1.0	-	-	n/a	n/a
Net (Loss)/Income Attrib. to Chi-Med	(7.3)	8.0	11.7	n/a	46%
EPS Attrib. to Company (Basic) (US\$)	(0.14)	0.15	0.20	n/a	34%
Accretion per share on redeemable NCI-Non-cash [4]	(0.48)	(0.79)	-		
EPS Attrib. to Ordinary Shareholders (Basic) <sup>[5]</sup>	(0.62)	(0.64)	0.20	n/a	n/a



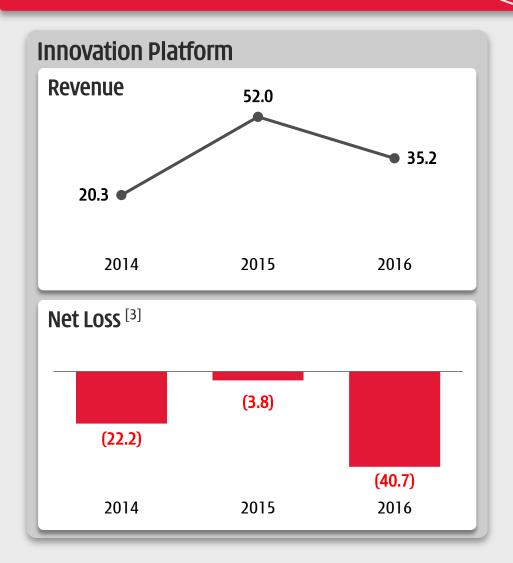
<sup>[1]</sup> Net (Loss)/Income = Net (Loss)/Income attributable to Chi-Med; [2] NSP = Nutrition Science Partners Limited; [3] SHPL = Shanghai Hutchison Pharmaceuticals Limited;

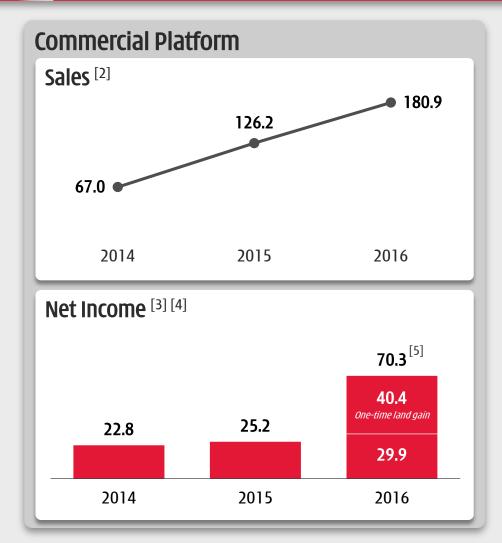
<sup>[4]</sup> Non-cash accretion relates to Mitsui's share in Innovation Platform, which was exchanged for Chi-Med shares in July 2015; [5] Including adjustment for accretion on redeemable non-controlling interests.

# Financial performance of main platforms



Sustainable biotech business model - \$170 million available cash[1]





(US\$ millions)

# Sufficient cash to fund pipeline well into 2019



Nasdaq listing, new bank facilities, land compensation & subsidies

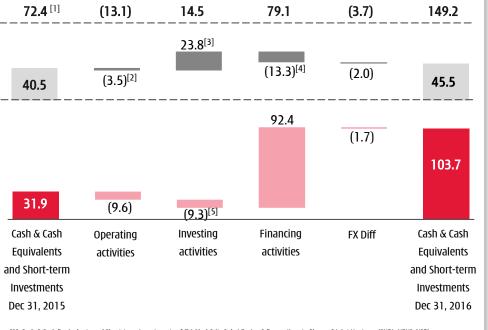
### **Chi-Med Group-level Cash Position:**

- 173.7 million available cash resources as at December 31, 2016 (Dec 31, 2015: \$38.8m).
  - **√** \$103.7m cash & cash equivalents & short-term investments<sup>[9]</sup> raised \$95.9m (net of costs) on Nasdaq in Mar 2016.
  - **◆ \$70m in unutilized banking facilities** from BAML, DB & HSBC held as at December 31, 2016 \$40m of which expired in Feb 2017<sup>[10]</sup>.
  - ✓ New \$70.0m bank facilities (unutilized) Set up new \$70.0m unsecured 18 month facilities with BAML/DB in Feb 2017.
- \$46.8 million in bank borrowings as at December 31, 2016 (December 31, 2015: \$49.8m).

#### JV-level Cash Position:

- \$91.0 million available cash as at December 31, 2016 (December 31, 2015: \$80.9m).
  - ✓ JVs have no bank borrowings.
  - √ ~\$72m cash from land compensation & subsidies received in 2016<sup>[11]</sup> ~\$40m dividend to Chi-Med Group level in H1 2017.

- Cash flow of Proportionate Share of Joint Ventures (SHPL<sup>[6]</sup>, HBYS<sup>[7]</sup>, NSP<sup>[8]</sup>).
- Proportionate Share of Cash & Cash Equivalents and Short-term Investments of Joint Ventures (SHPL, HBYS, NSP).
- Cash flow of Chi-Med & its Subsidiaries under Equity Accounting.
- Cash & Cash Equivalents and Short-term Investments of Chi-Med & its Subsidiaries.

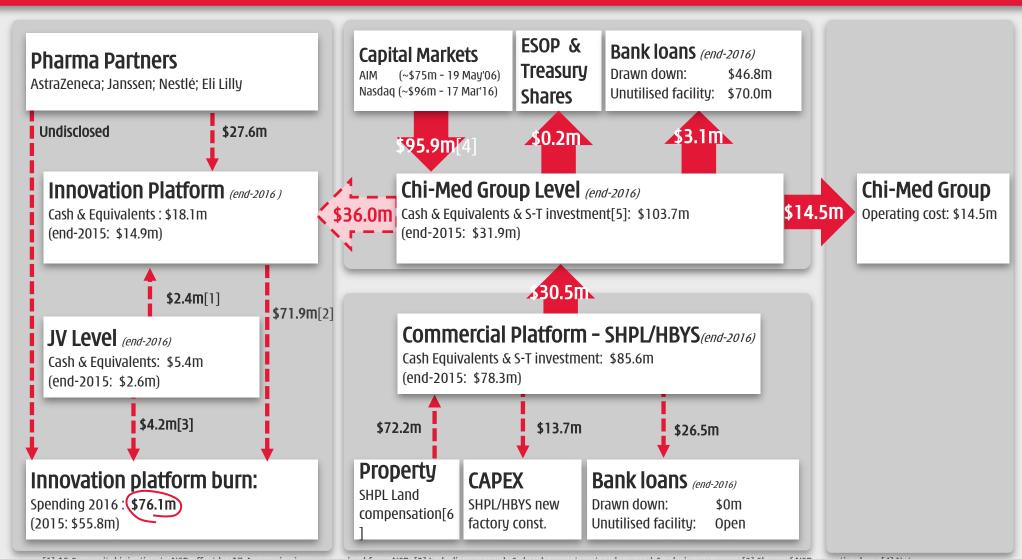


- [1] Cash & Cash Equivalents and Short-term Investments of Chi-Med & its Subsidiaries & Proportionate Share of Joint Ventures (SHPL, HBYS, NSP).
- [2] \$27.0m proportionate share of cash generated from operating activities less \$30.5m adjustment of dividend received in consolidation level.
- [3] \$0.1m proportionate share of cash used in investing activities offset with \$5.0m adjustment of capital injection to NSP in consolidation level and \$18.9m adjustment of net proceeds from Short-term Investments.
- [4] \$38.8m proportionate share of cash used in financing activities offset with a net total of \$25.5m adjustments of dividend received and NSP capital injection mentioned in items [2] and [3].
- [5] \$33.6m of cash used in investing activities offset with \$24.3m adjustment of net deposit in Short-term Investmen

# Inter-group cash flow



~\$103.7m in cash available (end-2016); \$70m in undrawn bank facilities



[1] \$5.0m capital injection to NSP offset by \$7.4m service income received from NSP; [2] Including research & development cost and general & admin. expenses; [3] Share of NSP operating loss; [4] Net proceeds: Gross proceeds deducted underwriting discounts and commissions, and other offering expenses; [5] Including \$24.3m short-term investment (over 3-month deposit) as at end of 2016; [6] Included cash received for SHPL land compensation and government subsidies in 2016.

# Risk-balanced pipeline & strategy



### **FIRST**

issues on high potential but difficult targets.

- Fix compound-related issues of failed first movers c-Met (renal tox.) & Syk (selectivity).
- Difficult novel kinase targets with deep body of evidence - FGFR (patient selection).
- Take fast action while others stuck in debate.
- Deep & DIVERSIFIED clinical pipeline.

### **BEST**

use world-class chemistry to design differentiated 2<sup>nd</sup> generation TKIs.

- No target related risk **VEGFR**, **EGFR** & **PI3Kδ**.
- Create 2<sup>nd</sup> generation TKIs w/ high selectivity & superior pharmacokinetic properties.
- A lot of room to optimize 1<sup>st</sup> generation TKIs tolerability, safety, efficacy.
- MULTIPLE fully funded pivotal studies - Not a binary proposition.

### **STRENGTHS**

Lower costs, huge team, & low-risk /fast clinical

- <u>leveraging China's</u> advantages.
- Large China patient population enables rapid
   lower risk development to proof-of-concept.
- Can afford to run ~330-person scientific team to create/manage diversified 8 asset portfolio.
- Practical, minimally dilutive, finance.

 SOLID CASH flow from Commercial Platform & global partners.

# Three collaborations have major aggregate financial impact









### ~\$1.2 billion in Partner payments to HMP/NSP<sup>[1]</sup>:

- \$118.5 million in upfront /milestone payments and equity injections as at December 31, 2016.
- **up to \$350 million** in further development and approvals milestones
- up to \$145 million in option payments.
- up to \$560 million in commercial milestones.
- customary tiered royalties on net sales.

### Clinical trial spending[2]:

- clinical costs for partnered drug candidates estimated at several hundred million US dollars.
- Partners to fund the vast majority of these clinical costs.

### Possible payment events in early 2017:

Savolitinib (AZD6094): Phase III initiation PRCC[3]

<sup>[1]</sup> Nutrition Science Partners Limited ("NSP") is the 50/50 joint venture between Nestlé Health Science ("Nestlé") and Chi-Med; [2] includes clinical and direct non-clinical costs.

<sup>[3]</sup> PRCC = papillary renal cell carcinoma.

### Sulfatinib - global potential



### Current approved treatments for NET remain somewhat limited

	S	omatostatin Based The	rapies		Kinase Inhibitor Therapi	<u>es</u>
	Sandostatin®	Somatuline Depot®	Lutathera®	Afinitor® (everolimus)	Sutent® (sunitinib)	Sulfatinib
Mechanism of Action	(octreotide) Somatostatin analogue	(lanreotide) Somatostatin analogue	( <sup>177</sup> Lu-Dotatate) <sup>[3]</sup> Somatostatin receptor targeting radiotherapy	mTOR inhibition	Inhibits multiple receptor tyrosine kinases	VEGFR/FGFR1 & CSF-1R inhibition
Mode of administration	Deep subcutaneous or Deep subcutaneous Suintravenous injection injection		Subcutaneous injection or intravenous injection	Oral tablet	Oral capsules	Oral tablet
Shelf-life	3 years	2 years	3 days (½ life)	3 years	3 years	
Primary Tumor Site						
Pancreas (6% NET)	×	*	×	$\checkmark$	$\checkmark$	$\checkmark$
Entire GI tract (67% NET)	*	$\checkmark$	×	$\checkmark$	×	$\checkmark$
with Mid-gut (20% NET)	$\checkmark$	√ (Ki67<10%)	$\checkmark$	$\checkmark$	×	$\checkmark$
Lung & Thymus (27% NET)	×	×	×	$\checkmark$	×	$\checkmark$
Other	×	×	×	*	×	$\checkmark$
	Sandostatin®/ Placebo	Somatuline Depot® / Placebo	Lutathera <sup>[4]</sup> / Sandostatin LAR 30mg	Afinitor® / Placebo	Sutent® / Placebo	Sulfatinib <sup>[2]</sup> (Ph.II ITT pop. N=81)
Median PFS (months)	14.3/6.0	NR / 18.0	Est. ~40.0 / 8.4 (mid-gut)	11.0 / 4.6 (pancreatic) 11.0 / 3.9 (lung & GI)	11.4 / 5.5	(19.4) (pancreatic) 13 <u>.6</u> (All non-pancreatic)
Hazard Ratio	0.34	0.47	0.21 (mid-gut)	0.35 (pancreatic) 0.48 (lung & GI)	0.42	
(p-value)	0.000072	⟨0.001	<i>&lt;0.001</i>	<0.001 (pancreatic) <0.001 (lung & GI)	⟨0.001	
Objective Response Rate [1]	2% / 2%	NR	18% / 3% (mid-gut)	5% / 2% (pancreatic) 2% / 1% (lung & GI)	9% / 0%	17.1% (pancreatic) 15.0% (All non-pancreatic)
Disease Control Rate [2]	69% / 40%	NR	95% / 76% (mid-gut)	73% / 51% (pancreatic) 81% / 64% (lung & GI)	72% / 60%	90.2% (pancreatic) 92.5% (All non-pancreatic)

[1] ORR = percent of patients with >30% tumor diameter shrinkage; [2] Sulfatinib Phase I: Intent to Treat ITT population = 21; patients evaluable for efficacy = 18; 3 patients withdrawn/lost to follow-up/AE); [3] DCR = percent of patients with tumor diameter growth <20%; [4] FDA action date December 28, 2016.

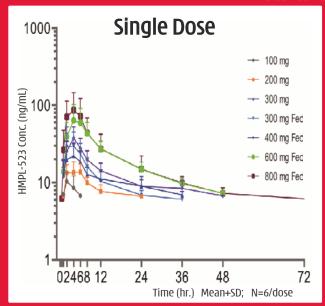
### HMPL-523 - Pharmacokinetic profile

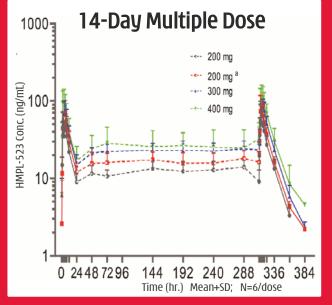


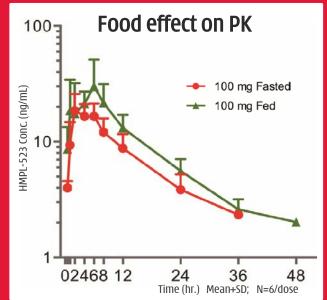
### Phase II dose of 300mg or less, once daily, for autoimmune disease

- A dose proportional increase of plasma exposure of HMPL-523 was observed.
- Exposure to HMPL-523 was increased 1.5 times when dosed in a fed condition with high-fat food. The elevated exposure could be a result of an increase in relative bioavailability.
- Preclinical models on HMPL-523 indicated a 10x drug exposure in tissue versus plasma.
- Of the 3 metabolites (M1, M2 and M3), only M1 reached plasma levels that could be characterised. The accumulation of M1 appeared greater over 14-day daily administration of HMPL-523 than that of the parent compound leading to 3 month toxicology study on the M1 metabolite which is expected to complete in H1 2017.

### **HMPL-523 CONCENTRATION-TIME PROFILE**



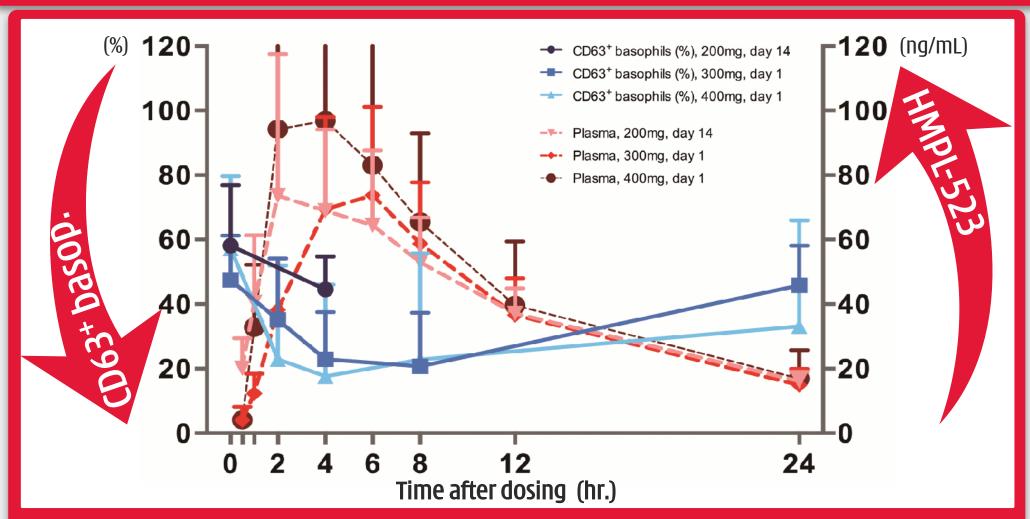




# HMPL-523 - Pharmacodynamic profile



Clear dose dependent inhibition of B-cell activation by HMPL-523



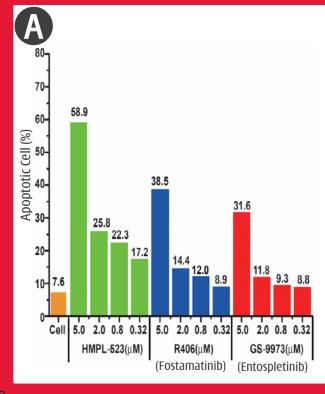
■ The EC<sub>50</sub> of HMPL-523 on the inhibition of anti-IgE-induced CD63+ expression in basophil was estimated to be 47.70 ng/mL

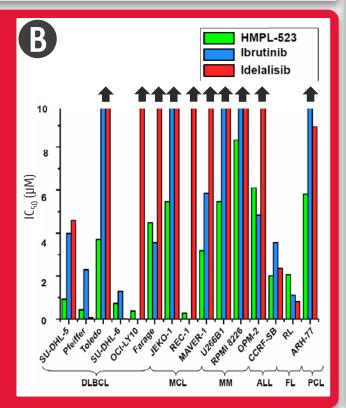
## HMPL-523 - hematological malignancies

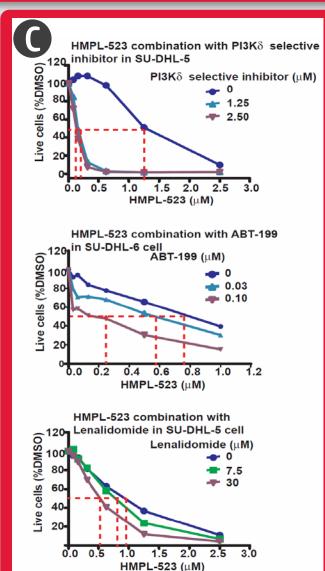


Pre-clinical superiority vs. both BTK/PI3Kδ TKIs as well as GS-9973 [1]

- A Syk inhibitors all showed a dose dependent increase in apoptotic rate (cell death) in REC-1 cells with HMPL-523 efficacy stand-out.
- B HMPL-523 inhibited cells survival in panel of human lymphoma & leukemia cells standout efficacy vs. ibrutinib (BTK) & idelalisib (PI3Kδ) inhibitors.
- Combination of HMPL-523 with other drugs (PI3Kδ TKI; ABT-199; Lenalidomide) promote cell killing in DLBCL through inducing apoptosis.



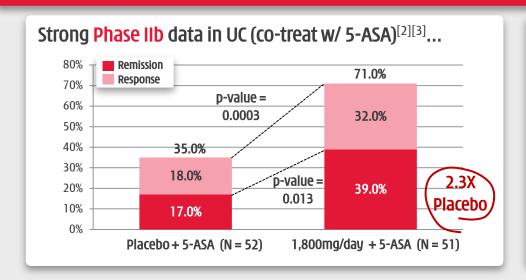


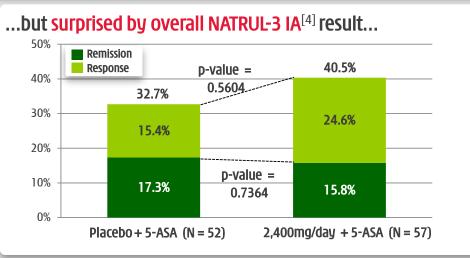


### HMPL-004 – Heavy pill burden/compliance issues

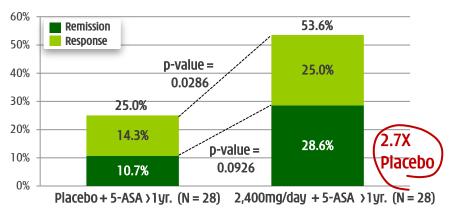


Reformulation - HM0046599 (>70% active) vs. HMPL-004 (~15% active)

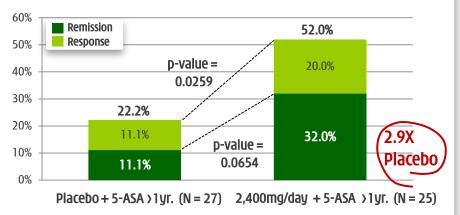








### ...particularly if difficult to treat patients stratified.



<sup>[1]</sup> Post-hoc analysis of IA: sub-group base sizes in these analyses are small and should be viewed for general indication purposes only; [2] UC = Ulcerative colitis;

<sup>[3] 1,800</sup>mg/day HMPL-004 plus Mesalamine (5-ASA) versus Mesalamine (5-ASA) alone (Placebo-arm); [4] IA = Phase III Interim Analysis conducted at ~1/3rd patient enrolment.

# Innovation Platform proxy peer group (1/2)



### HMP - A very deep pipeline and a very large organization/operation

		Mkt Cap		Ent.		201	7		Overview of pipeline assets						# of studies		
Name	7Mar'17	7Mar'16	7Mar'15	Value	Staff	Sales	EBITDA	Drug	Studies	Phase	Partner	drugs	Р3	POC	P		
Genmab	11,774	7,214	4,522	11,213	205	340	202	Ofatumumab	CLL, follicular lymphoma	Mktd, P3	Novartis	12	3	8			
								Ofatumumab (subcutaneous)	Relapsing remitting multiple sclerosis	P3	Novartis						
								Daratumumab	Double-refractory MM, relapsed & frontline MM, NHL, natura	Mktd, Reg., P3, 2x P2	,Janssen						
									killer / t-cell lymphoma, solid tumors	P1/2							
								Tisotumab vedotin	Solid cancers	P1/2	Seattle Genetics						
								HuMax-AXL-ADC	Solid cancers	P1/2	Seattle Genetics						
								AMG 714	Celiac disease	P2	Amgen						
								Teprotumumab	Graves' orbitopathy, diabetic macular edema	P2, P1	River Vision						
								HuMax-IL8, HuMax-TAC-ADC, JNJ-	Metastatic solid tumors, lymphoma, acute myeloid leukemia	, P1b, 4x P1	ADC, Bristol-Myers						
									8NSCLC, autoimmune disorder, acute myeloid leukemia		Squibb, Janssen						
Tesaro	9,499	1,842	2,177	8,845	446	96	(473)	Rolapitant IV (oral: Varubi)	CINV (oral and IV)	Mktd, Reg.	Opko	4	2	3			
								Niraparib	Ovarian maint., germline BRCAm+ breast, ovarian treat.	Reg., 2x P3, P2	Merck						
								Niraparib + Keytruda	Triple-negative breast cancer or ovarian cancer	P2	Merck						
								Niraparib + bevacizumab	Platinum-sensitive ovarian cancer (AVANOVA study)	P2	ENGOT						
								Niraparib + chemo, TSR-042 (PD-	Ewing's sarcoma, various tumor types	3x P1	AnaptysBio, SARC						
								mAb), TSR-022 (TIM-3 mAb)									
Exelixis	6,469	986	596	6,238	115	319	26	Cabometyx / Cometriq	Medullary thyroid cancer, adv. renal CC, adv. hepatocellular	Mktd, P3, 8xP2, 2xP	Ipsen	6	2	19			
								(Cabozantinib)	carcinoma, NSCLC, genitourinary tumors, & other								
								CS-3150	Hypertension	P3 (Japan)	Daiichi-Sankyo						
								Cobimetinib	CRC. NSCLC. melanoma. TNBC	P2, 3xP1b/2, P1b	Genentech						
								SAR245408	Adv. or recurr. endometrial cancer, ER/PR+ HER2- breast, lym	. P2, P1/2	Sanofi						
								SAR245409	NHL, glioblastoma, lymphoma, leukemia	P2. 3xP1b/2	Sanofi						
								XL888	Solid tumors	P1b, P1	-						
Galapagos	3,376	1,733	664	2,335	510	132	(71)	Filgotinib	RA, Crohn's (CD) , ulcerative colitis, small bowel CD	3xP3, P2	Gilead	7	3	4	- 3		
								GLPG1837	Cystic fibrosis	P2	AbbVie						
								GLPG1690	Idiopathic pulmonary disease	P2	-						
								GLPG2222	Cystic fibrosis	P2	AbbVie						
								GLPG1972, MOR106, GLPG2737	Osteoarthritis, inflammation, cystic fibrosis	3xP1	Servier, Morphosys						
Clovis	2,680	863	2,694	2,695	278	64	(222)	Rucaparib	Cancers: Ovarian treat./maint., prostate, triple negative	Approved, 3xP3, 6x		1	3	6			
									breast, breast, gastro esophageal, gynecological	P2, P1							
Juno	2,205	4,493	4,778	1,473	518	62	(332)	JCAR015	Acute lymphoblastic leukemia, NHL	P2	-	10	0	3	5		
								JCAR017	Pediatric acute lymphoblastic leukemia, adult NHL	P1	-						
								JCAR014	Chronic / acute lymphocytic leukemia, NHL	P1	-						
								JTCR016	AML, MDS, CML, NSCLC / mesothelioma	2xP1/2	-						
								JCAR018, BCMA, JCAR023, JCAR020	), Pediatric ALL / NHL, MM, pediatric neuroblastoma, ovarian,	6xP1	-						
								JCAR024, Lewis Y	NSCLC / breast, lung								

# Innovation Platform proxy peer group (2/2)



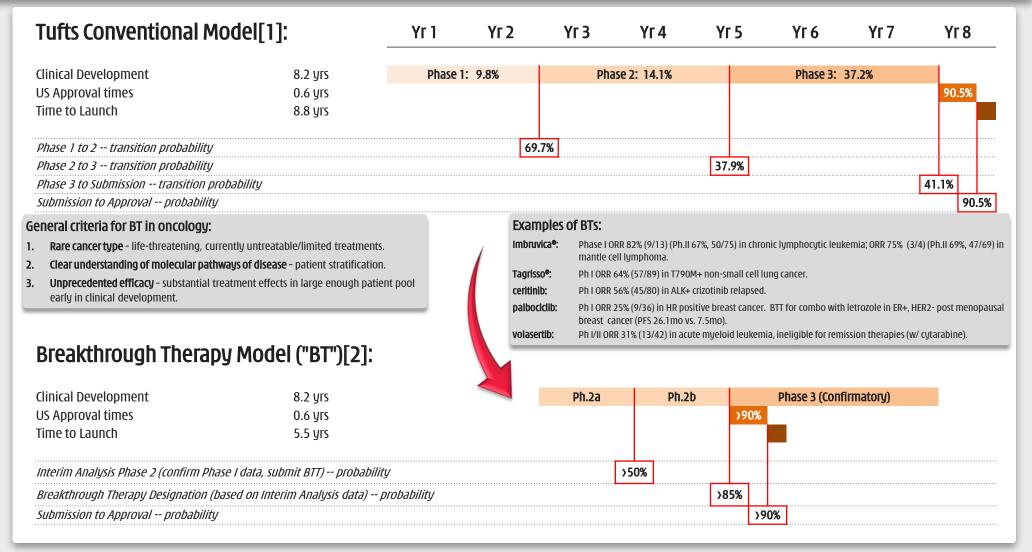
### HMP - A very deep pipeline and a very large organization/operation

		Mkt Cap		Ent.		20	17		Overview of pipeline assets (a)			# of	#	of stud	ies
Name	7Mar'17	7Mar'16	7Mar'15	Value	Staff	Sales	EBITDA	Drug	Studies	Phase	Partner	drugs	Р3	POC	P1
Agios	2,094	1,806	3,889	1,552	287	44	(299)	Enasidenib (AG-221)	R/R AML, frontline AML	P3, 2xP1/2, P1b	Celgene	4	3	6	3
								Ivosidenib (AG-120)	Frontline AML, R/R AML, solid tumors, cholangiocarcinom		-				
								AG-348	PK deficiency	P2	-				
								AG-881	Solid tumors	P1	Celgene				
Array	1,965	453	1,126	1,879	177	155	(76)	Binimetinib / MEK162	Melanoma, CRC	P3	-	7	2	3	2
								Encorafenib / LGX818	Melanoma, CRC	P3	-				
								Filanesib / ARRY-520	Multiple myeloma	P2	-				
								ARRY-797	Lamin A/C-related dilated cardiomyopathy	P2	-				
								ARRY-502	Asthma	P2	-				
								ARRY-382, ARRY-614	Solid tumors, myelodysplastic syndromes	2xP1	-				
Morphosys	1,856	1.203	2,139	1.699	278	72	(43)	MOR 208	CLL or small lymphocytic lym., diffuse large B-cell lym.	4x P2	-	3	0	5	1
	,	,		·			( )	MOR202	Multiple myeloma	P2	-				
								MOR107	Undisclosed	P1	-				
BeiGene	1,532	939	NA	1,347	318	6	(112)	BGB-3111; BGB-3111 + Ibrutinib	Waldenstrom's macro., relapsed or refractory MCL	P3, P2	-	4	1	7	1
								BGB-A317, -A317 + BGB-290, -	Advanced cancers, b-cell malignancies,	P1A/1B, 3xP1B, 2xP1A,	-				
								A317 + -3111, -290, -3111, BGB-	relapsed/refractory b-cell malignancies, b-cell lymphoid	P1					
								3111 + Obinutuzumab, BGB-283	malignancies						
Puma	1,315	1,729	7,432	1,086	156	24	(307)	Neratinib (PB272)	Adjuvant breast cancer, neoadjuvant BC, metastatic BC, metastatic BC, her2 BC metastatic	NDA,MAA, 2xP3, 8x P2	-	1	2	8	0
Ziopharm	842	1,250	1,535	886	36	7	(62)	Ad-RTS-IL-12 + veledimex	Locally adv. or met. breast can., recurrent or progressive	P2, P1b/2, P1	Intrexon	2	0	2	2
									GBM, pediatric brain tumor						
								CAR / cytokine product	Leukemia/lymphoma, AML	P1	Intrexon, MD Anders.				
Aduro	724	1,071	NA	377	143	31	(106)	CRS-207	Mesothelioma, ovarian cancer, pancreatic cancer	P2, P1b, P1/2	Incyte	4	0	3	3
								ADU-741, ADU-214, ADU-S100	Prostate cancer, lung cancer, multiple tumors	3xP1	Janssen, Novartis				
AVERAGE (13)	3,564	1,968	2,868									5	1	6	2
MEDIAN (13)	2,094	1,250	2,177									4	2	5	2
Chi-Med Inno	vation Pla	atform,			330	35-40 (	45)-(55)	Savolitinib	PRCC, CCRCC, NSCLC, gastric cancer	P3, P2b, 5xP2, 7xP1b	AstraZeneca	8	4	19	7
Hutchison Me	diPharma	a (HMP)				Ì	. , , ,	Fruquintinib	PRCC, CCRCC, NSCLC, gastric cancer Colorectal cancer, NSCLC, gastric cancer	2x P3, P1b. P1	Eli Lilly				
		` ′						Sulfatinib	NET. US bridging, thyroid cancer, biliary tract cancer	2x P3, 4xP2, P1	-				
								Epitinib	NSCLC, glioblastoma	P3. P2	-				
								Theliatinib	Solid tumors, esophageal cancer	P1b, P1	-				
								HMPL-523	RA. hematological cancers. immunology. lymphoma	4xP1	-				
								HMPL-689	Hematological cancers, lymphoma	2xP1	-				
								= 207	······································						

# Breakthrough Therapy Model







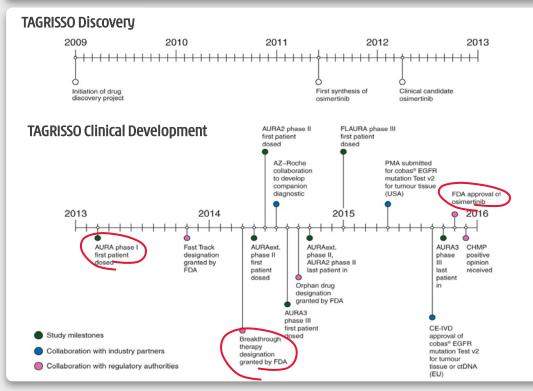
<sup>[1]</sup> Tufts Center for the Study of Drug Development (Feb 2010) - Transition probabilities for small molecule oncology drugs based on data of the 50 largest pharmaceutical companies 1993 through June 2009; [2] Hypothetical probabilities for BT estimated by Chi-Med - for general reference only, probabilities will vary dramatically based on scale/quality of Phase I data.

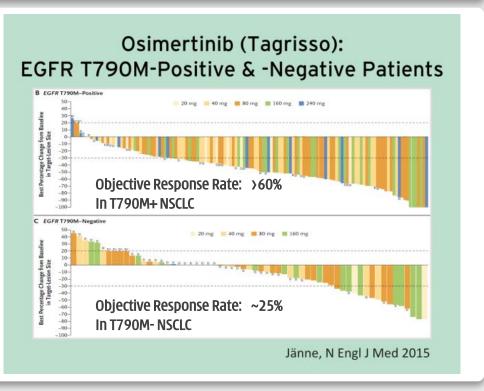
# AstraZeneca's Tagrisso®



Fastest U.S. FDA drug approval – just 2 yrs. 8 mo.

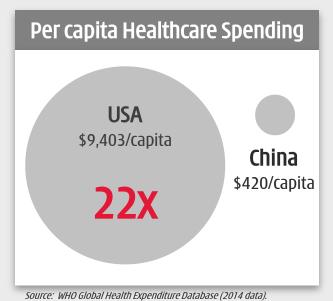
- Savolitinib has reported 55% Objective Response Rate (6/11 pts.) to-date in second line NSCLC (TATTON) if Phase IIb study re-affirms this we could follow the same accelerated approvals path taken by Tagrisso.
- Phase IIb study to complete in 2017 with ORR similar to TATTON we could target:
  - ✓ Potential Breakthrough Therapy designation application in 2017/2018.
  - ✓ Savolitinib submission for approval in 2018 and potential US FDA approval in 2019.

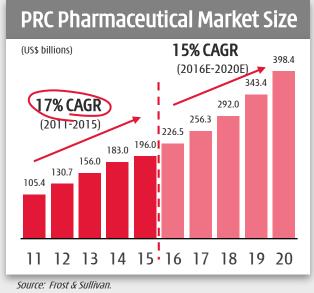


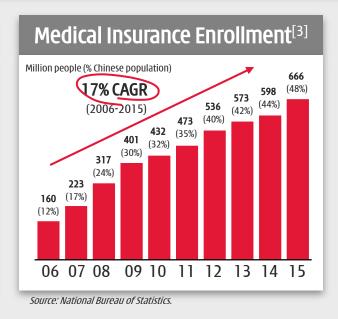


# China pharma market set to become the second largest globally in 2016/2017









- China pharmaceutical industry growth 17% CAGR<sup>[1]</sup> from 2011-2015 one of the highest rated industries in China with average P/E ratio of 42 for the 61 listed companies (next slide).
- Government healthcare spending grew 14% CAGR<sup>[2]</sup> from 2011 2015 and continues to increase rapidly Strategic priority.
- Expansion of State Medical Insurance Schemes<sup>[3]</sup> Link to increased drug reimbursement & sales.

# Targeted therapies – fastest growth & largest[1]



Pricing beyond reach of the 3.4 million new cancer patients/year in China

Global Oncology drug market<sup>[1]</sup>: \$112 billion

> China Oncology Market<sup>[2]</sup>: \$13 billion

China Pharmaceutical Market<sup>[3]</sup>: \$196 billion

Source: Frost & Sullivan; [1] 2015 global oncology market at ex-factory
price level: [3] 2015 china opcology market at wholesale price level:

	% Oncology Market <sup>[4]</sup>	Sub-Category	Share of Sub- category <sup>[4]</sup>	Product	Company	Est. Market Sales (\$m) <sup>[4]</sup>	Approx. patient cost/month (\$) <sup>[4]</sup>	12 mo. treatment (Est. # patients) <sup>[4]</sup>
Ī	20.9%	Targeted Therapies	19.3%	rituximab	Roche	443	16,780	2,200
			15.0%	trastuzumab	Roche	344	5,130	5,592
			14.2%	imatinib	Novartis	326	6,323	4,295
			8.5%	bevacizumab	Roche	195	6,251	2,601
			7.4%	erlotinib	Roche	170	3,108	4,554
			6.8%	gefitinib	AstraZeneca	156	2,730	4,764
			5.3%	cetuximab	BMS/BI	122	14,146	717
			4.6%	sorafenib	Bayer	106	8,329	1,056
			4.0%	bortezomib	Janssen	92	8,133	941
			14.9%	Other		342		
				Total Targeted 1		2,295		26,718
	20.4%	Anti-metabolites	29.1%	pemextred	Lilly/Hansoh	652		
			21.5%	capecitabine	Roche	482		
			20.4%	TS-1	Taiho/Qilu	457		
			16.6%	gemcitabine	Lilly/Hansoh	372		
			12.4%	Other		278		
				Total Anti-Meta		2,240		
	19.7%	Plant Alkaloids	49.3%	paclitaxel	BMS/Luye	1066		
			42.4%	docetaxel	Sanofi/Hengrui	916		
			8.4%	Other		181		
				Total Plant Alka		2,163		
	10.5%	DNA Damaging	46.5%	oxaplatin	Sanofi/Hengrui	546		
		agents	21.3%	temzolomide	Merck/Tasly	250		
			13.1%	nedaplatin		154		
			4.3%	carboplatin		51		
			14.8%	Other		174		
				Total DNA Dama		1,175		
	6.4%	Hormones	29.8%	letrozole	Novartis/Hengrui	209		
			23.0%	bicalutamide	AstraZeneca	162		
			19.5%	anastrozole	AstraZeneca	137		
			17.1%	exemestane	Pfizer/Qilu	120		
			10.6%	Other		74		
-				Total Hormones	5	703		

# CHI-MED

### China Commercial Platform has substantial value

- Chi-Med's Commercial Platform continues to perform well relative to our peer group.
- The market value, based on China Pharma PE multiples is approximately \$2.2-2.7 billion.<sup>[3]</sup> Given our share in the JVs, Chi-Med's share of this value is approximately \$1.0-1.3 billion.

			NET	SALES				NET INCO	ME		VALUA	ATION
	Code	2014	2015	LTM 2016 Jun	14-15 Growth	2014	2015	LTM 2016 Jun	14-15 Growth	LTM Margin	Market Cap.	P/E[2]
CHI-MED Commercial Platform Subsidiaries/JVs[1	]	465.4	518.9	560.0	11%	48.8	54.1	58.5	11%	10%	n/a	n/a
Tianjin Zhong Xin Pharma	600329	1,076.4	1,075.4	1,058.2	0%	57.6	69.5	70.7	21%	7%	1,720	30
Li Zhu Pharma	000513	842.1	1,005.5	1,105.7	19%	84.1	100.2	108.4	19%	10%	3,328	31
Shandong Dong E E Jiao		608.9	827.7	846.7	36%	208.4	248.8	257.6	19%	30%	5,281	21
Zhejiang Kang En Bai Pharma	600572	544.0	805.3	930.8	48%	110.5	76.5	47.1	-31%	5%	2,729	66
Kunming Pharma		625.8	746.6	808.5	19%	46.7	65.5	70.1	40%	9%	1,610	24
Guizhou Yi Bai Pharma	600594	479.5	501.6	522.0	5%	73.1	29.2	46.2	-60%	9%	1,976	42
Jin Ling Pharma	000919	421.0	489.3	525.3	16%	37.2	39.8	37.7	7%	7%	1,044	35
Jiangsu Kang Yuan	600557	389.3	428.4	439.6	10%	49.1	55.5	55.7	13%	13%	1,606	28
Jiang Zhong Pharma	600750	430.5	394.5	327.5	-8%	40.5	55.9	64.2	38%	20%	1,482	25
Zhuzhou Qian Jin Pharma	600479	333.3	371.6	397.2	12%	17.9	13.4	14.6	-25%	4%	801	50
Peer Group Weight Avg. (10 Comps. excl. Chi-Med		<b>575.1</b>	<b>664.6</b>	<b>696.2</b>	<b>16%</b>	<b>72.5</b>	<b>75.4</b>	<b>77.2</b>	<b>4%</b>	11%	<b>2,158</b>	<b>34</b>
All 61 Listed China Pharma. Companies Weight Av	erage	918.6	1008.3	1063.3	10%	68.4	80.4	89.1	18%	8%	2,784	42

Peer Group: 10 companies (excl. Chi-Med) selected as ALL listed and profitable mainland Chinese OTC/RX pharma manufacturing companies, with a focus on similar product types, and 2015 Net Sales in the ~\$350-1,100 million range.

<sup>[1]</sup> Total aggregate PRC domestic results of Chi-Med's 6 Commercial Platform companies (HBYS, SHPL, Hutchison Sinopharm, HHO, HHL, & HCPL), excluding discontinued operations;

<sup>[2]</sup> Price Earnings Ratio as at **January 6th, 2017**: Trailing Twelve Month PE weight averaged based on market capitalization;

<sup>[3]</sup> Peer group/China Pharma multiple of 34-42 x 2016 actual Net income after tax of \$63.3 million (excluding one-time property gain of \$80.8 million).

# SHPL old factory site surrender of land-use rights



Fully received \$113 million in cash compensation & subsidies (Feb 2017)



# 4.6 sq.km. new development zone 12km from CBD (re-zoned in 2014).

- "Smart City" new science & tech, commercial and residential area.
- SHPL old factory classified as Cat. 3 residential.



	Land Area (sq.m.)	Other Factors	Approx. Distance to CBD <sup>[1]</sup> (km)	Approx. Distance to Metro <sup>[2]</sup> (m)	Actual Compensation (US\$ million)	Compensation (\$/sq.m.)
★ SHPL Old Factory Plot	57,804	New Dev.	12.4	300	113.1	1,957
① Qing Pu Chemicals Plot	77,372	Nr. Airport	21.2	2,200	108.4	1,401
2 Shanghai Soap Factory Plot	62,846	Nr. River	8.0	500	122.6	1,951
3 Shanghai Electric (Fuels) Plot	27,091	Nr. River	11.4	2,000	89.1	3,290
4 Shen Bei Group Plot	4,976	Nr. River	3.3	300	34.5	6,928

# HBYS Plot 1&2 – 9km from Guangzhou city center



Property compensation expected in the range of ~\$120 million<sup>[2]</sup>

#### HBYS Plot 2 (26,700 sq.m. plot of land):

2.2 plot ratio, ~58,740 sq.m. of residential floor area. Estimated Auction Price[1]: \$123.4 million (\$2,100/sq.m.).



#### 163 Tong Bao Road (131,647 sq.m. plot of land):

Auction Date: November 24th 2014

~3.5 plot ratio, 460,765 sq.m. of residential floor area. Actual Auction Price: \$1,034 million (\$2,244/sq.m.).

#### 8-10 Tong Bao Road (65,055 sq.m. plot of land):

Auction Date: May 6th 2013

2.2 plot ratio, 143,121 sq.m. of residential floor area. Actual Auction Price[1]: \$305 million (\$2,132/sq.m.).

HBYS Plot 1 (59,400 sq.m. plot of land)



Tong He Metro Station (opened November 2010)

# New factories - triple capacity

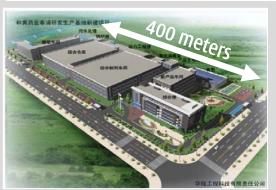
# CHI-

### JVs fund internally - \$139m of total \$142m (~98%) CAPEX spent

#### SHPL New Factory - SOP[1] Sep 2016

Feng Pu District, 78,000 sq.m. plot (~40km south of Shanghai city center). Approx. 3x designed capacity expansion (extraction & formulation).

Actual total CAPEX: \$102m











#### HBYS New Factory - SOP H1 2017

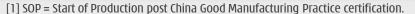
Bozhou, Anhui province (central China). 230,000 sq.m. plot. Approx. 3x extraction expansion & new formulation lines.

Estimated total CAPEX: \$40 m



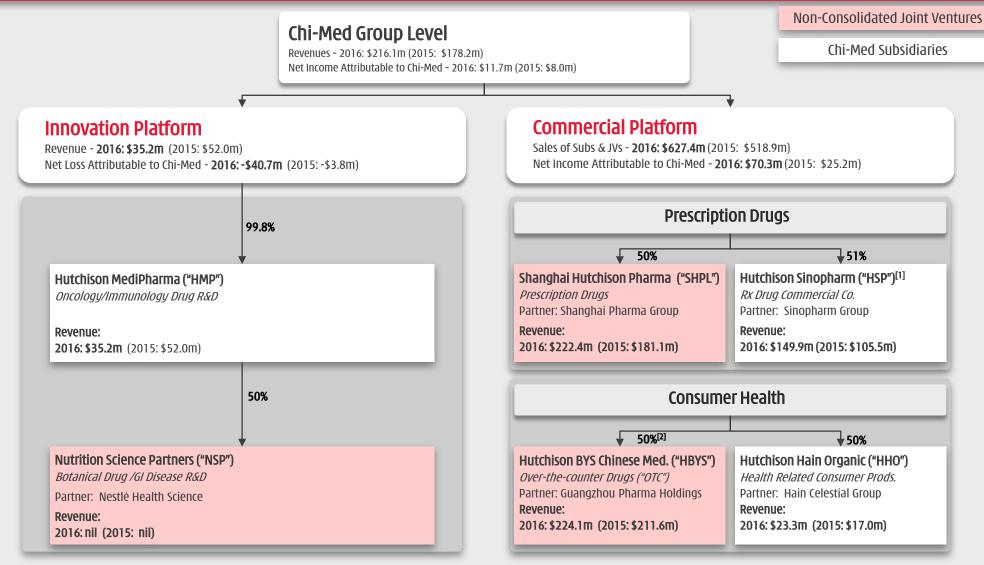






# Chi-Med Group structure - major entities





<sup>[1]</sup> Excluded HSP's ZLT business; [2] Held through an 80% owned subsidiary.



# Experienced pharma management team

POSITION	EXPERIENCE (yrs) Industry / Chi-Med	ROLE / BACKGROUND
CHRISTIAN HOGG, BSc, MBA Chief Executive Officer	Procter & Gamble 28 / 17	Led all aspects of the creation, implementation & management of Chi-Med's strategy, business & IPOs since 2000 start - incl. AZ, Lilly, Nestlé deals & est. of pharma business.
WEIGUO SU, PHD  EVP, Chief Scientific Officer	<b>Pfizer</b> 27/12	Created Chi-Med's R&D strategy, innovation platform & led all pipeline discovery; Director of Med Chem at Pfizer; Harvard Ph.D./post-doc under Nobel Laureate E. J. Corey.
JOHNNY CHENG, BEC, CA Chief Financial Officer	Bristol-Myers Squibb 27 / 8  KPING NESTIE	Former VP, Finance at BMS China; 8 years with Nestlé China heading finance & control in multiple businesses; KPMG & PWC in Australia & Beijing.
YE HUA, MD, MPH SVP, Clinical & Regulatory Affairs	NOVARTIS Celgene 18/3	Led Revlimid & Pomalyst global development in multiple myeloma; 15 yrs of global registrations incl. Humira, Zometa, Reclast, Femara, Cardioxane, Proleukin.
ZHENPING WU, PHD, MBA SVP, Pharmaceutical Sciences	Roche Pfizer 23/9	Leads all CMC development & manufacturing for Chi-Med's pipeline; Sr Director of PS at Phenomix; Director of Pharma Development at Pfizer San Diego; at Roche in Palo Alto.
MAY WANG, PHD SVP, Bus. Dev. & Strategic Alliances	Lilly 22/6	Leads alliance mgmt & BD for Chi-Med; long career in research, primarily biology, strategic alliance management, partnering & business development with Eli Lilly.
MARK LEE, BEng, MBA SVP, Corp. Finance & Development	CREDIT SUISSE 18/8	Focuses on strategic management, overall corporate operations & alliance support; Former US/UK banker advising & raising capital for major pharma & biotech.

- Management team comprised mainly of returnees averaging ~20 years in multinational pharma & biotech.
- Scientific leadership have participated in the discovery & development of global blockbusters.













