

### HUTCHISON CHINA MEDITECH

### 2017 Full Year Results

AIM/Nasdaq: HCM

March 12, 2018

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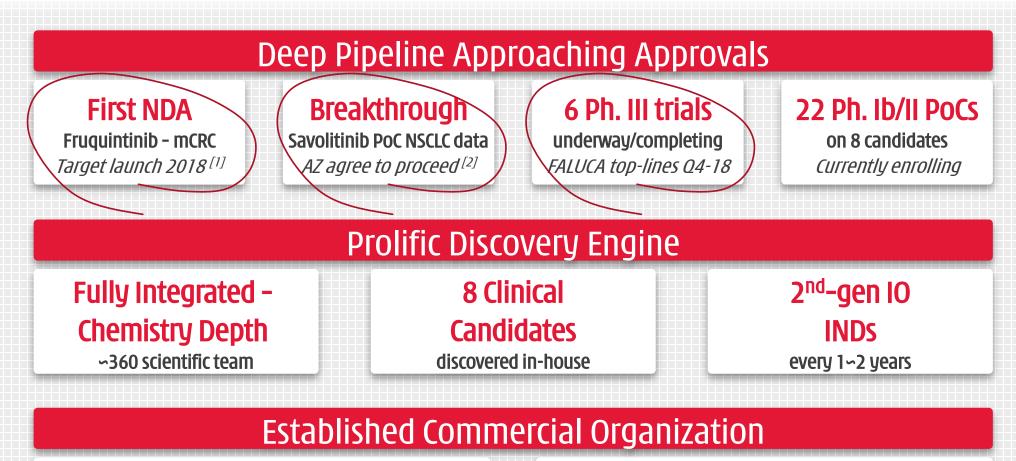
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### Chi-Med Highlights Momentum continues to build





### Pan-China Sales & Marketing

~2,300 medical reps

**Product Launch Ready** 

proven success in new indications

[1] Subject to China FDA approval; [2] Target to initiate global randomized chemo-doublet controlled study for savolitinib/Tagrisso® and multiple supporting studies in 2018; mCRC = metastatic colorectal cancer; AZ = AstraZeneca; FALUCA = Ph.III of fruguintinib monotherapy in 3<sup>rd</sup>-line non-small cell lung cancer; PoC = Phase Ib/II proof of concept study; IO = immuno-oncology

### 2017 Financial Results In line with guidance <sup>[1]</sup>

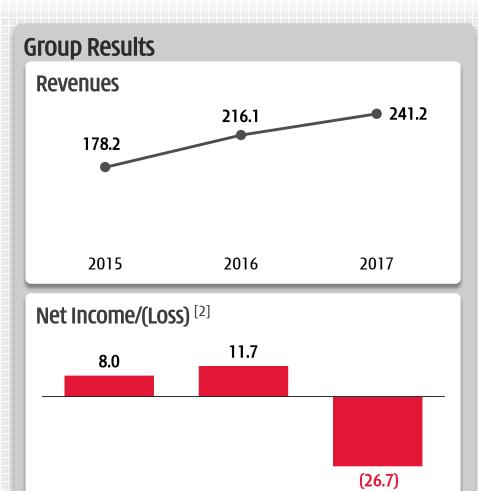


# Financial Summary

GROUP CONSOLIDATED REVENUES	178.2	216.1	241.2	225 - 240
Unconsolidated JV Revenues	392.7	446.5	472.0	480 - 500 [6]
NET (LOSS)/INCOME <sup>[2]</sup>				
INNOVATION PLATFORM	(3.8)	(40.7)	(51.9)	(45.0) - (55.0)
Base HMP Operations	(0.0)	(36.5)	(47.4)	
50% share of Nestle JV (NSP) <sup>[3]</sup>	(3.8)	(4.2)	(4.5)	
COMMERCIAL PLATFORM	25.2	29.9	37.5	32.0 - 34.0
Prescription Drugs Business	15.9	20.7	26.5	
Consumer Health Business	9.3	9.2	11.0	
GROUP COSTS	(13.4)	(17.9)	(14.8)	(18.0) - (19.0)
<b>GROUP COSTS</b> Administrative Expenses	<b>(13.4)</b> <i>(10.9)</i>	<b>(17.9)</b> <i>(12.6)</i>	<b>(14.8)</b> <i>(11.3)</i>	(18.0) - (19.0)
				(18.0) - (19.0)
Administrative Expenses	(10.9)	(12.6)	(11.3)	<b>(18.0) - (19.0)</b> 3.0 - 16.0
Administrative Expenses Interest/Tax	(10.9)	(12.6) (5.3)	(11.3) (3.5)	
Administrative Expenses Interest/Tax Land Compensation & Subsidies	(10.9) (2.5)	(12.6) (5.3) <b>40.4</b>	(11.3) (3.5) <b>2.5</b>	3.0 - 16.0
Administrative Expenses Interest/Tax Land Compensation & Subsidies Net Income/(Loss) Attrib. to Chi-Med	(10.9) (2.5) - 8.0	(12.6) (5.3) <b>40.4</b>	(11.3) (3.5) <b>2.5</b>	3.0 - 16.0

2016

2017



2016

2015

[1] Company guidance (July 31, 2017); [2] Net Income/(Loss) = Net Income/(Loss) attributable to Chi-Med; [3] NSP = Nutrition Science Partners Limited; [4] Non-cash accretion relates to Mitsui's share in Innovation Platform, which was exchanged for Chi-Med shares in July 2015; [5] Includes adjustment for accretion on redeemable non-controlling interests; [6] September 2017 divestment of HBYS subsidiary (Guanbao) - 2017 revenue \$38.6m (2016: \$45.0m), 2018 fully eliminated.

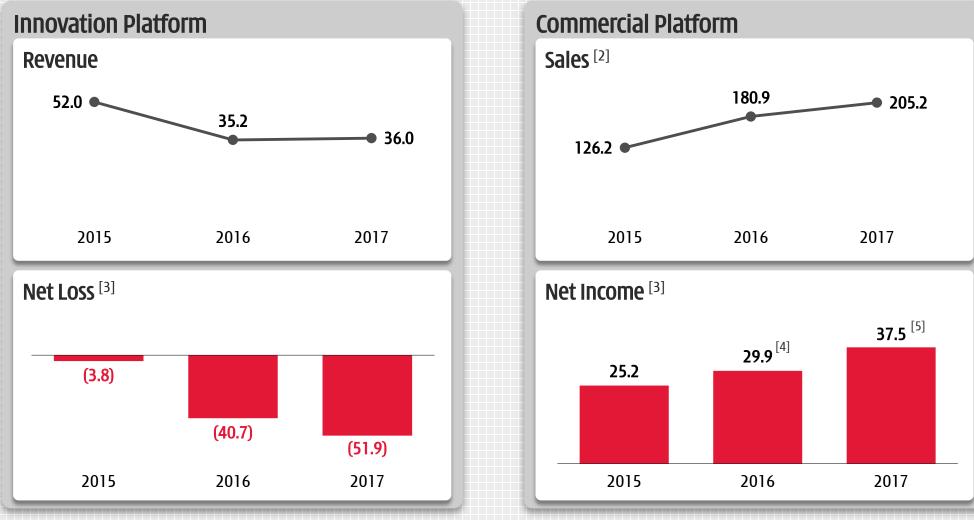
2017

Guidance

*(US\$ millions, except per share data)* 

### Financial performance of main platforms Sustainable model - \$88 million in innovation investment [1]





[1] Represents adjusted R&D expenses (non-GAAP). GAAP R&D expenses were \$75.5m in 2017 (2016: \$66.9m) – please see appendix "Non-GAAP Financial Measures and Reconciliation"; [2] Only includes sales of subsidiaries for Prescription Drugs and Consumer Health businesses – excludes joint ventures; [3] Adjusted Net Income/(Loss) = Adjusted Net Income/(Loss) attributable to Chi-Med (non-GAAP); [4] Excludes the share of one-time gain from SHPL land compensation of US\$40.4 million; [5] Excludes the share of a one-time gain from SHPL's R&D related subsidies of US\$2.5 million.



### Summary Balance Sheet & 2018 Guidance Strengthened cash position – supports higher innovation spend



(US\$ millions)

- 1. Chi-Med Group-level Cash Position
- **\$479.6 million available cash as at Dec 31, 2017** (Dec 31, 2016: \$173.7m).
  - ✓ \$358.3m cash & cash equivalents and short-term investments <sup>[1]</sup>
  - ✓ \$121.3m unutilized banking facilities from Scotiabank, BAML, DB & HSBC <sup>[2]</sup> held as at Dec 31, 2017.
- \$30.0 million in bank borrowings as at Dec 31, 2017 (Dec 31, 2016: \$46.8m). Weighted avg. total cost of borrowing on outstanding loan 2.7% (2016: 2.5%).
- \$292.7 million from follow-on offering in Oct 2017
- 2. JV-level Cash Position
  - \$67.0 million available cash as at Dec 31, 2017 (Dec 31, 2016: \$91.0m).
    - ✓ \$55.6m dividend to Chi-Med Group level in 2017.

#### 3. 2018 Guidance

- Innovation Platform Revenue up from potential fruquintinib launch/milestones. R&D expense up - expansion of clinical trials.
- Commercial Platform Revenue lower due to CFDA Two-Invoice policy & Guanbao divestment – no impact on Net Income growth.

Revenues	2017 Actual \$241.2	2018 Guidance \$155 - \$175
Innovation Platform		
Revenue Adjusted R&D expenses (non-GAAP) <sup>[3]</sup>	36.0 (88.0)	<u>40 - 50</u> (110) - <u>(120)</u>
Commercial Platform		
Sales (consolidated)	205.2	115 - 125 <sup>[4]</sup>
Sales of non-consolidated JVs	472.0	<b>460 - 480</b> <sup>[5]</sup>
Net Income		
On adjusted (non-GAAP) basis excluding one-time gains	37.5	41 - 43
One-time gains <sup>[6]</sup>	2.5	<u> </u>
Net Income	40.0	41 - 63
Chi-Med Group Costs		
Admin., interest, tax	(14.8)	(16) - (18)
Net (Loss)/Income Attributable to Chi-Med	(26.7)	(19) - (52)

[1] Short-term investments: 91-183 days deposits; [2] BAML = Bank of America Merrill Lynch, DB = Deutsche Bank, HSBC = Hong Kong Shanghai Banking Corporation; [3] R&D expenses, as adjusted (non-GAAP) excludes the actual or estimated impact of the revenue received from external customers of our Innovation Platform, which is reinvested into our clinical trials; [4] Impact of CFDA Two-Invoice policy - Hutchison Sinopharm no longer able to consolidate sales of certain third-party drugs (e.g. Seroquel®); [5] 2017 divestment of HBYS subsidiary (Guanbao) eliminates \$38.6 million in sales in 2018 vs. 2017; [6] Share of one-time gain from R&D related subsidies to SHPL in 2017 and potential land compensation from HBYS Plot 2 in 2018 guidance (dependent on Guangzhou government policy).

	nilestones for 2018 tions/clinical achievements on multiple candidates MED				
Savolitinib	<ol> <li>Initiation of global randomized, chemo-doublet controlled, study of savolitinib/ Tagrisso® combo in second-line NSCLC, along with multiple supporting studies;</li> <li>AZ decision on registration strategy for savolitinib/Tagrisso® in third-line NSCLC;</li> <li>AZ/Chi-Med decision on NSCLC pivotal study in China for savo /Iressa® combo;</li> <li>Molecular epidemiology study (n&gt;300) in PRCC <sup>[1]</sup> - possibly BTD <sup>[2]</sup> enabling.</li> </ol>				
Fruquintinib5. Potential NDA approval & launch in China in third-line CRC; ★ 6. Release top-line data for Phase III FALUCA study in third-line NSCLC. ★					
Epitinib	7. Initiate China Phase III study in first-line EGFRm NSCLC patients w/ brain mets. ★				
Sulfatinib	8. Initiate Phase IIa expansion study in NET patients in the U.S.				
HMPL-523 (Syk)	<ul> <li>9. Initiate dose expansion proof-of-concept studies in hematological cancer in both Australia &amp; China;</li> <li>10. Potential presentation of prelim. safety &amp; efficacy data from Phase I/Ib dose escalation expansion studies in hematological cancer. ★</li> </ul>				
HMPL-689 (PI3Kõ)	11. Initiate Phase Ib expansion studies in China in hematological cancer patients; 12. Present Phase I dose escalation data in Australian healthy volunteers.				

 $\frown$ 

[1] PRCC = Papillary renal cell carcinoma; [2] BTD = Breakthrough Therapy Designation.



### **Innovation Platform**

Near term: Driving for first product launches Mid-longer term: Building the pipeline for future growth

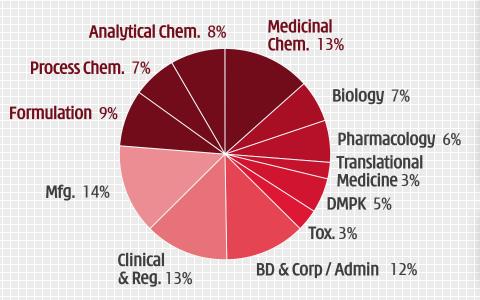


### Exceptional scale for pre-approval biotech Over 16 years with about \$500 million invested to-date



~360 SCIENTISTS & STAFF<sup>[1]</sup>

- ✓ 207 with advanced technical degrees
  ✓ 22 M.D.s
- ✓ 53 doctorate degrees



✓ 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 (4.3 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 16 years with stable financial support.

[1] Headcount as of December 31, 2017; Chem. = Chemistry; DMPK = Drug, Metabolism, & Pharmacokinetics; Tox. = Drug Safety Evaluation; Mfg = Manufacturing; Reg. = Regulatory; BD = Business Development; [2] CA Cancer J Clin 2016;66:115-132. 2016 American Cancer Society.

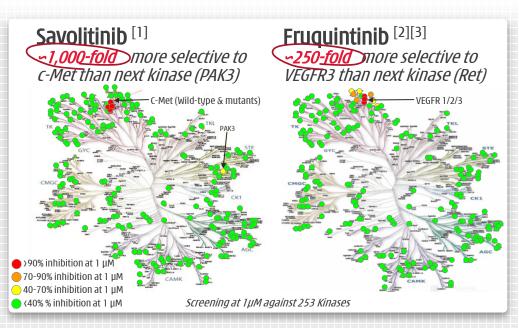
### **Chemistry is our edge** Seriously selective small molecules

CHI-(MED)

- 1. Fragment-based design of Novel Chemical Entities.
- Internally designed all 8 clinical drug candidates.
- Use of co-crystal structures.

10

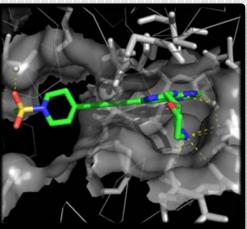
- Focus on small molecule interactions with tyrosine kinases - proteins/enzymes involved in cell signaling.
- 2. Total focus/discipline in designing and progressing drug candidates with superior kinase selectivity.
- Optimize binding to on-target protein, minimize offtarget protein binding.
- No off-target 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 Long-term use = prolonged/total target coverage = better efficacy MED

#### 3. Monotherapies – 1<sup>st</sup> generation TKIs not optimal for long-term use

- Multi-kinase TKIs major dose modifications due to off-target toxicities.
- Chi-Med's more selective TKIs designed for less dose modifications & discont.

EXAMPLES OF MONOTHERAPY APPROVED SMALL MOLECULE TKIS - targets (approval yr.)	2017 Sales	Recent Monotherapy Cancer Trial <sup>[2]</sup>	mPFS (months)	Dose Reductions	Discont. due to AEs	Total Discont- inuations
<b>Sutent<sup>®</sup> (sunitinib)</b> - VEGFR1,2,3, PDGFRβ, FLT3, CSF-1R, c-Kit, Ret (2006)	\$1.08b	1L ccRCC (CABOSUN) 1L ccRCC (COMPARZ)	5.6 9.5	49% 51%	22% 18%	38% 3 <u>3%</u>
<b>Nexavar<sup>®</sup>(sorafenib)</b> - RAF, VEGFR2, PDGFRβ, Flt3, c-Kit, FGFR1 (2005)	\$0.94b	2L RCC (AXIS)	5.7	54%	13%	23%
<b>Votrient<sup>®</sup> (pazopanib)</b> - VEGFR1,2,3, c-KIT, ITK, LCK, PDGFRα,β, FGFR1,3, c-Fms (2009)	\$0.81b	1L CCRCC (COMPARZ)	8.4	44%	23%	36%
<b>Inlyta® (axitinib)</b> - VEGFR1,2,3, PDGFRα, c-kit (2012)	\$0.34b	2L RCC (AXIS)	8.3	34%	8%	17%
Cabometyx <sup>®</sup> (cabozantinib) - AXL, c-Kit, FLT3, MET, RET, TIE-2, TrkB, VEGFR1,2,3 (2016)	\$0.35b	1L ccRCC (CABOSUN)	8.2	58%	21%	27%
Lenvima <sup>®</sup> (lenvatinib) - VEGFR1,2,3, Ret, PDGFR, c-Kit, FGFR1,2,3,4 (2015)	\$0.27b	2L ccRCC (Ph 2 reg.)	7.4	62%	25%	31%
<b>Stivarga® (regorafenib)</b> – VEGFR1,2,3, Raf, Ret, PDGFR, c-Kit (2012)	\$0.36b	≥3L CRC (CORRECT) ≥3L CRC (CONCUR China)	1.9 2.0	20% 23%	8% 14%	21%
savolitinib – c-Met (Ph II)		pRCC (JCO 2017)	6.2 (c-MET+)	13%	8%	14%
fruquintinib - VEGFR1,2,3 (FRESCO)		$\geq$ 3L CRC	3.7	24%	15%	19%
fruquintinib - VEGFR1,2,3 (Ph II)		3L NSCLC	3.8	13%	8%	11%
sulfatinib - VEGFR 1,2,3, FGFR1, CSF-1R (Ph II)		PNET, EP-NET	19.4, 13.4	25%	9%	19%
<b>epitinib</b> - EGFR (Ph I/II)		NSCLC w/brain mets		6%	N/D	N/D

## 4. Combination therapies proving to be a hard challenge

- Avg. 64% with grade 3-4 tox.
   vs. 37% in mono. trials.<sup>[1]</sup>
- ≤10 TKI+TKI or TKI+IO oncology combos FDA approved (as of YE 2017).<sup>[3]</sup>
  - ★ Drug-drug interactions.
  - ✗ Overlapping AEs.
- Keys to sustained combo use (i.e. minimize discont.):
  - Constituents must be highly tolerable.
  - ✓ Clear known AE profiles
     & careful management.

[1] Roda D et al. Clinical Cancer Research 2016 May 1;22(9):2127-32. [2] Sources: CABOSUN = Choueiri et al, J Clin Oncol. 2017 Feb 20;35(6):591-597; COMPARZ = Motzer et al, N Engl J Med. 2013 Aug 22;369(8):722-31; AXIS = Motzer et al, Lancet Oncol. 2013 May;14(6):552-62; lenvatinib Ph 2 = Motzer et al, Lancet Oncol. 2015 Nov;16(15):1473-82; CORRECT = Grothey et al, Lancet. 2013 Jan 26;381(9863):303-12; CONCUR China = Xu et al, "Efficacy and safety of regorafenib monotherapy in Chinese patients with previously treated metastatic colorectal cancer", CSCO 2014; savolitinib PRCC = Choueiri et al, J Clin Oncol. 2017 Sep 10;35(26):2993-3001; FRESCO = Li et al, J Clin Oncol. 2017 May 35(15\_suppl):3508-3508; fruquintinib NSCLC = Liu, ID4571, WCLC 2017; sulfatinib NET = Xu et al, #1697, ENETS 2017; epitinib NSCLC = Chi-Med data. [3] Approved TKI combos: HER2 inhibitor + HER2 inhibitor: BRAF inhibitor + MEK inhibitor: VEGFR inhibitor + MTOR inhibitor + CD20 inhibitor.

### **36 active or completing trials on 8 drug candidates** CHI-Four drug candidates in Ph.III, or about to start

Program	Target	Partner	Indication	Latest Status	Line	Target patient	Combo therapy	Site	Preclin.	Ph.I	Proof-of-concept	Pivotai/Ph.III
			1. Papillary renal cell carcinoma	Ph.III enrolling	1 <sup>st</sup>	c-Met-driven		Global				
			2. Papillary renal cell carcinoma	NCI Ph.II – savo vs. sunitinib vs. cabozan. vs. crizot.	All	All		US				
		≥	3. Papillary renal cell carcinoma	Ph.II enrolling	-	All	durvalumab (PD-L1)	UK/Sp				
		S.	4. Clear cell renal cell carcinoma	Ph.II enrolling	2 <sup>nd</sup>	VEGF TKI refractory		UK/Sp				
		stra	5. Clear cell renal cell carcinoma	Ph.II enrolling	2 <sup>nd</sup>	-	durvalumab (PD-L1)	UK/Sp				
		N	6. Non-small cell lung cancer	Ph.II enrolling; <b>planning next stage initiation 2018</b>	2 <sup>nd</sup>	EGFR TKI refractory		Global			i 📫	
Savolitinib	c-Met	<u>e</u>	7. Non-small cell lung cancer	Ph.II enrolling; pivotal decision pending	3 <sup>rd</sup>		Tagrisso® (T790M)	Global				
(AZD6094)	CMCC		8. Non-small cell lung cancer	Ph.II enrollment complete; pivotal under discussion	2 <sup>nd</sup>	EGFR TKI refractory	Iressa® (EGFR)	China			<u> </u>	
			9. Non-small cell lung cancer	Ph.II enrolling	1 <sup>st</sup>	c-Met-driven		China				
			10. Lung cancer	Ph.II enrolling	1 <sup>st</sup>	c-Met-driven		China				
			11. Gastric cancer	Ph.II enrolling		c-Met+		SK/PRC				
			12. Gastric cancer	Ph.II enrolling	2 <sup>nd</sup>	c-Met+	<b>docetaxel</b> (chemo)	SK				
			13. Gastric cancer	Ph.II enrolling	2 <sup>nd</sup>	c-Met O/E	docetaxel (chemo)	SK				
			14. Prostate cancer	CCTG Ph.II enrolling – umbrella trial	1 <sup>st</sup> /2 <sup>nd</sup>	c-Met-driven		Can				
			15. Colorectal cancer	Ph.III met all endpoints; NDA submitted Jun 2017	3 <sup>rd</sup>	All		China			I.	•
	VECED	Lilly	16. Non-small cell lung cancer	Ph.III fully enrolled; report top-line results late 2018	3 <sup>rd</sup>	All		China			n/a	⇒
Fruquintinib	VEGFR 1/2/3	(in China	17. Non-small cell lung cancer	Ph.II enrolling	1 <sup>st</sup>	All	Iressa®(EGFR)	China				
	1/2/5	(III CIIIIA Only)	18. Caucasian bridging	Ph.I enrolling	-	All comers		US				
			19. Gastric cancer	Ph.III enrolling	2 <sup>nd</sup>	All	<b>paclitaxel</b> (chemo)	China				
			20. Pancreatic NET	Ph.III enrolling	1 <sup>st</sup>	All		China				
			21. Non-pancreatic NET	Ph.III enrolling	1 <sup>st</sup>	All		China				
c	VEGFR/		22. Caucasian bridging	Ph.I enrolling	-	All comers		US				
Sulfatinib	CSF-1R/ FGFR1		23. Medullary thyroid ca.	Ph.II enrollment complete	2 <sup>nd</sup>	Radiotherapy ref.		China				
	FUFRI		24. Differentiated thyroid ca.	Ph.II enrollment complete	2 <sup>nd</sup>	Radiotherapy ref.		China				
			25. Biliary tract cancer	Ph.II enrolling	2 <sup>nd</sup>	Chemo ref.		China				
			26. Non-small cell lung cancer	Preparing for Ph.III; target initiation 2018	1 <sup>st</sup>	EGFRm+ brain mets		China				
Epitinib	EGFRm+		27. Glioblastoma	Ph.Ib/II enrolling	-	EGFR+		China				

### 6 pivotal trials underway with more preparing to start

Oncology

Immunology

Notes: Proof-of-concept = Phase Ib/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; EGFR+ = EGFR activating mutations; EGFR+ = EGFR gene amplification; EGFR wT = EGFR wild-type; 5ASA = 5-aminosalicylic acids; chemo = chemotherapy; c-Met+ = c-Met gene amplification; c-Met O/E = c-Met over-expression; FGFR = fibroblast growth factor receptor; CSF-1R = colony stimulating factor-1 receptor; NCI = U.S. National Cancer Institute; CCTG = 12 Canadian Cancer Trial Group; Aus = Australia; Can = Canada; SK = South Korea; PRC = People's Republic of China; Sp = Spain; UK = United Kingdom; US = United States; Global = >2 countries.

### Next wave of innovation now in proof-of-concept Four novel drug candidates in Phase I/II



Program	Target	Partner	Indication	Latest Status	Line	Target patient	Combo therapy	Site	Preclin.	Ph.I	Proof-of-concept	Pivotal/Ph.III
Theliatinib	EGFR WT		28. Solid tumors	Ph.I enrollment complete	-	All comers		China				
menatinib	LUIKWI		29. Esophageal cancer	Ph.Ib expansion enrolling	1 <sup>st</sup>	EGFR WT		China				
	_											
			30. Immunology	Ph.I complete; preparing for Ph.II	-	Healthy volunteers		Aus				
HMPL-523	Syk		31. Immunology	Ph.I dose escalation	-	Healthy volunteers		China				
	Зук		32. Hematological cancers	Ph.I enrolling	2 <sup>nd</sup> /3 <sup>rd</sup>	All comers		Aus				
			33. Lymphoma	Ph.I enrolling	-	All comers		China				
HMPL-689	ΡΙ3Κδ		34. Hematological cancers	Ph.I complete; preparing for Ph.II	-	Healthy volunteers		Aus				
	FISIO		35. Lymphoma	Ph.I enrolling	2 <sup>nd</sup> /3 <sup>rd</sup>	All comers		China				
HMPL-453	FGFR		36. Solid tumors	Ph.I enrolling	-	All comers		Aus				
	1/2/3		37. Solid tumors	Ph.I enrolling	-	All comers		China				
HM004-6599	NF-ĸB	Health	Ulcerative colitis (Induction)	HMPL-004 reformulation; IND submitted	2 <sup>nd</sup>	5ASA refractory		Aus/China				
	(TNF-α)	Science	Ulcerative colitis (Maintenance)	Await positive Ph.II in Ulcerative Colitis (Induction)	2 <sup>nd</sup>	5ASA refractory		China				
		Nestlē										
NSP DC2	TBD	Health Science	Immunology	IND end of 2018				China				
		ociciice										
Multiple	TBD		Oncology	Four small molecule/antibody programs in preclin.				TBD				

>3,500 subjects treated in studies (as of Dec 31, 2017); and >700 dosed in 2017.

Oncology

Immunology

Notes: Proof-of-concept = Phase Ib/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; EGFR+ = EGFR activating mutations; EGFR+ = EGFR gene amplification; EGFR WT = EGFR wild-type; 5ASA = 5-aminosalicylic acids; chemo = chemotherapy; c-Met+ = c-Met gene amplification; c-Met O/E = c-Met over-expression; FGFR = fibroblast growth factor receptor; CSF-1R = colony stimulating factor-1 receptor; NCI = U.S. National Cancer Institute; CCTG = 1 3 Canadian Cancer Trial Group; Aus = Australia; Can = Canada; SK = South Korea; PRC = People's Republic of China; Sp = Spain; UK = United Kingdom; US = United States; Global = >2 countries.

# 8-10 shots at pivotal success

First positive Ph.III outcome - fruquintinib in colorectal cancer



Est. Pivotal Read-out

**Breakthrough Therapy** 

					("BTD") potential	(if not BTD)
	Papillary renal cell carcinoma (MET-driven)	Pivotal Phase III	Global	Enrolling	Molecular epidemiology study MET as -ve prognostic H1-2018	2020
	NSCLC –2L 1 <sup>st</sup> Gen EGFR TKI refract, Tagrisso combo (MET+ , T790M+/-)	Pivotal Phase II/III <sup>[1]</sup>	Global	Randomized, chemo- doublet controlled study to init <u>iate</u> in 2018 <sup>[2]</sup>	ORR MET+/T790M+ 55% ORR MET+/T790M- 61%	2020
SAVO	NSCLC –3L 3 <sup>rd</sup> Gen EGFR TKI refract. Tagrisso combo (MET+)	Pivotal Phase III	Global	AZ pivotal study decision pending	ORR MET+ 33%	2021
	NSCLC –2L 1 <sup>st</sup> Gen EGFR TKI refract, Iressa combo (MET+, T790M-)	Pivotal Phase III	China	AZ/Chi-Med under discussion	ORR MET+/T790M- 52%	2021
	3L (or above) Colorectal cancer ("CRC")	Pivotal Phase III	China	Complete, <mark>Met All</mark> Endpoints, NDA submitted	$\checkmark$	March 3, 2017
FRUQ	3L Non-small cell lung cancer ("NSCLC")	Pivotal Phase III	China	Enrollment complete		04 2018 (Top-line results)
	2L Gastric cancer combo with Taxol	Pivotal Phase III	China	Enrolling		2020
	Pancreatic neuroendocrine tumors	Pivotal Phase III	China	Enrolling		2019
SULF	Non-pancreatic neuroendocrine tumors	Pivotal Phase III	China	Enrolling		2019
EPIT	1L EGFR-mutant NSCLC with brain metastasis	Pivotal Phase III	China	Initiating in 2018		2020

[1] Subject to the outcome of mature TATTON B and preliminary TATTON D data, and regulatory discussions; [2] In MET+, T790M- patients.



### Savolitinib (AZD6094) Potential first-in-class selective c-Met inhibitor

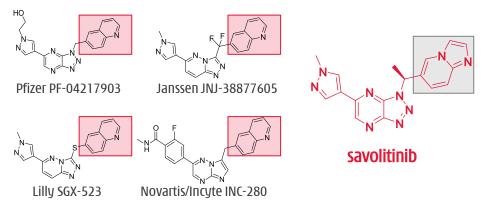


### Savolitinib (AZD6094)



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

- 1. Strong potential to become first selective c-MET inhibitor approved.
  - Clear clinical efficacy observed in non-small cell lung ("NSCLC"), kidney, gastric and colorectal cancers.
  - Partnered with AstraZeneca key comp. advantages in NSCLC (Tagrisso<sup>®</sup> 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 1<sup>st</sup>-gen c-Met compounds has dramatically reduced solubility and appeared to crystallize in the kidney resulting in obstructive toxicity.

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2. c-Met is aberrant in many tumor settings. [3]

		(C-MET)	New Cases (2015)		
Indication	Amplifi- cation	Mutation	Over- Expression	Global	China
Gastric	10%	1%	41%	1,034,000	679,100
Lung	8-10% [1]	8%	67%	1,690,000	733,300
Head & Neck		11%	46%	740,000	135,000
Colorectal	10%		65%	1,477,000	376,300
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	477,900
Prostate <sup>[4]</sup>			54-83%	1,100,000	60,300

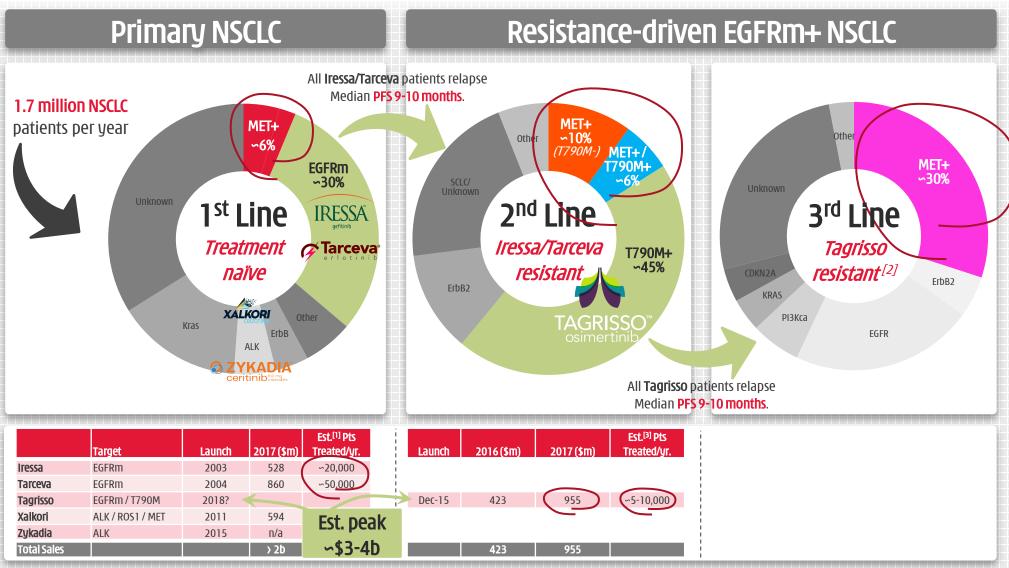
#### 4. AstraZeneca collaboration & 2016 amendment.

- \$20m received upfront (Dec 2011);
- \$120m in development/approvals milestones (\$25m received as of Dec 2017);
- Several hundred million in commercial milestones;
- Development costs: AZ pay 100% ex-China (excl. \$50m by Chi-Med) & 75% development cost in China (Chi-Med 25%);
- 14-18% tiered royalty ex-China; & 30% flat rate China royalty on all product revenues.

[1] Range includes (i) approximately 4% of c-Met+ naïve non-small cell lung cancer patients and (ii) 10 - 30% of EGFRm+ non-small cell lung cancer patients, which 15 to 20% develop EGFRm+ tyrosine kinase inhibitor resistance pathway as c-Met+; [2] Hereditary papillary renal cell carcinoma only; [3] Company estimates considering Frost & Sullivan data, National Central Cancer Registry of China and publicly available epidemiology data. [4] By IHC, c-Met overexpression in 54% of lymph node disease and 83% of bone metastases. Varkaris et al, Expert Opin Investig Drugs. 2011 Dec; 20(12): 1677–1684.

### Savolitinib Biggest opportunity is MET+ non-small cell lung cancer ("NSCLC")

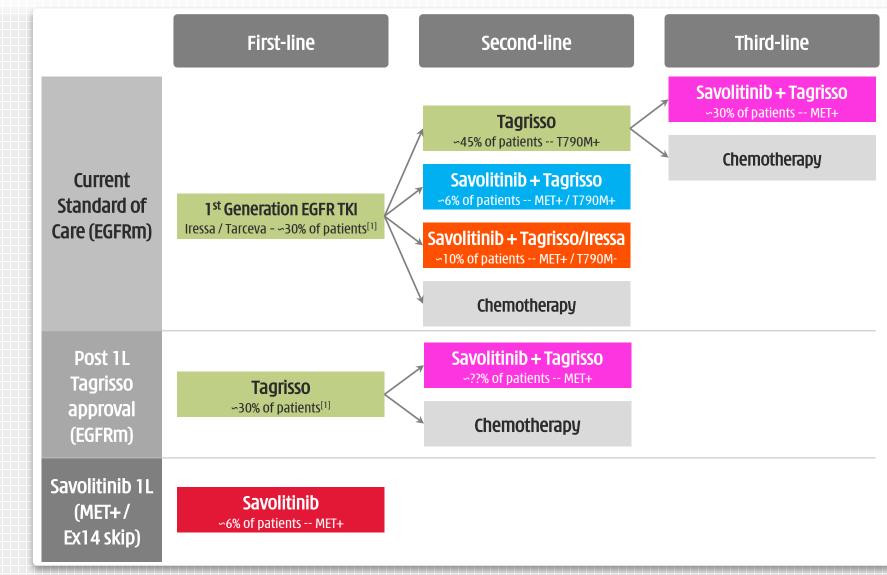


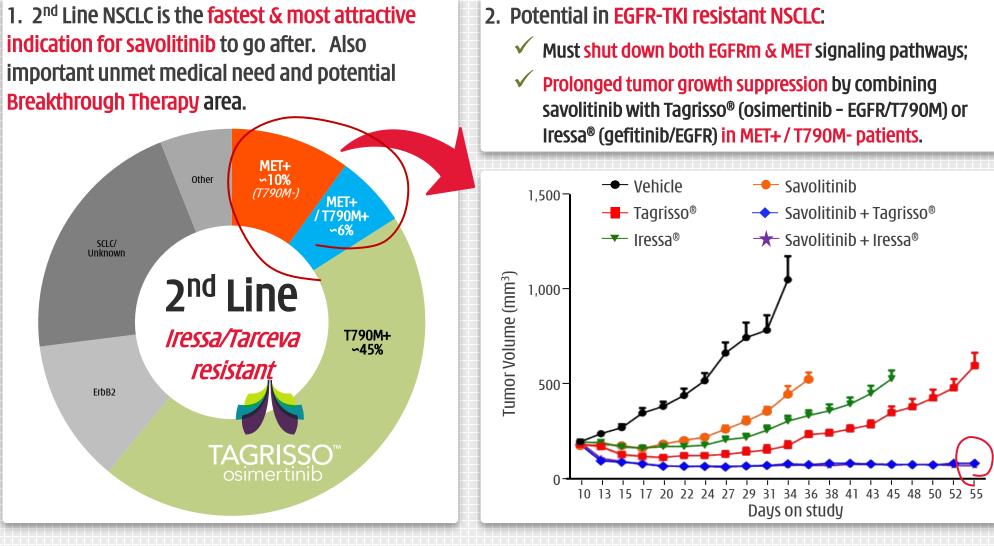


[1] General estimate based on mPFS ~9 mo. average cost/cycle ~\$2,500-3,000; [2] Primary drivers, based on aggregate rocelitinib/Tagrisso data published at 2016/2017 ASCO; [3] AstraZeneca 2016 /17 results.

### Savolitinib – NSCLC Five opportunities for savo in NSCLC





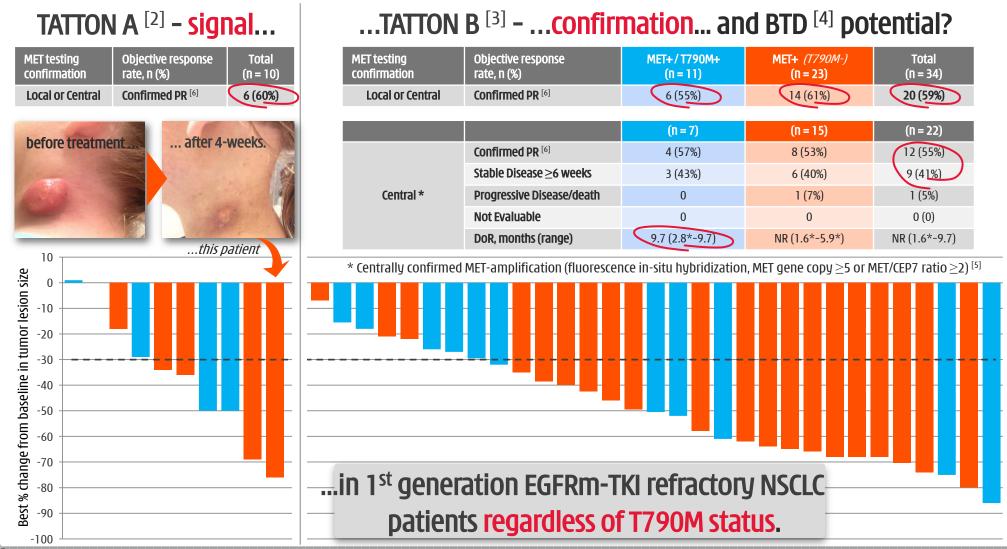


### Savolitinib – 2<sup>nd</sup> Line EGFRm NSCLC Very strong preclinical rationale for combination w/ EGFR-TKIs



### Savolitinib – 2<sup>nd</sup> Line NSCLC<sup>[1]</sup> combo w/ TAGRISSO<sup>\*\*</sup> TATTON A/B compelling – now starting next stage of development





20 [1] EGFRm NSCLC; [2] ESMO 2016 Galbraith - Novel Clinical Trials for Prec. Med.; [3] WCLC 2017 – Ahn M-J, et al. TATTON Phase Ib expansion cohort; [4] U.S. FDA Breakthrough Therapy designation potential; [5] Some local MET-status determined via IHC+3 in  $\geq$  50% of tumor cells),; [6] PR = Partial Response

#### Iressa<sup>®</sup> / savo combo in 1<sup>st</sup> gen. EGFRm-TKI refractory patients <sup>[2]</sup>...outstanding response in MET+/T790Min MET+/T790M- patients

MET testing confirmation	Objective response rate, n (%)	MET+ / T790M+ (n = 23)	MET+ <i>(T790M-)</i> (n = 23)	MET+ / T790M unk. (n = 5)	Total (n = 51)
	Confirmed PR <sup>[3]</sup>	2 (9%)	12 (52%)	2 (40%)	16 (31%)
Central *	SD $^{[4]} \ge$ 6 weeks	9 (39%)	7 (30%)	2 (40%)	18 (35%)
Central	PD <sup>[5]</sup> /death	7 (30%)	3 (13%)	0	10 (20%)
	Not Evaluable	5 (22%)	1 (4%)	1 (20%)	7 (14%)

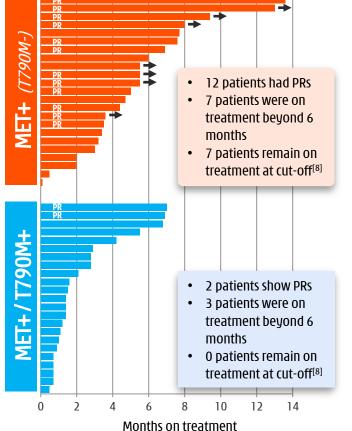
### ...vs. TATTON B data (savo / Tagrisso<sup>®</sup> combo) <sup>[6]</sup>

Objective response rate, n (%)	MET+/T790M+ (n = 11)	MET+ (7790M-) (n = 23)	MET+ / T790M unk. (n = 0)	Total (n = 34)
Confirmed PR <sup>[3]</sup>	6 (55%)	14 (61%)	0	20 (59%)
	(n = 7)	(n = 15)	(n = 0)	(n = 22)
Confirmed PR <sup>[3]</sup>	4 (57%)	8 (53%)	0	12 (55%)
SD $^{[4]} \ge$ 6 weeks	3 (43%)	6 (40%)	0	9 (41%)
PD <sup>[5]</sup> / death	0	1 (7%)	0	1 (5%)
Not Evaluable	0	0	0	0 (0)
	rate, n (%) Confirmed PR <sup>[3]</sup> Confirmed PR <sup>[3]</sup> SD <sup>[4]</sup> $\geq$ 6 weeks PD <sup>[5]</sup> / death	rate, n (%) $(n = 11)$ Confirmed PR [3] $6 (55\%)$ (n = 7)         Confirmed PR [3] $4 (57\%)$ SD [4] $\geq$ 6 weeks $3 (43\%)$ PD [5] / death $0$	rate, n (%)(n = 11)(n = 23)Confirmed PR $[3]$ 6 (55%)14 (61%)(n = 7)(n = 15)Confirmed PR $[3]$ 4 (57%)8 (53%)SD $[4] \ge 6$ weeks3 (43%)6 (40%)PD $[5]$ / death01 (7%)	rate, n (%)(n = 11)(n = 23)(n = 0)Confirmed PR $[3]$ 6 (55%)14 (61%)0(n = 7)(n = 15)(n = 0)Confirmed PR $[3]$ 4 (57%)8 (53%)0SD $[4] \ge 6$ weeks3 (43%)6 (40%)0PD $[5]$ / death01 (7%)0

\* Centrally confirmed MET-amplification (fluorescence in-situ hybridization, MET gene copy  $\geq$ 5 or MET/CEP7 ratio  $\geq$ 2)<sup>[9]</sup>.

[1] EGFRm NSCLC; [2] WCLC 2017Yang J-J, et al. A Ph.lb Trial of savolitinib plus gefitinib for patients with EGFR-mutant MET-amplified advanced NSCLC; [3] PR = Partial Response; [4] SD = Stable Disease; [5] PD = Progressive Disease; [6] WCLC 2017 - Ahn M-J, et al. TATTON Phase Ib exp. coho [7] DOR = Duration of Response; [8] Aug 21, 2017; [9] On TATTON B, some local MET-status determined via IHC+3 in  $\geq$  50% of tumor cells.

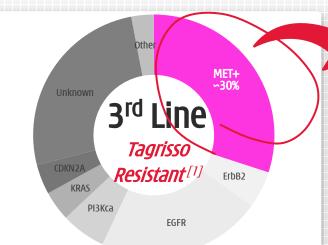
# VC TATTON D data (cavo / Taciaco® comba) [6]



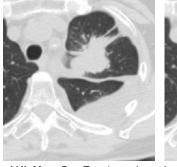


### Savolitinib – 2<sup>nd</sup> Line NSCLC<sup>[1]</sup> combo w/ IRESSA Compelling in MET+ / T790M-, pivotal decision under discussion

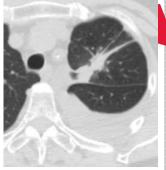
### Savolitinib - 3<sup>rd</sup> Line NSCLC<sup>[1]</sup> - A TAGRISSO resistant MET+ driven resistance in ~30% of patients



#### 3 out of 3 MET+ patients responded to savo/Tagrisso<sup>®</sup> combo.



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6 wks. on savo/Tag. Treatment LUL Mass Pre-Treatment

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	Y	-	T790M ND
4	L858R (de novo T790M)	2	Ŷ	<i>MET</i> amp, <i>EGFR</i> amp T790M (germline)	-
5	L858R	3	Y	T790wt, <i>EGFR</i> amp	T790M ND
6	L858R	4	Y	T790 WT	T790M ND
7	Del19	3	Y	-	T790M ND
8*	Del19	3		T790M/C797S	T790M/C797S
9	L858R	4	Y	T790 WT	-
10	Del19	3	Y	-	<i>PIK3CA</i> E545K, <i>PIK3CA</i> amp, T790M N
11	Del19	2	Y	<i>MET</i> amp, <i>EGFR</i> amp, T790 WT	T790M ND
12	Del19	2	Y	-	T790M/C797S
13	Del19	9		T790 WT	-
7	Del19	2	Y	T790 WT	T790M ND
م	Del19	1		T790 WT	FGFR1 D60N, FGFR1 amp, T790M NI
16	L858R	2		<i>MET</i> amp, T790 WT	MET, EGFR amp, T790M ND
17	L858R	3	Y	T790 WT	T790M ND
18	Del19 (de novo T790M)	3		SCLC, T790 WT	T790M ND, <i>EGFR</i> amp
19	Del19	3	Y	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	Y	-	T790M/C797S

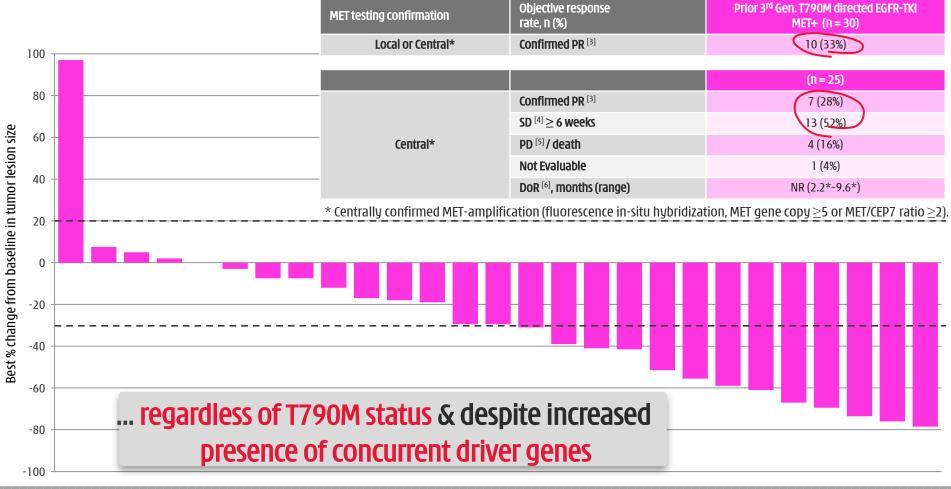
[1] Based on rocelitinib/Tagrisso data published at 2016/2017 ASCO; [2] In xenograft model H820, with EGFRm, T790M+ and MET CN gain. D'Cruz CM et al; #761 Preclinical data for changing the paradigm of treating drug resistance in NSCLC. Novel combinations of AZD6094, a selective MET inhibitor, and AZD9291 an irreversible, selective (EGFRm and T790M) EGFR TKI; American Association of Cancer Research Annual Meeting; April 19, 2015.



### Savolitinib – 3<sup>rd</sup> Line NSCLC<sup>[1]</sup> combo w/ simertinib No treatment options post Tagrisso<sup>®</sup> – pivotal decision pending

from accision penaing

...TATTON B<sup>[2]</sup> - ...promising efficacy in MET+ Tagrisso failure patients...



[1] EGFRM NSCLC; [2] WCLC 2017 – Ahn M-J, et al. TATTON Phase Ib expansion cohort; Waterfall plot based on evaluable patients (n=30): all patients dosed and with on-treatment assessment or discontinuation prior to first tumour assessment; Data cut-off 31 Aug 2017; [3] PR = Partial Response; [4] SD = Stable Disease; [5] PD = Progressive Disease; [6] DOR = Duration of Response

### Tolerability – savo plus IRESSA Or TAGRISSO TATTON D – 300mg QD dose potentially support long-term use



#### Efficacy / Tolerability analysis in $\ge 2^{nd}$ -Line NSCLC

				Effi	cacy	Disconti	nuations as %	Enrolled
US FDA Approval Date	Treatment / Control arms	Disease setting	n	ORR	Median PFS (mo.)	Due to AE	Withdrawn / Other	Total [5]
Non-Small	Cell Lung Cancer Treatment arms							
30-Mar-17	Tagrisso® (osimertinib)	<b>2L</b> EGFRi-refractory T790M+ NSCLC (AURA3)	279	71%	10.1	6%	6%	13%
29-Apr-14	<b>Zykadia</b> ® (ceritinib)	<b>2L</b> ALK+ NSCLC after Xalkori (single arm)	163	56%	6.9	10%	10%	20%
12-Dec-14	<b>Cyramza®</b> (ramucirumab) + <b>Taxotere</b> ®	2L NSCLC after plat-chemo	624	23%	4.5	15%	21%	37%
24-0ct-16	<b>Keytruda</b> <sup>®</sup> (pembrolizumab) 2mg/kg	<b>2L</b> PD-L1+ (TPS≥1%) NSCLC after plat-chemo (KEYNOTE-010)	345	18%	3.9	10%	26%	37%
2-0ct-15	<b>Keytruda®</b> (pembrolizumab) 10mg/kg	<b>2L</b> PD-L1+ (TPS≥1%) NSCLC after plat-chemo (KEYNOTE-010)	346	18%	4.0	9%	27%	36%
9-0ct-15	<b>Opdivo</b> ® (nivolumab)	<b>2L</b> NSCLC after plat-chemo	292	19%	2.3	15%	4%	20%
4-Mar-15	<b>Opdivo®</b> (nivolumab)	<b>2L</b> squ. NSCLC after plat-chemo	135	20%	3.5	12%	8%	20%
Non-Small	Cell Lung Cancer Control arms (aggregate .	/ weighted average)						
	<b>Chemo doublet</b> (platinum + pemetrexed)	<b>2L</b> NSCLC <i>(AURA3)</i>	136	31%	4.4	11%	17%	27%
	Taxotere® (docetaxel)	<b>2L</b> NSCLC <i>(REVEL; KEYNOTE-010; Opdivo x2)</i>	1,391	12%	3.5	13%	22%	36%
Savolitinib	)							
	<b>savolitinib</b> 600mg QD monotherapy <sup>[3]</sup>	All-lines Papillary RCC FOR REFERENCE ONLY	109 [1]	18%	6.2	9%	5%	14%
	savolitinib 600mg QD + Iressa® (gefitinib) [4]	≥ <b>2L</b> EGFRm+ c-MET+ T790M- NSCLC after 1st-gen EGFR TKI (expansion)	51 [2]	52%	ND	20%	14%	33%
	savolitinib 600mg QD + Tagrisso® [4]	$\geq$ <b>2L</b> EGFRm+ c-MET+ T790M-/+ NSCLC after 1 <sup>st</sup> -gen EGFR TKI (TATTON B)	34	59%	ND	- 30%	3%	33%
	savolitinib 600mg QD + Tagrisso® [4]	≥ <b>3L</b> EGFRm+ c-MET+ NSCLC after 3 <sup>rd</sup> -gen EGFR TKI (TATTON B)	30	33%	ND	50%	⊃%	55%

[1] PRCC Phase II - Efficacy data from MET+ patients (n=44), discontinuation data from late 2017 data cut-off; Tolerability data from all patients (n=109); [2] TATTON Study - Efficacy data for noted molecular subsets; Tolerability data from all patients (n=64); [3] September 2017 Journal of Clinical Oncology; [4] 2017 World Conference on Lung Cancer; ND = Not Disclosed; [5] Total discontinuations = Discontinuations NOT due to Disease Progression or Death.

#### Safety - savolitinib plus IRESA or TAGRISSO Gefitinib Adverse event profiles of combinations - manageable & tolerable

	<b>IPASS P</b> 1 <sup>st</sup> -Line EGI		
Grade ≥3 AEs, Preferred term, n (%)*	IPASS Iressa® (N=607)	IPASS carbo. + Taxol® (N=589)	≥ 2 <sup>nd</sup> -Line <sup>[2]</sup> Savo + Iressa® (N=51)
Any Grade ≥3 AE	29% (Gr. 3-4)	61% (Gr. 3-4)	17 (33%)
Vomiting	1 (<1%)	16 (3%)	
Rash or acne	19 (3%)	5 (1%)	
AST/ALT increase			8 (16%)
Nausea	2 (<1%)	9 (1%)	1 (2%)
Decreased appetite			
Fatigue			
Neutropenia	22 (4%)	387 (67%)	
ALP increased			11 (22%)
Neurotoxic effects	2 (<1%)	29 (5%)	
Anemia	13 (2%)	61 (11%)	
Leukopenia	9 (1%)	202 (35%)	
Thrombocytopenia			

FLAURA Phase III 1 <sup>st</sup> -Line EGFRm NSCLC						
Tagrisso® (N=279)	lressa® or Tarceva® (N=277)					
94 (34%)	124 (45%)					
0	4 (1%)					
3 (1%)	19 (7%)					
3 (1%)	37 (13%)					
0	0					
7 (3%)	5 (2%)					
2 (1%)	2 (1%)					
3 (1%)	3 (1%)					

#### AURA3 Phase III 2<sup>nd</sup>-Line EGFRm NSCLC $> 2^{nd}$ -Line<sup>[1]</sup> Chemo-doublet Tagrisso® (plat. + pemetrex.)Savo + Tagrisso<sup>®</sup> (N=279) (N=66) 63 (23%) 64 (47%) 33 (50%) 1 (<1%) 3 (2%) 5 (8%) 2 (1%) 4 (6%) 6 (2%) 2 (2%) 4 (6%) 2 (1%) 5 (4%) 3 (5%) 3 (1%) 4 (3%) 3 (5%) 3 (1%) 1 (1%) 3 (5%) 4 (1%) 16 (12%) 3 (5%) 2 (1%) 16 (12%) 5 (4%)

10 (7%)

1((1%))

Sources: [1] TATTON B – Figures where any grade AE  $\geq$ 10% patients. Ahn M-J, et al. Abstract #8985. Presented at the World Lung Cancer Congress (WCLC) 2017, Japan, October 2017; [2] Phase Ib/II study – Figures where any grade AE  $\geq$ 10% patients. Yang J-J, et al. Abstract #8995. Presented at WCLC 2017, Japan, October 2017.

AST = aspartate aminotransferase; ALT = alanine aminotransferase; ALP = alkaline phosphatase.

#### ALK RCC-47 PRCC model tumor volume (mm<sup>3</sup>) ASCO ANNUAL MEETING '16 Kalkori® 25mg/kg 1,000 Savolitinib 25mg/kg 500 MET Exon14 skipping: MET Exon14 skipping: Epidemiology of never-exposed to c-MET TKI ever exposed to c-MFT TKL posed to c-MFT TKI With concurrent Without concurrent No. of pts 27 34 c-MET amplification c-MET amplification Median OS 24.6 months 8.1 months 5.2 months Median OS 10.5 months 10 20 P=0.06 Days post treatment [1] Drilon A, Abstract 108 Efficacy and safety of crizotinib in patients with advanced MET Exon 14-altered non-small cell lung cancer; [2] ASCO 2017, Abstract 8511, Mark M. Awad et al.; [3] Paik, P.K., et al., Response to MET inhibitors in patients with stage arboring MET mutations causing exon 14 skipping. Cancer Discov, 2015. 5(8): p. 842-9.; [4] Schuller AG et al. "Regression in Papillary Renal Cell Carcinoma Patient-Derived Xenograft Models"

Savolitinib vs. Xalkori®

10x

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

Xalkori<sup>®</sup> (crizotinib) proof-of-concept in Exon 14 skip 1L NSCLC

Xalkori® (crizotinib)

19

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

IC<sub>so</sub>(nM)

1<sup>st</sup> Line

Treatment

naïve

IRESSA

Tarceva

**EBC1 Viability** 

26

EBC1 pMET 39 40x 293T MET (wild type) 79 11x

response (~50% ORR) but > 1/3<sup>rd</sup> of responses not durable (4/12)<sup>[1]</sup>.

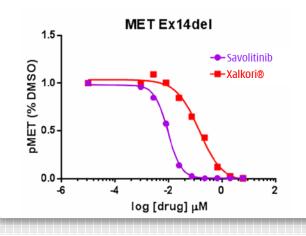




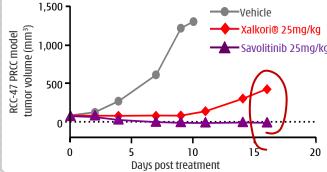
293	3T MET (Ex14del)	9	140	16X
2.	1 <sup>st</sup> line NSCLC – Xalkori® I	MET Exon14 sl	kipping - 2016 A	SCO – strong

Savolitinib





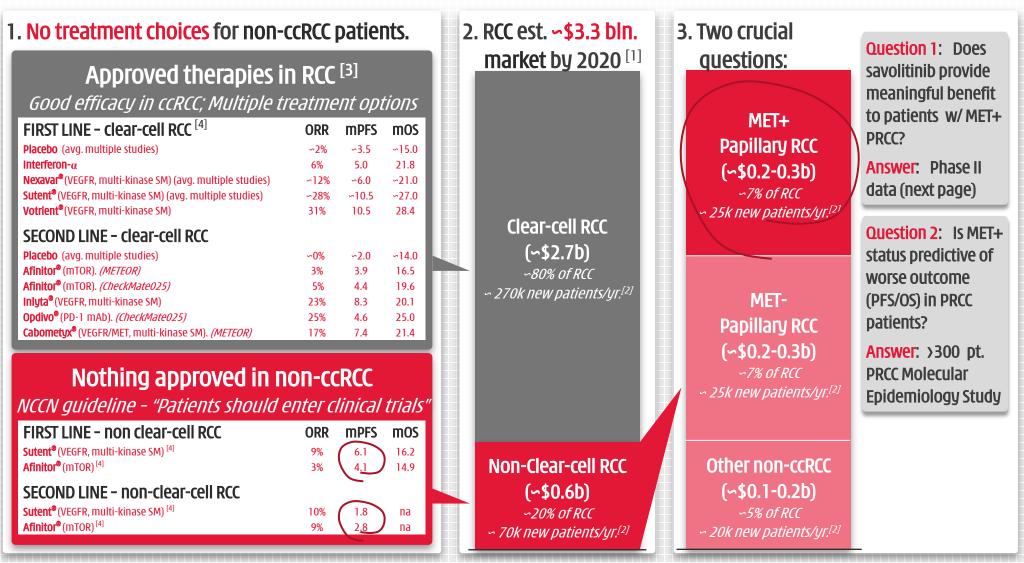
5. Durable tumor cell suppression for savolitinib but not for Xalkori<sup>®[4]</sup>.



### MET+ PRCC - unmet medical need

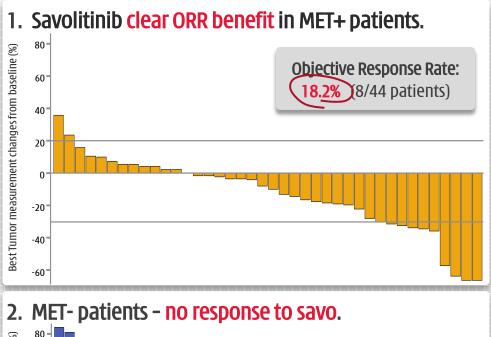
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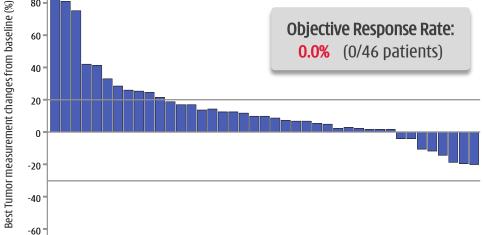




[1] Transparency Market Research, March 2015 - RCC (excl. non-RCC Kidney Cancer) global market size; [2] Frost & Sullivan, March 2016; [3] NCCN Guideline for kidney cancer. Version 3.2016, 05/26/16., RCC = renal cell carcinoma; [4] ORR = Objective Response Rate, mPFS = median Progression Free Survival, mOS = median Overall Survival

### Savolitinib – PRCC Phase II Clear efficacy & durable response in MET+ PRCC patients





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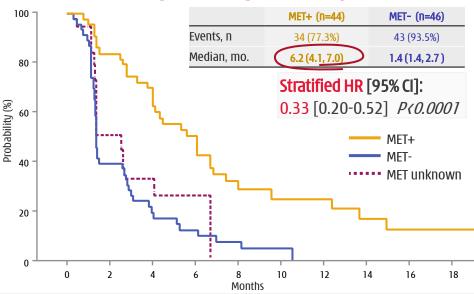
#### 3. Disease Control Rate ("DCR") – **big advantage** in MET+ with QCR 73.2% vs. MET- 28.2%.<sup>^</sup>

Tumor responses in the overall treatment population and by MET status

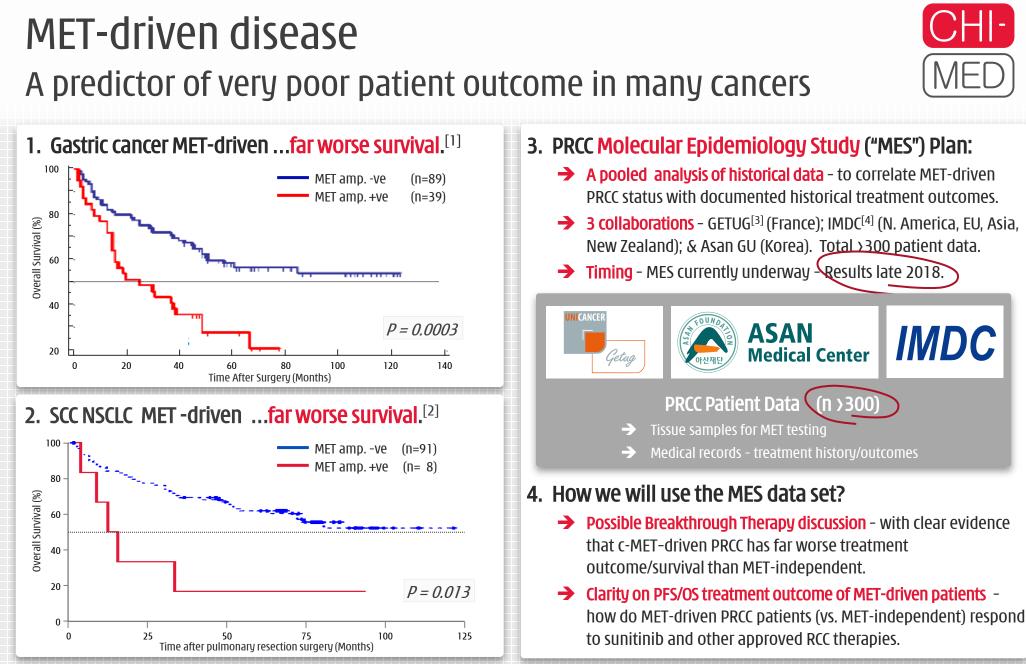
RECIST response, n (%)	MET+ (n=44)	MET- (n=46)	MET unknown (n=19)	Total (n=109)
Partial Response <sup>†</sup>	8 (18.2%)*	0 (0.0%)	0 (0.0%)	8 (7.3%)
Stable Disease	22 (50.0%)	11 (23.9%)	5 (26.3%)	38 (34.9%)
Progressive Disease	11 (25.0%)	28 (60.9%)	9 (47.3%)	48 (44.0%)
Not Evaluable	3 (6.8%)	7 (15.2%)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

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

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







[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 d'Étude des Tumeurs Urogénitales; [4] IMDC = International Metastatic Renal Cell Carcinoma Database Consortium.

### Savolitinib – PRCC Phase II

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

		PRCC PHASE II Savolitinib 1L/2L (n=109)	COMPARZ F Sunitinib 1L (n=548)	PHASE III <sup>[1]</sup> Pazopanib 1L (n=554)	METEOR PI Cabozantinib 2L (n=331)	HASE III <sup>[2]</sup> Everolimus 2L (n=322)	SINGLE-ARM PHASE III <sup>[3]</sup> Sunitinib 2L (n=106)	
MSKCC Risk Group	Favorable Intermediate Poor Missing	1 <u>4%</u> 45% 9% 32%	27 <u>%</u> 59% 9% 4%	2 <u>7%</u> 58% 12% 3%	45% 42% 12% 0%	46% 41% 13% 0%	5 <u>8%</u> 42% <sup>[6]</sup> 0%	→ Better safety data despite higher risk patient population: ✓ Only 14% "favorable" vs. 27-58%.
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 <sup>[4]</sup>	47%	77% <sup>[5]</sup>	76%[5]	68%	58%		
All Grade≥3 AEs with ≥5% incidence (AND selected savolitinib AEs for comparison) Hematologic Abnormalities Grade≥3 AEs with≥5% incidence:	Hypertension Fatigue Hand-foot-syndrome Diarrhea Neutropenia Thrombocytopenia	TR AES 0% 2% 0% 0% 0% 0% 0% 0% 0% (1%	TR AES 15% 17% 12% 8% 20% 24% 14% 6% 7%	TR AES 15% 11% 6% 9% 5% 4% 5% 1% 2%	All AEs 15% 9% 8% 11% 0% 0% 0% 0% 0% 5%	All AES 3% 7% (1% 2% 0% 0% 0% 0% 0% 16%	6% 11% 7% 16% 6%	Superior safety profile vs. other         TKIs - Most ≥3 G3 AEs ≈ 0-2%:         ✓       Hypertension: 0% vs. 6~17%.         ✓       Fatigue: 2% vs. 6~12%.         ✓       Diarrhea: 0% vs. ~10%.         ✓       Anemia: <1% vs. 7~16%.
Lab Abnormalities Grade≥3 AEs with≥5% incidence:	Increased ALT Increased AST Hypophosphatemia Hyponatremia Hypokalemia Hyperglycemia	5% 3% 0% 3% 0% 0%	4% 3% 9% 7% 1% 4%	17% 12% 4% 7% 3% 5%	2% 2% 4% 0% 5% <1%	<1% <1% 2% 0% 2% 5%		<ul> <li>≈ ALT/AST Increase: 3-5% vs. 0∽17%.</li> <li>✓ Other Lab Abnorm: 0% vs. ≤9%.</li> <li>Highly tolerable vs. other TKIs:</li> </ul>
Tolerability	Treatment discontinuation due to any AE <sup>[8]</sup> : Dose reduction due to AE:	8%	20% 51%	24% 44%	12% 62 <u>%</u>	11%	11%	<ul> <li>✓ Discontinued: 8% vs. 10~24%.</li> <li>✓ Dose reduction: 13% vs. 44-62%.</li> </ul>

[1] 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; [8] Early 2017 ASCO Genitourinary Cancers Symposium data cut-off.

### Savolitinib – Gastric cancer A major problem in east Asia – Japan, South Korea & China

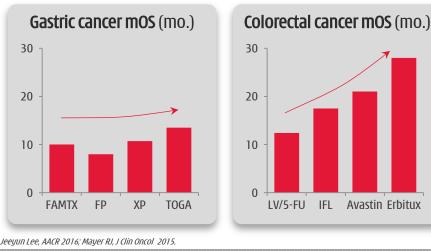


1. Gastric (stomach) cancer is the 5<sup>th</sup> most common cancer globally – 723,000 deaths/year.

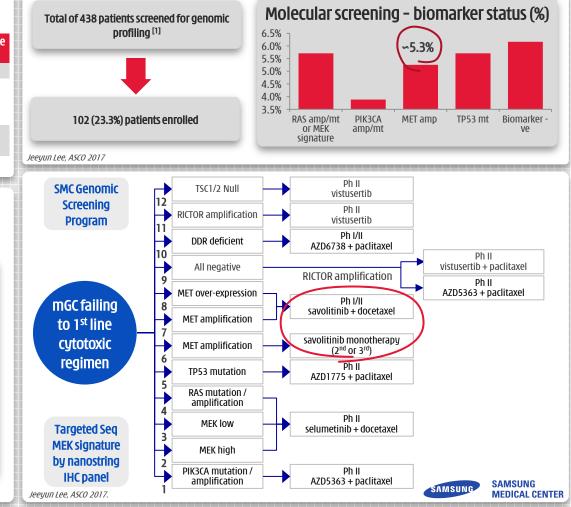
	Est. Age Standardised Rates (cases/100,000)	New cases ('000)	Deaths ('000)	5-year Prevalence ('000)
World	17.0	952	723	1,538
South Korea	41.8	22	17	32
Japan	29.9	38	29	56
China	22.7	405	325	594
EU-28	9.0	82	58	119
USA	6.8	21	12	32

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

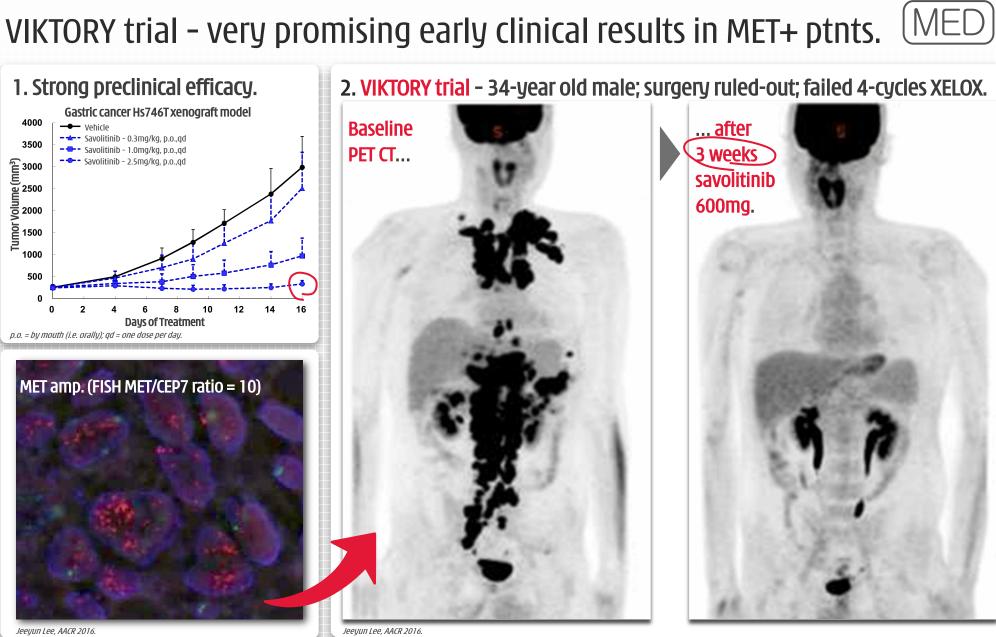
## 2. Little progress in gastric cancer<sup>[2]</sup> in improving overall survival ("OS") in first-line palliative setting.



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



[1] Since June 2014; [2] FAMTX = 5-FU + doxorubicin + methotrexate; FP = cisplatin + 5-FU; XP = capecitabine + cisplatin; TOGA = trastuzumab + chemo; LV/5-FU = leucovorin + 5-FU; IFL = irinotecan + 5-FU + leucovorin.



# Savolitinib – Gastric cancer



### Fruquintinib

Highly selective anti-angiogenesis inhibitor – Designed to be best-in-class relative to Stivarga<sup>®</sup> (regorafenib)



#### ✓ Validation of R&D approach - designed to only inhibit VEGFR1,2,3, 600 facilitating full target coverage & combinations.

- ✓ Pivotal Phase III in 3L CRC met all endpoints NDA submitted 02 '17.
- ✓ **Pivotal Phase III** in **3L NSCLC fully enrolled -** top-line result Q4 2018.
- ✓ **Pivotal Phase III Taxol<sup>®</sup> combo** in **2L gastric cancer initiated** Oct 2017.
- ✓ Phase II Iressa<sup>®</sup> combo in 1L EGFRm+ NSCLC early data at WCLC 2017.
- ✓ Phase I bridging in US initiated Dec 2017.

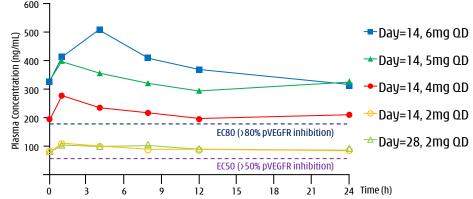
regulatory approval process underway.

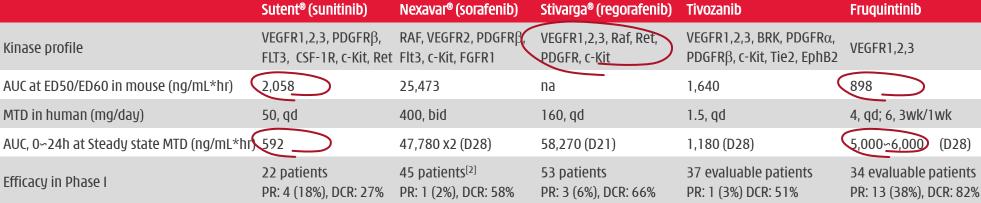
China GMP facility built & certification underway to support launch.

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

[1] Among small molecule tyrosine kinase inhibitors and to the best of Chi-Med's knowledge; [2] ( $\geq$ 100 mg bid); PR = Partial Response; DCR = Disease Control Rate.

#### 2. Only inhibits VEGFR – limits off-target toxicity & 1. Fruguintinib China NDA submission June 2017 allows for full & sustained target inhibition.



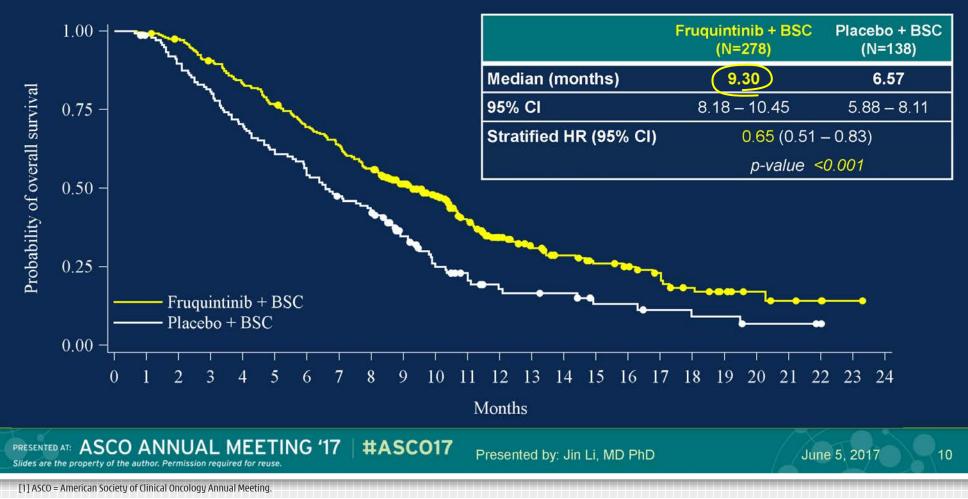




### **Fruquintinib – Third-line colorectal cancer** Best-in-class efficacy/safety – Ph.III FRESCO data ASCO 2017 <sup>[1]</sup>



**Overall Survival (Primary Endpoint)** FRESCO clearly succeeded in meeting the primary efficacy endpoint of OS





### Fruquintinib - FRESCO efficacy in 3L CRC

	Fruquintinib FRESCO Mainland China		Regorat	Regorafenib CONCUR		Regorafenib CONCUR		Regorafenib CORRECT	
Third-Line Metastatic Colorectal cancer			CONC						
			Chinese Patients (Mainland China, Hong Kong, Taiwan) <sup>[1]</sup>		Mainland China, Hong Kong, Taiwan, Vietnam, South Korea		Global		
Treatment arms	Fruquintinib	Placebo	Regorafenib	Placebo	Regorafenib	Placebo	Regorafenib	Placebo	
Patients (n)	278	138	112	60	136	68	505	255	
Complete Response, n (%) Partial Response, n (%) Stable Disease, n (%) <b>Disease Control Rate, n (%)</b> Median Progression Free Survival (mPFS) (mo.)	0.4% 4.3% 57.6% 62.2% +49 3.7 +1.9		0.0% 3.6% 40.2% 45.5% +38 2.0 +0		0.0% 4.4% 45.6% 51.5% +44 3.2 +1		0.0% 1.0% 42.8% 41.0% +26. 1.9 +0.2		
mPFS p-value	<0.0	<0.001		not published		<0.0001		<0.000001	
mPFS Hazard Ratio	0.2	0.26		0.32		0.31		0.49	
Median Overall Survival (mOS) (mo.) mOS p-value	<b>9.3 +2.7 6.6</b> (0.001 0.65		8.4 +2.2 6.2 not published		<b>8.8 +2.5 6.3</b> 0.0002		<b>6.4 +1.4 5.0</b> 0.0052		
mOS Hazard Ratio	0.6	5	0.56	)	0.55		0.77		

- 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 High VEGFR selectivity - lower off-target AEs & more tolerable



2	<b>J</b>			-		
Third-Line Metastatic Colorectal cancer	Fruquintinib FRESCO		Regorafenib CONCUR		Fruc	
<u>≥G3 AEs in &gt;4% of Patients</u>	Mainlar	nd China	Chinese Patients (Mainland China, Hong Kong, Taiwan) <sup>[1]</sup>		BIOC On-	
Treatment arms	Fruquintinib	Placebo	Regorafenib	Placebo	V	
Patients (n)	278	138	112	60	VI	
					VI	
≥G3 AE (Safety population)	61.1%	19.7%	69.6%	46.7%	Off-	
SAE (Safety population)	15.5%	5.8%	31.3%	26.7%	R	
VEGFR on-target related AEs:					FC	
Hypertension $\geq$ G3	21.2%	2.2%	12.5%	8.3%	(-	
	10.8%	0.0%	17.0%	0.0%	PI R/	
Hand Foot Syndrome (Palmar-plantar), ≥G3	10.0%	0.0%	17.0%	0.0%	B.	
Off-target (i.e. non-VEGFR) related AEs:					B	
Hypophosphatemia, ≥G3	0.0%	0.0%	8.0%	0.0%		
Hypokalemia, ≥G3	0.7%	0.7%	6.3%	0.0%	Dog	
Rash/desquamation, ≥G3	0.0%	0.0%	4.4%	0.0%	Reg	
Lipase increase, ≥G3	0.0%	0.0%	6.3%	1.7%	$\rightarrow$	
Hepatic function (Liver function) AEs:						
ALT increased, $\geq$ G3	0.7%	1.5%	7.1%	3.3%	$\rightarrow$	
AST increased, $\geq$ G3	0.4%	0.7%	8.9%	0.0%	ST	
Blood bilirubin increased, ≥G3	1.4%	1.5%	8.9%	8.3%	Ini	
NOTE: Baseline Characteristics Liver metastasis	66.5%	73.9%	na	na	$\mathbf{\hat{\lambda}}$	
Tolerability:					Se	
AE Leading to dose interruption	35.3%	10.2%	68.8%	25.0%	tr	
AE Leading to dose reduction	24.1%	4.4%	23.2%	0.0%	ar by	
AE Leading to treatment discontinuation	15.1%	5.8%	14.3%	6.7%	se	

		0.0%	8.0%	0.0%	
Pagarafanih liyer tayisity Plack		0.0%	6.3%	0.7%	
Regorafenib liver toxicity Black-		0.0%	4.4%	0.0%	
→ Increased liver function test mo		1.7%	6.3%	0.0%	
elevated) & remedial dose inte					
→ 3L CRC China - 65-75% liver met		3.3%	7.1%	1.5%	
STIVARGA (regorafenib) tablets, oral		0.0%	8.9%	0.7%	
Initial U.S. Approval: 2012	人	8.3%	8.9%	1.5%	
WARNING: HEPATOTOX See full prescribing information for comp Severe and sometimes fatal hepatotoxicity has be	·	na	na	73.9%	_
Severe and sometimes fatal hepatotoxicity ha					

ruquintinib far more selective than regorafenib					
BIOCHEMICAL ACTIVITY	Fruquintinib IC <sub>so</sub> (nmol/L)	Regorafenib IC <sub>so</sub> (nmol/L)			
On-Target Kinases:					
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				

#### -box warning:

- onitoring (weekly if rruption.
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plete boxed warning. een 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)

37 [1] Efficacy & safety of regorafenib monotherapy in Chinese patients with previously treated metastatic colorectal cancer: subgroup analysis of the CONCUR trial; R Xu

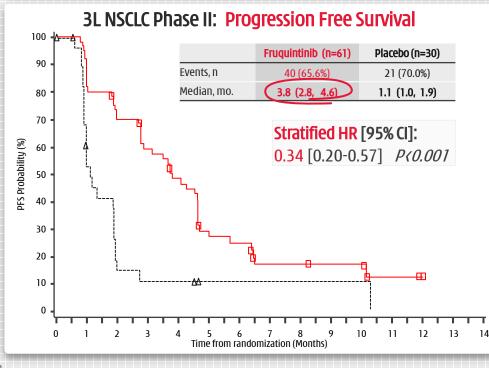
## **Fruquintinib – FALUCA Phase III in 3L NSCLC** Phase III enrolment complete (n=527); top-line results Q4 2018



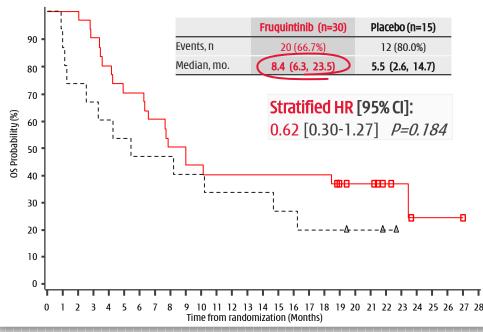
#### Non-small cell lung cancer ("NSCLC") Phase II PoC Results

- ✓ 91 <u>3<sup>rd</sup> line only</u> pts. **enrolled in ∽9 months** (Jun'14-Mar'15).
- Clearly met primary PoC endpoint of reduction in risk of progression.
- ✓ AEs consistent with the known safety profile and generally superior versus ≥3L colorectal cancer Phase III with lower >Gr.3 AEs (32.8% vs. 61.1%) and dose reductions (13.1% vs. 24.1%).

Phase III FALUCA study enrolment completed in February 2018.



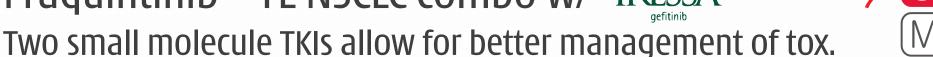
Patients, %	Fruquintinib (n=61)	Placebo (n=30)
All AEs, any grade	61 (100%)	27 (90.0%)
All AEs, grade $\geq$ 3	20 (32.8%)	6 (20.0%)
Hypertension, grade $\geq$ 3	5 (8.2%)	1 (3.3%)
Hand-foot syndrome ("HFS"), grade $\geq$ 3	3 (4.9%)	0
All other AEs, grade $\geq$ 3 (each)	≤2 (≤3.3%)	0
Leading to dose interruption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



#### 3L NSCLC Phase II: Overall Survival [1]

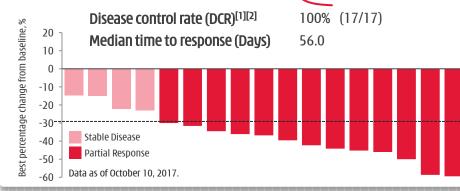
[1] EGFR Mutation positive (n=45)

### Fruquintinib - 1L NSCLC combo w/ IRESSA gefitinib





1. Promising efficacy in first-line –  $\frac{6\% \text{ ORR}}{13/17}$ .<sup>[1,2,3]</sup>



#### 2. Prelim. Safety data: fruquintinib vs. other VEGFRis.

Adverse Events ("AEs")	Iressa® or Tarceva® FLAURA <sup>[5]</sup> N = 277, n (%)	Avastin® + Tarceva® <sup>[6]</sup> N = 75, n (%)	Fruquintinib + Iressa® N = 26, n (%) <sup>[3]</sup>
All AEs, any grade	273 (98%)	≥74 (≥99%)	23 (89%)
All AEs, Grade ≥3	124 (45%)	68 (91%)	8 (31%)
AEs leading to death	6 (2%)	0 (0%)	0 (0%)
AEs leading to VEGFRi discontin.	NA	31 (41%)	1 (4%)
Grade ≥3 AEs:			
Liver function (e.g. ALT, AST incr.)	33 (12%)	6 (8%)	6 (23%)
Hypertension	NA	45 (60%)	1 (4%)
Proteinuria	NA	6 (8%)	1 (4%)
Rash	13 (5%)	19 (25%)	0 (0%)
Decreased appetite	22 (8%)	1 (1%)	NA

3. Combination of highly selective TKIs vs. MAbs: daily dose flexibility improves tolerability. This enables maintained drug exposure, leading to more durable response.<sup>[2,3]</sup>

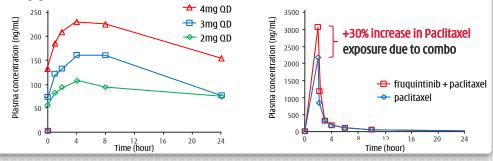
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0 28	56	84	112	140	168 1	96 224	2
Data as of Octobe	r 10. 2017.	D	uration of T	reatment (day	S)		

[1] Best tumor response for efficacy evaluable patients (patients who had both baseline and post-baseline tumor assessments); ORR = objective response rate; [2] Four PRs not yet confirmed at the time of data cut-off date; [3] Lu, S., et al, "A Phase II study of fruquintinib in combination with gefitinib in stage IIIb/IV NSCLC patients harboring EGFR activating mutations", ID 10907 IASLC 18<sup>th</sup> World Conference on Lung Cancer, Yokohama, Japan, October 15-18, 2017; [4] Drug discontinuation due to Grade 3 proteinuria and Grade 3 OTC prolonged; [5] Ramalingam S. et al, "LBA2\_PR Osimertinib vs standard of care (SoC) EGFR-TKI as first-line therapy in patients (pts) with EGFRm advanced NSCLC: FLAURA", ESMO 2017 Congress, Madrid, Spain, September 9, 2017; [6] Seto, T., et al, "erlotinib alone or with bevacizumab as first-line therapy in patients with advanced non-squamous non-small-cell lung cancer harbouring EGFR mutations (J025567); an open-label, randomised, multicenter, phase 2 study", The Lancet 2014, 15 (11) 1236-1244.

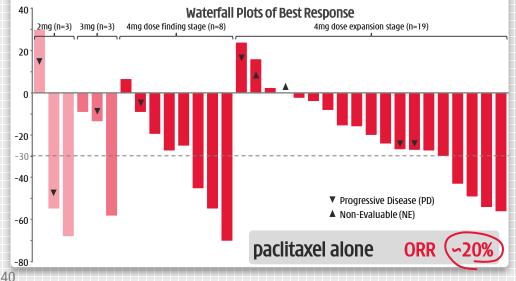
## Fruquintinib – Gastric combo with paclitaxel Phase III initiated October 2017



1. Dose proportional increase of fruquintinib AUC at steady state. Over 80% increase in paclitaxel drug exposure (mean AUC<sub>0-8</sub>) following multiple dose fruquintinib.



2. ORR of 36% (10/28) & DCR of 68% in efficacy evaluable pts. Fruquintinib 4mg,  $\geq 16$  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/m <sup>2</sup>			
	<b>Drug interruption</b>	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%)



## Sulfatinib

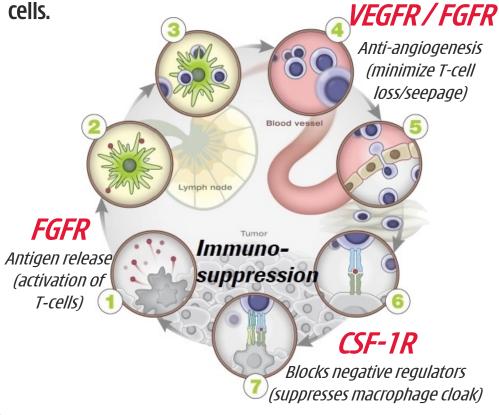
A highly active TKI with a unique angio-immuno Mechanism of Action



# Sulfatinib's unique angio-immuno kinase profile Multi-indication 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 production of macrophages (CSF-1R) which cloak cancer



47

#### Activity 1: Fast/first approval in China for all NET <sup>[1]</sup> 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); $\leq$ 2 prior systemic drugs.				
# of Sites	20-30 (China)				
# of Patients	~195				
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 / 2019 December 2015 / 2019				

#### Activity 2: Global development

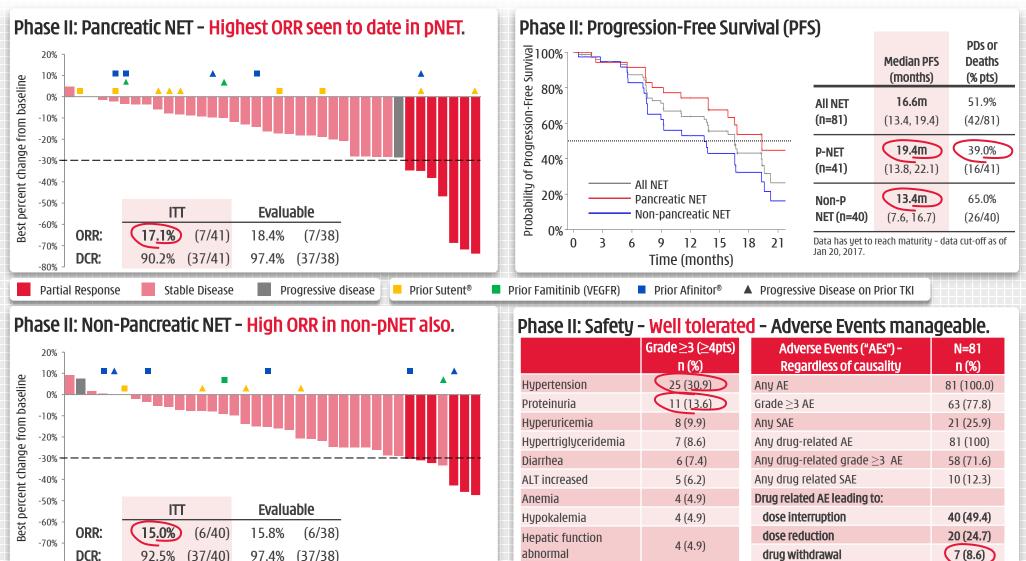
- U.S. Phase I bridging in Caucasian patients almost complete RP2D<sup>[3]</sup> expected to be similar to China – 300-400mg QD.
- U.S. Phase II in planning, focusing on areas of NET and solid tumors unmet medical need/BTD<sup>[4]</sup> opportunity.

#### 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<sup>[1]</sup>)* Efficacy in all NET; & patients who failed on Sutent<sup>®</sup>/Afinitor<sup>®</sup>





[1] ENETS = European Neuroendocrine Tumour Society. Data cut-off as of Jan 20, 2017. 43

(37/40)

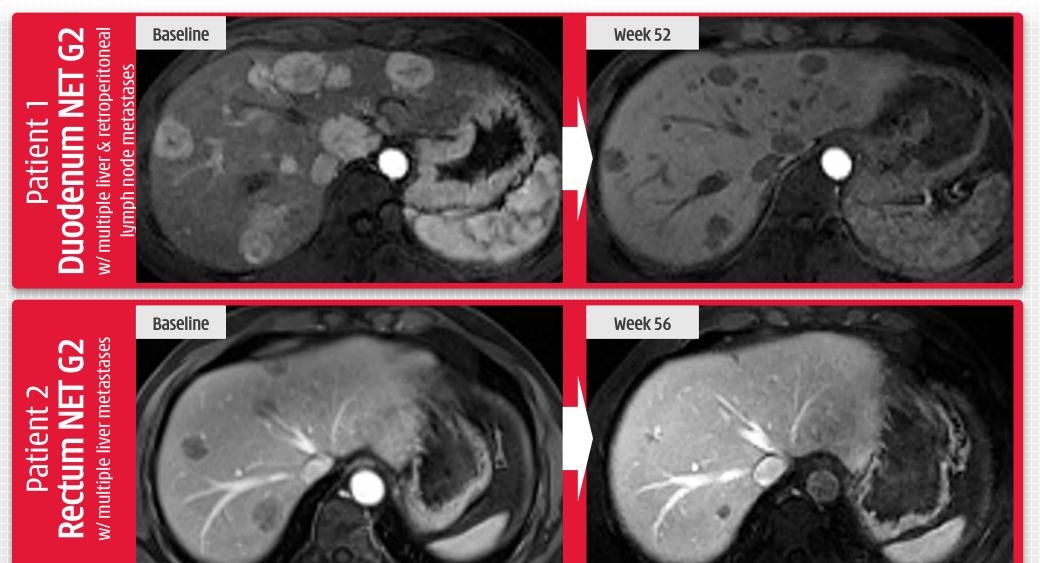
97.4%

92.5%

-80%

## Activity 1: China NET – Phase II *(ENETS 2017*<sup>[1]</sup>) Tumor devascularization & central necrosis







# Epitinib

*EGFR mutation kinase inhibitor that penetrates the blood-brain barrier Entering Phase III trials* 



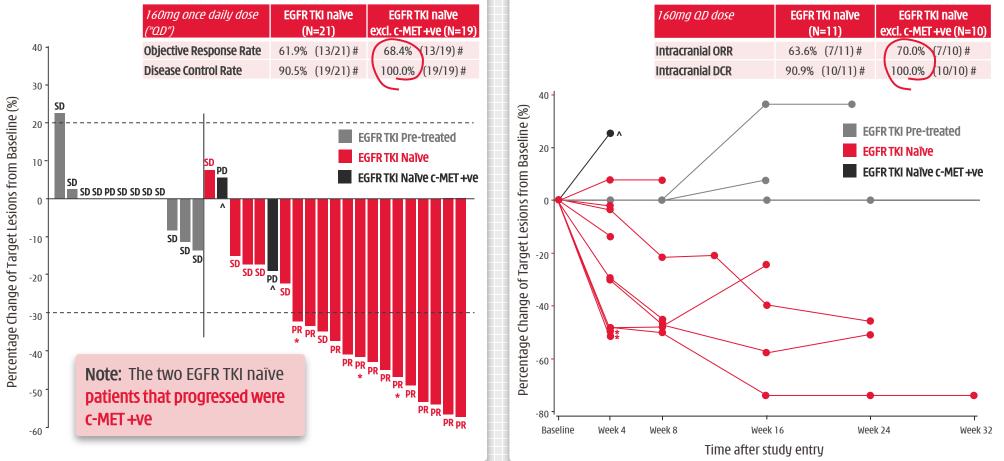
## **Epitinib – 70% response in NSCLC w/ brain mets**<sup>[2]</sup> Unmet medical need for ~50% of NSCLC patients w/ brain mets<sup>[1]</sup>



2. Phase Ib<sup>[2]</sup> - solid/durable efficacy in brain in EGFRm+

NSCLC patients with measurable brain mets (>10mm).

1. Phase Ib <sup>[2]</sup> – epitinib monotherapy in EGFRm+ NSCLC patients – <u>efficacy in lung</u> in-line with Iressa®/Tarceva®

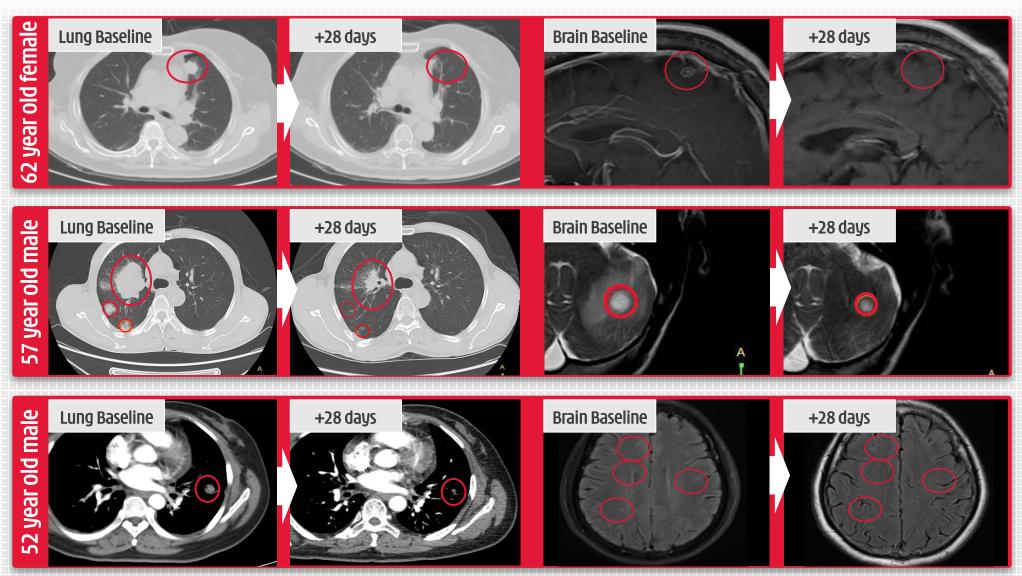


[1] Li B, Bao YC, Chen B, et al. Therapy for non-small cell lung cancer patients with brain metastasis. Chinese-German J Clin Oncol, 2014, 13: 483–488; [2] Dose expansion stage – data cut-off 20 Sept, 2016; \* Unconfirmed PR, due to no further assessment at cut-off date; # Includes both confirmed and unconfirmed PRs; ^ c-MET amplification/high expression identified

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## Epitinib – Strong PoC efficacy – 160mg QD dose



## **Epitinib – Safe & well tolerated** Pivotal Phase III study to initiate in 2018



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

<b>Dose Escalation Stage (n=35*)</b> (Drug related AEs reported >10%)			<b>Dose Expansio</b> (Drug related AE		
160mg QD dose	All Grades n (%)	Grade 3/4 n (%)	160mg QD dose	All Grades n (%)	Grade n (S
Skin rash	21 (60.0%)	1 (2.9%)	Skin rash	31 (83.8%)	2 (5.
Diarrhea	12 (34.3%)	-	Hyper-pigmentation	18 (48.6%)	1 (2.
AST increase	12 (34.3%)	1 (2.9%)	ALT increase	15 (40.5%)	7 (18
ALT increase	11 (31.4%)	1 (2.9%)	AST increase	15 (40.5%)	4 (10
Total bilirubin increase	10 (28.6%)	2 (5.7%)	ASP increase	11 (29.7%)	1 (2.
Stomatitis	5 (14.3%)	-	Diarrhea	10 (27.0%)	-
Exfoliative dermatitis	5 (14.3%)	-	Proteinuria	10 (27.0%)	-
Pruritus	5 (14.3%)	-	Total bilirubin increase	9 (24.3%)	1 (2.
Hyper-pigmentation	4 (11.4%)	-	Hyperuricemia	9 (24.3%)	2 (5.4
Gamma-GGT increase	4 (11.4%)	2 (5.7%)	Gamma-GGT increase	7 (18.9%)	4 (10
Conjugated bilirubin	4 (11.4%)	1 (2.9%)	Stomatitis	6 (16.2%)	-

#### 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 WCLC 2016.
  - ↗ China FDA Phase III clinical trial cleared in July 2016.
  - ↗ Finalizing dose in early 2018 (120mg vs.160mg QD), then initiating Phase III.
- EGFR gene amplified Glioblastoma (primary brain tumors):
   Phase Ib/II proof-of-concept underway.

## CASE STUDY – EGFR-TKI naïve patient

- Male, 46, diagnosed with Stage IV NSCLC adenocarcinoma (Exon21)
- Metastases in the brain, meninges, & bone
- 1<sup>st</sup>-line chemo naïve
- 120mg QD dosage

7%)

.9%

7%)

.4%) ).8%]  25 weeks (177 days) on treatment with clear response in multiple measurable (>10mm diameter) brain lesions



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[1] No Dose Limiting Toxicity ("DLT") was observed in any cohort; \* One patient did not join multiple dosing.

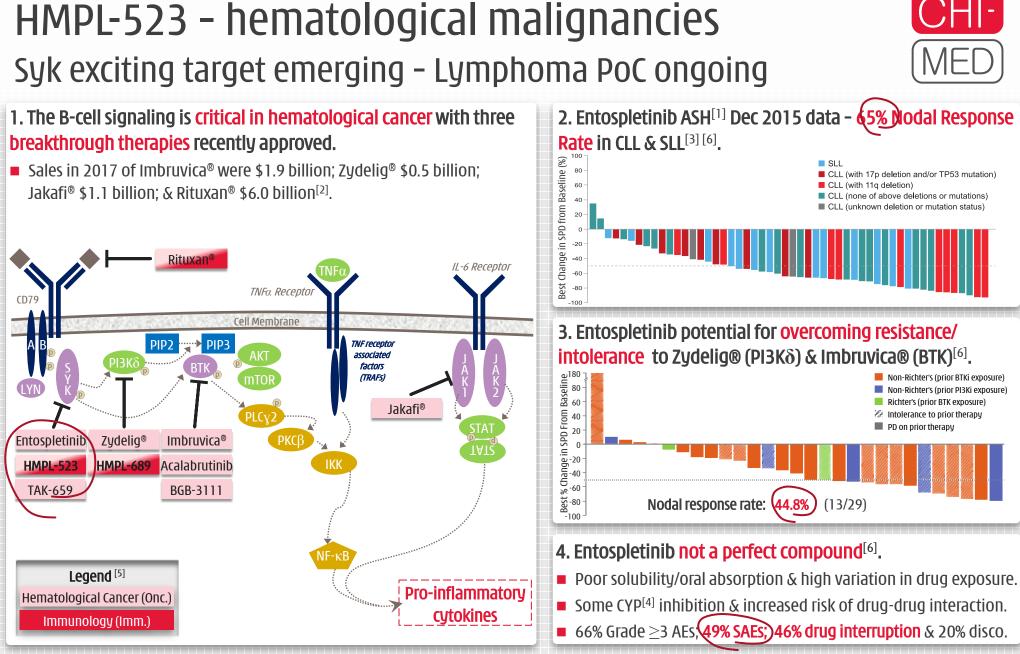
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## Additional Clinical Candidates HMPL-523 - potential first-in-class Syk inhibitor, Theliatinib, HMPL-689, HMPL-453 & HM0046599...





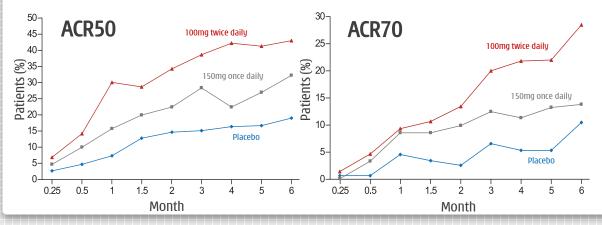


-[1] ASH = American Society of Hematology; [2] Rituxan<sup>®</sup> 2017 sales in oncology only; [3] Chronic lymphocytic leukemia ("CLL") & small lymphocytic lymphoma ("SLL"); [4] CYP3A4, CYP2D6 and CYP 1A2; [5] Approved Drug = ®; All others are clinical candidates; [6] Sharman et al, ASH Meetings 2015 & 2016.

## HMPL-523 – immunology potential Superior selectivity, better target coverage & efficacy vs. fosta.



**1. Fostamatinib good Phase II**<sup>[1]</sup> **RA**<sup>[2]</sup> **dose response**...



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

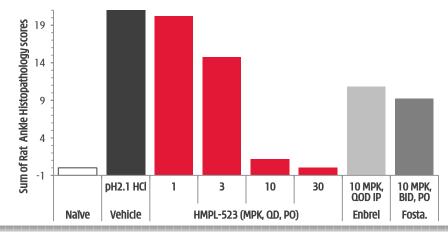
Percent of patients	Placebo (n = 153)	150mg QD (n = 152)	100mg BID (n = 152)
Diarrhea	3.0%	11.8%†	19.1%†
Upper respiratory infection	7.1	7.2	14.5 †
Jrinary tract infection	4.6	3.3	5.9
Nausea	4.6	5.9	4.6
Neutropenia	0.7	6.6 †	5.9 †
Headache	5.2	6.6	5.9
Abdominal pain	2.6	6.6 †	5.9 †
ALT > 3x ULN	2.0	3.9	3.9
Dizziness	2.0	2.6	4.6
Hypothyroidism	2.6	2.6	3.3
Cough	2.6	2.0	3.3
+ D / O OE for compariso	with placobo aro	ALT - ALT	aminotransfora

 $\dagger P < 0.05$  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.

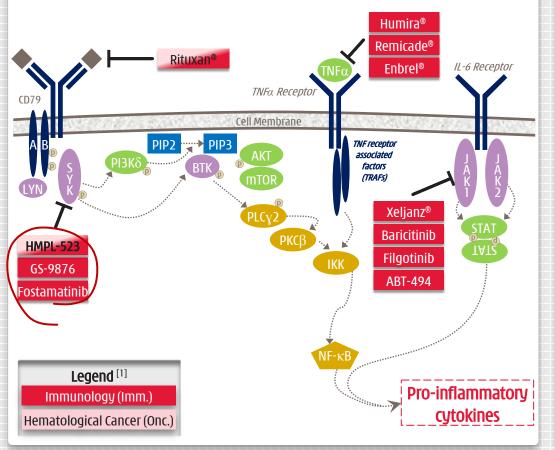


[1] Fostamatinib is a prodrug of the SYK inhibitor R406 - Phase II study data per N ENGL J MED 363;14; \*: HMPL data and Eun-ho Lee, 2011; \*\* Birth Defects Research (Part A) 2009, 85: 130-6; [2] RA = Rheumatoid Arthritis; [3] QD = one dose per day; BID = two doses per day; QOD = one dose every other day; PO = by mouth (i.e. orally); IP = by Intraperitoneal injection; Naïve = model score without induced arthritis.

## HMPL-523 – immunology potential Potential first-in-class Syk TKI in immunology – Ph.II in planning



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



2. RA expected to be a **\$45 billion**<sup>[2]</sup> market in 2020 with B-cell pathway; anti-TNF; & JAK the main focus.

(Methotrexate-IR: placebo adjusted)	ACR20	ACR50	ACR70	<b>2017 Sales</b> ( <b>\$ billion)</b> <sup>[3]</sup>
B-Cell receptor mAbs				
Rituxan® (24-Week)	33%	21%	11%	1.6
Anti-TNFα/NF-κB mAbs				
Humira® (24-Week)	33%	29%	18%	18.4
Remicade® (24-Week)	30%	22%	8%	6.3
Enbrel® (24-Week)	44%	36%	15%	7.9
JAK Inhibitors Small molecules				
Xeljanz® (24-Week)	25%	23%	13%	1.3
Xeljanz® (12-Week)	28%	21%	8%	1.5
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.

[1] Approved drug = (a); All other clinical candidates: mAb = antibody (extracellular); small molecule (intracellular); [2] Frost & Sullivan; [3] 2017 sales in immunology only.

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# Theliatinib – encouraging activity observed Potent & highly selective TKI – strong affinity to EGFRwt kinase



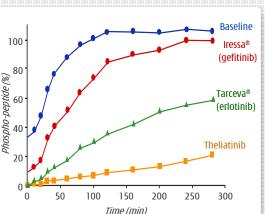
# 1. Major unmet medical need for wild-type EGFR activation tumors.

- 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.

	indennag in e		Iressa®, Tarceva®
Tumor Types	Wild-type: Gene Amplification	Wild-type: Over Expression	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- <u>84%</u>	42% (EGFR variant III)
			MAbs approved: Erbitux®, 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.



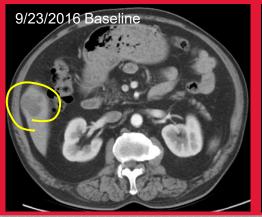
# 3. Esophageal cancer (EC): No effective treatment options.

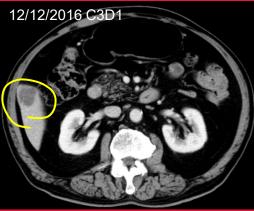
 Major issue in Asia with poor prognosis: 5 year survival 10-20%



### 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).





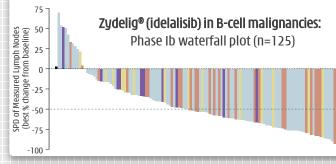
KIs = tyrosine kinase inhibitors; MAbs = monoclonal antibodies. [1] GLOBOCAN 2012 (http://globocan.iarc.fr/) and Chen W et al. Cancer statistics in China, 2015. CA Cancer J Clin. 2016; 66:115-132

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# HMPL-689 – Phase I Australia & China started Designed to be a best-in-class inhibitor of PI3K $\delta$



- 1. PI3K $\delta$  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  $\text{PI3K}\delta$  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. PI3K $\delta$ inhibitors being developed in a very broad range of indications.

Compound		Indication	Status	Issue	
Zydelig® (idelalisib) PI3Kô	Gilead	Chronic lymphocytic leukaemia, non-Hodgkin's lymphoma	Marketed	<b>High incidence of liver</b> <b>toxicity</b> seen with idelalisib (150mg bid)	
АМG-319 РІЗК <b>б</b>	Amgen	B-cell  lymphoma, non-Hodgkin's lymphoma, T-cell lymphoma, chronic lymphocytic leukaemia	Phase I Trial		
duvelisib		Relapsed or refractory chronic lymphocytic leukaemia / small lymphocytic lymphoma	Phase III Trial (NDA submitted)	<b>Need to spare PI3K</b> γ serious infection seen	
(IPI-145) ΡΙ3Κγ/δ	Verastem/ Infinity <sup>[1]</sup>	Refractory indolent non-Hodgkin lymphoma	Phase II Trial (NDA submitted)	with duvelisib due to strong immune	
		Peripheral T-cell lymphoma	Phase I Trial	suppression	
Aliqopa <sup>®</sup> (copanlisib) PI3Kα/δ	Bayer	Relapsed follicular B-cell non-Hodgkin lymphoma	Approved <sup>[2]</sup>	Serious and fatal infections and AEs	

#### 4. More potent / more selective than Zydelig<sup>®</sup>, duvelisib & Aliqopa<sup>®</sup>.

Enzyme IC <sub>50</sub> (nM)	HMPL-689	Zydelig®	duvelisib	Aliqopa®
РІЗКδ	0.8 (n = 3)	2	1	0.7
PI3Kγ (fold vs. PI3Kδ)	114 (142x)	104 <mark>(52x)</mark>	2 ( <u>2X</u> )	6.4 <b>(9x)</b>
PI3Kα (fold vs. PI3Kδ)	>1,000 (>1,250x)	866 <b>(433x)</b>	143 <b>(143x)</b>	0.5 (1X)
PI3K $\delta$ human whole blood CD63+	3	14	15	n/a
PI3Kβ (fold vs. PI3Kδ)	87 <mark>(109x)</mark>	293 <b>(147x)</b>	8 (8X)	3.7 <b>(5x)</b>

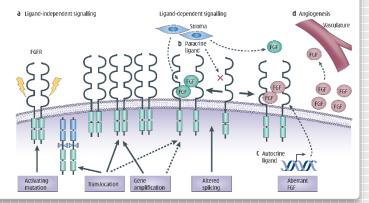
[1] AbbVie ended collaboration with Infinity in June 2016 following Phase II results in indolent non-Hodgkin's lymphoma. Duvelisib now licensed to Verastem; [2] Accelerated approval was granted based on ORR, and continued approval may be contingent upon verification and description of clinical benefit in a confirmatory trial.

## HMPL-453 – Phase I Australia & China underway Designed as first-in-class FGFR1/2/3 inhibitor



#### 1. FGFR genetic alterations are oncogenic drivers.

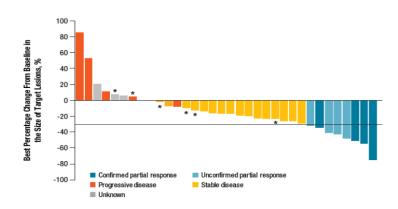
- FGF/FGFR signaling normally involved in embryonic development, tissue repair, angiogenesis, neuroendocrine and metabolism homeostasis.
- Multiple oncogenic driver genetic alterations in FGFR pathway: gene amplification, mutation, translocation, fusion, splicing, etc.



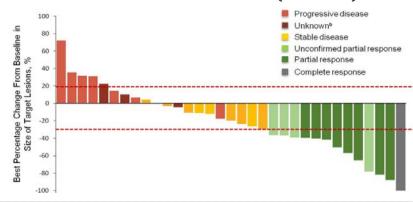
## 2. FGFR – diverse & complicated genetic changes with multiple tumor types harboring low incidence.

	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	<b>Gastric</b> (5~10%) <b>Breast</b> (4%)	Intra-hepatic biliary tract cancer (cholangiocarcinoma) (14%) Breast (n/a)	Endometrial (12~14%) Lung squamous (5%)			
FGFR3	<b>Bladder</b> (n/a) <b>Salivary adenoid cystic</b> (n/a)	Bladder (3~6%); Lung squamous (3%); Glioblastoma (3%) Myeloma (15~20%)	Bladder (60~80% NMIBC; 15~20 MIBC) Cervical (5%)			

- 3. Biliary Tract Cancer (cholangiocarcinoma) and bladder cancer have made much progress in clinic to date.
  - BGJ398 Phase II PoC in biliary tract cancer (2016 ASCO GI).



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



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## **China Commercial Platform** *Providing cash generation to fund R&D in Innovation Platform Established high-performance pan-China pharma sales organization*



# Chi-Med's Commercial Platform in China

## Long track record of commercial success – good source of cash



2 National house- hold name brands	Focus on largest disease categories	Major commercial & production scale	Leadership market shares	JVs with 3 leading China Pharmas			
上药牌	Most common disease diagnosed/treated in rural hospitals <sup>[1]</sup> :	<ul> <li>∽2,300 Rx &amp; ∽1,000 OTC sales</li> <li>people in over 300<sup>[2]</sup> cities &amp; towns in China.</li> </ul>	Market leader in the sub- categories/markets in which we compete <sup>[3]</sup> :	SPH 上海医药			
(夏夏)	Cold/Flu: 86%	Drugs in ~22,500 hospitals	SXBX pill: <sup>[4][5]</sup> ~15% Rx Cardiovascular TCM	SHANGHAI PHARMA			
	Cardiovascular: 78%	detailing ~98,000 doctors.	Banlangen: <sup>[6]</sup> ~53%				
(BEU)	Diabetes: 46%	Sold ~4.6 billion doses of	FFDS tablet <sup>[7]</sup> ~38%				
$\sim$	GI: 45%	medicine in 2017.	OTC Angina TCM	SINOPHARM			

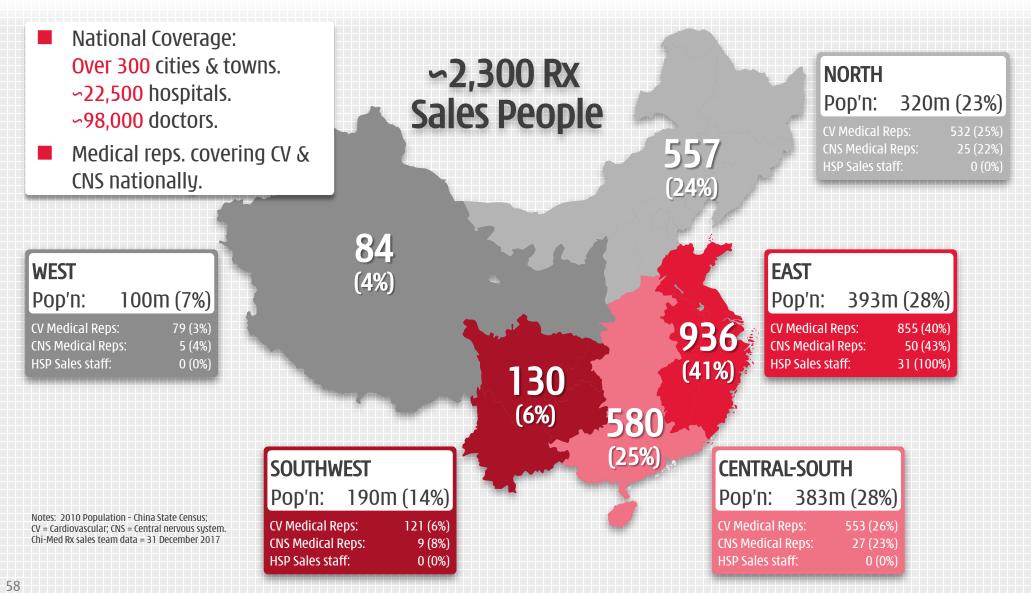
#### Commercial Platform Performance - 2003-2017<sup>[8][9]</sup>

					IFR	S							US GAAP			16-17
(US\$ millions)	03	04	05	06	07	08	09	10	11	12	13	14	15	16	17	Growth
Sales (Non-GAAP)	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	677.2	8%
Prescription Drugs	17.2	21.8	23.3	23.2	28.1	<i>39.5</i>	54.4	71.2	92.4	116.5	138.2	204.9	286.6	372.3	411.0	10%
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	266.2	4%
Total % Growth	n/a	27%	133%	56%	17%	31%	26%	20%	18%	29%	n/a	16%	11%	21%	8%	
Not (loss) (loss providentes) (Non CAAD)	(10.7)	(2.0)	2.2	17	11.5	147	21.5	27.0	20.1	22.1	20.7	40.0	F.4.1	<b>62 2</b> [11]	<b>77.3</b> <sup>[12]</sup>	22%
Net (loss)/income after tax (Non-GAAP)	(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	<b>63.3</b> <sup>[11]</sup>	11.3	22%
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	41.4	53.0	28%
Consumer Health	(10.3)	(4.9)	0.3	5.4	<i>9.3</i>	11.9	15.5	16.0	1 <i>5.9</i>	15.4	17.3	22.3	22.2	21.9	24.3	11%
% 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%	10.1%	11.4%	
					[10]	[10]	[10]	[10]	[10]	[10]	[10]	[10]	[10]	[11]		$\frown$
Net (loss)/income attrib. to Chi-Med	(5.7)	(3.7)	(0.5)	1.2	<b>4.5</b> <sup>[10]</sup>	<b>5.9</b> <sup>[10]</sup>	<b>9.3</b> <sup>[10]</sup>	<b>12.6</b> <sup>[10]</sup>	<b>13.6</b> <sup>[10]</sup>	<b>14.6</b> <sup>[10]</sup>	18.2 <sup>[10]</sup>	<b>22.8</b> <sup>[10]</sup>	<b>25.2</b> <sup>[10]</sup>	<b>29.9</b> <sup>[11]</sup>	37.5 [12]	2 <u>5%</u>
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	1 <i>5.9</i>	20.7	26.5	28%
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	<i>9.3</i>	9.2	11.0	20%
Total % growth	n/a	-35%	-86%	340%	275%	31%	58%	35%	8%	7%	n/a	26%	10%	19%	25%	

[1] Frost & Sullivan; [2] 300 cities & towns covered by Prescription Drug Business and 600 cities & towns including OTC business; [3] Frost & Sullivan 2017 market share data; [4] China coronary heart disease oral Chinese patented drugs market share; [5] She Xiang Bao Xin Pill ("SXBX pill"); [6] Banlangen Granules ("Banlangen") - OTC Antiviral; [7] Fu Fang Dan Shen tablets ("FFDS"); [8] 2003-2006 incl. disco. operation; [9] Prescription Drugs includes SHPL and Hutchison Sinopharm; and Consumer Health includes HBYS, HHO, HHL, and HCPL - please see appendix "Non-GAAP Financial Measures and Reconciliation"; [10] Continuing Operations; [11] Excluded the land compensation from SHPL of US\$80.8 million net income after tax and US\$40.4 million net income after tax and net income attributable to Chi-Med; [12] Excluded SHPL's R&D related subsidies of US\$5.0 million and \$2.5 million at income after tax and net income attributable to Chi-Med; [12] Excluded SHPL's R&D related subsidies of US\$5.0 million at \$2.5 million at income after tax and net income attributable to Chi-Med; [12] Excluded SHPL's R&D related subsidies of US\$5.0 million at \$2.5 million at income after tax and net income attributable to Chi-Med; [12] Excluded SHPL's R&D related subsidies of US\$5.0 million at income after tax and net income attributable to Chi-Med; [12] Excluded SHPL's R&D related subsidies of US\$5.0 million at income after tax and net income attributable to Chi-Med; [12] Excluded SHPL's R&D related subsidies of US\$5.0 million at income after tax and net income attributable to Chi-Med; [12] Excluded SHPL's R&D related subsidies of US\$5.0 million at income after tax and net income attributable to Chi-Med; [12] Excluded SHPL's R&D related subsidies of US\$5.0 million at income after tax and net income attributable to Chi-Med; [12] Excluded SHPL's R&D related subsidies of US\$5.0 million at income after tax and net income attributable to Chi-Med; [12] Excluded SHPL's R&D related subsidies of US\$5.0 million at income after tax and net income attrib

## A powerful Rx Commercial Platform in China.... Chi-Med management run all day-to-day operations





## ....and highly adaptable Sales of Seroquel<sup>®</sup> & Concor<sup>®</sup> up significantly since we took over

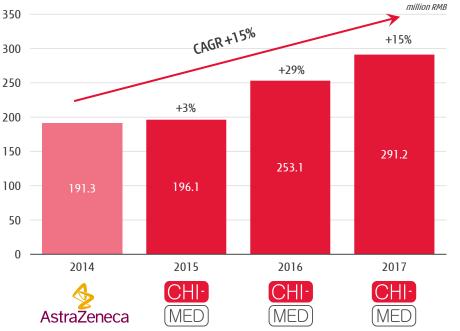


Seroquel XR 400 mg

Seroquel<sup>®</sup>, or quetiapine, is a second generation antipsychotic approved for the treatment of schizophrenia, bipolar disorder and as adjunct treatment of major depressive disorder. Chi-Med holds exclusive all China commercial rights – full service commercial role (fee-for-service<sup>[1]</sup>).

Took over from AZ Apr-2015<sup>[2]</sup>.

New team of ~120 CNS reps built from scratch.

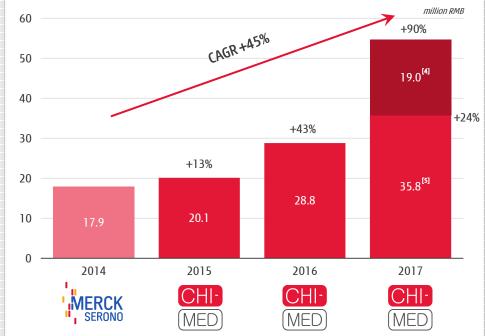


[1] In October 2017, as a result of the new CFDA Two-Invoice policy, the Seroquel® operating model changed to a "fee-for-service" model vs. the prior model in which Chi-Med consolidated the sales of Seroquel® -- the change has no impact on net income earned; [2] 2014 full year and Q1 2015 were managed by AstraZeneca. Chi-Med took over commercial function for Seroquel across all-China in Apr 2015.



Concor<sup>®</sup>, or bisoprolol hemifumarate, is a beta-blocker approved for the treatment of hypertension.

- Chi-Med runs six core territories w/ 360 mn. people - full service commercial role (fee for service).
- Took over from MS Jan-2015<sup>[3]</sup>.
- Leverages SHPL's existing >2,100
   CV medical reps.



[3] 2014 full year was managed by Merck Serono. Chi-Med took over commercial function for Concor in 3 original territories on fee-forservice basis in Jan 2015; [4] Sales into 3 new territories (Tianjin, Anhui and Jiangsu) were added from 2017: RMB 19.0 million; [5] 3 original territories (Shandong, Henan and Shanghai) contributed RMB 35.8 million in 2017 (+24.3%).

# Deep portfolio of household name drugs



Top 7 products represent 62% of sales<sup>[1]</sup> and 90% of gross profit<sup>D</sup> (MEC

Main Produc	ts <sup>[2]</sup> – SALES (Non-GAAP)	2011	2012	2013	2014	2015	2016	2017
·····································	<i>SXBX pill</i> Coronary artery disease (Rx) 15.4% National market share Patent expiry 2029	<b>79,438</b> +32%	102,215 <i>+29%</i>	<b>123,587</b> +21%	<b>138,848</b> +12%	<b>159,326</b> +15%	<b>195,371</b> <i>+23%</i>	<b>209,246</b> +7%
	<b>Banlangen granules</b> Anti-viral/flu (OTC) 53% National market share	57,278 +8%	<b>65,381</b> +14%	<b>72,300</b> +11%	<b>55,573</b> <i>-23%</i>	<b>54,793</b> - <i>1%</i>	<b>56,664</b> +3%	<b>59,898</b> +6%
	<b>FFDS tablet</b> Angina (OTC) 38% National market share	<b>57,001</b> - <i>3%</i>	<b>60,181</b> +6%	<b>69,996</b> +16%	<b>76,297</b> +9%	<b>60,154</b> -21%	<b>59,906</b> <i>0%</i>	<b>58,936</b> -2%
Seroquel XR and	<i>Seroquel tablets</i> Bi-polar/Schizophrenia (Rx) 6% National market share	n/a	n/a	n/a	n/a	21,131	<b>34,380</b> <i>+63%</i>	[3] <b>35,359</b> <i>+3%</i>
	<i>NXQ tablet</i> Cerebrovascular disease (Rx) Proprietary formulation	<b>3,741</b> +55%	<b>6,933</b> <i>+85%</i>	<b>10,142</b> +46%	<b>14,681</b> +45%	<b>17,581</b> +20%	<b>21,000</b> +19%	<b>20,408</b> - <i>3%</i>
	<i>KYQ granules</i> Periodontitis (OTC) >90% National market share	<b>15,412</b> +22%	16,351 +6%	<b>16,318</b> <i>0%</i>	<b>18,370</b> +13%	<b>17,051</b> - <i>7%</i>	<b>17,210</b> +1%	<b>17,620</b> +2%
	<b>Danning tablet</b> Gallbladder/stone (Rx) Patent expiry 2027	<b>9,914</b> +22%	<b>11,648</b> +17%	12,364 +6%	<b>13,822</b> +12%	<b>13,526</b> -2%	<b>9,041</b> <i>-33%</i>	<b>16,089</b> <i>+78%</i>

[1] Based on aggregate Non-GAAP sales (refer to page 57) and gross profit of consolidated subsidiaries and non-consolidated joint ventures of Commercial Platform, please see appendix "Non-GAAP Financial Measures and Reconciliation"; [2] Rx = prescription drug; OTC = over-the-counter drug; SXBX pill = She Xiang Bao Xin pill; FFDS tablet = Fu Fang Dan Shen tablet; NXQ tablet = Nao Xin Qing tablet; KYQ granules = Kou Yan Qing granules; Market Shares according to Frost & Sullivan or QuintilesIMS; [3] From October 2017, the majority of sales changed to a fee-for-service model due to the CFDA Two-invoice policy.

(US\$'000) (Growth % vs. Year Ago)



# **Appendices**



## Experienced pharma management team

POSITION	EXPERIENCE (yrs) Industry / Chi-Med	ROLE / BACKGROUND
CHRISTIAN HOGG, BSc, MBA <i>Chief Executive Officer</i>	Procter & Gamble 29/18	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> 28/13	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 29 / 10	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 <i>SVP, Clinical &amp; Regulatory Affairs</i>	NOVARTIS Celegene 19/4	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 24/10	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 <i>SVP, Bus. Dev. &amp; Strategic Alliances</i>	Lilly 24/8	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 <i>SVP, Corp. Finance &amp; Development</i>	Credit Suisse 19/9	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.



INCIVEK



ZOMETA



# A risk-balanced global-focused BioPharma

## Innovation Platform Deep late-stage pipeline

8 oncology drug candidates in 36 studies worldwide.

- 1<sup>st</sup> positive Ph.III result fruquintinib Launch 2018<sup>[1]</sup>
- ✓ 5 Phase III trials; with 4 more pending/in planning.
- ✓ ∽360-person Scientific Team.

## **Commercial Platform** *Solid cash flow from operations*

√ 23,300-persop China Sales Team (∽2,300 med. reps).

- ✓ To commercialize Innovation Platform drugs in China.<sup>[1]</sup>
- ✓ 2017 sales (non-GAAP)<sup>[2]</sup> up 8% to \$677.2 million.
- ✓ 2017 net income<sup>[3]</sup> up 25% to \$37.5 million.<sup>[4]</sup>

[1] If approved; [2] 2017 sales (non-GAAP) represents the sum of (i) the 2017 GAAP revenue from external customers of our commercial Platform (\$205.2 million), (ii) the 2017 revenue of our non-consolidated joint venture Shanghai Hutchison Pharmaceuticals Limited ("SHPL") (\$244.6 million) and (iii) the 2017 revenue of our non-consolidated joint venture Hutchison Whampoa Guangzhou Baiyunshan Chinese Medicine Company Limited ("HBYS") (\$227.4 million). SHPL and HBYS revenues are as reported in the audited consolidated financial statements of each of these companies which are prepared in accordance with IFRS; [3] Net income attributable to Chi-Med (non-GAAP); [4] Excludes the share of a one-time gain from SHPL's R&D related subsidies (\$2.5 million), while 2016's net income excludes the share of one-time gain from SHPL's R&D related subsidies (\$2.5 million), while 2016's net income excludes the share of one-time gain from SHPL's R&D related subsidies (\$2.5 million).



## Chi-Med Group structure - major entities

**Chi-Med Group Level** 

Revenues - 2017 \$241.2m (2016: \$216.1m)

Net Loss Attributable to Chi-Med - 2017: -\$26.7m (2016 net income: \$11.7m)

Non-Consolidated Joint Ventures

Chi-Med Subsidiaries

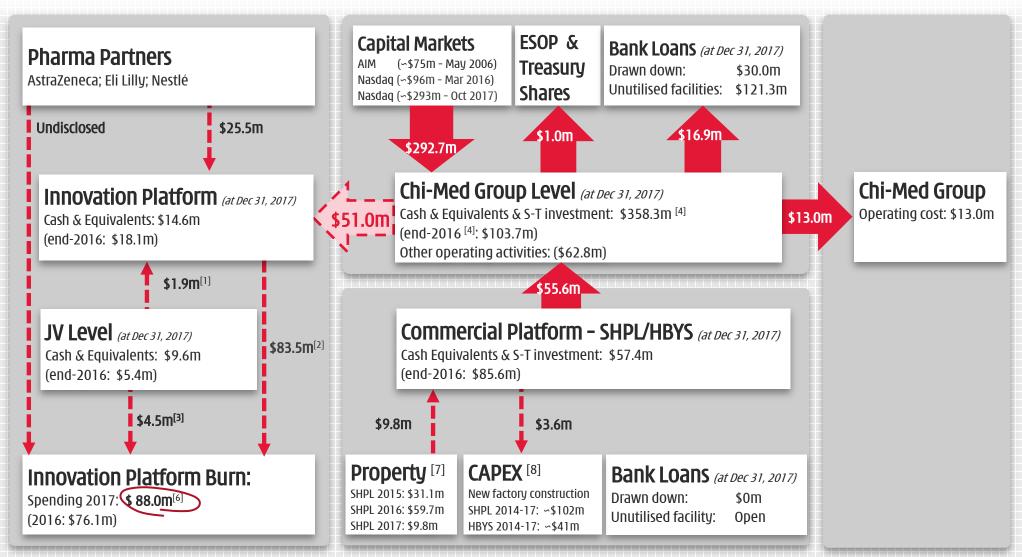
#### **Commercial Platform Innovation Platform** Sales of Subs & JVs - 2017: \$677.2m (2016: \$627.4m) Revenue - 2017: \$36.0m (2016: \$35.2m) Net Income Attributable to Chi-Med - 2017: \$37.5m (2016: \$29.9m) [1] Net Loss Attributable to Chi-Med - 2017: -\$51.9m (2016: -\$40.7m) **Prescription Drugs** 99.8% ★ 51% Shanghai Hutchison Pharma ("SHPL") Hutchison Sinopharm ("HSP")<sup>[2]</sup> Hutchison MediPharma ("HMP") Prescription Drugs Rx Drug Commercial Co. Oncology/Immunology Drug R&D Partner: Shanghai Pharma Group Partner: Sinopharm Group **Revenue: Revenue: Revenue:** 2017: \$244.6m (2016: \$222.4m) 2017: \$166.4m (2016: \$149.9m) 2017: \$36.0m (2016: \$35.2m) **Consumer Health** 50% 50% [3] ★ 50% Nutrition Science Partners ("NSP") Hutchison BYS Chinese Med. ("HBYS") Hutchison Hain Organic ("HHO") Botanical Drug /GI Disease R&D Over-the-counter Drugs ("OTC") Health Related Consumer Prods. Partner: Guangzhou Pharma Holdings Partner: Hain Celestial Group Partner: Nestlé Health Science **Revenue: Revenue: Revenue:** 2017: \$28.9m (2016: \$23.3m) 2017: \$227.4m (2016: \$224.1m) 2017: nil (2016: nil)

[1] Non-GAAP: excludes the share of land compensation and government subsidies from SHPL of US\$40.4million in 2016 and \$2.5million in 2017; [2] Excluded HSP's Zhi Ling Tong infant nutrition business; [3] Held through an 80% owned subsidiary.

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## **FY2017 Inter-group cash flow** \$358.3m cash (Dec 31, 2017); \$121.3m in undrawn bank facilities





[1] \$7.0m capital injection to NSP offset by \$8.9m service income received from NSP; [2] Including research & development cost and general & admin. expenses; [3] Share of NSP operating loss; [4] Including \$273.0m short-term investment (91-183 day deposit) as at end of 2017; [5] Cash received for SHPL land compensation (10% of total compensation) in Feb'17; [6] Please see appendix "Non-GAAP Financial Measures and Reconciliation" for a

Reconciliation of GAP to adjusted research and development expenses; [7] Cash received for SHPL land compensation; [8] CAPEX required to build new Shanghai (SHPL) and Bozhou (HBYS) factories.

65

(US\$ millions)

# Three collaborations have major aggregate financial impact



AstraZeneca





66

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

- \$135.5 million in upfront /milestone payments and equity injections as at December 31, 2017.
- **up to \$340 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<sup>[2]</sup>:

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

### Possible payment events in 2018:

- **Fruquintinib:** NDA approval for third line CRC.<sup>[3]</sup>
- Savolitinib: Start of Phase III in NSCLC.<sup>[4]</sup>

[1] 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.
 [3] CRC = Colorectal Cancer;
 [4] NSCLC = non-small cell lung cancer, and subject to regulatory discussions.



## Major market potential

		Pot. launch Year / Territory	Incidence (New pts./yr.) <sup>[1]</sup>	Approx. WAC <sup>[2]</sup> of various reference TKIs (US\$/month)	Median PFS (months) <sup>[3]</sup>	Potential Peak (US\$) <sup>[4]</sup> SalesNet Income
	Papillary renal cell carcinoma (c-Met-driven)	<b>2020/21</b> Global	~25,000	\$11,600 (Sutent®) \$10,500 (Afinitor®)	6.2 Ph.II (actual)	
	NSCLC –2L 1 <sup>st</sup> Gen EGFR TKI refract, Tagrisso combo (MET+ , T790M+/-)	<b>2020/21</b> Global	~35,000 - 40,000	\$15,100 (Tagrisso®)	TBD	
SAVO	NSCLC – 3L 3 <sup>rd</sup> Gen EGFR TKI refract. Tagrisso combo (MET+)	<b>2021</b> Global	TBD	\$15,100 (Tagrisso®)	TBD	
	NSCLC –2L 1 <sup>st</sup> Gen EGFR TKI refract, Iressa combo (MET+, T790M-)	2021 China	TBD	\$1,100 (Iressa®)	TBD	
	3L (or above) Colorectal cancer ("CRC")	2018 China	~50,000 - 60,000	\$14,000 (Regorafenib - global) \$2,870 (Apatinib - China off label)	3.7 Ph.II (actual)	<b>\$110-160m</b> @est. 20-25% penetration <sup>[5]</sup> <b>\$20-35m</b> @15-20% tier royalty/other
FRUQ	3L Non-small cell lung cancer ("NSCLC")	<b>2019</b> China	~60,000 - 70,000	No approved TKIs \$2,870 (Apatinib – China off label)	3.8 Ph.II (actual)	
	2L Gastric cancer combo with Taxol	<b>2020</b> China	~250,000 - 300,000	\$2,870 (Apatinib appr. 3L Gastric) \$1,810 (Apatinib NDRL <sup>[7]</sup> reimbursed)	3.7 Ph.II (actual)	
	Pancreatic neuroendocrine tumors	2019/20 China	∽5,000 - 6,000	\$11,000 (Sutent®/Afinitor® – global) \$5,500 (Somatuline ® – global)	19.4 Ph.II (actual)	
SULF	Non-pancreatic neuroendocrine tumors	2019/20 China	~50,000 - 60,000	\$11,000 (Sutent®/Afinitor® – global) \$2,190 (Afinitor® China NDRL) \$5,500 (Somatuline® – global)	13.4 Ph.II (actual)	
EPIT	1L EGFR-mutant NSCLC with brain metastasis	<b>2020/21</b> China	~30,000 - 40,000	\$15,100 (Tagrisso®) – <i>Brain pen.<sup>[6]</sup></i> \$1,100 (Iressa®) – <i>min. brain pen.</i> \$850 (Conmana®) – <i>min. brain pen.</i>	TBD	

[1] Addressable Patient Population = Company estimates considering Frost & Sullivan data, National Central Cancer Registry of China and publicly available epidemiology data; [2] WAC = Wholesaler Acquisition Cost; [3] Last published median Progression Free Survival ("PFS" or time to >20% tumor 67 growth) result for Chi-Med therapy (Chi-Med studies); [4] represents present company estimates; [5] Penetration = % of Addressable Patients treated for an average period equivalent to the median PFS; [6] Tagrisso approval in China expected in 2017; [7] NDRL = National Drug Reimbursement List.

## **Targeted therapies – fastest growth & largest**<sup>[1]</sup> Pricing beyond reach of the 10.2 million cancer patients in China



	% of Oncology Market <sup>[4]</sup>	Sub-Category	Share of Sub- category	Product	Company	Value Sales (\$m)	Approx. Monthly Pricing (\$)	12 mo. treatment (Est. # patients)
	23.0%	Targeted	19.5%	rituximab	Roche	327	13,090	2,090
		Therapies	14.9%	trastuzumab	Roche	250	4,500	4,640
			14.2%	imatinib	Novartis	238	6,320	3,140
			9.5%	gefitinib	AstraZeneca	160	2,730	4,870
Global Oncology 🔴 🔪 🚽			8.2%	bevacizumab	Roche	138	11,590	990
aiobai Olicology			7.4%	erlotinib	Roche	124	2,040	5,070
drug market <sup>[1]</sup> :			5.3%	cetuximab	BMS/BI	89	14,150	520
uluy markeu ···			4.6%	sorafenib	Bayer	77	7,250	890
			4.0%	bortezomib	Janssen	67	6,360	880
\$176 billion			12.4%	Other		208		
				Total Targeted T	herapies	1,679		23,080
	20.4%	Anti-metabolites	29.1%	pemetrexed	Lilly/Hansoh	433		
			21.5%	capecitabine	Roche	320		
			20.4%	TS-1	Taiho/Qilu	304		
			16.6%	gemcitabine	Lilly/Hansoh	247		
China Oncology			12.4%	Other		185		
Market <sup>[2]</sup> :				Total Anti-Meta	bolites	1,489		
\$7.3 billion	10.7%		40.20/	paclitaval	DMC/Luuro	700		
	19.7%	Plant Alkaloids	49.3% 42.4%	paclitaxel docetaxel	BMS/Luye	709		
			42.4% 8.4%	Other	Sanofi/Hengrui	609 120	Hiah-lev	el analysis i
			0.4%	Total Plant Alka	loids	1,438		
China 🔪 👘					10103	064,1	aeneral	reference o
	10.5%	<b>DNA Damaging</b>	46.5%	oxaplatin	Sanofi/Hengrui	356		
Pharmaceutical	1012/0	agents	21.3%	temzolomide	Merck/Tasly	163		
			13.1%	nedaplatin	i ici citi i cibig	100		
Market <sup>[3]</sup> :			4.3%	carboplatin		33		
			14.8%	Other		113		
				Total DNA Dama	aging Agents	767		
\$80 billion								
	6.1%	Hormones	29.8%	letrozole	Novartis/Hengrui	133		
			23.0%	bicalutamide	AstraZeneca	102		
			19.5%	anastrozole	AstraZeneca	87		
			17.1%	exemestane	Pfizer/Qilu	76		
ost & Sullivan; [1] 2016 global oncology market value sales;			10.6%	Other		47		
iina oncology market value sales;				Total Hormones	;	445		

## National Drug Reimbursement List Pricing ("NDRL") July'17 update – 15 new drugs in oncology<sup>[1]</sup> added to NDRL



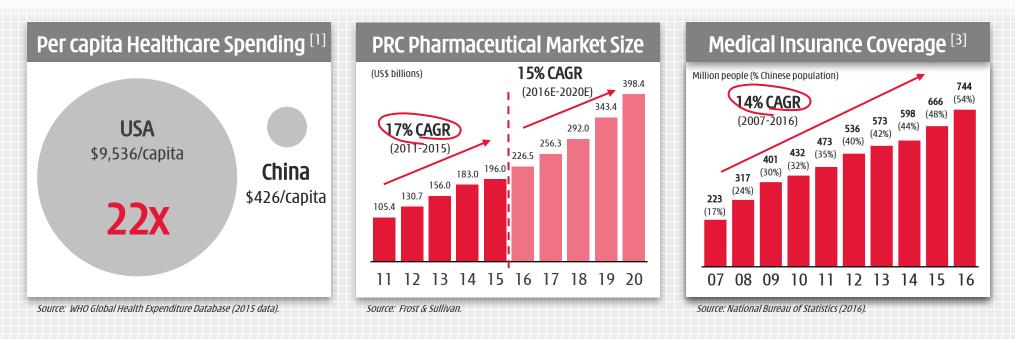
		U	nit Pricing (US\$	)[3]	Approximate Mor	nthly Pricing (U	JS\$) <sup>[3]</sup>	
Brand (generic)	Company	Dosage	Avg. Tender	Reimbursed $\Delta\%$	Dosage	Avg. Tender	Reimbursed	Indication coverage
Herceptin® (trastuzumab)	Roche	440mg:20ml	\$3,298.81	\$1,125.93 <b>-66%</b>	Breast: 4mg/kg wk 1, 2mg/kg weekly. <sup>[2]</sup>	\$4,500	\$1,540	Breast: Her2+; Her2+ meta. Her2+ late-stage meta. gastric.
Avastin® (bevacizumab)	Roche	100mg:4ml	\$772.74	\$296.00 <b>-62%</b>	10mg/kg 0.2W.	\$11,590	\$4,440	Late-stage meta. CRC or advanced non-squamous NSCLC.
<b>TheraCIM<sup>©[4]</sup></b> (nimotuzumab)	Biotech Pharma	50mg:10ml	\$435.26	\$251.85 <b>-42%</b>	100mg weekly.	\$3,730	\$2,160	Combo with radiotherapy for EGFR+ Stage III/IV nasopharyngeal carcinoma.
<b>Rituxan®</b> (rituximab)	Roche	500mg:50ml <sup>[2]</sup>	\$2,544.74	\$1,228.15 <b>-52%</b>	375 mg/m² weekly.	\$13,090	\$6,320	Restorative or resistant follicular central type lym.; CD20+ stage III-IV follicular NHL, CD20+ DLBCL.
Tarceva® (erlotinib)	Roche	150mg <sup>[2]</sup>	\$68.15	\$28.89 <b>-58%</b>	150mg QD.	\$2,040	\$870	Advanced NSCLC with limited EGFR gene mutation.
Nexavar® (sorafenib)	Bayer	0.2g	\$60.44	\$30.07 <b>-50%</b>	400mg BID.	\$7,250	\$3,610	Unresectable RCC. Unresectable HCC. meta. Diff. thyroid after radio-iodine therapy.
Tykerb® (lapatinib)	GSK	250mg	\$17.63	\$10.37 <b>-41%</b>	1,500mg QD.	\$3,170	\$1,870	Adv./meta. breast cancer with Her2 O/E, after anthracycline, paclitaxel, trastuzumab.
AiTan® (apatinib)	Hengrui	425mg <sup>[2]</sup>	\$47.85	\$30.22 <b>-37%</b>	850mg QD.	\$2,870	\$1,810	3L gastric adenocarcinoma or esophageal junction with adenocarcinoma.
Velcade® (bortezomib)	١&١	3.5mg <sup>[2]</sup>	\$1,873.78	\$906.07 <b>-52%</b>	1.3mg/m² quartic every 3 wks.	\$6,360	\$3,080	Myeloma; recurring or refractory mantle cell lymphoma.
EnDu® (rh-endostatin)	Simcere	15mg	\$132.15	\$93.33 <b>-29%</b>	7.5mg/m² iv QD 2-wks- on / 1-week-off.	\$2,110	\$1,490	Late-stage NSCLC.
Epidaza® (chidamide)	Chipscreen	5mg	\$81.48	\$57.04 <b>-30%</b>	30mg QD, 2x per wk.	\$4,190	\$2,930	2L+ Recurring or refractory peripheral T-cell lymphoma (PTCL).
<b>Zytiga®</b> (abiraterone)	۲%۱	250mg	\$45.63	\$21.48 <b>-53%</b>	1,000mg QD.	\$5,480	\$2,580	Metastatic or ovariectomized prostate cancer.
Faslodex® (fulvestrant)	AstraZeneca	250mg:5ml	\$806.81	\$355.56 <b>-56%</b>	500mg per month.	\$1,610	\$710	Advanced ER/PR+ breast can., failing aromatase inhibitor.
Afinitor® (everolimus)	Novartis	5mg <sup>[2]</sup>	\$36.44	\$21.93 <b>-40%</b>	10mg QD.	\$2,190	\$1,320	Adv. RCC after sunitinib or sorafenib. Adv./meta. pancreatic NETs. Tuberous sclerosis with renal angiomyolipoma.
<b>Revlimid</b> (lenalidomide)	Celgene	25mg <sup>[2]</sup>	\$413.93	\$163.26 <b>-61%</b>	25mg QD 3-wks-on / 1-wk-off.	\$9,310	\$3,670	2L+ Recurring myeloma.

Source: Ministry of Human Resources and Social Security (MOHRSS); Yaozhi; BofA Merrill Lynch Global Research

[1] Excluding 3 botanical oncology drugs; [2] Reference SkU or reference recommended dosage for monthly pricing calculation; [3] Calculation assumes an exchange rate of CN¥6.75 per US\$1; [4] Marketed as Tai Xin Sheng® in China.

# China pharma market has become the second largest globally since 2016





- China pharmaceutical industry growth 17% CAGR from 2011-2015 one of the higher rated industries in China with average P/E ratio of 36 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 Link to increased drug reimbursement & sales.

<sup>[1]</sup> Current health expenditure by revenues of health care financing schemes (in current US\$ per capita); [2] National Bureau of Statistics of China; [3] Urban Basic Medical Care Insurance – total persons covered at year-end CAGR = Compound annual growth rate

## 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 \$2.6 2.8 billion.<sup>[1]</sup> Given our share in the JVs, Chi-Med's share of this value is approximately \$1.2 1.4 billion.

			NET SALES			NET I	NCOME		VALUAT	ION <sup>[3]</sup>
	Code	2016 Jan-Jun	2017 Jan-Jun	16-17 1H Growth	2016 Jan-Jun	2017 Jan-Jun	16-17 1H Growth	2017 1H Margin	Market Cap.	P/E
CHI-MED Commercial Platform Subsidiaries/JVs <sup>[2]</sup>		331.9	357.0	8%	47.9	51.9	8%	15%	n/a	n/a
Tianjin Zhong Xin Pharma	600329	486.0	447.2	-8%	37.5	41.3	10%	9%	1,350	30
Li Zhu Pharma	000513	566.5	639.9	13%	64.0	82.4	29%	13%	5,358	34
Shandong Dong E E Jiao	000423	400.4	439.3	10%	124.8	135.1	8%	31%	6,151	23
Zhejiang Kang En Bai Pharma	600572	456.6	350.3	-23%	44.4	58.0	31%	17%	2,769	29
Kunming Pharma	600422	374.2	408.6	9%	37.4	32.4	-13%	8%	1,111	26
Guizhou Yi Bai Pharma	600594	258.4	292.2	13%	26.1	29.7	14%	10%	1,111	25
Jin Ling Pharma	000919	270.7	256.2	-5%	19.1	18.4	-4%	7%	626	31
Jiangsu Kang Yuan	600557	224.9	249.0	11%	28.9	29.1	1%	12%	1,182	25
Zhuzhou Qian Jin Pharma	600479	191.4	226.4	18%	5.4	8.1	49%	4%	790	26
ZhangZhou Pian Zai Huang	600436	143.8	262.4	83%	46.3	63.3	37%	24%	6,885	48
Peer Group Weighted Avg. (10 Comps. excl. Chi-Med)		337.3	357.2	6%	43.4	49.8	15%	14%	2,733	33
All 61) isted China Pharma. Companies Weighted Average		567.1	624.8	10%	51.1	57.0	19%	9%	3,251	36

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 2017E Net Sales in the ~\$400-1,300 million range.

Source: Company data, Deutsche Bank, FactSet

[1] Peer group/China Pharma multiple of 33x-36x 2017 actual Net income after tax of \$77.3 million (excluding SHPL's R&D related subsidies of US\$5.0 million at net income after tax);

- [2] Total aggregate PRC domestic results of Chi-Med's 6 Commercial Platform companies (HBYS, SHPL, Hutchison Sinopharm, HHO, HHL, & HCPL), excluding discontinued operations and land compensation from SHPL;
- [3] Market Capitalization and Price Earnings Ratios as at February 27th, 2018: Trailing Twelve Month PE weight averaged based on market capitalization.

## **Innovation Platform proxy peer group (1/2)** A very deep pipeline and a very large organization/operation



	Mkt	Cap (Feb 1	<u>6)</u>	Ent.		a second seco	Overview of pipeline assets			_ # 0f _	#	of stud	ies	
ame	2018	2017	2016	Value <sup>[1]</sup>	Staff	Drug	Studies	Phase	Partner	drugs	P3	P2		
enmab	11,126	12,415	6,177	10,284	257	Arzerra (ofatumumab)	CLL, FL	Mktd, P3	Novartis	12	13	6		
						Ofatumumab (subcutaneous)	Relapsing multiple sclerosis	2xP3	Novartis					
						Darzalex (daratumumab)	MM, amyloidosis, NKT-cell lym., myelodysplastic syndromes, solid tumors	Mktd, Reg., 9xP3, 3xP2, 5xP1	Janssen					
						Teprotumumab (RV001)	Graves' orbitopathy (thyroid eye disease)	P3	Horizon					
						Tisotumab vedotin	Solid tumors	1xP2, 2xP1/2	Seattle Genetics					
						HuMax-AXL-ADC, HexaBody-DR5/DR5	Solid tumors	1xP1/2 (ongoing), 1xP1/2 (to start in 2018)						
						DuoBody-CD3xCD20	Hematological malignancies	P1/2 (to start in 1H2018)						
						AMG 714	Celiac disease	2xP2	Amgen					
						ADCT-301, JNJ-61186372, JNJ-63709178, JNJ-64007957	Lym., AML, ALL, NSCLC, R/R MM	5xP1	ADC, Janssen					
telixis 8,633 6,488 959 8,266 372 Cabometyx / Cometriq (cabozantinib) Cotellic (cobimetinib) Esaxerenone (CS-3150) SAR245408 (XL147) SAR245409 (XL765)	Thyroid cancer, advanced renal CC, adv. hepatocellular carcinoma, NSCLC, genitourinary tumors, endometrial cancer, breast cancer & others	Mktd, 2xP3, 14xP2, 5xP1	6	7	22									
		Metastatic or unresectable locally advanced melanoma, CRC, BC, pancreatic cancer	Mktd, 3xP3, 2xP2, P1											
							Hypertension, diabetic nephropathy	2xP3	Daiichi Sankyo					
							Variety of cancer indications	P2	Sanofi					
				SAR245409 (XL765)	NHL, glioblastoma, lym., BC, leukemia, combos w/ Treanda, Rituxan	5xP2	Sanofi							
	XL888	XL888	BRAF V600 Mutation-Pos advanced melanoma, Malignant melanoma	2xP1										
eiGene	6,199	1,618	225	5,614	900	BGB-3111; BGB-3111 + Gazyva	WM, 1L CCL, R/R MCL, R/R CLL, R/R DLBCL, R/R FL	2xP3, 4xP2		5	6	7		
						BGB-A317	3GB-A317 2L NSCLC, 1L hepatocellular carcinoma, R/R Hodgkins lym. 2L+ UC 4xP3, 2xP							
						BGB-290	3L gBRCA+ ovarian cancer	P1, P2						
						BGB-283	BRAF and RAS mutated solid tumors	2xP1						
						BGB-A317 + BGB-290; BGB-A317 + BGB-3111	Solid tumors; B-cell malignancies	2xP1						
	s       8,633       6,488       959       8,266       372       Cabometyx / Cometriq (cabozantin Cotellic (cobimetinib)         Esaxerenone (CS-3150)       SAR245408 (XL147)       SAR245408 (XL147)         SAR245409 (XL765)       XL888         ve       6,199       1,618       225       5,614       900       BGB-3111; BGB-3111 + Gazyva BGB-3117         BGB-290       BGB-3317       BGB-290       BGB-283       BGB-290; BGB-A317 + BGB-290; BGB-A317 + BGB-290; BGB-A317 + BGB-290; BGB-A333 + CC-122	BGB-290 +(RT/)Chemo; BGB-A333 +/- BGB-A317	Solid tumors, glioblastoma	3xP1										
						((-122	R/R DLBCL, NHL	P1						
							NSCLC	P1	Mirati					
llapagos	5,209	3,002	1,897	3,490	578	Filgotinib	RA, CD, UC, small bowel CD, Fistulizing CD, Sjogren's, ankylosing spondylitis, psoriatic arthritis, cutaneous lupus, lupus nephropathy, uveitis	3xP3, 8xP2	Gilead	11	3	12		
	'2222; '2222 + Kalydeco '2451 + '2222 + '2737; '3067 + '2222 + '2737; '3067 + '2222 + '3221;	Cystic fibrosis	2xP2, 3xP1	AbbVie										
						GLPG1690; '1205; ,3499	Idiopathic pulmonary disease	P2	-					
						GLPG1972; MOR106	Atopic dermatitis, Osteoarthritis	2xP1	Servier, Morphosys					

Source: Company data, FactSet, public filings

[1] As of February 16, 2018

Key: CLL = chronic lymphocytic leukemia; Lym. = lymphoma; NHL = Non-Hodgkin's Lymphoma; AML = acute myeloid leukemia; ALL = acute lymphoblastic leukemia; WM = Waldenstrom's macroglobulinemia; MCL = mantle cell lymphoma; FL = follicular lymphoma; DLBCL = diffuse large B-cell lymphoma; RA = Rheumatoid Arthritis; MM = Multiple Myeloma; CC = Cell Carcinoma; NSCLC = Non-Small Cell Lung Cancer; BC = Breast Cancer; CRC = colorectal cancer; CD = Crohn's disease; R/R = relapsed / refractory; Mktd = Marketed; Reg. = Under Registration

(\$ millions unless otherwise stated)

## Innovation Platform proxy peer group (2/2) A very deep pipeline and a very large organization/operation



	Mkt	Cap (Feb 1	6)	Ent.		and the second	Overview of pipeline assets			# Of	#	<u>of stud</u>	es
Name	2018	2017	2016	Value <sup>[1]</sup>	Staff	Drug	Studies	Phase	Partner	drugs	P3	P2	P
Agios	4,329	2,105	1,497	3,905	382	Idhifa; + Vidaza; + (7+3)	R/R AML, frontline AML	Mktd., P3, 2xP2	Celgene	5	4	3	_
						Ivosidenib; + Vidaza; + (7+3);	Frontline AML, R/R AML, cholangiocarcinoma, low grade glioma	Reg., 3xP3,	-				
						+ AG-881		5xP1					
						AG-348	PK deficiency	P2					
						AG-270		P1	-				
						AG-881	Low grade glioma	P1	Celgene				
Array <sup>[2]</sup>	3,526	1,889	406	3,212	209	ARRY-797	LMNA-related DCM	P2	-	2	0	2	
						ARRY-382	Solid tumors	P2	-				
Tesaro	3,263	9,750	1,530	2,882	446	Varubi (IV and oral)	CINV (oral and IV)	Mktd, Reg.	Opko	5	1	5	
						Zejula (niraparib); + anti-PD-1	Ovarian cancer maintenance, ovarian cancer treatment, NSCLC	Mktd, Reg., P3, 2xP2	Merck				
						Niraparib + Pembrolizumab	Triple-negative BC or ovarian cancer (TOPACIO study)	P2	Merck				
						Niraparib + Bevaciumab	Ovarian cancer, 1L ovarian cancer maintenance	2xP2	Roche				
							Advanced NSCLC, advanced or metastatic cancer, SCCL, Ewing's sarcoma, various		AnaptysBio, SARC				
		TSR-033	tumor types										
Loxo 3,129 1,0	1,057	345	2,757	47	Larotrectinib (LOXO-101)	Cancers Harboring Alterations of TRK	Reg., 2xP2, 2xP1	Bayer	3	0	2		
						LOXO-292	Cancers Harboring Alterations of RET	P1					
						LOXO-195	Next-Gen TRK inhibitor for potential acquired resistance	P1	Bayer				
Morphosys <sup>[2]</sup>	2,761	1,533	1,084	2,533	351	MOR208	CLL, SLL, DLBCL	P3, 2xP2		3	1	3	_
						MOR202	Multiple myeloma	P2					
						MOR107	Undisclosed	P1					
Clovis	2,654	2,739	746	2,308	278	Rubraca (rucaparib); + nivolumab; + atezolizumab		1	4	3	_		
							prostate, triple negative BC, BC, gastro esophageal, gynecological	4xP3, 3xP2, P1					
Puma	2,516	1,375	1,371	2,410	308	Neratinib (PB272)	Adjuvant BC, neoadjuvant BC, metastatic BC, metastatic BC wit brain met., met.	Mktd., P3, 8xP2		1	1	8	
							her2 BC						
AVERAGE	4,805	3,997	1,476	4,333	375					5	3	7	
Median	3,526	2,105	1,084	3,212	351					5	3	5	
nnovation					∽360	Savolitinib	PRCC, CCRCC, NSCLC, gastric cancer, lung cancer, prostate cancer	P3, 13xP2	AstraZeneca	8	6	18	1
Platform						Fruquintinib	CRC, NSCLC, caucasian bridging, gastric cancer	Reg., 2xP3, P2, P1	Eli Lilly				
						Sulfatinib	Pancreatic and non-pancreatic NETs, Caucasian bridging, meduliary thyroid	2xP3, 3xP2, P1					
							cancer, differentiated thyroid cancer, biliary tract cancer						
						Epitinib	NSCLC, glioblastoma	P1b/2					
						Theliatinib	Solid tumors, esophageal cancer	P1b, P1					
						HMPL-523	RA, hematological cancers, immunology, lym.	2xP1b, P1					
						HMPL-689	Hematological cancers, lym.	2xP1					
						HMPL-453	Solid tumors	2xP1					

Source: Company data, FactSet, public filings

[1] As of February 16, 2018

[2] Only non-partnered products included for Morphosys and Array. Array also owns two products in phase 3 (Binimetinib and Encorafenib) in which Array maintains US and Canadian rights.

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lymphoma; FL = follicular lymphoma; DLBCL = diffuse large B-cell lymphoma; RA = Rheumatoid Arthritis; MM = Multiple Myeloma; CC = Cell Carcinoma; NSCLC = Non-Small Cell Lung Cancer; BC = Breast Cancer; CRC = colorectal cancer;

CD = Crohn's disease; R/R = relapsed / refractory; Mktd = Marketed; Reg. = Under Registration

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# Non-GAAP Financial Measures and Reconciliation (1/2)



Reconciliation of Adjusted Research and Development Expenses (Page 5 and Page 65):

Reconciliation of Top 7 products' Gross Profit as Percentage of Aggregated Gross Profit for Commercial Platform (Page 60):

	2017	2016		2017
Research and development expenses	(75.5)	(66.9)	Sales of goods — third parties and related parties	205.2
Plus: Innovation Platform — administrative and other expenses	(8.0)	(5.0)	Less: Costs of sales of goods — third parties and related parties	(175.8)
	(0.0)	(5.0)	Consolidated gross profit	29.4
Plus: Equity in earnings of equity investees — NSP and other	(4.5)	(4.2)	Plus: Gross profit — HBYS and SHPL	269.8
			Adjusted gross profit	299.2
Plus: Innovation Platform — interest income	0.0	0.0		
			Top 7 products gross profit	267.8
Adjusted research and development expenses	(88.0)	(76.1)	% of Top 7 products to adjusted gross profit	90%

# Non-GAAP Financial Measures and Reconciliation (2/2)



#### Reconciliation of Non-GAAP Sales and Non-GAAP Net (loss)/income after tax<sup>[1]</sup>

Prescription Drugs: includes our Consolidated subsidiary (Hutchison Sinopharm) and Non-consolidated joint venture (SHPL);
 Consumer Health: includes our Consolidated subsidiaries (HHO, HHL and HCP) and Non-consolidated joint venture (HBYS).

					IFR	5							US GAAP			16-17
(US\$ millions)	03	04	05	06	07	08	09	10	11	12	13	14	15	16	17	Growth
Sales (Non-GAAP)	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	677.2	8%
Prescription Drugs	17.2	21.8	23.3	23.2	<i>28.1</i>	<i>39.5</i>	54.4	71.2	92.4	116.5	<i>138.2</i>	204.9	286.6	372.3	411.0	10%
- Consolidated subsidiary	-	-	-	-	-	-	-	-	-	-	-	50.2	105.5	149.9	166.4	11%
- Non-consolidated joint venture	17.2	21.8	23.3	23.2	28.1	39.5	54.4	71.2	92.4	116.5	138.2	154.7	181.1	222.4	244.6	10%
Consumer Health	4.7	6.1	41.8	<i>78.2</i>	90.9	116.3	142.6	165.2	186.2	244.2	264.1	260.5	232.3	255.1	266.2	4%
- Consolidated subsidiaries	4.7	6.1	<i>9.3</i>	<i>8.9</i>	3.7	5.5	7.0	14.1	14.9	15.5	16.5	16.8	20.7	31.0	38.8	25%
- Non-consolidated joint venture	-	-	32.5	<i>69.3</i>	87.2	110.8	135.6	151.1	171.3	228.7	247.6	243.7	211.6	224.1	227.4	1%
Total Sales Growth	n/a	27%	133%	56%	17%	31%	26%	20%	18%	29%	n/a	16%	11%	21%	8%	
Net (loss)/Income after tax (Non-GAAP)	(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	63.3 <sup>[3]</sup>		22%
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	41.4	53.0	28%
- Consolidated subsidiary	-	-	-	-	-	-	-	-	-	-	-	0.1	0.6	1.6	2.4	50%
- Non-consolidated joint venture	(0.4)	1.3	1.9	1.3	1.9	2.8	6.0	11.9	14.2	17.7	22.4	26.4	31.3	<i>39.8</i>	50.6	27%
Consumer Health	(10.3)	<b>(4.9)</b>	0.3	5.4	<i>9.3</i>	11.9	15.5	16.0	<i>15.9</i>	15.4	17.3	22.3	22.2	21.9	24.3	11%
- Consolidated subsidiaries	(10.3)	(4.9)	(2.9)	(2.4)	0.2	-	0.8	1.0	(0.4)	(1.1)	0.1	1.5	0.8	1.5	3.5	>100%
- Non-consolidated joint venture	-	-	3.2	7.8	<i>9.1</i>	11.9	14.7	15.0	16.3	16.5	17.2	20.8	21.4	20.4	20.8	2%
% Margin	<i>-48.9%</i>	-12.9%	3.4%	6.6%	9.4%	9.4%	10.9%	11.8%	10.8%	9.2%	<i>9.9%</i>	10.5%	10.4%	10.1%	11.4%	
					<b>Fa1</b>	<b>Fa1</b>	Fe ]	541	F= 1	F93	F=1	F=1	Fe1	F=1	543	
Net (loss)/income attrib. to Chi-Med	(5.7)	(3.7)	(0.5)	1.2	<b>4.5</b> <sup>[2]</sup>	<b>5.9</b> <sup>[2]</sup>	9.3 <sup>[2]</sup>	12.6 <sup>[2]</sup>	13.6 <sup>[2]</sup>	14.6 <sup>[2]</sup>	18.2 <sup>[2]</sup>	<b>22.8</b> <sup>[2]</sup>	25.2 <sup>[2]</sup>	<b>29.9</b> <sup>[3]</sup>	37.5 <sup>[4]</sup>	25%
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	<i>15.9</i>	20.7	26.5	28%
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	11.0	20%
Net (loss)/income attrib. to Chi-Med growth	n/a	-35%	-86%	340%	275%	31%	58%	35%	8%	7%	n/a	26%	10%	19%	25%	

[1] 2003–2006 incl. disco. operation; [2] Continuing Operations; [3] Excludes one-time gains associated with land compensation and R&D subsidies received by SHPL of US\$80.8 million in net income after tax to SHPL and US\$40.4 million net income attributable to Chi-Med; [4] Excludes one-time gains associated with R&D subsidies received by SHPL of US\$5.0 million net income to SHPL and \$2.5 million at net income after tax and net income attributable to Chi-Med;



## HUTCHISON CHINA MEDITECH

Thank you