

HUTCHISON CHINA MEDITECH

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Use of Non-GAAP Financial Measures - Certain financial measures used in this presentation are based on non-GAAP financial measures. Please see the appendix slides titled "Non-GAAP Financial Measures and Reconciliation" for further information relevant to the interpretation of these financial measures and reconciliations of these financial measures to the most comparable GAAP measures.

Agenda



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Building a global science-focused biopharma company from a powerful base in China...





Global Innovation

- 5 clinical drug candidates in US/EU development
- Building global clinical development footprint
- World-class >420 person scientific team

China Oncology



 Major market potential driven by regulatory reforms & high unmet medical need in oncology

• Elunate[®] (Fruquintinib capsules) first ever homegrown cancer drug launched in China^[1]

8 oncology assets in China development



Existing China Business

- Cash generative China Commercial Platform
- Platform for future innovative drug launches



Important milestones in Chi-Med's evolution





Proven innovation & commercial operations

Management Team	Industry / Chi-Med (years)
Mr. CHRISTIAN HOGG, BSc, MBA Chief Executive Officer	Procter & Gamble 30 / 19
Dr. WEIGUO SU, PHD EVP, Chief Scientific Officer	Pfizer 29 / 14
Mr. JOHNNY CHENG, BEC, CA Chief Financial Officer	Bristol-Myers Squibb Sin Nestle 30 / 11
Dr. ZHOU JUN JIE, MD, MBA General Manager, SHPL	28 / 18
Dr. MAREK KANIA, MD, MBA SVP, Chief Medical Officer, US	Lilly 25/1
Dr. ZHENPING WU, PHD, MBA SVP, Pharmaceutical Sciences	(Roche) 25 / 11
Mr. CHEN HONG, BSc, MBA SVP, Chief Commercial Officer	Bristol-Myers 21 / 9
Dr. MAY WANG, PHD SVP, Bus. Dev. & Strategic Alliances	Lilly 25/9
Mr. MARK LEE, BEng, MBA SVP, Corp. Finance & Development	Credit Suisse 20 / 10
Mr. ENRICO MAGNANELLI, BA, MBA Head of International Operations	🧭 GILEAD 20/1

Integrated Innovation Organization^[1]



Commercial Team & Joint Ventures [1]

ommercial Team (subsidiaries):

>200 staff covering:

- Drug distribution operations; &
- New Oncology Business Dept.

50/50 Joint Ventures: >2,500 Rx medical sales reps.;

>950 person OTC sales team; &

>1,500 staff in two major factories

Portfolio Summary: (1) Eight self-discovered assets; (2) multiple early- & registration-stage studies in a wide range of indications; (3) marketed drugs portfolio in China



Dose Finding / Safety Run-In	Proof-of-Concept	Registration	Marketed
Fruquintinib + Tyvyt (PD-1)	Savo / Savo + Imfinzi (CALYPSO)	Savo + Tagrisso (SAVANNAH)	Elunate (Fruquintinib capsules)
Solid Tumors ⁽¹⁾	x2: PRCC & ccRCC	2L/3L Tagrisso-refractory MET+ NSCLC	≥3L Colorectal cancer
Surufatinib + Tuoyi (PD-1)	Savolitinib (VIKTORY)	Savolitinib	SXBX^[3] Pills
Solid Tumors ⁽¹⁾	MET+ Gastric cancer	MET Exon 14 deletion NSCLC	Coronary artery disease
HMPL-523 (Syk)	Savolitinib (CCGT 1234B)	Fruquintinib + Taxol (FRUTIGA)	>10 other Rx / OTC drugs
Indolent NHL ^{[1] [2]}	MET+ Prostate cancer	2L Gastric cancer	
HMPL-689 (PI3Kδ)	Fruquintinib	Surufatinib (SANET-p)	
Indolent NHL ^[1]	3L/4L Colorectal cancer ^[1]	Pancreatic NET	
Fruquintinib + Tyvyt (PD-1)	Surufatinib	Surufatinib (SANET-ep)	
Solid tumors ^[1]	2L Pancreatic NET	Non-Pancreatic NET	
Fruquintinib + genolimzumab (PD-1)	Fruquintinib + Iressa	Surufatinib	
Solid tumors	1L EGFRm+ NSCLC	2L Biliary Tract cancer	
Surufatinib + Tuoyi (PD-1) Solid tumors	HMPL-523 B-cell malignancies; ITP [1]		Global Innovation
Surufatinib + HX008 (PD-1) Solid tumors ⁽¹⁾	HMPL-523 + azacitidine AML		China Oncology
HMPL-453 (FGFR1/2/3) Solid tumors	HMPL-689 Indolent NHL	[1] In planning / imminent: [2] Proof-of-concept in Australia: [3] SXBX =	She Xiang Bao Xin (cardiovascular).
	Epitinib Glioblastoma	Targets: Savolitinib = MET; Fruquintinib = VEGFR1/2/3; Surufatinib = VEGF Epitinib = EGFRm in the brain; Theliatinib = EGFR wild-type; HMPL-453 = Indications: NHL = Non-Hodgkin's Lymphoma; NET = Neuroendocrine tur ITP = Immune thrombocytopenia: NSCLC = Non-small cell lung carcer.	R1/2/3 / FGFR1 / CSF-1R; HMPL-523 = Syk; HMPL-689 = PI3K&; FGFR1/2/3. mors; RCC = Renal cell carcinoma; AML = Acute myeloid leukemia;

Major targets/news flow in 2019





Targets: Savolitinib = MET; Fruquintinib = VEGFR1/2/3; Surufatinib = VEGFR1/2/3 / FGFR1 / CSF-1R; HMPL-523 = Syk; HMPL-689 = PI3K8. Indications: NHL = Non-Hodgkin's Lymphoma; NET = Neuroendocrine tumors; RCC = Renal cell carcinoma; NSCLC = Non-small cell lung cancer.

Global clinical drug portfolio (1/2)





MET = mesenchymal epithelial transition receptor, VEGFR = vascular endothelial growth factor receptor, NSCLC = non-small cell lung cancer, RCC = renal cell carcinoma, PRCC = papillary RCC, CRC = colorectal cancer; [1] Efficacy Evaluable Patients. Data cut-off: Oct. 10, 2017; [2] Dosed to-date = patients in all clinical trials (treatment & placebo).

Global clinical drug portfolio (2/2)





[1] Dosed to-date = patients in all clinical trials (treatment & placebo); [2] American Society of Hematology. Blood, vol. 132 no. Suppl 1 5324 (Nov 2018); VEGFR = vascular endothelial growth factor receptor, FGFR1 = fibroblast growth factor receptor 1, CSF-1R = colony stimulating factor-1 receptor, Syk = spleen tyrosine kinase, PI3K\delta = Phosphatidylinositol-3-Kinase delta, pNET = pancreatic neuroendocrine tumors, ep-NET = non-pancreatic neuroendocrine tumors, AML = acute myeloid leukemia, FL = follicular lymphoma, CLL = chronic lymphocytic leukemia, SLL = small lymphocytic leukemia.





Highlights & Strategies – Global Innovation Pushing the envelope on our most valuable assets

One of China's largest & most prolific discovery platforms in oncology





Global step-change innovation

• Multiple potential first-in-class assets



Kinase selectivity – enable combos

• Dial out off-target toxicity & address TKI resistance



Building broad range of assets against novel targets

• 2nd generation I/O & expanding to mAbs





Attack cancer from multiple angles at same time

Immune Desert Insufficient T cell response

- Chemotherapies
- Vaccines
- CAR-T (pro-inflammatory strategies)
- TCB's

Antigen Release

- Aberrant genetic drivers
- Targeted therapies (small molecule & antibody)



Excluded Infiltrate Inadequate T cell homing

- Anti-angiogenics
- Stromal targets
- Chemokines
- Vaccines

Inflamed

Inactivated T cell response

- Immunotherapies (address negative regulators)
- Vaccines

Need combinations of potent, yet tolerable drugs against specific targets

Our advanced medicinal chemistry provides superior selectivity & safety profiles...





Savolitinib

∽1,000 times more selective to c-MET than next kinase (PAK3) ^[5]

> Screening at 1µM against 253 Kinases



ELUNATE Fruquintinib Capsules ~250 times more selective to

VEGFR3 than next non-VEGFR kinase (Ret) ^[6]

	Disconti	nuations as %	Enrolled
Non-small cell lung cancer (NSCLC)	Due to AE	Withdrawn / Other	Total ^[1]
Monotherapy – Tagrisso® / savolitinib			
Tagrisso® (osimertinib)	6%	6%	13%
Savolitinib 600mg QD PRCC (for reference only – not NSCLC) ^[2]	9%	5%	14%
Combination - Tagrisso® + savolitinib			
savolitinib 600mg QD + Tagrisso® [3]	29%	6%	35%
Approved treatments in NSCLC			
Zykadia ® (ceritinib)	10%	10%	20%
Cyramza ® (ramucirumab) + Taxotere ®	15%	21%	37%
Keytruda® (pembrolizumab) 2mg/kg	10%	26%	37%
Opdivo ® (nivolumab)	15%	4%	20%
Chemo doublet (platinum + pemetrexed)	11%	17%	27%
Taxotere® (docetaxel)	13%	22%	36%

3 rd -Line Metastatic CRC	FRESCO Study Mainland China		CONC (China, Hi	UR Study K, Taiwan) ^[4]
Treatment arms	Elunate®	Placebo	Stivarga ®	Placebo
VEGFR on-target related AEs:				
Hypertension ≥G3	21.2%	2.2%	12.5%	8.3%
Hand-Foot Syndrome (Palmar-plantar), ≥G3	10.8%	0.0%	17.0%	0.0%
Off-target (i.e. non-VEGFR) related AEs:				
Hypophosphatemia, ≥G3	0.0%	0.0%	8.0%	0.0%
Hypokalemia, ≥G3	0.7%	0.7%	6.3%	0.0%
Rash/desquamation, ≥G3	0.0%	0.0%	4.4%	0.0%
Lipase increase, ≥G3	0.0%	0.0%	6.3%	1.7%
Hepatic function (Liver function) AEs:				
ALT increased, ≥G3	0.7%	1.5%	7.1%	3.3%
AST increased, ≥G3	0.4%	0.7%	8.9%	0.0%
Blood bilirubin increased, ≥G3	1.4%	1.5%	8.9%	8.3%
Tolerability:	\frown		\frown	
AE Leading to dose interruption	35.3%	10.2%	68.8%	25.0%

[1] Total discontinuations = Discontinuations NOT due to Disease Progression or Death; [2] September 2017 Journal of Clinical Oncology; [3] 2019 AACR # CT032, CT033; [4] Efficacy & safety of regorafenib monotherapy in Chinese patients with previously treated metastatic colorectal cancer: subgroup analysis of the CONCUR trial; R Xu; [5] W. Su, et al, 2014 American Association of Cancer Research; [6] Sun et al., Cancer Biology & Therapy 15:12, 1635--1645; December 2014.

...**Superior safety allows for combinations** TKI + TKI combos to address acquired resistance





- MET amplification is the most common resistance mechanism for Tagrisso[®].
- Requires addition of MET inhibitor savolitinib – in combo with Tagrisso[®].

RESULTS: CANDIDATE ACQUIRED RESISTANCE MECHANISMS WITH OSIMERTINIB (n=91)*

- No evidence of acquired EGFR T790M
- The most common resistance mechanisms were *MET* amplification and EGFR C797S mutation • Other mechanisms included *HER2* amplification, *PIK3CA* and *RAS* mutations





- C481S or PLC_γ are the most common resistance mechanisms for Imbruvica[®].
- Invalidating BTK inhibitor requires a possible Syk, PI3Kδ &/or BTK TKIs.



Savolitinib Biggest opportunity is MET+ NSCLC





[1] Primary drivers, based on aggregate rocelitinib/Tagrisso data published at 2016/2017 ASCO; [2] AstraZeneca 2016/17/18/1019 results and company estimates.

Savolitinib – 2L NSCLC^[1] combo w/ Simertinib TATTON B Study at AACR 2019



...TATTON B^[2] - ...promising efficacy in MET+ T790M- Iressa/Tarceva failure patients

2L post Iressa®/ Tarceva®





Best response after treatment with savolitinib and Tagrisso	# pts	% Enrolled (n=46)	% Efficacy Evaluable (n=43)		
Complete or partial response	24	52%	56%		
Stable disease (≥6 weeks)*	16	35%	37%		
Progressive disease	3	7%	7%		
Not evaluable	3	7%	-		
Time to response, median (IQ range)		43 days (40-4	43)		
Duration of response, median (IQ range)	7.1 months (4.1 - 10.7)				



[1] EGFRm NSCLC; [2] AACR 2019 - Sequist, *et al.* TATTON Phase Ib expansion cohort: Osimertinib plus savolitinib for patients (pts) with EGFR-mutant, MET-amplified NSCLC after progression on prior epidermal growth factor receptor (EGFR) tyrosine kinase inhibitor (TKI); * Includes four patients with unconfirmed partial responses.

Savolitinib – 2L/3L NSCLC^[1] combo w/ simertinib TATTON B Study at AACR 2019



...TATTON B^[2] - ...promising efficacy in MET+ Tagrisso failure patients...

2L/3L post Tagrisso®





Best response after treatment with savolitinib and Tagrisso	# pts	% Enrolled (n=48)	% Efficacy Evaluable (n=39)		
Complete or partial response	12	25%	31%		
Stable disease (≥6 weeks)*	21	44%	54%		
Progressive disease	6	13%	15%		
Not evaluable	9	19%	-		
Time to response, median (IQ range)		46 days (43-1	51)		
Duration of response, median (IQ range)	9.7 months (5.5 - NC)				



[1] EGFRm NSCLC; [2] AACR 2019 - Sequist, *et al.* TATTON Phase Ib expansion cohort: Osimertinib plus savolitinib for patients (pts) with EGFR-mutant, MET-amplified NSCLC after progression on prior epidermal growth factor receptor (EGFR) tyrosine kinase inhibitor (TKI); * Includes five patients with unconfirmed partial response.

SAVANNAH Study Encouraging TATTON data – led to the initiation of SAVANNAH





SAVANNAH (NCT03778229)

Phase II single-arm study:

- ➢ Global − N. & S. America, Eur., & Asia.
- Primary endpoint ORR.
- Secondary endpoints: PFS, OS, DOR & percent change in tumor size.
- > Primary data completion est. 2021.

• Weight-based dosing regimen:

- TATTON D exploring lower savo dose in order to maximize long-term tolerability for combo.
- > TATTON D enrollment complete.

• ORCHARD study:

- Post FLAURA Platform study offering targeted treatments for all patients expect high enrollment.
- > MET+ patients prioritize to SAVANNAH.

PRCC – unmet medical need Lower response rates to treatments





[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 4.2019, April 25, 2019) category 1 options, RCC = renal cell carcinoma; [4] ORR = Objective Response Rate, mPFS = median Progression-Free Survival, mOS = median Overall Survival, NR = not reached; For approved subgroup of patients; [5] only approved for patients with intermediate or poor risk RCC; [6] Not in guideline; FDA approved on May 15, 2019.



Savolitinib + Imfinzi[®] combination



Immunotherapy combinations... our assets are ideal TKI combo partners for immunotherapy





Multiple global immunotherapy combo deals...



[1] Source: 1. B. Rini et al, for the for the KEYNOTE-426 Investigators, NEJM 2019 Feb 16. doi: 10.1056/NEJMoa1816714, Pembrolizumab plus Axitinib versus Sunitinib for Advanced Renal-Cell Carcinoma; 2. D.F. McDermott et al, ASCO 2018 #4500, Pembrolizumab monotherapy as first-line therapy in advanced clear cell renal cell carcinoma (accRCC): Results from cohort A of KEYNOTE-427; [2] BTD = Breakthrough Therapy Designation.

Australia & China Phase I/Ib studies Extensive Ph.I dose escalation study now complete in Australia & China (total n=60); **Complete** Stage I: dose escalation "3 + 3" each dose cohort RP2D^[1] determined & large Ph. lb until disease • Australia: Relapsed/refractory Studied HMPL-523 N = 33progression, dose expansion study, total hematologic malignancy 100-1,000mg QD & death. 200-400mg BID in **n=192**, underway in 13 active • China: Relapsed/refractory mature B intolerable N = 27 13 dose cohorts lymphoma toxicity, etc. sites in Australia & China; Phase I/Ib data set currently >110 patients; Stage II: dose expansion ...Now enrolling US IND application cleared by FDA Relapsed or refractory, measurable & U.S./E.U. Phase I imminent; disease – multiple arms: until disease Aus Chronic lymphocytic leukemia progression, Plan to initiate China registration N = 40600mg QD Small lymphocytic lymphoma death, studies in 2019. China intolerable Mantle cell lymphoma N = 152toxicity, etc. • Follicular lymphoma Diffuse large B-cell lymphoma (PRC)

HMPL-523 (Syk) in hematological cancer Australia & China – large Ph.Ib expansion. US/EU Ph.I imminent

5 assets in global development ...US/EU clinical & regulatory team fully operational



Program	Treatment	Indication	Target patient	Study name	Sites	Dose finding / safety run-in	Proof-of-concept		Registration
	Savolitinib + Tagrisso®	NSCLC	2L/3L EGFRm; Tagrisso [®] ref.; MET+	SAVANNAH	Global	Oxnard/Ahn – DF/SMC			
	Savolitinib + Tagrisso®	NSCLC	2L EGFRm; EGFR TKI ref.; MET+	TATTON	Global	Oxnard – Dana Farber			Full Ph.II data at
	Savolitinib + Imfinzi [®] (PD-L1)	Papillary RCC	All	CALYPSO	UK/Spain	Powles – Queen Mary's			AACK API 2019
Savolitinib	Savolitinib + Imfinzi [®] (PD-L1)	Clear cell RCC	VEGFR TKI refractory	CALYPSO	UK/Spain	Powles – Queen Mary's			
MET	Savolitinib	Gastric cancer	MET+	VIKTORY	South Korea	Lee – Samsung Med. Ctr			Prelim. PoC at
	Savolitinib + Taxotere®	Gastric cancer	MET+	VIKTORY	South Korea	Lee – Samsung Med. Ctr [1]			ASCO GU FED 2019
	Savolitinib + Taxotere®	Gastric cancer	MET over expression	VIKTORY	South Korea	Lee – Samsung Med. Ctr [1]			
	Savolitinib	Prostate cancer	MET+	CCGT 1234B	Canada	Kolinsky/Muk'jee/Ong/Chi			Prelim. PoC
	_								11110 2019
Fruquintinib	Fruquintinib	Colorectal cancer	3L/4L; Stivarga [®] /Lonsurf [®] ref./intol.		US	Eng /Desari - MD And. [2]		Plan	ning US/EU registr.
VEGFR 1/2/3	Fruquintinib + Tyvyt® (PD-1)	Solid tumors	1L		US	In planning			study based on
	_							F	RESCO/US Ph.Ib
	Surufatinib	Pancreatic NET	2L; Sutent [®] /Afinitor [®] refractory		US	Dasari/Yao – MD Anderson		Plan	nina US/FU reaistr
FGFR1/2/3; FGFR1; CSF-1R	Surufatinib + Tuoyi [®] (PD-1)	Solid tumors				In planning		stu	dy based on China
	_								Ph.II/US Ph.Ib
HMPL-523	HMPL-523	Indolent NHL			Australia	N/A		clah	
Syk	HMPL-523	Indolent NHL			US	Fowler - MD Anderson [3]		GIOD	al Ph.I/Poc data-set
HMPL-689	HMPL-689	Healthy volunteers			Australia			Data-	set now emerging
РІЗКЪ	HMPL-689	Indolent NHL			US	Gnosn/Cohen-Levine/Emory[3]		in C	hina Ph.I (n ~31)

[1] Further patient enrolment directed to savolitinib monotherapy arm due to the high efficacy observed; [2] in U.S., in E.U. Tabernero - Vall d'Hebron & Sobrero - Genova; [3] In planning.

Note: MET = mesenchymal epithelial transition receptor, VEGFR = vascular endothelial growth factor receptor, EGFRm = epidermal growth factor receptor mutation, FGFR1 = fibroblast growth factor receptor 1, CSF-1R = colony stimulating factor-1 receptor, Syk = spleen tyrosine kinase, PI3K8 = Phosphatidylinositol-3-Kinase delta, NSCLC = non-small cell lung cancer, RCC = renal cell carcinoma, NHL = Non-Hodgkin's Lymphoma, AACR = American Association of Cancer Research annual meeting, ASCO GU = American Society of Clinical Oncology Genitourinary Cancer Symposium, PoC = Proof of Concept.

What is next from discovery? Differentiated assets against multiple targets to emerge 2019-22



Priming & activations aOX40 4-1BB

Antigen release

- MET (savolitinib)
- EGFR (epitinib/theliatinib)
- Syk (HMPL-523)
- PI3Kδ (HMPL-689)
- FGFR (HMPL-453)

• ERK

RIP1KIDH



<u>Anti-angiogenesis</u>

- VEGFR (fruquintinib)
- VEGFR/FGFR (surufatinib)
- FGFR (HMPL-453)

Negative regulators

- Treg (HMPL-689)
- CSF-1R (surufatinib)
- IDOi
- AhRi
- TIM3
- **TCBs**

Pre-clinical - small molecule
 Pre-clinical - antibody

Creating highest-quality range of assets against novel targets for use in combos

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Global Innovation Main targets for 2019-2021

Aim for Savolitinib / Tagrisso[®] combo approval & launch

Build out US/EU development operation

 US/EU C&R operation set up in Florham Park, NJ in 2018; expected to reach ~30 staff by end 2019



- Fruq (ex-China) & suru registration studies & exploration of combos with PD-1s;
- Syk & PI3K δ registration studies & exploration of combos with other TKIs

S Aim to move ~1 novel drug candidate into global development per year









CHI-China oncology - ~24% of world's cancer patients^[1] MED



Industry's attention turning to unmet medical need in China oncology

- Regulatory reforms in China addressing low SoC [2]
- Major investment inflow



Chi-Med is a first mover

- Elunate[®] launch in 3L mCRC; First ever in China^[3]
- Deep pipeline 8 clinical drug candidates with 5 registration studies underway/set to start in China



Major commercial opportunity

• National Drug Reimbursement; Medical coverage



China now world's 2nd largest pharma market ...investment, approvals & access all accelerating rapidly







PRC Healthcare VC/PE Funds^[2]



Number of Priority Review NDAs ^[3]



Medical Insurance Coverage^[1]



Improved Access since 2017

- 128 western drugs added to NRDL;
- Further 17 oncology drugs added to NRDL in Oct 2018 (15 in Jul 2017);
- Essential drug list expanded from 520 to 685 molecules. Including oncology.

Source: McKinsey

[1] Urban Basic Medical Care Insurance (for both employees & residents) - total persons covered at year-end. National Bureau of Statistics (2017); 2017-18 extrapolated based on growth in coverage of urban employees (no data for urban residents only after 2016); [2] Funds raised; [3] NDA = New Drug Application. Note: CAGR = Compound annual growth rate.

Cancer is a major unmet need in China ... investments in launches/access starting to have an impact















上市会

ELUNATE Fruquintinib Capsules

95% CI(月 8.18-10.4

5 88-8 11

Launched - Nov. 25, 2018

100SIE 伏特》组 曾備着°+BS 日間和+BSC -0.831 P < 0.001 9.30月 1.87个月

First ever oncology drug discovered & launched in China [1]







3rd-line colorectal cancer ("CRC")

1. Epidemiology







3. Latest status

- Launch of Elunate[®] underway & doing well
- In 5 weeks in Nov/Dec 2018: Revenues of \$3.3m from product purchases (manufacturing); & royalty of \$0.3m (15% of ~\$2.0m external sales);
- > Encouraging month-to-month growth trajectory.



Lilly amendment – Dec 2018 Secures long-term commercial potential



- **Chi-Med will pay full cost of any future development in China.** In return, Chi-Med gains:
- Freedom to operate in selecting & pursuing any future indications in China;
- Materially higher milestones & royalties upon launch in new LCI;
- Freedom to collaborate with any third-party in clinical development; and
- **Possible promotion rights in 30-40% of China for Elunate**[®]. Not expected before 2021, until then, Lilly responsible for all launch & commercialization costs in China. If we assume promotion rights, we will receive service fees, which we expect to be net income accretive.

	Original 2013 Agreement	Amendment (Dec 2018)
LCI ^[1] Development Costs – Paid by Lilly	70%	0%
LCI Development Costs – Paid by Chi-Med	30%	100%
LCI Regulatory Approval Milestones – Paid to Chi-Med ^[2]	12.5	20.0
Royalty Payments – Paid to Chi-Med ^[3]	15 - 20%	15 - 29%
Co-Promotion Rights in China (% of provinces) Co-Promotion Service Fees – paid to Chi-Med (% Net Sales)	0% 0%	30 - 40% Not disclosed

More control & higher long-term economics on bestin-class asset

[1] LCI = Life Cycle Indication; [2] Lifecycle Indication - China - per LCI, up to 3 LCIs; [3] On Total Molecule Sales in China triggered upon launch of 1st LCI.

Savolitinib – MET Exon 14 deletion NSCLC China's lead MET inhibitor



1. Competitive landscape outside China:

			Treatment Line	Ν	Investigator (ORR	95% CI	BIC	R ORR	95% CI
Capmatinib	selective	ESMO 2018 #LBA52	2/ <u>3L</u>	69	42.0% (29/	/69)	30.2%, 54.5%	39.1%	(27/69)	27.6%, 51.6%
(Novartis/ Incyte)	MET	ESMO 2018 #LBA52	1L	25	68.0% (17/2	/25)	46.5%, 85.1%	72.0%	(18/25)	50.6%, 87.9%
Tepotinib (Merck Serono)	selective MET	WCLC 2018 #12896	35% 1L, 65% ≥2L	40	57.5% (23/-	/40)	40.9%, 73.0%	42.9%	(12/28) ^[1]	24.5%, 62.8%
Xalkori®	multi-	WCLC 2018 #13453	38% 1L	65	32% (21/6	65)	21%, 45%		na	na
(Pfizer)	kinase	WCLC 2018 #12937	Median 1L (1L-4L)	25	na		na	40%	(10/25)	21%, 61%

2. Xalkori[®] a multi-kinase TKI – probably will be the first approval in MET Exon14 deletion pts outside China.

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


Savolitinib – MET Exon 14 deletion NSCLC Potential China NDA submission in 2020



- 4. Savolitinib aims to be 1st approved drug in China in MET Exon14 deletion NSCLC:^[1]
 - Expected fully enrolled in H2 2019.
 - Primary data expected in H1 2020.
 - Early CDE^[3] discussion potential accelerated approval.
- 2-3% of NSCLC est. incidence of ~10,000 new patients / year in China.
 Well over 400 screened to date.



6. Encouraging preliminary, midstudy China data at AACR 2019^[2]

- 41 pts; 31 pts efficacy evaluable.
- Promising antitumor activity.
- Rapid, durable tumor response observed.
- Anti-tumor activity observed in brain mets.
- Savolitinib generally well tolerated; most related TEAEs were grade 1 or 2.



[1] The trial is focused on patients with MET Exon 14 mutation who have failed prior systemic therapy, or are unable to receive chemotherapy, however the target patient population is intended to be all MET Exon 14 mutation patients.

[2] Data cut-off Feb. 26, 2019. Lu S et al, CT031 - Preliminary efficacy and safety results of savolitinib treating patients with pulmonary sarcomatoid carcinoma (PSC) and other types of non-small cell lung cancer (NSCLC) harboring MET exon 14 skipping mutations. Presented at American Association of Cancer Research Annual Meeting 2019, Atlanta, GA, Mar. 31, 2019.

[3] Center for Drug Evaluation of the National Medicinal Products Administration of China

Surufatinib – China NET – 2x Ph. III interims in 2019 Efficacy in all NET & patients who failed on Sutent®/Afinitor® [1]





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

8 assets in China development ... frug launched – savo/suru NDAs & Syk/PI3K δ PoC ahead



Program	Treatment	Indication	Target patient	Study name	Sites	Dose finding / safety run-in	Proof-of-concept	Registration
	Savolitinib	NSCLC	MET Exon 14 deletion		China	Lu Shun – SH Chest Hosp.		n ∽60
Savolitinib	Savolitinib + Iressa®	NSCLC	2L EGFRm; Iressa [®] ref.; MET+		China	Wu Yilong – GD General		Launchod
MET	Savolitinib	Gastric cancer	MET+		China	Shen Lin – BJ Univ. Tumor		Nov 2018
	Fruquintinib	Colorectal cancer	≥3L; chemotherapy refractory	FRESCO	China	Li Jin – Fudan Univ.		
	Fruquintinib + Taxol®	Gastric cancer	2L	FRUTIGA	China	Xu Ruihua – Sun Yat Sen		Interim
Fruguintinib	Fruquintinib	NSCLC	3L; chemotherapy refractory	FALUCA	China	Lu Shun – SH Chest Hosp.		Early 2019
VEGFR 1/2/3	Fruquintinib + Iressa®	NSCLC	1L EGFRM		China	Lu Shun – SH Chest Hosp.		Dublich 2010
	Fruquintinib + genolimzumab (PD-1)	Solid tumors			China	In planning		Publish 2019
	Fruquintinib + Tyvyt® (PD-1)	Solid tumors			China	In planning		Interim
	Surufatinib	Pancreatic NET	All	SANET-p	China	Xu Jianming – #5 Med. Ctr.		Late
Surufatinih	Surufatinib	Non-Pancreatic NET	All	SANET-ep	China	Xu Jianming - #5 Med. Ctr.		2019
VEGFR 1/2/3;	Surufatinib	Biliary Tract cancer	2L; chemotherapy refractory		China	Xu Jianming – #5 Med. Ctr.		Interim
FGFR1; CSF-1R	Surufatinib + Tuoyi [®] (PD-1)	Solid tumors			China	Shen Lin - BJ Univ. Tmr.		Edily
	Surufatinib + HX008 (PD-1)	Solid tumors			China	In planning		2017
	HMPL-523 + azacitidine	Acute Myeloid Leuke.	1L		China	Wang/Qi – CN Hem. Hosp.		
HMPL-523	HMPL-523	B-cell malignancies	All		China	Multiple leads by sub-types		
Зук	HMPL-523	ITP	All		China	Yang – CN Hem. Hosp. [1]		Planning China Ph.II/III
	HMPL-689	Indolent NHL			China	Cao/Zhou - Fudan/ Tongii		in several iNHL types
ΡΙ3Κδ								Ph.Ib data now n >110
	Faitinib		FC FDm with brain motastasis		Chipp			Data-set emerging in
E pitinid Eger	Epitinib		EGER gono amplified		China	Ving Mag - SH Huashan		China Ph.I (n ~31)
Lank	Epitilliv	GIIUDIdSLUIIId	EGER GEHE dilipilited		Clilla	Tilly Mau - SH Huasilali		
Theliatinib	Theliatinib	Esophageal cancer	EGFR over expression		China	Shen Lin – BJ Univ. Tumor [2]		
EGFR wt								
HMPI -453	HMPL-453	Solid tumors			China	Xu Ruihua – SYS		
FGFR 1/2/3								

[1] In planning; [2] Discontinued. ITP = immune thrombocytopenic purpura; PoC= proof of concept

China Oncology Main targets for 2019-2021



Sestablish Elunate[®] as the best-in-class VEGFR TKI in China market

- Work with Lilly to maximize penetration & sales performance;
- Aggressively expand PD-1 combination collaborations & broader LCI program

3 Launch our un-partnered oncology drugs

- Target surufatinib NDA in neuroendocrine tumors potentially in late 2019;
- Expand Oncology Commercial Org. from current ∽30 people to ∽200 by end 2020

Savolitinib NDA in MET Exon 14 NSCLC potentially in early 2020

Progress development pipeline

- Syk & PI3K δ into registration studies & aim to establish PoC for epitinib, theliatinib & FGFR;
- Aim for 2-3 further novel drug candidates into early development by 2021





Existing China business





Chi-Med spent 17 years building China commercial presence

- Valuable know-how in operating within the complex medical system in China
- Clear operating synergies with our novel oncology assets
- China operations/JVs have generated
 >\$500 million in Net Income since 2005
- China pharma industry grew at circa. 15% CAGR over last 15 years & set to continue ^[1]
 - Aging population; rapid urbanization; economic development

Chi-Med's Commercial Platform in China Integrated platform built from ground up



2 National House-Hold Name Brands



Major Commercial & Production Scale

2,500 RX & >950 OTC sales people in over 320 ^[1] cities & towns in China.

Drugs in ~24,900 hospitals detailing ~108,000 doctors.

Sold ~4.8 billion doses of medicine in 2018.

Leadership Market Shares

Market leader in the subcategories/markets in which we compete ^[2]:

SXBX pill: ^{[3][4]}	∽17%
Rx Cardiovascular TCM	
Banlangen: ^[5]	∽ 5 4%
OTC Anti-viral /flu TCM	
FFDS tablet: ^[6]	∽ <mark>38%</mark>
OTC Angina TCM	

JVs with 3 Major China Pharmas









Chi-Med's Commercial Platform in China Proven track record of success – important source of cash



Sales (Non-GAAP) [1][2] (US\$ millions) 2003-2018 (AGR.+26% 06 07 08 09 10 11 12 13 14 15 16 17 18

Net Income/(Loss) attrib. to Chi-Med



[1] 2003-2006 incl. disco. Operation; [2] Excluding Guanbao (from 2011 until divested in Sep 2017); [3] Based on aggregate Non-GAAP net income / (loss) of consolidated subsidiaries and non-consolidated joint ventures of Commercial Platform, please see appendix "Non-GAAP Financial Measures and Reconciliation".

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



...highly adaptable commercial platform 3rd party products – sales of Seroquel[®] & Concor[®] up significantly





Seroquel®, 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^{[1][2]}).

Luye acquisition. **Chi-Med retain rights through 2025 if we hit sales targets**. 2018 target RMB354m or +22% & +15% p.a. thereafter.



[1] In Oct 2017, as a result of the new NMPA Two-Invoice System 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 material 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 April 2015.



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

- Chi-Med runs **nine core territories covering ~600m people** – full service commercial role (fee-for-service).
- Took over from MS Jan-2015 ^[3].
- Leverages SHPL's existing >2,300 cardiovascular 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-for-service basis in Jan 2015.

Existing China Business Plans for 2019-2021





Continue organic growth

 Focus on proprietary prescription drug products. Mid- to long-term target of high single-digit percentage growth.

Build out synergies with China Oncology Organization

Strategically evaluate potential for M&A

- Expand the scope & scale of our joint ventures
- Continue to evaluate potential for divestment of certain non-strategic assets





3 Historical Financial Results and 2019 Guidance

2018 Operating Highlights

CHI-MED

Fruquintinib (Elunate®)

- Received China NDA approval for fruquintinib & launched in Nov 2018 for colorectal cancer;
- > Completed an agreement with Lilly to amend the original 2013 license & collaboration agreement.

Savolitinib

- Initiated two studies with potential for registration in lung cancer;
- > Presented Phase II data of Imfinzi[®] / savolitinib combo in papillary renal cell carcinoma.

Hematological malignancies

- > Australia & China Phase Ib expansion in lymphoma for HMPL-523 (Syk) & HMPL-689 (PI3K δ);
- Cleared U.S. IND applications (523/689). U.S. and E.U. clinical development set to start in H1 2019.

Immunotherapy combinations

Signed 4 co-development collaborations for fruquintinib & surufatinib PD-1 antibodies.

Global clinical development

Expansion of U.S. & international C&R operations. 5 Chi-Med drug candidates in global development.

2018 Financial results



	2016	2017	2018
GROUP REVENUES Unconsolidated JV Revenues ^[1]	216.1 401.5	241.2 433.3	214.1 491.5
SEGMENT NET INCOME/(LOSS) ^[2]			
INNOVATION PLATFORM	(40.7)	(51.9)	(102.4)
COMMERCIAL PLATFORM	29.9	37.5	41.4
<i>Prescription Drugs Business Consumer Health Business</i>	20.7 9.2	26.5 11.0	32.1 9.3
Chi-Med Group Costs	(17.9)	(14.8)	(13.8)
Land Comp. & Subsidies	40.4	2.5	-
GROUP NET INCOME/(LOSS) [2]	11.7	(26.7)	(74.8)
EPS Attrib. to Ord. S-H (Basic) (US\$)	0.20	(0.43)	(1.13)





2018 Financial results – Innovation Platform



\$26.9m revenues from Lilly:

- CRC approval milestone & service fees (\$23.3m);
- Last 5 weeks of 2018 Elunate[®] manufacturing revenue & royalty (\$3.6m).

R&D expenses of \$142.2m (non-GAAP):

- Development of 8 drug candidates (5 in U.S./International);
- Established GMP small molecule manufacturing (formulation) in China;
- Expanded U.S./International C&R operation in New Jersey.





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2018 Financial results - Commercial Platform



Net income up +10% to \$41.4m (non-GAAP):

- SXBX pill (cardiovascular Prescription drug) sales up +11% to \$233.1m;
- Seroquel[®] & Concor[®] service fees up +61% to \$21.2m.

China Two-Invoice System implemented:

- HSP sales lower due to move to fee-for-service model - from revenue consolidation - on some 3rd party drugs; No impact on net income;
- Restructure of Prescription Drugs distrib./logistics network under SHPL.



Cash position & 2019 guidance \$420 million in cash resources ^[1]

Cash Position

- \$301 million cash / cash equiv. / ST inv.^[2]
- \$119 million additional unutilized banking facilities ^[3]
- \$42 million additional cash in **Commercial JVs**
- **\$27 million** in bank borrowings ✓ Avg. cost 2.8%



	2019 Guidance	
Research & Development Expenses	(160) - (200)	
Adj. (non-GAAP) Group Net Cash Flows [4]	(120) - (150)	

Innovation Platform:

- Elunate[®] revenues ramp-up in coming years gradual start in 2019;
- Increase in R&D investment. U.S./E.U. expansion.

Commercial Platform

- China reforms ^[5] could narrow 2019 growth before seeing mid- to long-term benefit;
- RMB 5% weaker vs. US\$ than first half 2018.

[1] Including cash, cash equivalents, short-term investments & unutilized banking facilities; [2] Short-term investments: deposits over 3 months; [3] From Scotiabank, Bank of America Merrill Lynch, Deutsche Bank, Hong Kong Shanghai Banking Corporation; [4] Adjusted (non-GAAP) Group net cash flows excluding financing activities; [5] Two-Invoice System leading to change in SHPL distribution/logistics network & 4+7 Quality Consistency Evaluation System affects some of Hutchison Sinopharm's third-party products.

Adj.

CHI-MED

Objectives for existing assets 2019-2021



- NDA submission for savolitinib combo with Tagrisso[®]
- Expand savolitinib Exon14d development global
- 2 compounds to enter registration studies in 2020, surufatinib & fruquintinib
- Proof-of-concept achieved on both Syk & PI3Kδ compounds



- Establish Elunate[®] as best-in-class VEGFR TKI
- 2 further NDAs by mid 2020, savolitinib Exon14d NSCLC & surufatinib NET
- 2 more compounds into registration trials by 2020, Syk & PI3Kδ
- Expanded life cycle development on all assets, incl. PD-1 combos

Existing China Business

- Cash generative China Commercial Platform
- Platform for future innovative drug launches
- Opportunity for strategic exit













Savolitinib (AZD6094) Potential first-in-class selective MET inhibitor

Savolitinib (AZD6094)



Potential first-in-class selective MET inhibitor

2. MET is aberrant in many tumor settings. [7]

		New Cases (2018)			
Indication	Amplifi-cation	Mutation	Over- Expression	Global	China
Gastric	10%	1%	41%	1,033,700	442,300
Non-small Cell Lung Cancer	4%/16%/30% [1]	2% [2]	39%	1,779,800	737,400
Head & Neck	17-39%	11% [3]	46% [4]	887,700	137,000
Colorectal	10%	3%	65%	1,801,000	426,700
Papillary Renal Cell Carcinoma	64%	70-100% [5]	55%	45,400	3,700
Clear Cell Renal Cell Carcinoma	54%	NA	35%	281,300	57,500
Esophagus	8%	NA	92%	572,000	271,600
Prostate	NA	NA	54/83% [6]	1,276,100	99,300

4. AstraZeneca collaboration & 2016 amendment.

- \$20m received upfront (Dec 2011);
- \$120m in development/approvals milestones (\$25m received as of Dec 2018);
- 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%);
- From 9% up to 18% tiered royalty ex-China ^[8] & 30% flat rate China royalty on all product revenues.

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[1] MET amplification in non-small cell lung cancer patients occurs in approximately 4% of patients not previously exposed to systemic therapies and in approximately 16% to 30% of patients with acquired resistance to EGFR inhibitors; [2] MET Exon 14 skipping mutation only; [3] Oropharynx squamous cell cancer only; [4] Head and neck squamous cell cancer only; [5] Type 1 papillary renal cell carcinoma only; [6] MET expression is increased with progression of prostate cancer, which is 54% of lymph node metastases and 83% of bone metastases; [7] Company estimates considering Frost & Sullivan data, National Central Cancer Registry of China and publicly available epidemiology data; [8] Base royalty of 9%-13%. Additional 5% royalty subject to approval in the papillary renal cell carcinoma (PRCC) indication, for a total of 14%-18% tiered royalty. After total agregate sales of savolitinib have reached 55bn, the royalty will step down over a two-year period, to an ongoing royalty rate of 10.5% to 14.5%.

- 1. Strong potential to become first selective MET inhibitor approved in certain indications.
 - Clear clinical efficacy observed in non-small cell lung ("NSCLC"), kidney, gastric and colorectal cancers.
 - Partnered with AstraZeneca key comp. advantages in NSCLC (Tagrisso® combo) & biomarker testing.
- 3. Savolitinib design eliminates renal toxicity first
 generation of selective MET inhibitors encountered ~900 patients involved in clinical studies to date.



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

Savolitinib Biggest opportunity is MET+ NSCLC





[1] Primary drivers, based on aggregate rocelitinib/Tagrisso data published at 2016/2017 ASCO; [2] AstraZeneca 2016/17/18/1019 results and company estimates.

Savolitinib – MET Exon 14 deletion NSCLC Prelim data at AACR; Potential China NDA submission in 2020



1. Savolitinib aims to be first approved drug in China in MET Exon14 deletion NSCLC:

- Preliminary, mid-study China Phase II data^[1] presented at AACR 2019;
- Primary data completion expected in 2020;
- Study continues to enroll patients;
- 2-3% of NSCLC estimated incidence of ~10,000 new patients / year in China.

Preliminary interim data encouraging

- Promising antitumor activity;
- Rapid, durable tumor response observed;
- Anti-tumor activity observed in brain;
- Savolitinib generally well tolerated; most of the related TEAEs were grade 1 or 2.

Best response from baseline in efficacy # Stable disease (increase of the sum of the target tumors was less than 5mm): * PR unconfirmed vet

2. Xalkori[®] a multi-kinase TKI – probably will be the first approval in MET Exon14 deletion pts outside China.

	Savolitinib IC ₅₀	Xalkori® IC ₅₀	Savolitinib vs. Xalkori®	Source	Line of treatment	N	Investigator ORR	95% CI
EBC1 Viability EBC1 pMET	2nM 1	19nM 39	10x 40x	WCLC 2018 #13453	38% 1L	65	32% (21/65)	21%, 45%
293T MET (wild type) 293T MET (Ex14del)	7 9	79 140	11x 16x	WCLC 2018 #12937	[Median 1 (range 0-4)]	25	na	na

evaluable patients (%)



Savolitinib – 2L EGFRm NSCLC Very strong preclinical rationale for combination w/ EGFR-TKIs



Savolitinib – 2L NSCLC^[1] combo w/ IRESSA[®] compelling in MET+ / T790M-, next step under discussion

Savo / Iressa[®] combo in 1st gen. EGFRm-TKI refractory patients^[2]...outstanding response in MET+ / T790M-

WCLC 2017	MET+ / T790M+ (n = 23)	MET+ <i>(1790M-)</i> (n = 23)	MET+ / T790M unk (n = 5)
Confirmed response	2 (9%)	12 (52%)	2 (40%)
Stable disease≥ 6 weeks	9 (39%)	7 (30%)	2 (40%)
Progressive disease / death	7 (30%)	3 (13%)	0
Not Evaluable	5 (22%)	1 (4%)	1 (20%)
MET status all centrally confirmed.			

...vs. TATTON B data (savo / Tagrisso[®] combo)^[3]

	MET+ / T790M+ (n = 11) WCLC 2017 ^[2]	MET+ <i>(T790M-)</i> (n = 46) AACR 2019 ^[3]			
Confirmed response	6 (55%)	24 (52%)			
Stable disease≥ 6 weeks	NA (43% central confirm.)	16 (35%)			
Progressive disease / death	NA (0 central confirm.)	3 (7%)			
Not EvaluableNA (0 central confirm.)3 (7%)					
MET status locally or centrally confirmed.					

...Iressa[®] combo - <u>6</u> Duration of Response in MET+ / T790M- patients



[1] EGFRm NSCLC; [2] WCLC 2017 - Yang J-J, et al. A Ph.Ib Trial of savolitinib for patients with EGFR-mutant MET-amplified advanced NSCLC; [3] AACR 2019 - Sequist, et al. TATTON Phase Ib expansion cohort: Osimertinib plus savolitinib for patients (pts) with EGFR-mutant, MET-amplified NSCLC after progression on prior epidermal growth factor receptor (EGFR) tyrosine kinase inhibitor (TKI); [4] PR = Partial Response; [5] Aug 21, 2017.



Savolitinib – EGFR TKI Refractory NSCLC MET the main resistance mechanism for Tagrisso[®] 1L failure



Analysis from **plasma samples from FLAURA patients** who progressed or discontinued Tagrisso[®] (osimertinib) treatment. Frequency of MET amplification may be higher in tissue samples.

RESULTS: CANDIDATE ACQUIRED RESISTANCE MECHANISMS WITH OSIMERTINIB (n=91)*

- No evidence of acquired EGFR T790M
- The most common resistance mechanisms were *MET* amplification and EGFR C797S mutation
 - Other mechanisms included HER2 amplification, PIK3CA and RAS mutations



Ramalingam SS et al, "Mechanisms of acquired resistance to first-line osimertinib: preliminary data from the phase III FLAURA study", ESMO 2018 Congress, October 19, 2018.

Savolitinib – EGFR TKI Refractory NSCLC CHMET also the main resistance mechanism for Tagrisso[®] \geq 2L failure MEC

Analysis from **plasma samples from AURA3** patients who progressed or discontinued Tagrisso[®] (osimertinib) treatment. Frequency of MET amplification may be higher in tissue samples.

Acquired resistance mechanisms post-osimertinib (n=73)

Summary

- Acquired *EGFR* mutations: 21%
- MET amp*: 19%
- Cell cycle gene alterations: 12%
- HER2 amp*: 5%
- PIK3CA amp* / mutation: 5%
- Oncogenic fusion: 4%
- BRAF V600E: 3%





*Amplification events may be underrepresented in plasma analyses amp, amplification

Savolitinib – 2L/3L NSCLC^[1] – TAGRISSO[®] resistant MET+ driven resistance in ~30% of patients



3 out of 3 MET+ patients responded to savo/Tagrisso[®] combo.





LUL Mass Pre-Treatment 6 wks. on sa

Та	grisso® resista	a <mark>nt</mark> tissu	IA analysis 🛯 🖉 🖉	SSACHUSETTS NERAL HOSPITAL ANCER CENTER			
Pt	EGFR mutation	# Prior Therapies	Prior 3 rd 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 ND		
11	Del19	2	Y	MET amp, EGFR amp, T790 WT	T790M ND		
12	Del19	2	Y	_	T790M/C797S		
13	Del19	9		T790 WT	-		
	Del19	2	Y	T790 WT	T790M ND		
د	Del19	1		T790 WT	FGFR1 D60N, FGFR1 amp, T790M ND		
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		MET amp, EGFR 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		
(-) Testi	-) Testing not performed: EGFR - Epidermal Growth Factor Receptor: TKI- Turosine Kinase Inhibitor: amp - amplification: WT - wild tune: ND - not detected						

Safety & tolerability



Tagrisso[®] & savo both highly selective/tolerable monotherapies (MED

				Eff	ficacy	Discont	inuations as %	5 Enrolled
US FDA Approval	Treatment	Disease setting	n	ORR	Median PFS (mo.)	Due to AE	Withdrawn / Other	Total ^[5]
Monot	therapy – Tagrisso® / savolitini	b						
30-Mar-17	Tagrisso® (osimertinib)	2L EGFRi-refractory T790M+ NSCLC (AURA3)	279	71%	10.1	6%	6%	13%
	savolitinib 600mg QD monotherapy [3]	All-lines Papillary RCC FOR REFERENCE ONLY NOT NSCLC	109 [1]	18%	6.2	9%	5%	14%
Combi	nation – Tagrisso® + savolitini	b						
	savolitinib 600mg QD + Iressa® (gefitinib) [2]	≥ 2L EGFRm+ MET+ T790M- NSCLC after 1 st -gen EGFR TKI (expansion)	51	52%	ND	20%	14%	33%
	savolitinib 600mg QD + Tagrisso® [3]	≥ 2L EGFRm+ MET+ T790M-/+ NSCLC after 1 st -gen EGFR TKI (TATTON B)	46	56%	ND	37%	9%	46%
	savolitinib 600mg QD + Tagrisso® [4]	EGFRm+ MET+ NSCLC after 3 rd -gen EGFR TKI (TATTON B)	48	31%	ND	21%	4%	25%
Approv	ed treatments in NSCLC							
29-Apr-14	Zykadia® (ceritinib)	2L ALK+ NSCLC after Xalkori (single arm)	163	56%	6.9	10%	10%	20%
12-Dec-14	Cyramza® (ramucirumab) + Taxotere®	2L NSCLC after plat-chemo	624	23%	4.5	15%	21%	37%
24-0ct-16	Keytruda® (pembrolizumab) 2mg/kg	2L PD-L1+ (TPS≥1%) NSCLC after plat-chemo (KEYNOTE-010)	345	18%	3.9	10%	26%	37%
2-0ct-15	Keytruda® (pembrolizumab) 10mg/kg	2L PD-L1+ (TPS≥1%) NSCLC after plat-chemo (KEYNOTE-010)	346	18%	4.0	9%	27%	36%
9-0ct-15	Opdivo® (nivolumab)	2L NSCLC after plat-chemo	292	19%	2.3	15%	4%	20%
4-Mar-15	Opdivo® (nivolumab)	2L squ. NSCLC after plat-chemo	135	20%	3.5	12%	8%	20%
2008	Chemo doublet (platinum + pemetrexed)	2L NSCLC (AURA3)	136	31%	4.4	11%	17%	27%
1999	Taxotere® (docetaxel)	2L NSCLC (REVEL; KEYNOTE-010; Opdivo x2 aggregate total)	1,391	12%	3.5	13%	22%	36%

Tagrisso® + savo combo tolerable even in late-stage ≥3L patients

[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] WCLC 2017 #8995; [3] AACR 2019 CT032; 43 efficacy evaluable patients, 46 safety evaluable patients; ECOG = 0 in 30% of patients; [4] 2019 AACR CT033; 39 efficacy evaluable patients, 48 safety evaluable patients; ECOG = 0 in 50% of patients; [5] Total discontinuations NOT due to Disease Progression or Death; ND = Not Disclosed.

Safety - savolitinib plus IRESSA® or TAGRISSO gefitinib Or Adverse event profiles of combinations - manageable & tolerable MEC

	IPASS P 1 st -Line EGI		
Grade ≥3 AEs, Preferred term, n (%)*	IPASS Iressa® (N=607)	IPASS carbo. + Taxol® (N=589)	≥ 2 nd -Line ^[2] 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 st -Line EGFRm NSCLC				
Tagrisso® (N=279)	Iressa® 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 2 nd -Line E	Phase III GFRm NSCLC	
Tagrisso® (N=279)	Chemo-doublet (plat. + pemetrex.) (N=136)	≥ 2 nd -Line ^[1] Savo + Tagrisso® (N=94)
63 (23%)	64 (47%)	43 (46%)
1 (<1%)	3 (2%)	4 (4%)
2 (1%)		2 (2%)
6 (2%)	2 (2%)	4 (4%)
2 (1%)	5 (4%)	3 (3%)
3 (1%)	4 (3%)	3 (3%)
3 (1%)	1 (1%)	5 (5%)
4 (1%)	16 (12%)	4 (5%)
2 (1%)	16 (12%)	
	5 (4%)	
1 (<1%)	10 (7%)	

Sources: [1] TATTON B - Figures where any grade AE \geq 15% patients; AACR 2019 CT032 and CT032

[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.

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



2. MET- patients – no response to savo.



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

Tumor responses in the overall treatment population and by MET status

RECIST response,	MET+ (n=44)	MET- (n=46)	MET unknown (n=19)	Total (n=109)
		(11-40)	(11-12)	
Partial Response [†]	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.[†]Unconfirmed responses excluded. ^ Evaluable patients.

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





6



Highest selectivity delivers better tolerability

		PRCC PHASE II	COMPARZ P	HASE III ^[1]	METEOR PH	IASE III ^[2]	SINGLE-ARM PHASE III ^[3]	
		Savolitinib 1L/2L (n=109)	Sunitinib 1L (n=548)	Pazopanib 1L (n=554)	Cabozantinib 2L (n=331)	Everolimus 2L (n=322)	Sunitinib 2L (n=106)	
MSKCC Risk Group	Favorable Intermediate Poor Missing	14% 45% 9% 32%	27% 59% 9% 4%	27% 58% 12% 3%	45% 42% 12% 0%	46% 41% 13% 0%	58% 42% ^[6] 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 ^[4]	47%	77% ^[5]	76%[5]	68%	58%		
All Grade≥3 AEs with ≥5% incidence (AND selected savolitinib AEs for comparison)	Hypertension Fatigue Hand-foot-syndrome Diarrhea	TRAES 0% 2% 0% 0%	TRAES 15% 17% 12% 8%	TRAES 15% 11% 6% 9%	<u>All AEs</u> 15% 9% 8% 11%	All AEs 3% 7% <1% 2%	6% 11% 7%	Superior safety profile vs. other TKIs - Most \geq 3 G3 AEs \approx 0-2%: \checkmark Hypertension: 0% vs. 6~17%.
Hematologic Abnormalities Grade≥3 AEs with≥5% incidence:	Neutropenia Thrombocytopenia Lymphocytopenia Leukopenia Anemia	0% 0% 0% (1%	20% 24% 14% 6% 7%	5% 4% 5% 1% 2%	0% 0% 0% 5%	0% 0% 0% 16%	16% 6% 6%	 ✓ 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%	4% 3% 9% 1% 4%	17% 12% 4% 7% 3% 5%	2% 2% 4% 0% 5% <1%	<1% <1% 2% 0% 2% 5%		 ALI/AST Increase: 3-5% vs. 0∽17%. ✓ Other Lab Abnorm: 0% vs. ≤9%. Highly tolerable vs. other TKIs:
Tolerability	Treatment discontinuation due to any AE ^[7] : Dose reduction due to AE:	8% 13%	20%	24%	12%	11%	11%	 ✓ Discontinued: 8% vs. 10∽24%. ✓ Dose reduction: 13% vs. 44-62%.

[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. TRAEs = Treatment-Related Adverse Events; [7] 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 5th most common cancer globally - 782,700 deaths/year

	New cases ('000)	Deaths ('000)	5-year Prevalence ('000)
World	1034	783	1,590
South Korea	38	8	104
Japan	116	49	298
China	442	318	604
EU-28	133	56	195
USA	26	11	41

overall survival ("OS") in first-line palliative setting.



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





Savo potential not only in NSCLC... ...highly promising efficacy in MET+ gastric cancer (...& kidney)





MET amp. +ve (n=39)

120

140

100



[1] mOS = median overall survival post surgery.

P = 0.0003

40

60

80

Time After Surgery (Months)

20

20





Fruquintinib best-in-class VEGFR TKI Cutting off blood flow a \$16+bn market in ~30 tumor settings



	Drug	FDA Approved Indications				
Company	(INN Name)	Indication	Үеаг	2016 Sales		
		2L bevacizumab-pretreated mCRC	2013			
	Avastin® (Bevacizumab)	1/2L mCRC	2004			
		1L non-sq NSCLC	2006			
		2L GBM	2009	_		
Roche		1L ccRCC	2009	\$6,890m		
		1L Cervical Ca.	2014			
		1L Ovarian Ca.	2018			
		1/2L platinum-sensitive Ovarian Ca.	2016			
		2/3L platinum-resistant Ovarian Ca.	2014			
		2L GIST	2006	-		
	Cutopt®	≥1L pNET	2011			
Pfizer	, Sutent® (Sunitinib)	adjuvant RCC	2017	\$1,049m		
		1L RCC	2007			
		≥2L cytokine-ref. ccRCC	2006			
Boehringer Ingelheim	Vargatef® Ofev® (Nintedanib)	2L adeno-NSCLC (by EMA)	2014	\$1,076m ^[1]		
	Nexavar® (Sorafenib)	≥1L RCC	2005			
Bayer		1L HCC	2007	\$788m		
		lodine-ref. DTC	2013			
Novartis	Votrient®	1/2L RCC	2009	¢878m		
Novarus	(Pazopanib)	2L STS	2012	#020III		
	Cyramza® (Ramucirumab)	2L GC	2014	\frown		
Lilly		2L NSCLC	2014	\$821m		
		2L mCRC	2015			
Fxelixis/	kis/ Cometriq® Cabometyx®	\geq 1L MTC	2012			
Insen		1L ccRCC	2017	\$783m		
	(Cabozantinib)	≥2L ccRCC	2016			
_	Stivarga®	3L mCRC	2012	\frown		
Bayer	(Regorafenib)	2L GIST	2013	\$348m		
	(2L HCC	2017			
Pfizer	Inlyta® (Axitinib)	2L ccRCC	2012	\$298m		

	Drug	FDA Approved Indications		
Company	(INN Name)	Indication	Year	2010 30103
Merck/	Lenvima®	lodine-ref. DTC	2015	
Eisai	(Lenvatinib)	2L ccRCC	2016	\$575m
	(,	1L HCC	2018	
Hengrui	AiTan® (Apatinib)	3L GC (by CFDA)	2015	\$255m
Sanofi	Zaltrap® (Ziv-Aflibercept)	2L mCRC	2012	\$101m
Simcere	Endu® (rh-Endostatin)	≥1L NSCLC (by CFDA)	2005	NA
Sanofi	Caprelsa® (Vandetanib)	≥1L MTC	2011	NA
Aveo	Fotivda® (Tivozanib)	1/2L ccRCC (by EMA)	2017	NA
Sino Biopharm	FocusV® (Anlotinib)	3L NSCLC (by CFDA)	2018	NA
VEGF Produ	iction	Blood vessel grov	vth	
Tumor			► Tum	or growth -
			WWW C	- V
			Me	etastases

Note: * Active indications in US as of July 3, 2018. Some indications have been approved for frontline therapy. Sources: FDA approved label; Medtrack; Corporate annual reports; D. Ri batti, Oncotarget 2017 8(24) 38080-1, Sales for anti-angiogenic drugs. [1] 2017 sales, includes sales for idiopathic pulmonary fibrosis
Fruquintinib – 24hr full target coverage

The most selective VEGFR inhibitor in clinical trials globally ^[1]



- ✓ Validation of R&D approach designed to only inhibit VEGFR1,2,3, facilitating full target coverage & combinations.
- ✓ Approval and launch for 3L CRC.
- ✓ Pivotal Phase III Taxol[®] combo in 2L gastric cancer initiated Oct 2017.
- ✓ Phase II Iressa[®] combo in 1L EGFRm+ NSCLC early data at WCLC 2017.
- ✓ Phase I in solid tumors in US initiated Q4 2017.
- ✓ China GMP **facility built and certified** to support launch.
- ✓ PD-1 combination collaborations.

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



3. Selectivity and potency superior to competitors' drugs.

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

[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.



Fruquintinib – 3L/4L colorectal cancer Develop in US/EU for rego/TAS-102 ref./intol. patients^[1]



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



[1] ASCO = American Society of Clinical Oncology Annual Meeting.

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Better tolerability = Better efficacy

Fruquintinib		Regorat	fenib	Regorafenib		Regorafenib		
Third-Line Metastatic Colorectal cancer	FRES	FRESCO CONCUR Mainland China Chinese Patients (Mainland China, Hong Kong, Taiwan) ^[1]		CONCUR		CONCUR		ECT
	Mainland			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 (%)	0.4%	0.0%	0.0%	0.0%	0.0%	0.0%	0.0%	0.0%
Partial Response, n (%)	4.3%	0.0%	3.6%	0.0%	4.4%	0.0%	1.0%	0.4%
Stable Disease, n (%)	57.6%	12.3%	40.2%	6.7%	45.6%	7.4%	42.8%	14.5%
Disease Control Rate, n (%)	62.2% +4%	2) 12.3%	45.5% +38	.8 6.7%	51.5% 🖽	.1 7.4%	41.0% +26.	D 14.9%
Median Progression-Free Survival (mPFS) (mo.)	3.7 +1.9	1.8	2.0 +0,	3 1.7	3.2 +1,	5 1.7	1.9 +0.2	1.7
mPFS p-value	<0.0	01	not publ	ished	<0.0	001	<0.000	001
mPFS Hazard Ratio	0.2	6	0.32	2	0.3	31	0.4	9
			6		6			
Median Overall Survival (mOS) (mo.)	9.3 +2.	6.6	8.4 +2	2 6.2	8.8 +2	6.3	6.4 +14	5.0
mOS p-value	<0.0	01	not publ	ished	0.00	002	0.00	52
mOS Hazard Ratio	0.6	5	0.56		0.5	5	0.7	7

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

ELUNATE[®] Fruquintinib Capsules



	ELUNATE [®] Fruquintinib Capsules	Stivarga [®] (regorafenib) tablets
BIOCHEMICAL ACTIVITY	IC ₅₀ (nmol/L)	IC ₅₀ (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	7
PDGFRβ	>10,000	22
RAF-1	>10,000	2.5
B-RAF	>10,000	28
B-RAF ^{V600E}	>10,000	19

Stivarga[®] liver toxicity black-box warning:

→ Increased liver function test monitoring (weekly if elevated) & remedial dose interruption.

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

> WARNING: HEPATOTOXICITY See full prescribing information for complete boxed warning.

- Severe and sometimes fatal hepatotoxicity has been observed in clinical trials. (5.1)
- Monitor hepatic function prior to and during treatment. (5.1)
- 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)

	ELUNATE®		Stiva (regorafenil	arga [®] 1) tablets
3 rd -Line Metastatic Colorectal cancer	FRESCO Mainland	Study China ^[1]	CONCUE (Mainland China	R Study 1, HK, Taiwan) ^[2]
Treatment arms	Elunate®	Placebo	Stivarga [®]	Placebo
Patients (n)	278	138	112	60
≥G3 AE (Safety population)	61.1%	19.7%	69.6%	46.7%
SAE (Safety population)	15.5%	5.8%	31.3%	26.7%
VEGFR on-target related AEs:				
Hypertension ≥G3	21.2%	2.2%	12.5%	8.3%
Hand-Foot Syndrome (Palmar-plantar), ≥G3	10.8%	0.0%	17.0%	0.0%
Off-target (i.e. non-VEGFR) related AEs:				
Hypophosphatemia, ≥G3	0.0%	0.0%	8.0%	0.0%
Hypokalemia, ≥G3	0.7%	0.7%	6.3%	0.0%
Rash/desquamation, ≥G3	0.0%	0.0%	4.4%	0.0%
Lipase increase, ≥G3	0.0%	0.0%	6.3%	1.7%
Hepatic function (Liver function) AEs:				
ALT increased, ≥G3	0.7%	1.5%	7.1%	3.3%
AST increased, ≥G3	0.4%	0.7%	8.9%	0.0%
Blood bilirubin increased, ≥G3	1.4%	1.5%	8.9%	8.3%
Tolerability:				
AE Leading to dose interruption	35.3%	10.2%	68.8%	25.0%
AE Leading to dose reduction	24.1%	4.4%	23.2%	0.0%
AE Leading to treatment discontinuation	15.1%	5.8%	14.3%	6.7%

Elunate[®] higher selectivity; lower off-target toxicity; superior tolerability

FALUCA – Third-line NSCLC



FALUCA Phase III

- 527 NSCLC (3rd-line) patients enrolled;
- Topline results released Nov 2018;
- Anticipate presenting full data set and analysis at scientific conference in 2019.

Phase II Study (reported May 2015)

- 91 NSCLC (3rd-line) patients enrolled;
- Clearly met Primary Endpoint: mPFS vs. placebo;
- AEs consistent & more tolerable than \geq 3L CRC ^[2].

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%)

FALUCA Phase III – Topline Results

- Did not achieve Primary Endpoint of median Overall Survival;
- Clearly met all Secondary Endpoints: mPFS; ORR; DCR; & Duration of Response vs. placebo^[1];
- **AEs consistent** with those observed in prior clinical studies.



Phase II – Median PFS

Fruquintinib – 1L NSCLC combo w/ IRESSA[®] gefitinib Two small molecule TKIs allow for better management of tox.





2. Prelim. safety data: fruquintinib vs. other VEGFRis.

Adverse Events ("AEs")	Iressa® or Tarceva® FLAURA ^[5]	Avastin®+ Tarceva®[6]	Fruquintinib + Iressa®
	N = 277, n (%)	N = 75, n (%)	N = 26, n (%) ^[3]
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. ^[2,3]

PR	PR	PR		PR		PR		✦
PR	PR	PR		PR		PR	+	
SD		SD		→				
PR	PR	PR	+					
SD		PR 🟓	•					
PR	PR	<u>→</u>	•					
PR	PR	PR 🔿	•					
SD		SD →	•					
PR	PR	→ ·						
PR	PR [4	4]						
PR								
SD	_							
РК								
חח			5 m	a fruauint	tinib + 25	0ma li	ressa®	
	•		4m	a fruquini		oma li	rocca®	
			4111	g nuquin	11110 + 25	Unign	lessa	
→ ×			3 m	g fruquint	tinib + 25	Omg li	ressa®	
→			fruo	nuintinih :	and Iressa	e inte	rrunte	h
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→			PR Par	tial respo	nse ^[2]			
→			SD Stal	ble diseas	e			
→				atmont co	ntinuina			
PR 🔶					mununiy			
PR 🔶								
→								
28 56	84	112	140	168	196	224	25	52
Data as of October 10, 2017.	D	uration of	Treatment (da	ays)				

[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; mAb = Monoclonal Antibody; [3] Lu, S., et al., "A Phase II study of fruquintinib in combination with gefitinib in stage IIIb/IV NSCLC patients harboring EGR activating mutations", ID 10907 IASLC 18th 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) EGR-TKI as first-line therapy in patients (pts) with EGRm 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 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 (bc) with EGR-TKI as first-line therapy.", an open-label, randomised, multicenter, phase 2 study", The Lancet 2014, 15 (11) 1236-1244.

Fruquintinib – Gastric combo with paclitaxel Phase III initiated Oct 2017 – Interim analysis early 2019



1. Dose proportional increase of fruquintinib AUC at steady state. Over 30% increase in paclitaxel drug exposure (mean AUC₀₋₈) following multiple dose fruquintinib.



2. ORR of 36% (10/28) & DCR of 68% in efficacy evaluable pts. Fruquintinib 4mg, ≥ 16 wk. PFS of 50% & ≥ 7 mo. OS of 50%.



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 ²				
	Drug interruption	Drug reduction			
Dose modification with Fruquintinib N (%)	2 (10.5%)	2 (10.5%)			
Dose modification with Paclitaxel N (%)	5 (26.3%)	1 (5.3%)			

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

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



VEGFR / immunotherapy (PD-1s) combinations



Potent two prong attack - Anti-angiogenesis + activated T-cell response

Source: 1. B. Rini et al, Lancet Oncol 2013 14(12) 1233-42, Axitinib with or without dose titration for first-line metastatic renal-cell carcinoma: a randomised double-blind phase 2 trial; 2. D.F. McDermott et al, ASCO 2018 #4500, Pembrolizumab monotherapy as first-line therapy in advanced clear cell renal cell carcinoma (accRCC): Results from cohort A of KEYNOTE-427; 3, . B. Rini et al, for the for the KEYNOTE-426 Investigators, NEJM 2019 Feb 16. doi: 10.1056/NEJMoa1816714, Pembrolizumab plus Axitinib versus Sunitinib for Advanced Renal-Cell Carcinoma.

Fruquintinib & surufatinib both unique VEGFR TKISideal VEGFR combination partners for immunotherapy



TKI	15	st Generatio	n	2	2 nd Generation		Next Generation		
Selectivity		Multiple targets			Relatively selective		Highly selective	Selective angio-immuno kinase inhibitor	
Inhibitors	Sunitinib	Sorafenib	Anlotinib	Tivozanib	Lenvatinib	Axitinib	Fruquintinib	Surufatinib ^[1]	
Status	Launched	Launched	Launched	Launched	Launched	Launched	Launched	Ph. IIIs ongoing	
VEGFR1 (nM)	2	26	27	30	22	3	33	2	
VEGFR2 (nM)	9	90	0.2	6.5	4	7	25	24	
VEGFR3 (nM)	19	20	0.7	15	5	1	0.5	1	
Phos-KDR (nM)	10	30	0.1-1	0.16	0.8	0.2	0.6	2	
Other kinases (IC50 < 100nM)	PDGFR _α PDGFRβ c-Kit Flt3 Ret CSF-1R	Raf-1 b-raf Flt3 P38 c-Kit Ret	PDGFR _α PDGFR _β FGFR1-4 c-Kit	PDGFR _α PDGFRβ EphB2 c-Kit Tie2	PDGFR _α PDGFR _β FGFR1-4 Ret c-Kit	PDGFR _α PDGFR _β c-Kit	none	CSF-1R FGFR1 FLT3 TrkB	
Patent Expiration					2021/10/19 (US7253286B2)	2025/04/29 (US6534524B1)	2029 (without extension)	2030 (without extension)	

Fruquintinib is uniquely selective – unlike other TKIs with off-target toxicity
 Surufatinib inhibits TAM^[2] production – amplifying PD-1 induced immune response



Chi-Med immunotherapy collaborations



5 PD-1/PD-L1 combos underway/in planning on savo, fruq & suru







Surufatinib Highly active TKI with unique angio-immuno activity

Surufatinib's unique angio-immuno kinase profile CH Multi-indication global development program, initially for NETs^[1]

Surufatinib's unique angio-immuno kinase profile & MoA^[1] activates & enhances the body's immune system, namely T-cells, via VEGFR/FGFR while inhibiting the production of macrophages (CSF-1R) which cloak cancer cells.



VEGFR / FGFR

FGFR Antigen release (activation of T-cells)



Aiming for fast/first approval in China for all NET^[1] patients

Pancreatic NET ("P-NET") & Non-Pancreatic NET ("EP-NET")

- SANET-p & SANET-ep active in 25 China sites.
- Primary endpoint median PFS.
- Target Interim Analysis in 2019
 SANET-ep in H1 2019 & SANET-p in H2 2019.
- Enrolment expected to complete late 2019 / early 2020.
- Potential launch in China in late 2020 / 2021 first un-partnered oncology asset for Chi-Med.



Biliary Tract Cancer ("BTC")

- Clear unmet medical need a few agents being tested in 2L BTC but standard of care not yet established.
- Phase II PoC^[2] initiated in early 2017.
- Phase II/III pivotal study in BTC in China initiated H1 2019.

U.S. Development Expanding

- Phase I dose escalation study in the U.S. completed (N=29), 5 dose cohorts (50-400mg QD), established
 300mg. QD as RP2D (same as China).
- U.S. Phase Ib/II study in P-NET & BTC initiated July 2018.
- PD-1 combination collaborations.

Surufatinib – China NET – Phase II *(ENETS 2017*^[1]) Tumor devascularization & central necrosis





Patient 2 w/ multiple liver metastases











[1] Rituxan[®] 2018 sales in oncology only; [2] Approved Drug = @; All others are clinical candidates; [3] ASH = American Society of Hematology; [4] Chronic lymphocytic leukemia ("CLL") & small lymphocytic lymphoma ("SLL"); [5] Sharman et al, ASH Meetings 2015 & 2016; [6] CYP3A4, CYP2D6 and CYP 1A2.

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





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

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

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

1. Fostamatinib good Phase II^[1] RA^[2] dose response...

Selectivity	HMPL-523 IC ₅₀ (nM)	fostamatinib IC ₅₀ (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; GI = Gastrointestinal; 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

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**^[2] market in 2020 with B-cell pathway; anti-TNF; & JAK the main focus.

(Methotrexate-IR: placebo adjusted)	ACR20	ACR50	ACR70	2018 Sales (\$ billion) ^[3]
B-Cell receptor mAbs				
Rituxan® (24-Week)	33%	21%	11%	1.6
Anti-TNFα/NF-κB mAbs				
Humira® (24-Week)	33%	29%	18%	19.9
Remicade [®] (24-Week)	30%	22%	8%	5.3
Enbrel® (24-Week)	44%	36%	15%	6.9
JAK Inhibitors Small molecules				
Xeljanz® (24-Week)	25%	23%	13%	1.8
Xeljanz® (12-Week)	28%	21%	8%	1.0
baricitinib 4mg QD (12-Week)	30%	28%	14%	n/a
filgotinib 100mg BID (12-Week)	35%	40%	23%	n/a
ABT-494 24mg QD (12-Week)	32%	24%	18%	n/a
Syk Inhibitor Small molecule				
fostamatinib 100mg BID (24-Week)	32%	24%	18%	n/a

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

HMPL-689 – Phase I Australia & China ongoing Designed to be a best-in-class inhibitor of PI3K δ



1. PI3K δ now a proven target.

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



2. PI3K δ inhibitors being developed in a very broad range of indications.

Compound		Indication	Status	Issue
Zydelig® (idelalisib) PI3K&	Gilead	Chronic lymphocytic leukaemia, non-Hodgkin's lymphoma	Marketed	High incidence of liver toxicity seen with idelalisib (150mg bid)
AMG-319 ΡΙ3Κδ	Amgen	B-cell lymphoma, non-Hodgkin's lymphoma, T-cell lymphoma, chronic lymphocytic leukaemia	Phase I Trial	
Copiktra®		Relapsed or refractory chronic lymphocytic leukaemia / small lymphocytic lymphoma	Approved	Need to spare PI3Ky serious infection seen &
(duvelisib) PI3Kγ/δ	Infinity ^[1]	Relapsed or refractory follicular lymphoma	Approved ^[2]	associated with a boxed warning for 4 fatal and/or
		Peripheral T-cell lymphoma	Phase II enrolling	serious toxicities
Aliqopa [®] (copanlisib) PI3Kα/δ	Bayer	Relapsed follicular B-cell non-Hodgkin lymphoma	Approved ^[2]	Serious and fatal infections and AEs

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.

4. More potent / more selective than Zydelig[®], Copiktra[®] & Aliqopa[®].

HMPL-689	Zydelig®	Copiktra®	Aliqopa®
0.8 (n = 3)	2	1	0.7
114 (142x)	104 <mark>(52x)</mark>	2 (2X)	6.4 (9x)
>1,000 (>1,250x)	866 <mark>(433x)</mark>	143 (143x)	0.5 (1X)
3	14	15	n/a
87 (109x)	293 (147x)	8 (8X)	3.7 (5x)
	HMPL-689 0.8 (n = 3) 114 (142x) >1,000 (>1,250x) 3 87 (109x)	HMPL-689 Zydelig® 0.8 (n = 3) 2 114 (142x) 104 (52x) 1,000 (>1,250x) 866 (433x) 3 14 87 (109x) 293 (147x)	HMPL-689Zydelig®Copiktra®0.8 (n = 3)21114 (142x)104 (52x)2 (2x)>1,000 (>1,250x)866 (433x)143 (143x)3141587 (109x)293 (147x)8 (8x)

[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 trials.







Epitinib – 70% response in NSCLC w/ brain mets^[1] Unmet medical need. Investment case under review.



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



2. Phase Ib ^[1] - solid/durable <u>efficacy in brain</u> in EGFRm+ NSCLC patients with measurable brain mets (>10mm).



[1] Dose expansion stage - data cut-off September 20, 2016; [2] 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; * Unconfirmed PR, due to no further assessment at cut-off date; # Includes both confirmed and unconfirmed PRs; ^ MET amplification/high expression identified.



Epitinib – Strong PoC efficacy – 160mg QD dose





Epitinib - Safe & well tolerated

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

Dose Escalatio (Drug related AE	n Stage (n= Es reported	:35*) →10%)	Dose Expansion Stage (n=37) (Drug related AEs reported >10%)						
160mg QD dose	All Grades n (%)	Grade 3/4 n (%)	160mg QD dose	All Grades n (%)	Grade 3/4 n (%)				
Skin rash	21 (60.0%)	1 (2.9%)	Skin rash	31 (83.8%)	2 (5.4%)				
Diarrhea	12 (34.3%)	-	Hyper-pigmentation	18 (48.6%)	1 (2.7%)				
AST increase	12 (34.3%)	1 (2.9%)	ALT increase	15 (40.5%)	7 (18.9%)				
ALT increase	11 (31.4%)	1 (2.9%)	AST increase	15 (40.5%)	4 (10.8%)				
Total bilirubin increase	10 (28.6%)	2 (5.7%)	ASP increase	11 (29.7%)	1 (2.7%)				
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 (2.7%)				
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 (10.8%)				
Conjugated bilirubin	4 (11.4%)	1 (2.9%)	Stomatitis	6 (16.2%)	-				

4. 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
- 1st-line chemo naïve
- 120mg QD dosage
- 25 weeks (177 days) on treatment with clear response in multiple measurable (>10mm diameter) brain lesions



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Theliatinib Potent & highly selective TKI – strong affinity to EGFRwt kinase



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

- EGFR TKIs are less effective in solid tumors with wild-type EGFR activation (gene amplification & protein over expression).
- Ph.Ib study in esophageal cancer short-term response & stable disease observed. Does not warrant continued development as monotherapy. Consider potential immunotherapy combo.

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

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

- 5-10-fold more potent than Tarceva[®].
- 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).





HMPL-453 – Phase I in China ongoing Designed as best-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	Gastric (5∽10%) Breast (4%)	Intra-hepatic biliary tract cancer (cholangiocarcinoma) (14%) Breast (n/a)	Endometrial (12~14%) Lung squamous (5%)
FGFR3	Bladder (n/a) Salivary adenoid cystic (n/a)	Bladder (3~6%); Lung squamous (3%); Glioblastoma (3%) Myeloma (15~20%)	Bladder (60~80% NMIBC; 15~20 MIBC) Cervical (5%)

- 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).









Chi-Med Group Structure - Major Entities



[1] Excluding Guanbao (divested in Sep 2017); [2] Non-GAAP: excludes the share of government subsidies from SHPL of \$2.5million in 2017; [3] Excluded HSP's Zhi Ling Tong infant nutrition business, revenue from prescription drug business has decreased by 20% as a result of the Chinese government's implementation of the new Two-Invoice System ("TIS"), pursuant to which we had converted to earning service fees from the commercialization of certain third-party products instead of recognizing the gross sales from these products in our revenue as we had done prior to implementation of TIS in October 2017; despite the TIS change, service fees (non-GAAP) earned from the key third-party product, anti-psychotic Seroquel®, grew rapidly, up 51% to \$17.2 million (2017; \$11.4m); [4] Held through an 80% owned subsidiary.

FY2018 Inter-group cash flow \$301.0m cash (Dec 31, 2018); \$119.3m in undrawn bank facilities





[1] \$8.0m capital injection to NSP offset by \$6.9m service income received from NSP; [2] Including research & development cost and general & admin. expenses; [3] Share of NSP operating loss; [4] Including \$214.9m short-term investment (deposits over 3 months) as at end of 2018; [5] Please see appendix "Non-GAAP Financial Measures and Reconciliation" for a Reconciliation of GAAP to adjusted research and development expenses.



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 median PE multiples is approximately \$1.7 1.8 billion.^[1] Given our share in the JVs, Chi-Med's share of this value is approximately \$0.8 0.9 billion.

		NET SALES				NET I	NCOME		VALUATION ^[4]	
	Code	2017 Jan-Jun	2018 Jan-Jun	17-18 1H Growth	2017 Jan-Jun	2018 Jan-Jun	17-18 1H Growth	2018 1H Margin	Market Cap.	P/E
CHI-MED Commercial Platform Subsidiaries/JVs ^[2]		328.0 ^[3]	360.3	10%	51.9	55.1	6%	15%	n/a	n/a
Tianjin Zhong Xin Pharma	600329	451.3	470.2	4%	41.6	47.6	14%	10%	1,699	22
Li Zhu Pharma	000513	645.8	689.6	7%	83.2	102.1	23%	15%	3,619	21
Shandong Dong E E Jiao	000423	443.3	451.1	2%	136.4	130.5	-4%	29%	4,519	15
Zhejiang Kang En Bai Pharma	600572	353.6	540.3	53%	58.6	83.1	42%	15%	3,201	24
Kunming Pharma	600422	412.4	511.4	24%	32.7	27.7	-15%	5%	831	18
Guizhou Yi Bai Pharma	600594	294.9	285.9	-3%	30.0	26.2	-13%	9%	723	18
Jin Ling Pharma	000919	258.5	236.4	-9%	18.6	17.4	-6%	7%	533	31
Jiangsu Kang Yuan	600557	251.3	278.7	11%	29.3	30.8	5%	11%	1,137	19
Zhuzhou Qian Jin Pharma	600479	228.5	225.0	-2%	8.1	12.1	49%	5%	572	13
ZhangZhou Pian Zai Huang	600436	264.8	363.2	37%	63.9	91.7	44%	25%	9,681	62
Peer Group Median (10 Comps. excl. Chi-Med)		324.2	407.1	26%	37.2	39.2	6%	10%	1,418	20
All 61 Listed China Pharma. Companies Median		258.5	278.7	8%	29.3	31.6	8%	11%	1.137	21

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 2018E Net Sales in the ~\$400-1,400 million range.

Source: Company data, Deutsche Bank, FactSet

[1] Peer group/China Pharma multiple of 20X-21X 2018 actual Net income after tax of \$83.6 million; [2] Total aggregate PRC domestic results of Chi-Med's 6 Commercial Platform companies (HBYS, SHPL, Hutchison Sinopharm, HHO, HHL & HCPL); [3] Excluding Guanbao (US\$ millions) (divested); [4] Market Capitalization and Price Earnings Ratios as at March 1st, 2019: Trailing Twelve Month PE weighted averaged based on market capitalization.

Deep portfolio of household name drugs



Top 7 products represent 69% of sales^[1] and 89% of gross profit^[1]

Main Product	ts ^[2] – SALES (Non-GAAP)	2012	2013	2014	2015	2016	2017	2018	
「小田」の	SXBX pill Coronary artery disease (Rx) 17% National market share Patent expiry 2029	102,215 <i>+29%</i>	123,587 +21%	138,848 <i>+12%</i>	159,326 +15%	195,37 1 <i>+23%</i>	209,246 +7%	233,096 +11%	
	Banlangen granules Anti-viral/flu (OTC) 54% National market share	65,381 +14%	72,300 +11%	55,573 <i>-23%</i>	54,793 -1%	56,664 +3%	59,898 +6%	62,585 +4%	
	FFDS tablet Angina (OTC) 38% National market share	60,181 +6%	69,996 +16%	76,297 +9%	60,154 -21%	59,906 <i>0%</i>	58,936 -2%	56,342 -4%	
	<i>NXQ tablet</i> Cerebrovascular disease (OTC) Proprietary formulation	6,933 <i>+85%</i>	10,142 +46%	1 4,681 +45%	17,581 +20%	21,000 +19%	20,408 - <i>3%</i>	37,250 +83%	
Seroquel XR and	<i>Seroquel tablets</i> Bi-polar/Schizophrenia (Rx) 6% National market share	n/a	n/a	n/a	21,131	34,380 +63%	35,359 +3%	29,211 ^[3] - <i>17%</i>	
	<i>KYQ granules</i> Periodontitis (OTC) >90% National market share	16,351 +6%	16,318 <i>0%</i>	18,370 <i>+13%</i>	17,051 <i>-7%</i>	17,210 +1%	17,620 +2%	19,329 <i>+10%</i>	
	Danning tablet Gallbladder/stone (Rx) Patent expiry 2027	11,648 <i>+17%</i>	12,364 +6%	13,822 <i>+12%</i>	13,526 -2%	9,041 <i>-33%</i>	16,089 <i>+78%</i>	17,378 +8%	

(US\$'000)

(Growth % vs. Year Ago)

[1] Based on aggregate Non-GAAP Sales 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-thecounter 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 Two-invoice policy. Net service fee increased by 51% to \$17.2m in 2018 (2017: \$11.4m).

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



Reconciliation of Adjusted Group net cash flows and Adjusted Group net cash flows excluding financing activities:

Reconciliation of Top 7 products' Gross Profit as Percentage of Aggregated Gross Profit for Commercial Platform:

				2018
Cach and each aquivalents and short term investments	2018	2019 Guidanc	Revenue from external customers	172.0
cdsir driu cdsir equivalents driu short-term investments at end of year	301.0	150-180 [1	– commercial platform	172.9
Less: cash and cash equivalents and short-term		(2.2.2)	Less: Costs of goods and services	(142.4)
investments at beginning of year	(358.3)	(300)	Gross profit – commercial platform	30.5
Adjusted Group net cash flows	(57.3) (120)		Add: Gross profit – HBYS and SHPL	306.1
	(37.3)	(120) (130)	Adjusted gross profit	336.6
Add: Net cash used in financing activities for the year	8.2	[1		
Adjusted Group net cash flows excluding financing			Top 7 products gross profit	298.1
activities	(49.1)	(120) - (150)	% of Top 7 products to adjusted gross profit	89%
Reconciliation of Adjusted Service Fees for Seroquel:			Reconciliation of Adjusted Research and Development Expenses:	
		2018 20 1	2018	2017
Revenue – Seroquel		29.2 35	Segment operating loss – Innovation Platform (102.6)	(52.0)
Less: Cost of goods - Seroquel	((12.0) (24.	Less: Segment revenue from external customers (41.2)	(36.0)
			- Innovation Platform	(50.0)
Adjusted services fees for Seroquel		17.2 11	Add: Costs of goods - third parties 1.6	-
			Adjusted R&D expenses (142.2)	(88.0)

[1] For the purposes of this reconciliation, 2019 guidance for net cash used in or generated from financing activities for the year is not provided and as such, cash and cash equivalents and short-term investments at the end of year excludes the effect of any net cash used in or generated from financing activities for the year.

(US\$ millions unless otherwise stated) 104

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



Reconciliation of Non-GAAP Sales and Non-GAAP Net (loss)/income after tax^[1]

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).

		IFRS									US GAAP				17-18		
(US\$ millions)	03	04	05	06	07	08	09	10	11	12	13	14	15	16	17	18	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	664.4	-2%
Prescription Drugs	17.2	21.8	23.3	23.2	28.1	39.5	54.4	71.2	92.4	116.5	138.2	204.9	286.6	372.3	411.0	408.5	-1%
- Consolidated subsidiary	-	-	-	-	-	-	-	-	-	-	-	50.2	105.5	149.9	166.4	132.8	-20%
- 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	275.7	13%
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	255.9	-4%
- Consolidated s ubsidiaries	4.7	6.1	9.3	<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	40.1	3%
- Non-consolidated joint venture	-	-	32.5	69.3	87.2	110.8	135.6	151.1	171.3	228.7	247.6	243.7	211.6	224.1	227.4	215.8	-5%
Total Sales Growth	n/a	27%	133%	56%	17%	31%	26%	20%	18%	29%	n/a	16%	11%	21%	8%	-2%	
- GuanBao divested in Sept'2017	-	-	-	-	-	-	-	-	(11.4)	(50.5)	(51.6)	(49.7)	(40.7)	(45.0)	(38.6)	0.0	n/a
Adjusted Consumer Health	4.7	6.1	41.8	78.2	90.9	116.3	142.6	165.2	174.8	193.7	212.5	210.8	191.6	210.1	227.6	255.9	12%
- Adjusted Non-consolidated joint venture	0.0	-	32.5	69.3	87.2	110.8	135.6	151.1	1 <i>59.9</i>	178.2	196.0	194.0	1 <i>70.9</i>	179.1	188.8	215.8	14%
Adjusted Sales (Non-GAAP)	21.9	27.9	65.1	101.4	119.0	155.8	197.0	236.4	267.2	310.2	350.7	415.7	478.2	582.4	638.6	664.4	4%
Total Adjusted Sales Growth	n/a	27%	133%	56%	17%	31%	26%	20%	13%	16%	13%	19%	15%	22%	10%	4%	
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[3]	77.3[4]	83.6	8%
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	63.9	21%
- Consolidated subsidiary	-	-	-	-	-	-	-	-	-	-	-	0.1	0.6	1.6	2.4	4.1	74%
- 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	<i>59.8</i>	18%
Consumer Health	(10.3)	(4.9)	0.3	5.4	9.3	11.9	15.5	16.0	15.9	15.4	17.3	22.3	22.2	21.9	24.3	1 <i>9.</i> 7	-19%
- 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	2.8	-20%
- Non-consolidated joint venture	-	-	3.2	7.8	9.1	11.9	14.7	15.0	16.3	16.5	17.2	20.8	21.4	20.4	20.8	16.9	-19%
% 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%	12.6%	
Net (loss)/income attrib. to Chi-Med	(5.7)	(3.7)	(0.5)	1.2	4.5 ^[2]	5.9 ^[2]	9.3 ^[2]	12.6 ^[2]	13.6 ^[2]	14.6 ^[2]	18.2 ^[2]	22.8 ^[2]	25.2 ^[2]	29.9 ^[3]	37.5 ^[4]	41.4	10%
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	32.1	21%
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	9.3	-16%
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%	10%	

[1] 2003-2006 incl. disco. operation; [2] Continuing Operations; [3] Excludes the land compensation in SHPL of \$80.8 million from net income after tax and \$40.4 million from net income attributable to Chi-Med for 2016; [4] Excludes SHPL's R&D related subsidies of \$5.0 million from net income after tax and \$40.4 million from net income attributable to Chi-Med for 2016; [4] Excludes SHPL's R&D related subsidies of \$5.0 million from net income after tax and \$40.4 million from net income attributable to Chi-Med for 2016; [4] Excludes SHPL's R&D related subsidies of \$5.0 million from net income after tax and \$2.5 million from net income attributable to Chi-Med for 2017.

National Reimbursement Drug List Pricing ("NRDL") July'17 update – 15 new drugs in oncology^[1] added to NRDL



		U	nit Pricing (US\$)	[3]		Approximate Mor	nthly Pricing (L	JS\$) ^[3]	
Brand (generic)	Company	Dosage	Avg. Tender	Reimbursed	Δ %	Dosage	Avg. Tender	Reimbursed	Indication coverage
Herceptin® (trastuzumab)	Roche	440mg:20ml	\$3,298.81	\$1,125.93	-66%	Breast: 4mg/kg wk 1, 2mg/kg weekly. ^[2]	\$4,500	\$1,540	Breast: Her2+; Her2+ meta. Her2+ late-stage meta. gastric.
Avastin® (bevacizumab)	Roche	100mg:4ml	\$772.74	\$296.00	-62%	10mg/kg 0.2W.	\$11,590	\$4,440	Late-stage meta. CRC or advanced non-squamous NSCLC.
TheraCIM ^{®[4]} (nimotuzumab)	Biotech Pharma	50mg:10ml	\$435.26	\$251.85	-42%	100mg weekly.	\$3,730	\$2,160	Combo with radiotherapy for EGFR+ Stage III/IV nasopharyngeal carcinoma.
Rituxan® (rituximab)	Roche	500mg:50ml ^[2]	\$2,544.74	\$1,228.15	-52%	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 ^[2]	\$68.15	\$28.89	-58%	150mg QD.	\$2,040	\$870	Advanced NSCLC with limited EGFR gene mutation.
Nexavar® (sorafenib)	Bayer	0.2g	\$60.44	\$30.07	-50%	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	-41%	1,500mg QD.	\$3,170	\$1,870	Adv./meta. breast cancer with Her2 O/E, after anthracycline, paclitaxel, trastuzumab.
AiTan® (apatinib)	Hengrui	425mg ^[2]	\$47.85	\$30.22	-37%	850mg QD.	\$2,870	\$1,810	3L gastric adenocarcinoma or esophageal junction with adenocarcinoma.
Velcade® (bortezomib)	١%١	3.5mg ^[2]	\$1,873.78	\$906.07	-52%	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	-29%	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	-30%	30mg QD, 2x per wk.	\$4,190	\$2,930	2L+ Recurring or refractory peripheral T-cell lymphoma (PTCL).
Zytiga® (abiraterone)	١&١	250mg	\$45.63	\$21.48	-53%	1,000mg QD.	\$5,480	\$2,580	Metastatic or ovariectomized prostate cancer.
Faslodex® (fulvestrant)	AstraZeneca	250mg:5ml	\$806.81	\$355.56	-56%	500mg per month.	\$1,610	\$710	Advanced ER/PR+ breast can., failing aromatase inhibitor.
Afinitor® (everolimus)	Novartis	5mg ^[2]	\$36.44	\$21.93	-40%	10mg QD.	\$2,190	\$1,320	Adv. RCC after sunitinib or sorafenib. Adv./meta. pancreatic NETs. Tuberous sclerosis with renal angiomyolipoma.
Revlimid (lenalidomide)	Celgene	25mg ^[2]	\$413.93	\$163.26	-61%	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.

National Reimbursement Drug List Pricing ("NRDL") Oct'18 update – 17 new drugs in oncology added to NRDL



			Unit Pricing (US\$) ^[2]		Approximate Monthly F				
Brand (generic)	Company	Dosage	Avg. Tender	Reimbursed	Δ %	Dosage ^[1]	Avg. Tender	Reimbursed	Indication coverage	
Focus V [®] (anlotinib)	Sino Biopharn	n 12mg	\$127	\$70	-45%	12mg QD (2 wks-on/1-wk-off)	\$1,783	\$981	3L NSCLC	
Oncaspar [®] (pegaspargase)	Hengrui	5ml:3750 IU	\$560	\$429	-23%	\leq 2ml every 14 days	\$1,231	\$943	1L ALL	
Vidaza [®] (azacitidine)	Celgene	100mg	\$378	\$152	-60%	1 st cycle: 75mg QD for 7 days; 4wk cycle. After 2 cycles increase dose to 100mg, min of 4-6 cycles	\$14,022	\$5,636	Refractory anemia (RA) or RA with ringed sideroblasts (RARS), RA with excess blasts (RAEB / RAEB-T), and chronic myelomonocytic leukemia (CMMoL)	
Inlyta [®] (axitinib)	Pfizer	5mg	\$99	\$30	-70%	5mg BID	\$5,957	\$1,787	2L Advanced renal cell carcinoma	
Tagrisso [®] (osimertinib)	AstraZeneca	80mg	\$253	\$73	-71%	80mg QD	\$7,597	\$2,201	EGFR TKI refractory T790M+ NSCLC	
Ninlaro [®] (ixazomib)	Takeda	4mg	\$3,234	\$710	-78%	4mg on Days 1, 8, 15 (28 day cycle)	\$12,934	\$2,839	2L Multiple myeloma	
Xalkori [®] (crizotinib)	Pfizer	250mg	\$123	\$37	-70%	250mg BID	\$7,407	\$2,245	Locally adv. or meta. ALK+ or ROS1+ NSCLC	
Gilotrif [®] (afatinib)	Boehringer	40mg	\$116	\$29	-75%	40mg QD	\$3,483	\$863	NSCLC with EGFR	
Tasigna [®] (nilotinib)	Novartis	200mg	\$39	\$14	-65%	400mg BID	\$4,645	\$1,635	CML	
Votrient [®] (pazopanib)	Novartis	200mg	\$66	\$23	-65%	800mg QD	\$7,891	\$2,348	RCC	
Sutent [®] (sunitinib)	Pfizer	12.5mg	\$49	\$22	-55%	GIST & RCC: 50mg QD pNET: 37.5mg QD	\$5,544 \$4,455	\$2,498 \$2,007	RCC, GIST, pNET	
Stivarga [®] (regorafenib)	Bayer	40mg	\$52	\$28	-46%	160mg QD	\$6,216	\$3,384	Meta. CRC, GIST, HCC	
Zykadia [®] (certinib)	Novartis	150mg	\$108	\$28	-74%	450mg QD	\$9,699	\$2,564	NSCLC	
Zelboraf [®] (vemurafenib)	Roche	240mg	\$30	\$16	-47%	960mg BID	\$7,252	\$3,868	Melanoma	
Erbitux [®] (cetuximab)	Merck	100mg	\$571	\$186	-67%	400mg/m2 initial dose, 250mg weekly	\$10,446	\$3,074	Colorectal cancer, head and neck cancer	
Sandostatin LAR [®] (octreotide)	Novartis	20mg	\$1,169	\$835	-29%	20mg Q4W	\$1,169	\$835	GEP-NENS	
Imbruvica [®] (ibrutinib)	JNJ	140mg	\$78	\$27	-65%	MCL: 560mg QD CLL & WM: 420mg QD	\$9,324 \$6,993	\$3,263 \$2,447	MCL, CLL/SLL	

Source: Ministry of Human Resources and Social Security (MOHRSS); Yaozhi; China Merchants Securities Research; Citi Global Research. [1] Reference SKU or reference recommended dosage for monthly pricing calculation; [2] Calculation assumes an exchange rate of CN¥6.95 per US\$1.





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Thank you