

#### HUTCHISON CHINA MEDITECH

## **2019 Full Year Results**

March 3, 2020 Nasdaq/AIM: HCM





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*Use of Non-GAAP Financial Measures* - This presentation includes certain 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.

# Building a global science-focused biopharma company from an established base in China



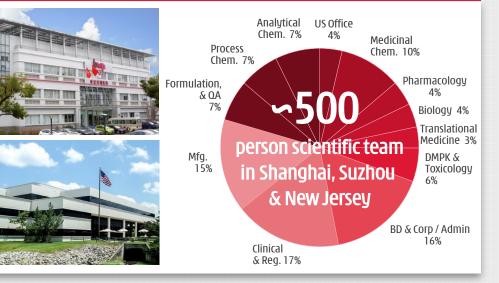




## Proven innovation & commercial operations



### Integrated Innovation Organization<sup>[1]</sup>



### Commercial Team & Joint Ventures [1]

Commercial Team (subsidiaries):

- ∽220 staff covering:
- Drug distribution & marketing operations; &
- New Oncology Business Dept.

50/50 Joint Ventures:

∽2,300 Rx medical sales reps.;

∽900 person OTC sales team; &

>1,500 staff in two major factories.



## Building blocks in place

Savolitinib for NSCLC & more	<ul> <li>Tagrisso<sup>®</sup> combo. Further global reg. studies.</li> <li>1<sup>st</sup> NDA submission (China)<sup>[1]</sup> &amp; data presentation.</li> </ul>	P7
Surufatinib for NET	<ul> <li>NET China NDA submissions.<sup>[2]</sup></li> <li>U.S./Europe/Japan global reg discussions.</li> </ul>	P19
Elunate <sup>®</sup> for CRC (fruquintinib)	<ul> <li>Global late-stage development (FRESCO2).<sup>[3]</sup></li> <li>Maximizing China access.</li> </ul>	P26
Organization & Other Candidates	<ul> <li>Oncology China sales team.</li> <li>Lymphoma China &amp; U.S. progress.</li> <li>PD-1 combos. HMPL-306 (IDH1/2).</li> </ul>	P33
Financials	<ul> <li> <ul> <li>                  2019 Rev.: \$205m. Net Loss: \$106m.             </li> <li>                  Available cash: &gt;\$300m (Dec-19)<sup>[4]</sup> + \$110m (Jan-20)<sup>[5]</sup>.                  </li> <li>                  2020 net cash flow guidance: \$(140)-(160)m.<sup>[6]</sup> </li> </ul> </li> </ul>	P39

[1] Planned for H1 2020; [2] Non-pancreatic NET NDA accepted and pancreatic NET NDA being prepared; [3] Study initiation planned for mid-2020; [4] Available cash resources = 12/31/2019 cash + cash equivalents + short-term investments + unutilised banking facilities; [5] Net proceeds from Jan 2020 Offering; [6] Adjusted non-GAAP Group net cash flows excluding financing activities.

## Portfolio summary Multiple waves of innovation – progressing rapidly

Indolent NHL







## AstraZeneca

## AstraZeneca and Chi-Med Harnessing the power of Chinese Innovation



# Savolitinib – global partnership with AstraZeneca MET-driven development strategies



A. INITIAL APPROVAL OF MONOTHERAPY - ESTABLISH SINGLE AGENT

**A2. PAPILLARY RCC** ~8% RCC. No biomarker therapies approved. A1. EXON14 MUTATION NSCLC 2~3% 1<sup>st</sup> line. First in China.

## **B. SOLIDIFY KEY COMBINATION OPPORTUNITIES WITH TKIS & IO**

**B2. PD-L1 COMBINATION** Preliminary signal with Imfinzi<sup>®</sup>. Exploring further. **B1. POST-EGFR TKI NSCLC** ~30% Tagrisso®-resistant pts. (Tag. 2019 \$3.2bn, #1 globally).

## **C. EXPLORATORY DEVELOPMENT**



## Highly selective MET inhibitor Current development status





Indication	Treatment	Target Patient	Study Name	Dose Finding / Safety Run-in	Proof-of-concept	Registration
	<b>Savolitinib</b> + Tagrisso	2L/3L EGFRm; Tagrisso ref.; MET+	SAVANNAH			
1 NSCLC	Savolitinib	MET Exon 14 deletion				
2	Savolitinib + Iressa	2L EGFRm; Iressa ref.; MET+				
	Savolitinib	MET+ Papillary RCC	SAVOIR			
2 Kidney	<b>Savolitinib</b> + Imfinzi (PD-L1)	Papillary RCC *	CALYPSO			
	<b>Savolitinib</b> + Imfinzi (PD-L1)	Clear cell RCC <sup>*</sup>	CALYPSO			
	Savolitinib	MET+ Gastric cancer *	VIKTORY			
3 Prostate, Colorectal	Savolitinib	MET+ Gastric cancer				
	Savolitinib	MET+ Prostate cancer *	CCTG I234B			
	Savolitinib	MET+ Colorectal cancer *				

## 1. Prime position in NSCLC – 2 ongoing registration-intent studies:

- MET Exon 14 del NSCLC completed enrollment. NDA submission in early 2020;
- Savo/Tagrisso® combo interim analysis mid-2020. Complete enrollment by end-2020.

## 2. Kidney cancer – Renewed global development strategy:

Savo monotherapy - ~60 pt. mature SAVOIR data. Actively evaluating restart in MET-driven PRCC;

Savo/Imfinzi<sup>®</sup> combo – Preliminary signal showing durable efficacy and tolerability.

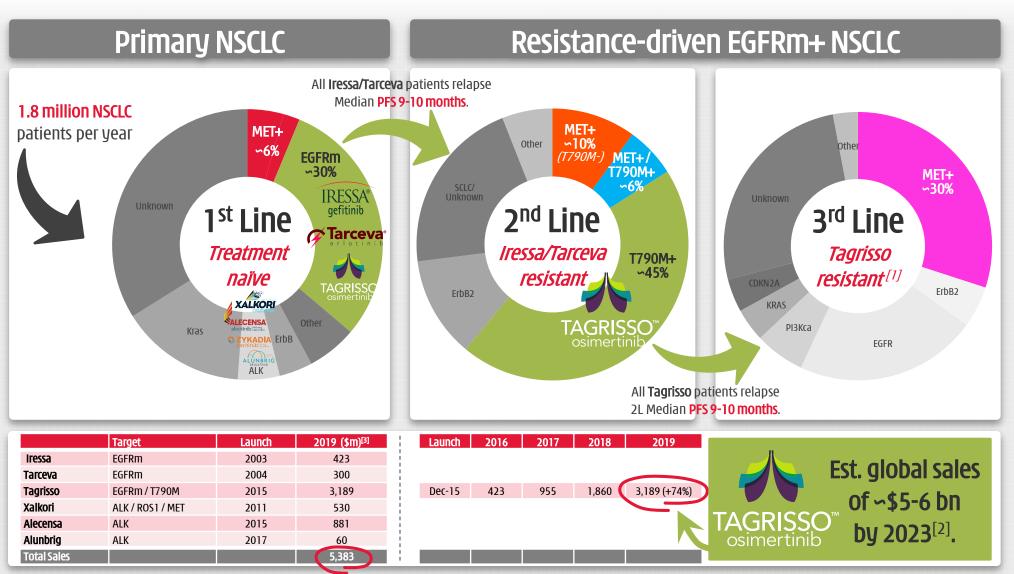
3. Gastric cancer & other exploratory studies:



Monotherapy in gastric - 50% ORR in VIKTORY;

## Savolitinib Biggest opportunity is MET+ NSCLC





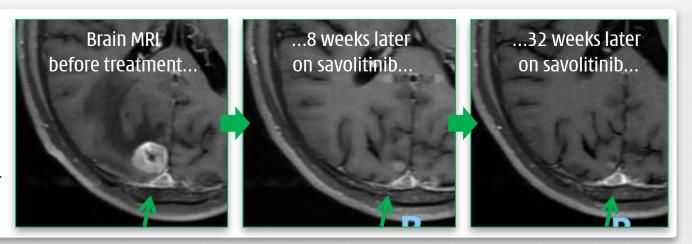
[1] Primary drivers, based on aggregate rocelitinib/Tagrisso data published at 2016/2017 ASCO; [2] Research estimates; [3] company annual reports and Frost & Sullivan.

## Savolitinib – MET Exon 14 deletion NSCLC<sup>[1]</sup> Potential China NDA submission in 2020<sup>[2]</sup>



#### 1. Encouraging MET Exon14d NSCLC study China data at CSCO 2019<sup>[3]</sup>

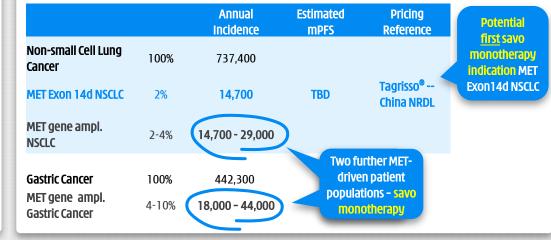
- SO pts; 36 pts efficacy evaluable.
- Promising antitumor activity.
- $\bigcirc$ Rapid, durable tumor response observed.
- Anti-tumor activity observed in brain mets.
- Savolitinib generally well tolerated; most related 1 TEAEs were grade 1 or 2.



#### 2. MET Exon14d NSCLC potential NDA filing 2020<sup>[2]</sup>

2019		2020
G-	<b>≻</b> 0	0-0→
Mar 31, '19 – Oral AACR Pres. <sup>[4]</sup> • 41 patient data	Topline results	<ul> <li>Potential NDA submission</li> <li>CDE<sup>[5]</sup> discussion</li> <li>Final results &amp; potential</li> </ul>
<b>Jun-Jul '19</b> registration s enrolled (n~6	study fully	NDA submission <ul> <li>Incl. global safety data</li> </ul> Mid- '20 - Presentation at
		a scientific conference

#### 3. Savolitinib monotherapy China market opportunity



[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] We expect that the Phase II study of savolitinib in MET Exon 14d NSCLC would, if successful, be sufficient to support NDA submission; [3] Data cut-off April 10, 2019, Lu S et al, Abstract #5707, presented at the 22<sup>nd</sup> Annual Meeting of the Chinese Society of Clinical Oncology, in Xiamen, China on Sept 20, 2019; [4] Data cut-off Feb. 26, 2019. Lu S et al, Abstract #CT031, presented at American Association of Cancer Research Annual Meeting 2019, Atlanta, GA, Mar. 31, 2019; [5] Center for Drug Evaluation of the National Medicinal Products Administration of China.

## TAGRISSO<sup>TH</sup> + Savo in EGFR TKI refractory NSCLC TATTON B & D data - efficacy



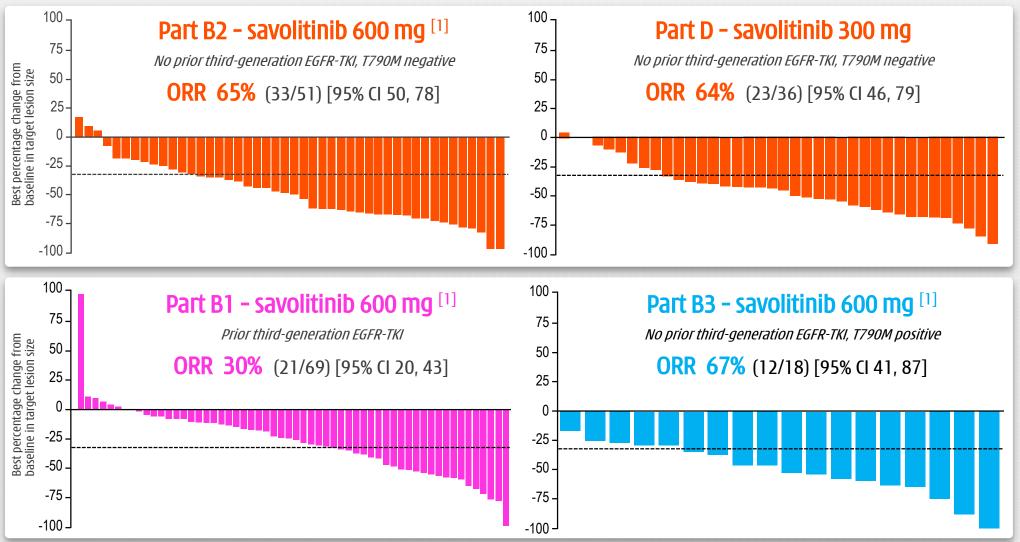
	(	<b>TATTON Part D</b> osimertinib 80 mg + savolitinib 300 mg		
	<b>Part B1 (n=69)</b> Prior third-generation EGFR-TKI	<b>Part B2 (n=51)</b> No prior third-generation EGFR-TKI (T790M negative)	Part B3 (n=18) No prior third-generation EGFR-TKI (T790M positive)	<b>Part D (n=36)</b> No prior third-generation EGFR-TKI (T790M negative)
<b>Objective response rate</b> ,* % [95% CI] Complete response, % Partial response, %	<b>30%</b> [20, 43] 0 30%	<b>65%</b> [5 <u>0,</u> 78] 0 65%	<b>67%</b> [41, 87] 0 67%	<b>64%</b> [46, 79] 0 64%
Non-response, % Stable disease (≥ 6 weeks) Progressive disease Not evaluable	45% 10% 14%	24% 6% 6%	33% 0 0	28% 3% 6%
Disease control rate, #% [95% CI]	<b>75%</b> [64, 85]	<b>88%</b> [76, 96]	<b>100%</b> [81, 100]	<b>92%</b> [78, 98]
Median DoR, months [95% CI]	<b>7.9</b> [4.0, 10.5]	<b>9.0</b> [6.1, 22.7]	12.4 [2.8, NR]	<b>8.0</b> [4.5, NR]
Median PFS, months [95% CI]	<b>5.4</b> [4.1, 8.0]	<b>9.0</b> [5. <u>5, 11.9]</u>	11.0 [4.0, NR]	<b>9.1</b> [5.4, 12.9]

## No reduction in efficacy with 300mg savo – SAVANNAH converted to 300mg dose

[1] Most patients were enrolled to Part B1, B2, B3 on 600 mg savolitinib, prior to weight-based dosing implementation, but following a protocol amendment in response to a safety signal of hypersensitivity, the final 21 patients enrolled in Part B were dosed with savolitinib by body weight as follows: patients who weighed  $\leq$ 55 kg (n=8) received 300 mg daily and those weighing >55 kg (n=13) received 600 mg daily; Best response data are for patients who had an opportunity to have two follow-up scans; \* Complete or partial response confirmed at  $\geq$ 4 weeks; # Disease control rate = confirmed complete response + stable disease at  $\geq$ 5 weeks; CI, confidence interval; NR, not reached.

Sequist LV, Han JY, Ahn MJ, et al. Osimertinib plus savolitinib in patients with EGFR mutation-positive, MET-amplified, non-small-cell lung cancer after progression on EGFR tyrosine kinase inhibitors: interim results from a multicentre, open-label, phase 1b study. Lancet Oncol. 2020; S1470-2045(19)30785-5. doi:10.1016/S1470-2045(19)30785-5.

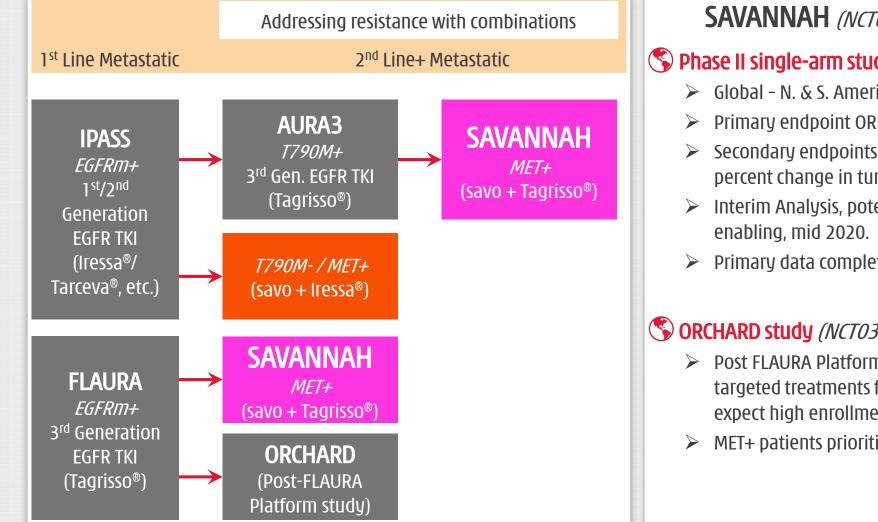




ORR = Objective Response Rate; EGFR = Epidermal Growth Factor Receptor; TKI = Tyrosine Kinase Inhibitor; [1] Most patients were enrolled to Part B1, B2, B3 on 600 mg savolitinib, prior to weight-based dosing implementation, but following a protocol amendment in response to a safety signal of hypersensitivity, the final 21 patients enrolled in Part B were dosed with savolitinib by body weight as follows: patients who weighed  $\leq$ 55 kg (n=8) received 300 mg daily and those weighing >55 kg (n=13) received 600 mg daily.

## TAGRISSO<sup>TH</sup> + Savo in EGFR TKI refractory NSCLC SAVANNAH - global registration intent study





#### **SAVANNAH** (*NCT03778229*)

#### S Phase II single-arm study:

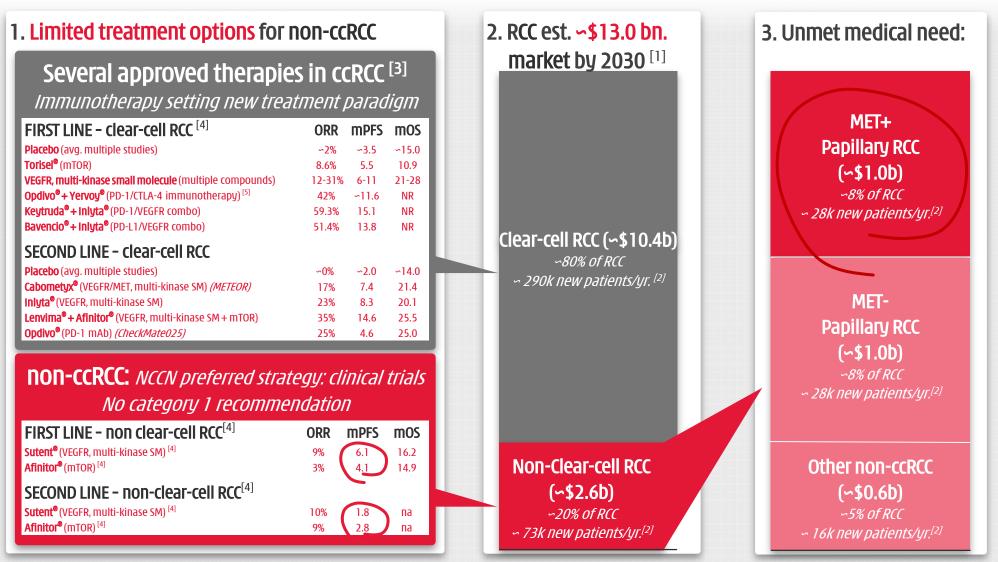
- Global N. & S. America, Eur., & Asia.
- Primary endpoint ORR.
- Secondary endpoints: PFS, OS, DoR & percent change in tumor size.
- Interim Analysis, potentially BTD
- Primary data completion est. 2021.

#### **S ORCHARD study** (*NCT03944772*):

- 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

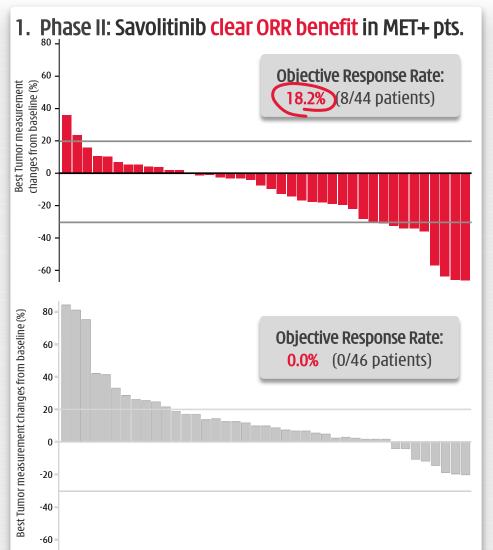




[1] Frost & Sullivan; [2] Frost & Sullivan, based on US incidence mix and global incidence rate in 2018; [3] NCCN Guideline for kidney cancer (Version 1.2020, June 7, 2019) preferred or category 1 options, RCC = renal cell carcinoma; [4] ORR = Objective Response Rate, mPFS = median Progression-Free Survival, mOS = median Overall Survival, NR = not reached; For approved subgroup of patients; [5] only approved for patients with intermediate or poor risk RCC.

## **Restart Savolitinib in PRCC** SAVOIR Phase III data planned mid-year<sup>[1]</sup>



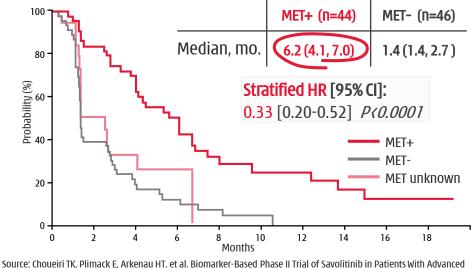


## 3. Phase II: Disease Control Rate ("DCR") – advantage in MET+ with OCR 73.2% vs. MET- 28.2%.^

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

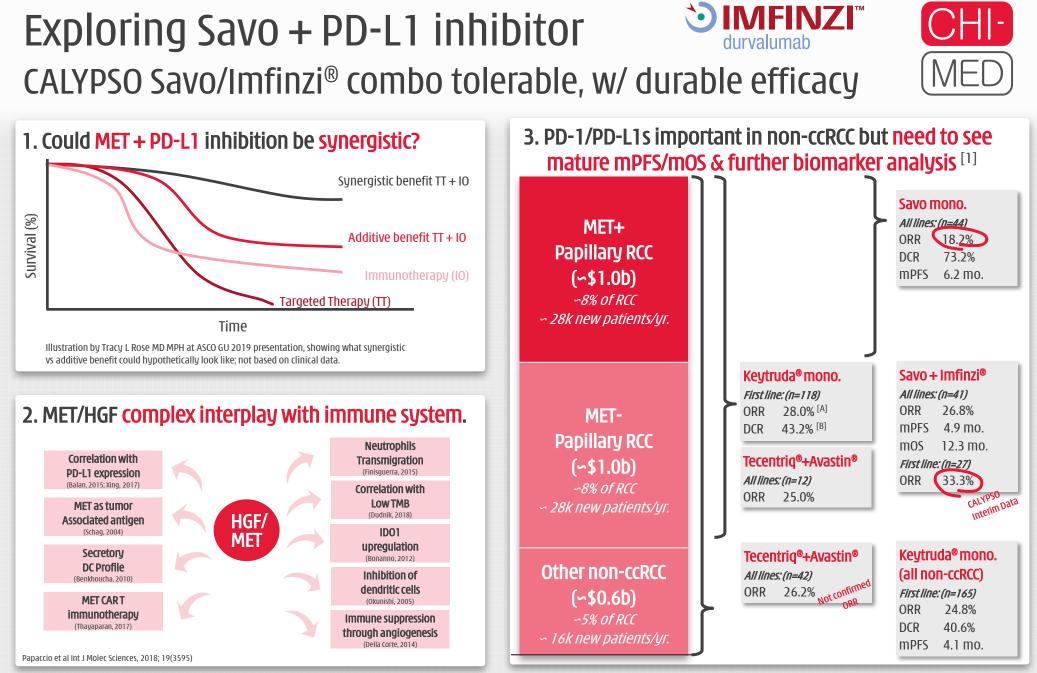
\* P=0.002 versus MET-independent subgroup (Fisher exact test). Responses assessed according to RECIST version 1.1.<sup>†</sup> Unconfirmed responses excluded. ^ Evaluable patients.

#### 4. Phase II: Median PFS – **big advantage in MET+ pts**.



Source: Choueiri TK, Plimack E, Arkenau HT, et al. Biomarker-Based Phase II Trial of Savolitinib in Patients With Advanced Papillary Renal Cell Cancer. J Clin Oncol. 2017;35(26):2993–3001. doi:10.1200/JC0.2017.72.2967

[1] In late 2018, enrollment was terminated in SAVOIR, a global Phase III registration study of savolitinib monotherapy compared with sunitinib monotherapy in MET-positive PRCC. Data from the approximately 60 patients randomized in SAVOIR prior to termination has matured during 2019 and will be presented at an upcoming scientific conference in mid-2020.

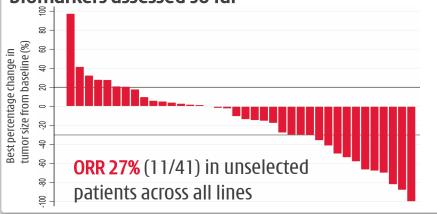


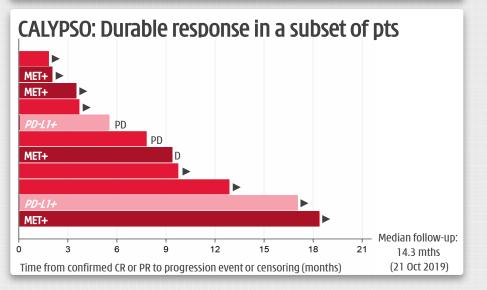
[1] CALYPSO: Suárez C et al. J Clin Oncol 38, 2020 (suppl 6; abstr 619); Keytruda mono - Keynote 427 cohort B [A] ESMO 2019, [B] ASCO GU 2019; Tencentriq; ORR = Objective Response Rate; DCR = Disease Control Rate; mPFS = median Progression-Free Survival; mOS = median Overall Survival.

# Savo + durva in PRCC (CALYPSO) Study expansion underway, alongside further MET



CALYPSO: Encouraging response independent of biomarkers assessed so far

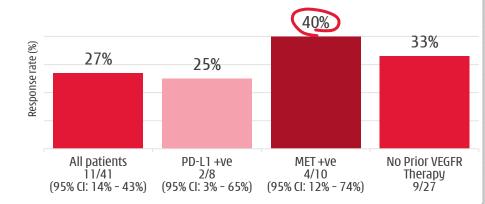




## CALYPSO: MET +ve results to be confirmed based on genetic alterations (40% ORR based on IHC $\geq$ 3)

durvalumat

MFINZI™



#### **CALYPSO: next steps**

- S Further assessment of biomarkers (6 not assessable)
  - Only MET+ overexpression assessed to date (10/41 positive, 25/41 negative);
  - MET+ gene amplification / other MET aberrations to evaluate.

## Section 2017 Secti

C. Suarez CALYPSO (PRCC cohort) ASCO GU 2020; Abstract 619. IIT = Investigator Initiated Study; ORR = Objective Response Rate; DCR = Disease Control Rate; mPFS = median Progression-Free Survival; mOS = median Overall Survival; PD = progressive disease; D = discontinued; PD-L1+ defined as: >25% immune component with SP263 Ab; MET +ve is defined as: >3+ in > 50% tumor cells with IHC.



#### Mechanism of Action

<u>Anti-angiogenesis</u>: cut off <u>blood flow to tumor</u> (VEGFR/FGFR).

Immunotherapy: inhibit expression of tumorassociated macrophages which cloak cancer cells from T-cell attack (CSF-1R).

2

**Surufatinib** 

Tumor-associated macrophages

T-cells

Angiogenesis

## **Surufatinib – wholly owned worldwide** Global development strategies



## A. BROAD APPROVAL FOR NET - ESTABLISH SINGLE AGENT

A3. BILIARY TRACT CANCER Poor prognosis pts. A2. GLOBAL NET REGISTRATION 1<sup>st</sup> targeted therapy with pan-NET success in Phase III. A1. 1<sup>ST</sup> CANCER PRODUCT LAUNCH NDA under review. Own oncology commercial org.

## **B. SOLIDIFY KEY COMBINATION OPPORTUNITIES WITH IO**

**B1/B2. PD-1 COMBINATIONS** Multiple PD-1s approach. Very broad potential. MOA synergy, especially CSF-1R and PD-1.

## **C. EXPLORATORY DEVELOPMENT**



## Unique VEGFR & CSF-1R inhibitor **Current development status**



Indication	Treatment	Treatment		Target Patient		Dose Finding / Safety Run-in	Proof-of-concept	Registration	
	Surufatinib		NET						
1 NET	Surufatinib		Pancreatic NET		SANET-p				
2 Surufatinib			Non-Pancreatic NET		SANET-ep				
3 Biliary TC	Surufatinib	fatinib 2L; chemo ref. biliary tract car		ncer					
	Surufatinib + Tuoyi (PD-1)		Solid tumors			*			
4 PD-1 Combo	<b>Surufatinib</b> + Tuoyi (PD-1)		Solid tumors						
Combo	Surufatinib + Tyvyt (PD-1)		Solid tumors			*			
		2. Pre			3. Initiated Phase II/III in biliary tract cancer (March		4. PD-1 combos progressing		
		consul					😋 Tuoyi® (Junshi PD-1)		
子 2 positive Phase IIIs –		and Ja	pan:	2019):		combo: Completed Pha			
torminat	ad aarly following	💽 11 C			Intorim -	nalucic in	in China: initiated Dhace II		

- terminated early following positive interim analyses (met mPFS primary endpoint):
- 😋 SANET-ep (June 2019) NDA accepted Nov 2019;
- 😋 SANET-p (January 2020) NDA preparations underway.
- (S) U.S. Phase Ib/II monotherapy in solid tumors initiated;
- **NET** enrollment complete.

- 😋 Interim analysis in 2020.
- in China; initiated Phase II in multiple solid tumors;
- **Tyvyt<sup>®</sup>** (Innovent PD-1) combo: in planning;
- **(S)** U.S. Phase Ib/II PD-1 combo study to start by mid-2020.

## **G1/2 Advanced NET**<sup>[1]</sup> *(Ki-67 Index 0-20)* Global opportunity in lung/other NETs & China wide-open

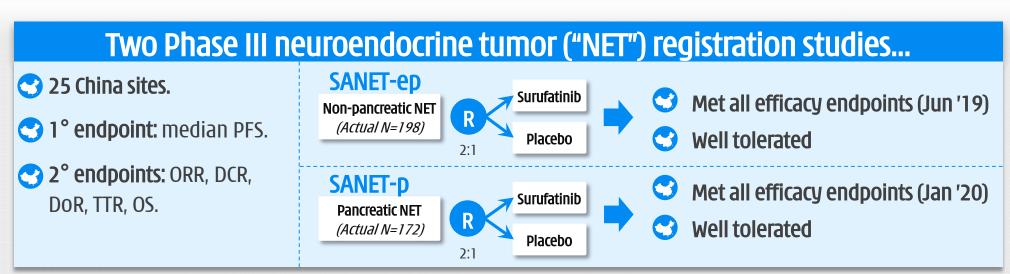


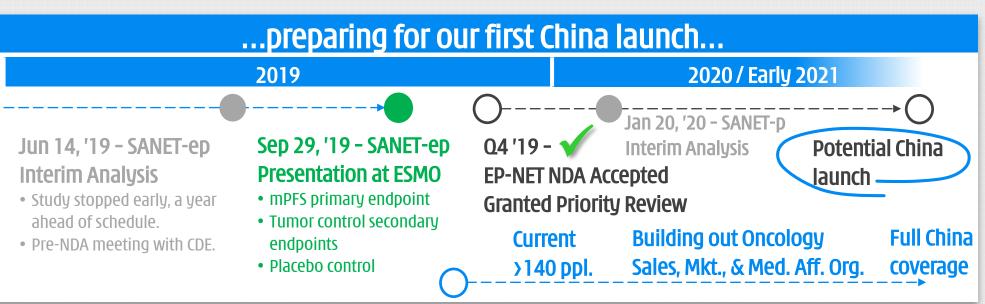
Site		est. %	Octreotide	Lanreotide	<sup>177</sup> Lu-Dotatate	Streptozocin	Sunitinib	Everolimus	Surufatinib
Disease status			Treatment naïve	Stable disease	Progressed in past 3 yrs.	Historical	Progressed in past 12 mo.	Progressed in past 6 mo.	Progressed in past 12 mo.
	Stomach	7%		CLARINET <sup>[2]</sup>	Historical Ph. II SSR over expression			RADIANT-4 <sup>[3]</sup>	SANET-ep
	Small bowel / appendix	9%	PROMID	CLARINET <sup>[2]</sup>	NETTER-1			RADIANT-4 <sup>[3]</sup>	SANET-ep
GI Tract	Colon & Rectum	31%		CLARINET <sup>[2]</sup>	Historical Ph. II SSR over expression			RADIANT-4 <sup>[3]</sup>	SANET-ep
Pancreas		6%		CLARINET <sup>[2]</sup>	Historical Ph. II SSR over expression	Historical	PHASE III	RADIANT-3 <sup>[3]</sup>	SANET-p Met primary endpt. (PFS)
Lung		20%						RADIANT-4 <sup>[3]</sup>	SANET-ep
Other	Other	∽17%							SANET-ep
	Unknown Primary	∽10%						RADIANT-4 <sup>[3]</sup>	SANET-ep

[1] Yao ESMO 2019; [2] CLARINET approved only for Ki-67 Index <10 (i.e. est. ~50% of G1/G2); [3] Everolimus approved in non-Functional NET (~60% pNET; 90% Lung NET; majority mid-gut/small bowel NET); [4] RADIANT-3 – Progressed in past 12 months.

China

Global (ex-China)



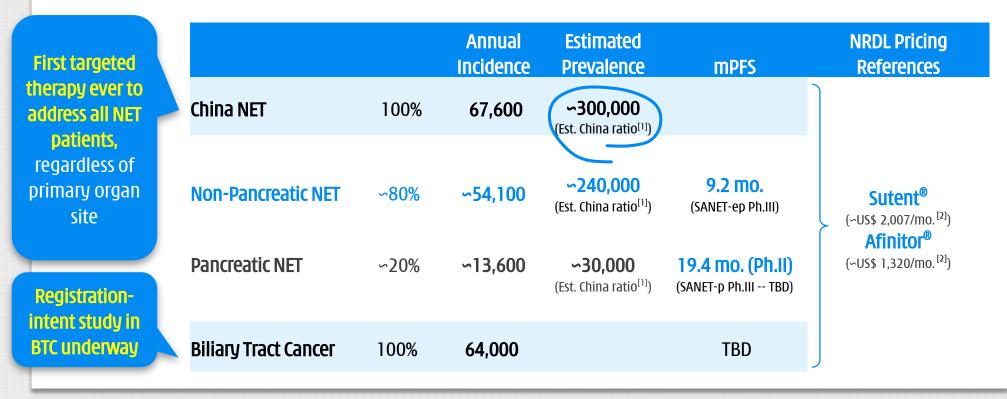


## Surufatinib in NETs Potentially our first un-partnered oncology drug launch

## Surufatinib – China NET Two positive Ph. III studies unblinded a year ahead of schedule

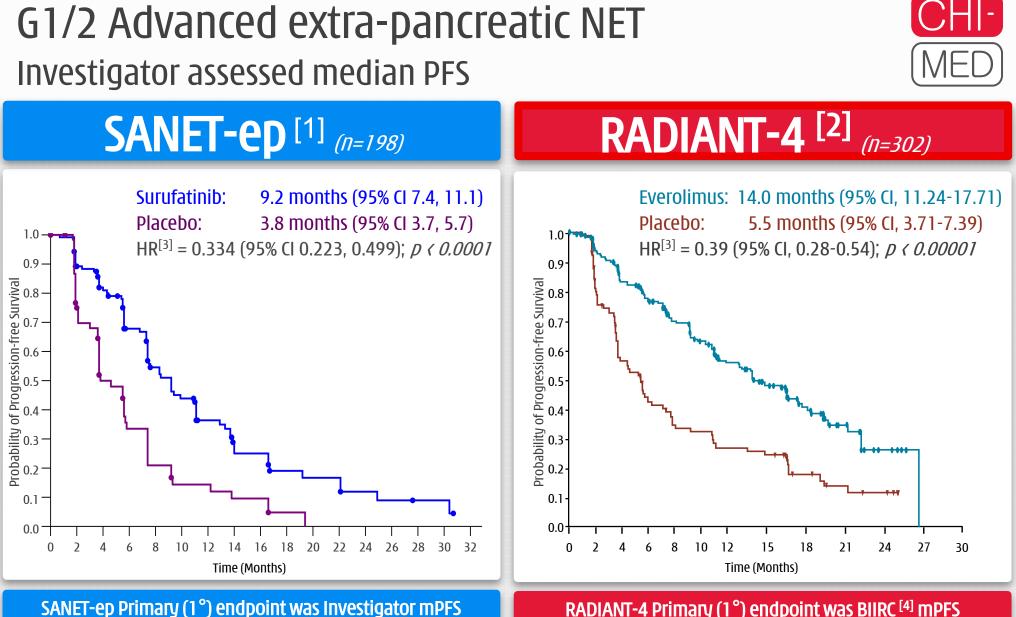


## **Epidemiology –** *China NET & BTC patient populations*



## NET is major unmet medical need in China – with long treatment duration

[1] Source: Frost & Sullivan. Current estimated Prevalence to Incidence ratio in China at 4.4, lower than U.S. 7.4 ratio due to lower access to treatment options; [2] NRDL pricing references calculations assume exchange rate of RMB6.74 per US\$1.



BIIRC<sup>[4]</sup> mPFS for supportive analysis not 1° or 2° endpoint

RADIANT-4 Primary (1°) endpoint was BIIRC <sup>[4]</sup> mPFS Investigator mPFS not 1° or 2° endpoint

[1] ESMO 2019 LBA#76; [2] Yao et al. "Everolimus for the treatment of advanced, non-functional neuroendocrine tumours of the lung or gastrointestinal tract (RADIANT-4)" Lancet. 2016 Mar 5;387(10022):968-977. doi: 10.1016/S0140-6736(15)00817-X. Epub 2015 Dec 17; [3] P-value is obtained from the stratified one-sided log-rank test; Hazard ratio is obtained from stratified Cox model; CI, confidence interval; HR, hazard ratio; [4] BIIRC = Blinded Independent Image Review Committee (Central).



# 3 Elunate<sup>®</sup> (fruquintinib capsules)

## ELUNATE highly selective VEGFR1/2/3 inhibitor Development strategies

## A. ESTABLISH BEST-IN-CLASS VEGFR INHIBITOR - SAFETY, TOLERABILITY, EFFICACY

#### A2. GLOBAL CRC REGISTRATION Maximize value through Phase III in the US, Europe & Japan. A1. BROAD ACCESS IN CRC Bolster excellent product profile with better accessibility for pts.

## **B. SOLIDIFY KEY COMBO OPPORTUNITIES WITH IO, TKIS & chemo**

**B2/B3. PD-1 COMBINATIONS** "Clean" profile enhances tolerability. Multiple PD-1s approach. **B1. "UPSTREAM" POTENTIAL** 2x~5x more pts in earlier lines. e.g. 2L gastric (+chemo).

## **C. EXPLORATORY DEVELOPMENT**



## **ELUNATE** highly selective VEGFR1/2/3 inhibitor **Current development status**



	Indication	Treatment	Target Patient	Study Name	Dose Finding / Safety Run-in	Proof-of-concept	Registration
1	Colorectal	Fruquintinib	Colorectal cancer ("CRC")	FRESCO2			
2	CUIVIECIAI	Fruquintinib	≥3L; chemotherapy ref. CRC	FRESCO			
3	Gastric	Fruquintinib + Taxol	2L gastric cancer	FRUTIGA			
	NSCLC	F <b>ruquintinib</b> + Iressa	1L EGFRm NSCLC				
	Collid	<b>Fruquintinib</b> + Tyvyt (PD-1)	Solid tumors		*		
4	Solid Tumors	<b>Fruquintinib</b> + Tyvyt (PD-1)	Solid tumors				
		Fruquintinib + genolimzumab (PD-1)	Solid tumors				

- 1. Elunate<sup>®</sup> China commercialization:
- China 1<sup>st</sup> product sales by Lilly: \$18m in 2019;
- NRDL inclusion from Jan 1,
  - 2020: Elunate<sup>®</sup> now the most attractive approved therapy in 3L CRC in China in terms of price, efficacy and safety;

Jan-Feb 2020 sales: \$6.6m<sup>[1]</sup>.

- 2. Prep. for global CRC registration trial (unpartnered):
- Section 22 [2] completed with U.S. FDA in Feb 2020;
- Seurope & Japan EOP2 mtgs. planned shortly;
- Study to start shortly after regulatory interactions;

Phase Ib/II in CRC ex-China enrollment completed.

- 3. FRUTIGA Phase III in 2L gastric cancer:
- 1<sup>st</sup> interim analysis in April 2019 - study continued without changes;
- ✓ 2<sup>nd</sup> interim analysis in 2020;
- On track to complete enrollment in H2 2020.

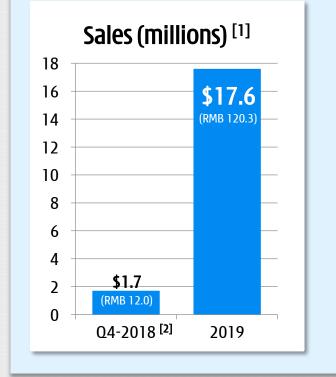
#### 4. PD-1 combos:

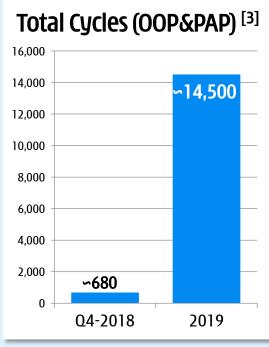
- Tyvyt<sup>®</sup> (Innovent PD-1): approaching completion of Phase I in China;
- 😋 Genolimzumab (Genor PD-1): Phase I underway.

## **ELUNATE** Fruquintinib Capsules **FY2019 performance**



#### Elunate<sup>®</sup> Performance





#### Chi-Med Revenue (US\$ million)

Total HCM Revenue	\$3.6M	\$10.8m
Royalty	\$0.3m	\$2.7m
Manufacturing <sup>[4]</sup>	\$3.3m	\$8.1m
	Q4-2018	2019



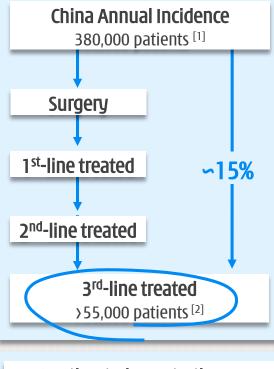
### Elunate<sup>®</sup> early progress – PAP worked but NRDL will provide greater access

[1] Royalties to Chi-Med in Q4 2018 and FY 2019 of \$0.261m and \$2.653m, respectively; at the lowest tier royalty rate of 15%, this implies net sales from Eli Lilly to third parties of \$1.7m and \$17.6m, respectively; at RMB:US\$ exchange rate of 6.87:1 and 6.83:1, respectively, this implies RMB sales of 12.0m and 120.3m, respectively; [2] Elunate® launched in Q4 2018; [3] Treatment cycle = 28 day, i.e. assume three x 7 capsule 5mg packs per cycle or five x 21 capsule 1mg packs per cycle; OOP = Out of pocket payment; PAP = Patient access program; [4] Sales of Elunate® manufactured by Chi-Med to Eli Lilly.

# ELUNATE<sup>®</sup> NRDL - 2020 accessible pricing



#### Epidemiology



#### 2019 estimated penetration:

- ~14,500 cycles used (OOP & PAP);
- Average 5 months per patient;
- ~3,000 patients paid for Elunate;
- Representing ~5% penetration.

## National Reimbursed Drug List (NRDL)

#### Effective Jan 1, 2020:

- 8 newly listed oncology drugs, including Elunate®
- NRDL reimburses 50-70% of patient costs under urban scheme

Out-of-pocket	costs for 3L CRC	Urban Med. Insur.	Non-UMI
Patients per cy	Icle (all US\$) <sup>[3]</sup>	Scheme (UMI)	
Population		317m	1,053m
<i>% China</i>		<i>23%</i>	<i>77%</i>
<b>Elunate®</b>	Pre-NRDL	3,260	3,260
(fruquintinib)	Post-NRDL	1,180	1,180
<b>Stivarga®</b> (regorafenib)	<b>3L CRC Pts OOP</b> Pre-NRDL Post-NRDL	<b>350 - 600</b> 4,490 2,450	1,180 4,490 2,450
	<b>3L CRC Pts OOP</b>	730 - <u>1,220</u>	2,450

## **2020 post NRDL:** Jan-Feb Sales – \$6.6 million <sup>[4]</sup>

[1] W. Chen, R. Zheng et al, CA Cancer J Clin. 2016 Mar-Apr;66(2):115-32. Cancer Statistics in China, 2015. doi:10.3322/caac.21338. Epub 2016 Jan 25; [2] Frost & Sullivan; [3] RMB:USD exchange rate 6.73:1.00.; OOP = Out of pocket payment; PAP = Patient access program; [4] January-February 2020 In-market sales of Elunate® to third-parties, as provided by Lilly and unaudited.



## Efficacy advantage



	FRESCO <sup>[1]</sup> Mainland China		CONCUR Chinese Patients (Mainland China, Hong Kong, Taiwan) <sup>[2]</sup>		CONC	UR	CORRECT	
Third-Line Metastatic Colorectal cancer					Mainland China, Hong Kong, Taiwan, Vietnam, South Korea		Global	
Treatment arms	Elunate®	Placebo	<b>Stivarg</b> a <sup>®</sup>	Placebo	<b>Stivarga<sup>®</sup></b>	Placebo	<b>Stivarga</b> <sup>®</sup>	Placebo
Patients (n)	278	138	112	60	136	68	505	255
Objective Response Rate, n (%)	4.7%	0.0%	3.6%	0.0%	4.4%	0.0%	1.0%	0.4%
Disease Control Rate, n (%)	62.2% +4	9.9 12.3%	45.5% +38	8 6.7%	51.5% +44.	1 7.4%	41.0% +26	.1 14.9%
Median Progression-Free Survival (mPFS) (mo.)	3.7 +1	. <mark>9</mark> 1.8	2.0 +0.	<b>3</b> 1.7	3.2 +1.5	1.7	1.9 +0.	2 1.7
Median Overall Survival (mOS) (mo.)	9.3 +2	2.7 6.6	8.4 +2.	2 6.2	8.8 +2.5	6.3	6.4 +1	4 5.0

Advantage for Elunate<sup>®</sup> efficacy vs. Stivarga<sup>®</sup> in Chinese metastatic **CRC** patients;

#### C Advantage for Elunate<sup>®</sup> post **VEGF/EGFR** targeted therapy

- mOS: 7.69 mo. vs. 5.98 mo. placebo (HR 0.63 & p-value 0.012)
- mPFS: 3.65 mo. vs. 1.84 mo. placebo (HR 0.24 & p-value < 0.001)

Overall Survival subgroup analysis by Prior Treatment [1]	
	Jazard Datio

100% Avastin prior use

		Hazard Ratio (95% Cl)	p-value
Overall		0.65 (0.51, 0.83)	<0.001
with prior anti-VEGF therapy		0.68 (0.45, 1.03)	0.066
without prior anti-VEGF therapy		0.60 (0.45, 0.80)	<0.001
with prior anti-VEGF or anti-EGFR therapy		0.63 (0.46, 0.90)	0.012
without prior anti-VEGF or anti-EGFR therapy		0.63 (0.43, 0.86)	0.003
	0 0.5 1.0 1.5	2.0	
	Favors Fruquintinib Favors Place	bo	

[1] Effect of Fruquintinib vs Placebo on Overall Survival in Patients with Previously Treated Metastatic Colorectal Cancer: The FRESCO Randomized Clinical Trial; [2] Efficacy & safety of regorafenib monotherapy in Chinese patients with previously treated metastatic colorectal cancer: subgroup analysis of the CONCUR trial; R Xu.

## ELUNATE<sup>®</sup> Fruquintinib Capsules



	ELUNATE <sup>®</sup>	Stivarga <sup>®</sup> (regorafenib) tablets
BIOCHEMICAL ACTIVITY	IC <sub>50</sub> (nmol/L)	IC <sub>50</sub> (nmol/L)
On-Target Kinases:		
VEGFR1	33	13
VEGFR2	35	4.2
VEGFR3	0.5	46
Off-Target Kinases:		
Ret	128	1.5
FGFR1	181	202
c-kit	458	7
PDGFRβ	>10,000	22
RAF-1	>10,000	2.5
B-RAF	>10,000	28
B-RAF <sup>V600E</sup>	>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 Sivarga for hepatotoxicity as manifested

by elevated liver function tests or hepatocellular necrosis, depending upon severity and persistence. (2.2)

	ELUNATE <sup>®</sup>			
3 <sup>rd</sup> -Line Metastatic Colorectal cancer	FRESCO Study Mainland China <sup>[1]</sup>		CONCUR Study (Mainland China, HK, Taiwan) <sup>[2]</sup>	
Treatment arms	Elunate®	Placebo	<b>Stivarga</b> <sup>®</sup>	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 $\geq$ 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:			$\frown$	
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%

Toxicity limitations of Stivarga<sup>®</sup>

## Elunate<sup>®</sup> superior safety - advantage especially for liver mets patients







## **350 person dedicated oncology commercial team** Building on >15 yrs Rx commercial knowhow in mainland China



#### To cover ~1,300 hospitals across China

- Establishing dedicated oncology commercial team to cover ~95% of initial market opportunity;
- Fully in-place & in-training by Q3 2020;
- Includes sales reps, sales mgrs., product mktg., medical mktg., distribution,

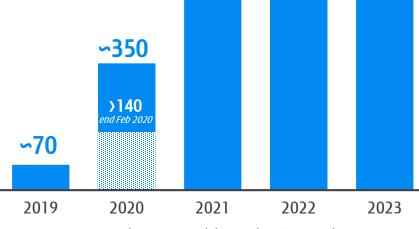
training, mgmt.;

30 provinces / municipalities to be covered at launch (>90% already covered).

#### Full suru launch team in place by mid-2020

- All key senior roles are already in-place;
- Vast majority of new staff from successful China oncology companies;
- >140 staff already on board;
- Plan to expand oncology team to 900+ by end-2023 to support future product launches.





## Next wave of innovation Development strategies and current status



Program	Treatment	Target Patient	Sites	Dose Finding / Safety Run-in	Proof-of-concept	Registration
HMPL-523 Syk	HMPL-523	Indolent NHL	Australia			
	HMPL-523	Indolent NHL	US			
	HMPL-523	B-cell malignancies	China			
	HMPL-523	ITP	China			
Η <b>ΜΡΙ-689</b> ΡΙ3Κδ	HMPL-689	Healthy volunteers	Australia			
	HMPL-689	Indolent NHL	US			
	HMPL-689	Indolent NHL	China			
	HMPL-453	Solid tumors	China			
FGFR 1/2/3						

1. Non-Hodgkin's lymphoma (China):

China Phase Ib dose expansions of both HMPL-523 and HMPL-689;

China registration study decisions in 2020.

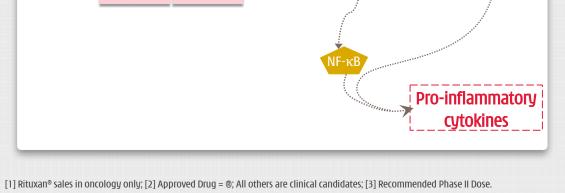


- S 20 Phase I sites in U.S. and Europe now enrolling;
- Multiple dose cohorts completed for both HMPL-523 and HMPL-689.

- 3. HMPL-453:
- Phase II study in advanced malignant mesothelioma in China set to initiate.

#### 4. HMPL-306 and others:

Phase I for our 9<sup>th</sup> inhouse discovered asset (IDH 1/2 dual inhibitor) set to initiate.



## HMPL-523 (Syk) & HMPL-689 (PI3K $\delta$ ) Exciting targets emerging – our next wave of innovation

TNF receptor associated

factors

(TRAFS)

Jakafi®

IL-6 Receptor

# The B-cell signaling is critical in hematological cancer with three breakthrough therapies recently approved.

 2019 sales: Imbruvica<sup>®</sup> \$5.7bn; Zydelig<sup>®</sup> \$0.1bn; Jakafi<sup>®</sup> \$2.8bn; & Rituxan<sup>®</sup> \$4.8bn <sup>[1][2]</sup>.

TNFα Recept

**Cell Membrane** 

AKT

PLC<sub>V</sub>2

РКСВ

**Rituxan**®

Imbruvica®

Brukinsa®

PIP2

HMPL-689 Calquence®

Zydelig®

umbralisib

B-Cell Receptor

mivavotinib

HMPL-523

**CD79** 

#### HMPL-523 (Syk inhibitor)

#### Large Phase Ib expansion in Australia & China

- Ph.I dose escalation complete in Australia & China (N>60) -RP2D<sup>[3]</sup> determined;
- Large Ph. Ib dose expansion study (N>200), underway in ~30 active sites in Australia & China;
- US/EU Phase I/Ib enrolling, with 12 sites.

#### HMPL-689 (PI3Kδ inhibitor)

#### Phase I/Ibs in China, US & EU ongoing

Designed to be a best-in-class inhibitor of  $\mbox{PI3K}\delta$ 

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

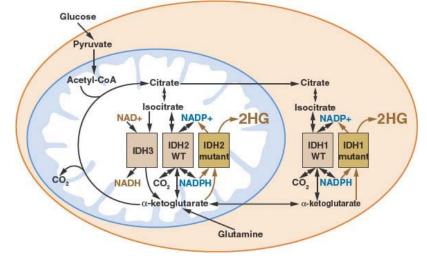
Phase I/Ib data will inform China registration study decisions on HMPL-523 & -689 in 2020.



## HMPL-306 – Phase I in China set to initiate Designed as potential best-in-class IDH 1/2 inhibitor



- The IDH family converts isocitrate to α-KG via oxidative decarboxylation, an important process for normal cellular metabolism.
- Mutant IDH1/2 catalyze the reaction of α-KG to 2-HG, leading to accumulation of 2-HG in tumor cells;
- IDH inhibitors could restore 2-HG levels to normal physiological levels, induce tumor cell differentiation and ultimately stop tumor cell progression;
- Mutant IDH isoform switching, either from cytoplasmic mutant IDH1 to mitochondrial mutant IDH2, or vice versa, as a mechanism of acquired resistance to IDH inhibition.



[1] Amary et al., 2011; Paschka et al., 2010; Yan et al., 2009; Fujii T et al., 2016, Abstract 3101, AACR 2016.

2. Unmet medical need and potential indications – IDH1/2 mutations are frequent genetic alterations in AML, glioma and various solid tumors.

Tumor	% IDH Mutation <sup>[1]</sup>					
	Total	IDH1-R132	IDH2-R140	IDH2-R172		
Brain tumor						
Grade 2 and 3 glioma	60-80%	60-80%	0%	1%		
Secondary gliomablastoma	70%	70%	0%	1%		
Hematopoietic tumor						
Acute myelocytic Leukemia (AML)	15-25%	5-10%	5-15%	0-5%		
Myelodysplastic syndrome (MDS)	10%	5%	5%	0%		
Angioimmunoblastic T-cell lymphoma	26%	0%	1%	25%		
Solid tumor						
Chondrosarcoma	55%	40%	0%	15%		
Osteosarcoma	25%	0%	0%	25%		
Cholangiocarcinoma	22%	20%	0%	2%		
Giant cell tumors of bone	80%	0%	0%	80%		

3. HMPL-306 is a potent IDH1/2 dual inhibitor.

- IDH1 & 2 mutations are validated targets with approval of ivosidenib (IDH1) and enasidenib (IDH2) in R&R AML;
- HMPL-306 provides comparable efficacy in preclinical model while wider safety window;
- The higher penetration of blood-brain barrier with HMPL-306 makes exploring IDHm glioma attractive.

## What is next from discovery? Differentiated assets against multiple targets

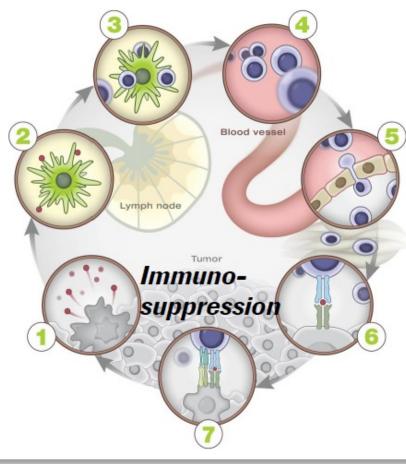


# Priming & activations aOX40 4-1BB

### Antigen release

- MET (savolitinib)
- EGFR (epitinib/theliatinib)
- Syk (HMPL-523)
- PI3Kδ (HMPL-689)
- FGFR (HMPL-453)
- IDH 1/2 (HMPL-306)
- ERK

## • RIP1K



### <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





## **5 2019 Financial Results, Cash Position & Guidance**

## 2019 Financial results

Global Innovation

China Oncology

China Commercial



	2018	2019	Growth	at CER <sup>[2]</sup> (Non-GAAP)
GROUP REVENUES Unconsolidated JV Revenues	<b>214.1</b> 491.5	<b>204.9</b> 487.5	<b>-4%</b> -1%	<b>-1%</b> +3%
SEGMENT NET INCOME/(LOSS) [1]				
INNOVATION PLATFORM <sup>[3]</sup>	(104.4)	(133.2)	-28%	-33%
COMMERCIAL PLATFORM Prescription Drugs Business <sup>[3]</sup> Consumer Health Business	<b>43.4</b> 34.1 9.3	<b>47.4</b> 37.5 9.9	<b>+9%</b> +10% +7%	<b>+13%</b> +14% +12%
Chi-Med Group Costs	(13.8)	(20.2)	-46%	-46%
<b>GROUP NET LOSS</b> <sup>[1]</sup> EPS Attrib. to Ord. S-H (Basic) (US\$) <sup>[4]</sup>	<b>(74.8)</b> (0.11)	(106.0)	-42%	-46%

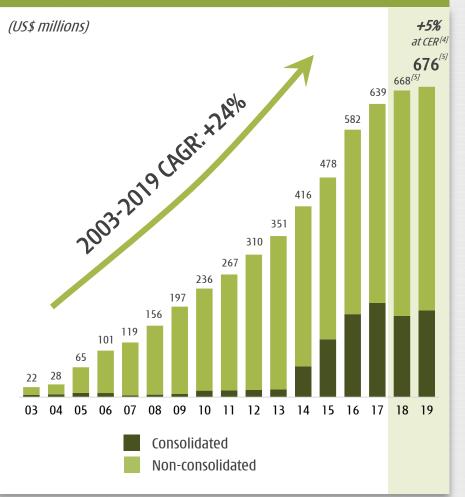
(US\$ millions, except per share data)

[1] Net Income / (Loss) attributable to Chi-Med; [2] at CER = at Constant Exchange Rate, which is a non-GAAP financial measure used to present period-to-period comparisons without the effects of currency movements by retranslating the current period's performance at the previous period's foreign currency exchange rates. Please refer to the slides titled "Non-GAAP Financial Measures and Reconciliation" for more information and a reconciliation of these measures to the most comparable GAAP measure; [3] In 2019, the results of innovative medicines developed by the Innovation Platform have been reallocated from Innovation Platform to Commercial Platform- Prescription Drugs business. 2018 information has been revised for comparison purpose; [4] EPS was adjusted retroactively to take into account the share split which each ordinary share has subdivided into 10 ordinary shares effective from May 30, 2019.

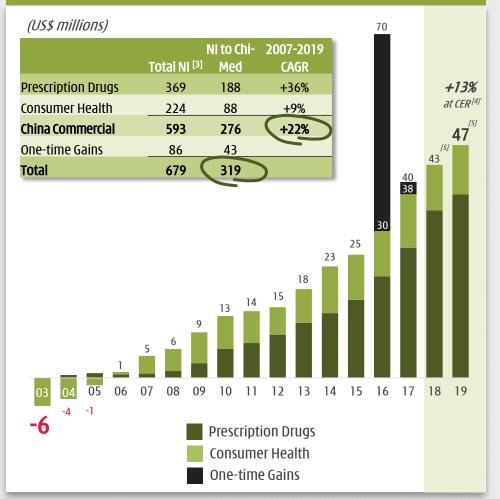
## Chi-Med's Commercial Platform in China Proven track record, \$319 million in net income since inception



### Revenues (Non-GAAP) [1][2]



### 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"; [4] at Constant Exchange Rate (at CER), which is a non-GAAP financial measure used to present period-to-period comparisons without the effects of currency movements by retranslating the current period's performance at the previous period's foreign currency exchange rates. Please refer to the slide titled "Non-GAAP Financial Measures and Reconciliation" for more information and a reconciliation of these measures to the most comparable GAAP measure; [5] In 2019, the results of innovative medicines developed by the Innovation Platform have been reallocated from Innovation Platform to Commercial Platform - Prescription Drugs business. 2018 information has been revised for comparison purpose. 4

## Cash position & guidance >\$300m available cash (Dec-19)<sup>[1]</sup> + \$110m raised (Jan-20)<sup>[3]</sup>



Cash Position (est. at end Dec 2019)

- \$217 million cash / cash equiv. / ST inv. <sup>[2]</sup>
- \$110m net raised on Nasdaq (Jan 2020)<sup>[3]</sup>

### **\$120m**

additional unutilized banking facilities <sup>[4]</sup>

## **\$63m**

additional cash in JVs

### \$27m in bank borrowings

	Ļ
Global Innovation	
China	

Oncology

(US\$ millions)	<b>2019</b> Guidance <sup>[5]</sup>	2019 Actual	2020 Guidance
Adj. (non-GAAP) Innovation Platform Segment Operating Loss	(130) - (170)	(149.3)	(180) - (210)
Adj. (non-GAAP) Group Net Cash Flows excluding financing activities	(90) - (120)	(82.3)	(140) - (160)

### Performance in-line with 2019 guidance:

> Better cash flow from one-time investing activity<sup>[6]</sup>;

### Increased cash use in 2020:

- Global registration studies start on suru & fruq;
- > Capital investment in small molecule facility;
- > Commercial Platform continued cash flow growth;
- > No material impact from COVID-19 outbreak.

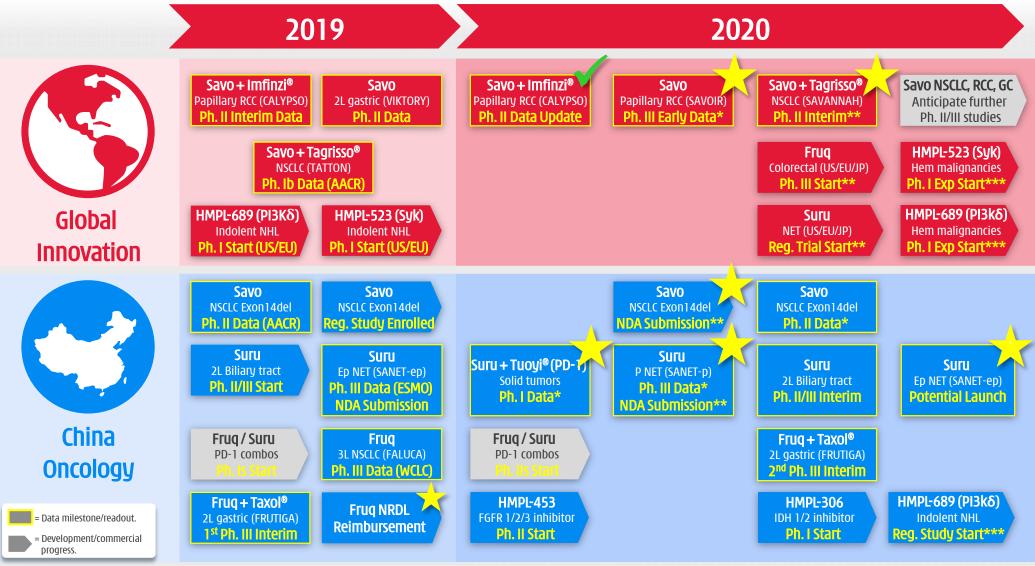
[1] Including cash, cash equivalents, short-term investments & unutilized banking facilities; [2] Short-term investments: deposits over 3 months; [3] Net proceeds of \$110.1m from NASDAQ follow-on offering: Total gross proceeds of \$118.3m netting off with underwriters' commission, legal and professional fees of \$8.2m; [4] From Bank of America Merrill Lynch, Deutsche Bank, HSBC; [5] 2019 Financial Guidance update on July 30, 2019; [6] In Dec 2019, we acquired our joint venture partner's 50% shareholding in Nutrition Science Partners, after which Nutrition Science Partners has become our subsidiary and the Group consolidated its financial position, which contributed net cash inflow of \$8.7m.







## Potential 2020 upcoming events



\* submission to scientific conference; \*\* subject to regulatory interaction; \*\*\* subject to supportive data; Targets: Savolitinib = MET; Fruquintinib = VEGFR1/2/3; Surufatinib = VEGFR1/2/3 / FGFR1 / CSF-1R; HMPL-523 = Syk; HMPL-689 = PI3K\delta; Indications: NHL = Non-Hodgkin's Lymphoma; NET = Neuroendocrine tumors; RCC = Renal cell carcinoma; NSCLC = Non-small cell lung cancer.



2020 Targets		MED		
Suru Launch	<ul> <li>Chi-Med's first unpartnered oncology drug launch</li> <li>Oncology commercial team targeting ~300-350 staff</li> </ul>			
Savo Breakout	SAVANNAH (w/Tagrisso®) interim	SAVOIR PRCC data & strategy		
ELUNATE® NRDL	<ul> <li>NRDL Jan 2020 - broad China access</li> <li>Establish Elunate<sup>®</sup> as best-in-class VEGFR TKI</li> </ul>			
US & EU C&R Team	<ul> <li>Fruq &amp; Suru global Phase IIIs starting</li> <li>HMPL-523 (Syk) &amp; HMPL-689 (PI3Kδ) global development</li> </ul>			
M&A (In 2020 & beyond)	<ul> <li>Add large molecule development capability/assets</li> <li>Non-core commercial assets</li> </ul>			





HUTCHISON CHINA MEDITECH

Thank you





## (A1a) R&D Strategy and Portfolio Overview

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





## **Global step-change innovation**

• Aiming for multiple potential first-in-class assets



## Kinase selectivity – enable combos

• Limit off-target toxicity & address TKI resistance



# Discovery of broad range of assets against novel targets





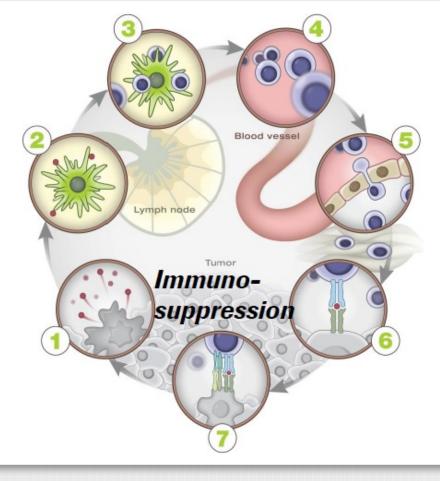
## Attack cancer from multiple angles at same time

### 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

Note: Adapted from Chen DS et al. Oncology Meets Immunology: The Cancer-Immunity Cycle. Immunity, Volume 39, Issue 1, 1 - 10.

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

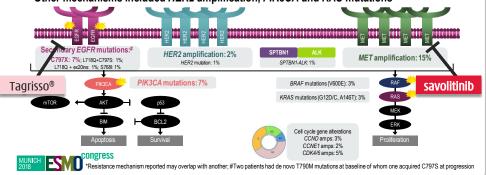




- MET amplification is the most common resistance mechanism for Tagrisso<sup>®</sup>.
- Requires addition of MET inhibitor savolitinib – in combo with Tagrisso<sup>®</sup>.

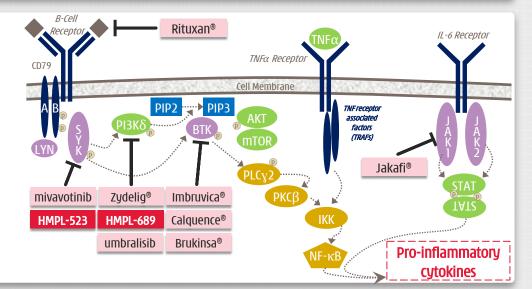
### RESULTS: CANDIDATE ACQUIRED RESISTANCE MECHANISMS WITH OSIMERTINIB (n=91)\*

- 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



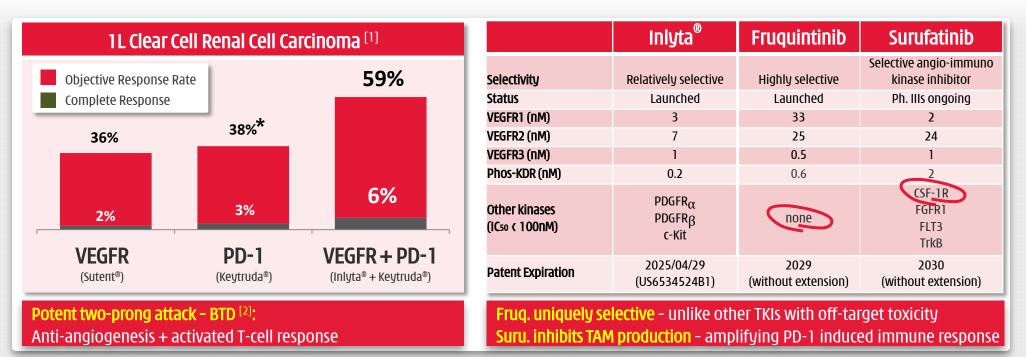


- C481S or PLCγ are the most common resistance mechanisms for Imbruvica<sup>®</sup>.
- Invalidating BTK inhibitor requires a possible Syk, PI3Kδ &/or BTK TKIs.

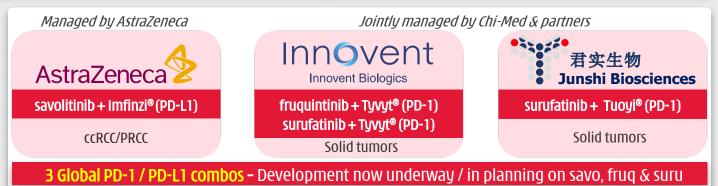


## Immunotherapy combinations... assets potentially ( ideal TKI combo partners for immunotherapy





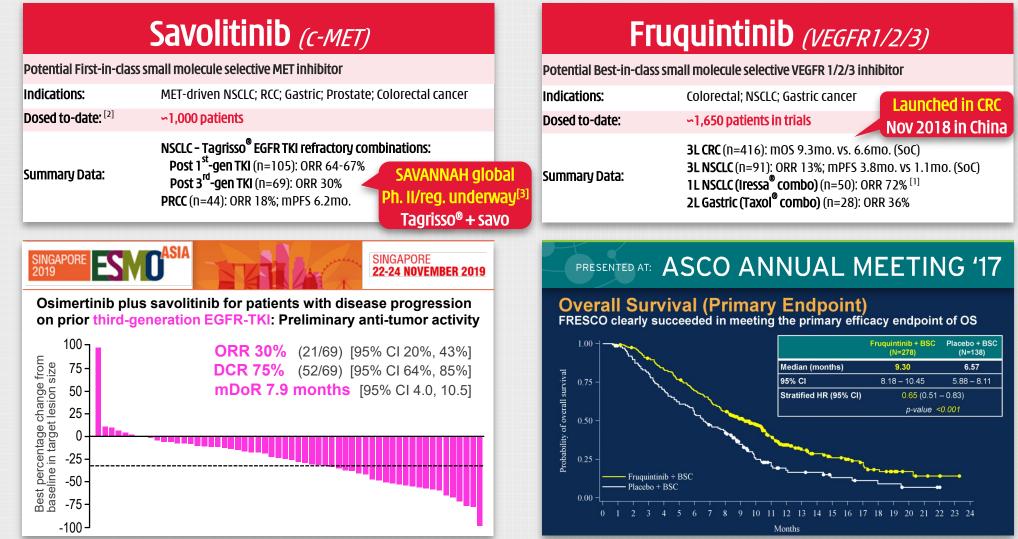
Multiple global immunotherapy combo deals...



[1] Sources: (i) 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; (ii). 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; \* ORR =38.2% for all PD-L1 expression combined positive scores (CPS) - ORR=50.0% for CPS 21 pts, ORR=26.4% for CPS (1 pts.; [2] BTD = Breakthrough Therapy Designation.

## Global clinical drug portfolio (1/2)

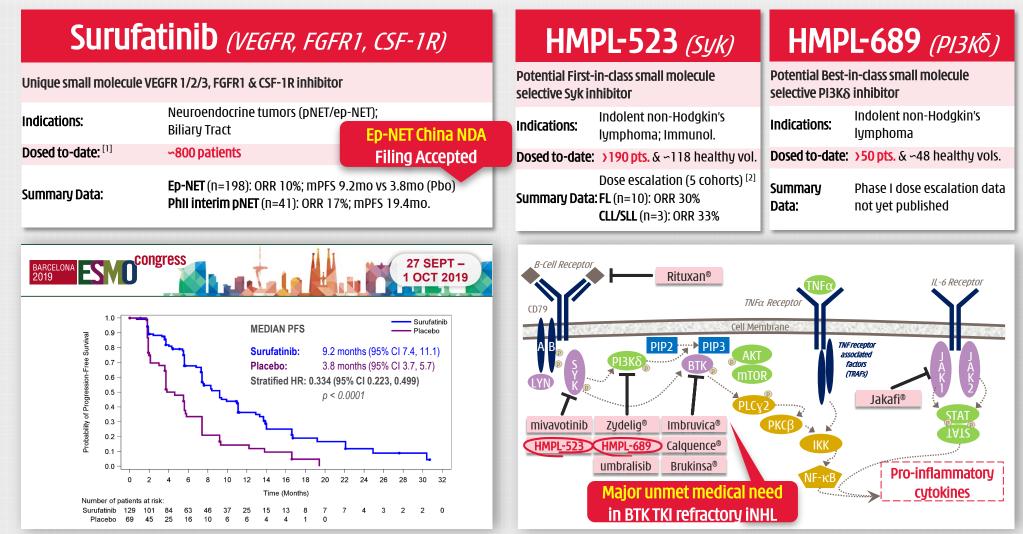




MET = mesenchymal epithelial transition receptor, VEGFR = vascular endothelial growth factor receptor, NSCLC = non-small cell lung cancer, RCC = renal cell carcinoma, PRCC = papillary RCC, CRC = colorectal cancer; [1] Lu, S., et al, "Phase II Study of Fruquintinib plus Gefitinib in Stage IIIb/IV NSCLC Patients Harboring EGFR Activating Mutations", #4780 ESMO Asia, Singapore, Nov 23, 2019; [2] Patients in all clinical trials (treatment & placebo); [3] Phase II registration intent study subject to regulatory discussions.

## 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, Syk = spleen tyrosine kinase, Pl3K\delta = Phosphatidylinositol-3-Kinase delta, pNET = pancreatic neuroendocrine tumors, ep-NET = non-pancreatic neuroendocrine tumors, FL = follicular lymphoma, CLL = chronic lymphocytic leukemia, SLL = small lymphocytic leukemia.

## 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 <sup>®</sup> ref.; MET+	SAVANNAH	Global	Oxnard/Ahn – DF/SMC		
	Savolitinib	Papillary RCC	MET+	SAVOIR	Global	Choueiri – Dana-Farber		
	Savolitinib + Imfinzi® (PD-L1)	Papillary RCC *	All	CALYPSO	UK/Spain	Powles – Queen Mary's		Interim PoC at
Savolitinib MET	Savolitinib + Imfinzi® (PD-L1)	Clear cell RCC *	VEGFR TKI refractory	CALYPSO	UK/Spain	Powles – Queen Mary's		ASCO GU Feb 2020
	Savolitinib	Gastric cancer *	MET+	VIKTORY	S Korea	Lee – Samsung Med. Ctr		PoC published in
	Savolitinib	Prostate cancer *	MET+	CCTG I234B	Canada	Kolinsky/Muk'jee/Ong/Chi		Can. Discovery Oct 2019
	Savolitinib	Colorectal cancer *	MET+		US	Strickler – Duke Uni		
								Planning US/EU registr.
Fruquintinib	Fruquintinib	Colorectal cancer	Stivarga <sup>®</sup> /Lonsurf <sup>®</sup> ref./intol.	FRESCO2	US	Eng /Desari – MD And. [1]		study based on FRESCO / US Ph. Ib
VEGFR 1/2/3	Fruquintinib + Tyvyt <sup>®</sup> (PD-1)	Solid tumors				In planning		
								Planning US/EU registr.
Surufatinib VEGFR 1/2/3;	Surufatinib	NET	Refractory		US	Dasari/Yao – MD Anderson		study based on China Ph.III / US Ph. Ib
FGFR1; CSF-1R	<b>Surufatinib</b> + Tuoyi <sup>®</sup> (PD-1)	Solid tumors				In planning		
								US & EU Phase I/Ib study
HMPL-523	HMPL-523	Indolent NHL			Australia			enrolment underway
Syk	HMPL-523	Indolent NHL			US			
								US & EU Phase I/Ib study
HMPL-689	HMPL-689	Healthy volunteers			Australia			enrolment underway
ΡΙ3Κδ	HMPL-689	Indolent NHL			US	Ghosh/Cohen-Levine/Emory		

[1] in U.S., in E.U. Tabernero - Vall d'Hebron & Sobrero - Genova; \* Investigator initiated trials (IITs).

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, PI3K\delta = Phosphatidylinositol-3-Kinase delta, NSCLC = non-small cell lung cancer, RCC = renal cell carcinoma, NET = neuroendocrine tumors; NHL = Non-Hodgkin's Lymphoma; ASCO GU = American Society of Clinical Oncology Genitourinary Cancer Symposium; PoC = Proof of Concept.

## **8 assets in China development** ...fruq launched – savo/suru NDAs & Syk/PI3Kδ PoC ahead



Program	Treatment	Indication	Target patient	Study name	Sites	Dose find / safety run-in	Proof-of-concept	Registration	Fully Enrolled
Savolitinib	Savolitinib	NSCLC	MET Exon 14 deletion		China	Lu Shun – SH Chest Hosp.			NDA H1'20
MET	<b>Savolitinib</b> + Iressa®	NSCLC	2L EGFRm; Iressa <sup>®</sup> ref.; MET+		China	Wu Yilong – GD General			Launched
	Savolitinib	Gastric cancer	MET+		China	Shen Lin – BJ Univ. Tumor			Nov 2018
	Fruquintinib	Colorectal cancer	≥3L; chemotherapy refractory	FRESCO	China	Li Jin – Fudan Univ.			Interim OK
_	Fruquintinib + Taxol®	Gastric cancer	2L	FRUTIGA	China	Xu Ruihua – Sun Yat Sen			April 2019
Fruquintinib VEGFR 1/2/3	Fruquintinib + Iressa®	NSCLC	1L EGFRM		China	Lu Shun – SH Chest Hosp.		-	ESMO Asia
	<b>Fruquintinib</b> + Tyvyt <sup>®</sup> (PD-1)	Solid tumors			China	Bai Yuxian - Harbin Med. U.			Nov 2019
	<b>Fruquintinib</b> + genolimzumab (PD-1)	Solid tumors			China	Li Jin – Fudan Univ.			Met Primary
	Surufatinib	Pancreatic NET	All	SANET-p	China	Xu Jianming - #5 Med. Ctr.			Endpt (PFS)
Surufatinib	Surufatinib	Non-Pancreatic NET	All	SANET-ep	China	Xu Jianming - #5 Med. Ctr.			Jan 2020
VEGFR 1/2/3;	Surufatinib	Biliary tract cancer	2L; chemotherapy refractory		China	Xu Jianming – #5 Med. Ctr.			NDA accepted
FGFR1; CSF-1R	<b>Surufatinib</b> + Tuoyi <sup>®</sup> (PD-1)	Solid tumors			China	Shen Lin – BJ Univ. Tmr.			Nov 2019
	<b>Surufatinib</b> + Tyvyt <sup>®</sup> (PD-1)	Solid tumors			China	In planning			
HMPL-523	HMPL-523	B-cell malignancies	All		China	Multiple leads by sub-types		Pha	ase I/Ib data <b>to</b>
Syk	HMPL-523	ITP	All		China	Yang – CN Hem. Hosp.		info	orm registration
HMPL-689	HMPL-689	Indolent NHL			China	Cao/Zhou – Fudan/ Tongji			decisions
ΡΙ3Κδ									data <b>to inform</b>
Epitinib	Epitinib	NSCLC	EGFRm with brain metastasis		China	Wu Yilong – GD General		registrat	cion decisions
EGFR	Epitinib	Glioblastoma	EGFR gene amplified		China	Ying Mao – SH Huashan			
Theliatinib	Theliatinib	Esophageal cancer	EGFR over-expression		China	nı			
EGFR wt		. 5							
HMPL-453	HMPL-453	Solid tumors			China	Xu Ruihua – SYS			Phase II set to
FGFR 1/2/3									initiate





## China Oncology Opportunities

Next-gen oncology drugs to meet major needs in China

## China oncology - ~24% of world's cancer patients<sup>[1]</sup> 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<sup>®</sup> launch in 3L mCRC; First ever in China<sup>[3]</sup>
- Deep pipeline 8 clinical drug candidates with 5 registration studies underway/set to start in China



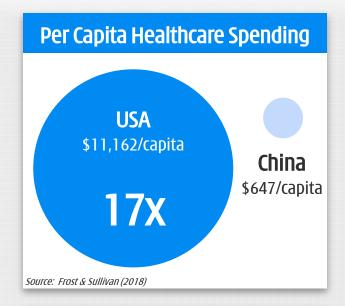
## Major commercial opportunity

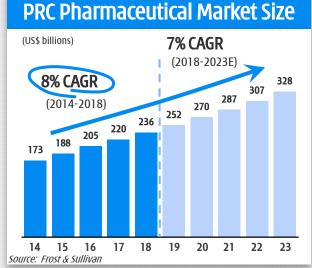
• National Drug Reimbursement; Medical coverage



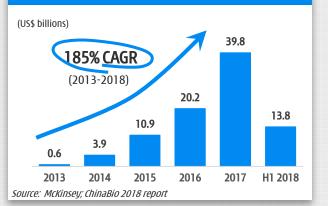
# China now world's 2<sup>nd</sup> largest pharma market ...investment, approvals & access all accelerating rapidly



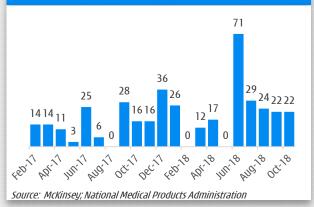




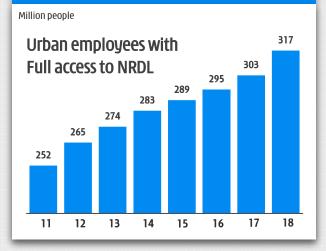
#### PRC Healthcare VC/PE Funds<sup>[2]</sup>



### Number of Priority Review NDAs<sup>[3]</sup>



### Medical Insurance Coverage<sup>[1]</sup>



### **Improved Access since 2017**

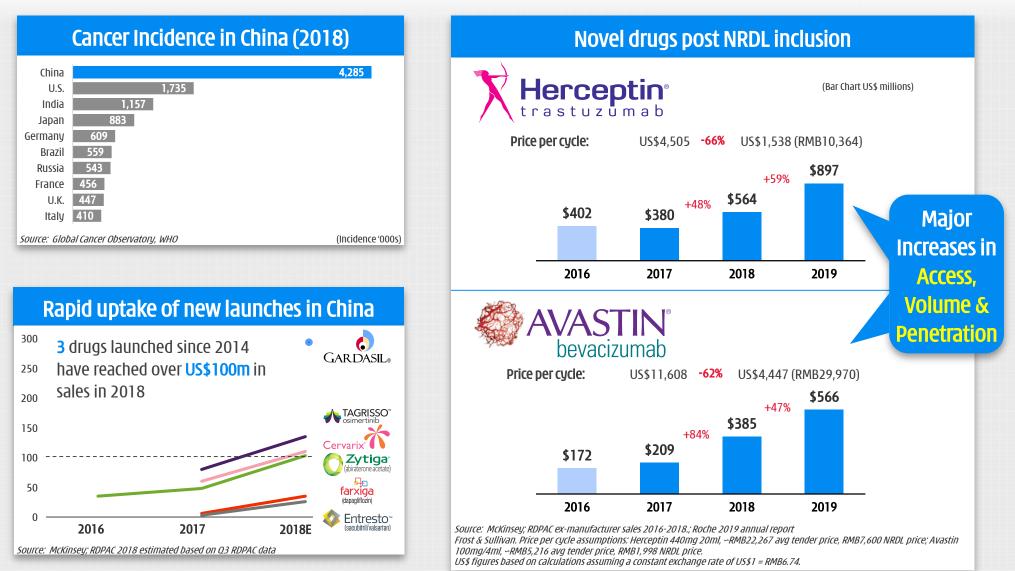
- 128 western drugs added to NRDL;
- Further 17 oncology drugs added to NRDL in Oct 2018 (15 in Jul 2017);
- Essential drug list expanded from 520 to 685 molecules. Including oncology.

Source: McKinsey

[1] Urban Basic Medical Care Insurance (for both employees & residents) - total persons covered at year-end. National Bureau of Statistics (2017); includes rural residents from 2017 and beyond; [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









Cash generation & China commercial know-how / infrastructure



## **China Commercial**

### Chi-Med spent 19 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
   >\$260 million dividends since inception

### China pharma industry grew at ∽10% CAGR over last 15 years <sup>[1]</sup>

• 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

2,400 RX & ~900 OTC sales people in over 330 <sup>[1]</sup> cities & towns in China.

Drugs in >25,200 hospitals detailing ~82,000 doctors.

Sold ~4.7 billion doses of medicine in 2019.

#### Leadership Market Shares

Market leader in the subcategories/markets in which we compete <sup>[2]</sup>:

SXBX pill: <sup>[3][4]</sup>	∽18%
Rx Cardiovascular TCM	
Banlangen: <sup>[5]</sup>	∽ <b>5</b> 4%
OTC Anti-viral /flu TCM	
FFDS tablet: <sup>[6]</sup>	∽ <mark>38%</mark>
OTC Angina TCM	

### JVs with 3 Major China Pharmas





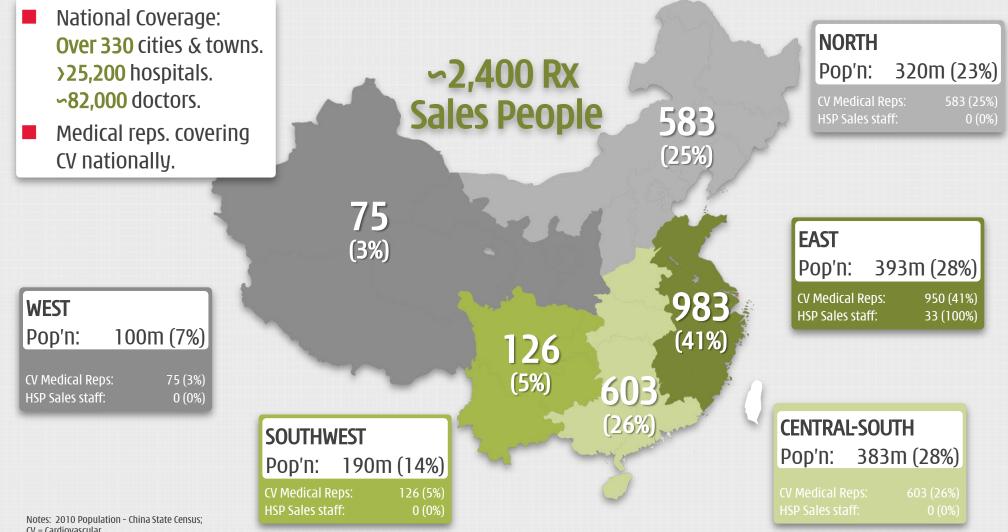


63



[1] 330 cities & towns covered by Prescription Drug Business and 600 cities & towns including OTC business; [2] Frost & Sullivan 2018 market share data, except SXBX pill which is 2019 data; [3] China coronary heart disease oral Chinese patented drugs market share; [4] She Xiang Bao Xin Pill ("SXBX pill") - RX Coronary artery disease; [5] Banlangen Granules ("Banlangen") - OTC Antiviral; [6] Fu Fang Dan Shen tablets ("FFDS") - OTC Angina.

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



CV = Cardiovascular Chi-Med Rx sales team data = December 31, 2019







**Product Candidate Details** Further details on each drug candidate







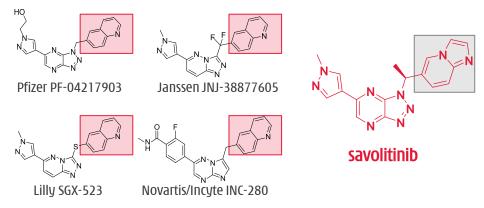
Savolitinib Potential first-in-class selective MET inhibitor

## Savolitinib



## Potential first-in-class selective MET inhibitor

- 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.
- Savolitinib design eliminates renal toxicity first generation of selective MET inhibitors encountered – ~1,000 patients involved in clinical studies to date.



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

2. MET is aberrant in many tumor settings.<sup>[7]</sup>

		New Cases (2018)			
Indication	Amplification	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 December 2019);
- 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 <sup>[8]</sup> & 30% flat rate China royalty on all product revenues.

67

[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 aggregate sales of savolitinib have reached \$5bn, the royalty will step down over a two-year period, to an ongoing royalty rate of 10.5% to 14.5%.

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

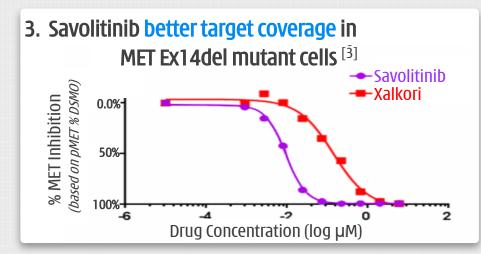


### 1. Competitive landscape outside China:

			Treatment Line	N	BICR <sup>[1]</sup> ORR	95% CI
Capmatinib	coloctivo MET	ASCO 2019 #9004	2/ <u>3L</u>	69	<b>40.6%</b> (28/69)	28.9%, 53.1%
(Novartis/ Incyte)	selective MET	ASCO 2019 #9004	(IL)	28	<b>67.9%</b> (19/28)	47.6%, 84.1%
<b>Tepotinib</b> (Merck Serono)	selective MET	ASCO 2019 #9005	39% 1L, 61% ≥2L	51	<b>45.1%</b> (23/51)	31.1%, 59.7%
Xalkori®	multi kinaca	WCLC 2018 #13453	38% 1L	65	<b>32.3%</b> (21/65) <sup>[2]</sup>	<b>21%, 45%</b> <sup>[2]</sup>
(Pfizer)	multi-kinase	WCLC 2018 #12937	Median 1L (1L-4L)	25	<b>40.0%</b> (10/25)	21%, 61%

## 2. Xalkori<sup>®</sup> a multi-kinase TKI – selective MET inhibitors reporting better response – superior selectivity.

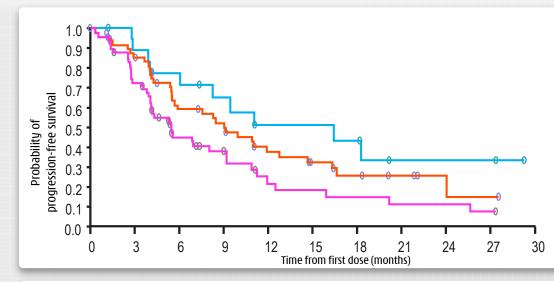
	Savolitinib IC <sub>50</sub>	Xalkori® IC <sub>50</sub>	Savolitinib vs. Xalkori®
EBC1 Viability	2nM	19nM	10x
EBC1 pMET	1	39	40x
293T MET (wild type)	7	79	11x
293T MET (Ex14del)	9	140	16x



[1] BICR = blinded independent central review; [2] Investigator reviewed data (not BICR); [3] Paik, P.K., et al., Response to MET inhibitors in patients with stage IV lung adenocarcinomas harboring MET mutations causing exon 14 skipping. Cancer Discov, 2015. 5(8): p. 842-9.

## TATTON B & D data - PFS

## Tagrisso<sup>®</sup> + savolitinib in EGFR TKI refractory NSCLC



12

9

15

18

Time from first dose (months)

21

24

27

1.0

0.9

0.8

0.7

0.6 0.5

0.4

0.3 0.2 0.1 0.0

0

3

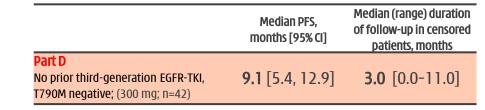
progression-free survival

Probability of

	Median PFS, months [95% Cl]	Median (range) duration of follow-up in censored patients, months
<b>Part B1</b> Prior third-generation EGFR-TKI; (600 mg <sup>[1]</sup> ; n=69)	<b>5.4</b> [4.1, 8.0]	<b>2.6</b> [0.0-27.3]
Part B2 No prior third-generation EGFR-TKI, T790M negative; (600 mg <sup>[1]</sup> ; n=51)	<b>9.0</b> [5.5, 11.9]	<b>10.1</b> [0.0-27.5]
<b>Part B3</b> No prior third-generation EGFR-TKI, T790M positive; (600 mg <sup>[1]</sup> ; n=18)	11.0 [4.0, NR]	<b>14.7</b> [1.2-29.3]

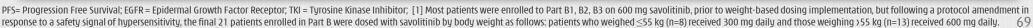
#### Progression data had a maturity of 62%.

Patients who had not progressed or died at the time of analysis were censored at the time of the latest date of assessment from their last evaluable RECIST assessment. Circles indicate censored observations.



#### Progression data had a maturity of 40%.

Patients who had not progressed or died at the time of analysis were censored at the time of the latest date of assessment from their last evaluable RECIST assessment. Circles indicate censored observations.



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## TAGRISSO<sup>T</sup> + Savo in EGFR TKI refractory NSCLC TATTON B & D data - AEs & tolerability



Event, n (%)	<b>All Part B (n=138)</b> osimertinib 80 mg <b>+ savolitinib 600 mg</b> <sup>[1]</sup>	<b>Part D (n=42)</b> osimertinib 80 mg + <b>savolitinib 300 mg</b> <sup>[1]</sup>
Any AE	135 (98)	39 (93)
Any AE possibly related to savolitinib	115 (83)	25 (60)
AE grade $\geq$ 3	79 (57)	16 (38)
AE possibly causally related to study treatment leading to discontinuation of:		
Savolitinib	38 (28)	9 (21)
Osimertinib	14 (10)	2 (5)
Any AE leading to death	6 (4)	2 (5)
Any SAE	62 (45)	11 (26)

[1] Most patients were enrolled to Part B1, B2, B3 on 600 mg savolitinib, prior to weight-based dosing implementation, but following a protocol amendment in response to a safety signal of hypersensitivity, the final 21 patients enrolled in Part B were dosed with savolitinib by body weight as follows: patients who weighed  $\leq$ 55 kg (n=8) received 300 mg daily and those weighing >55 kg (n=13) received 600 mg daily. Part D data are preliminary, therefore, for osimertinib, the mean actual treatment exposure was 8.5 months vs 6.1 months for Parts B and D, respectively, and 7.1 months vs 4.9 months for Parts B and D, respectively; Sequist LV, Han JY, Ahn MJ, et al. Osimertinib plus savolitinib in patients with EGFR mutation-positive, MET-amplified, non-small-cell lung cancer after progression on EGFR tyrosine kinase inhibitors: interim results from a multicentre, open-label, phase 1b study. Lancet Oncol. 2020; S1470-2045(19)30785-5. doi:10.1016/S1470-2045(19)30785-5

# TATTON B & D data - AEs & SAEsTAGRISSOMost common $AEs^{[1]}$ independent of causality & SAEs ( $\geq$ 3%)<sup>[2]</sup>



All Part B (n=138)		Part D (n=42)			All Part B (n=138)		Part D (n=42)		
AE*, n (%)	All grades	Grade≥3	All grades	Grade $\geq$ 3	AE*, n (%)	All grades	Grade $\geq$ 3	All grades	Grade $\geq$ 3
Nausea	67 (49%)	4 (3%)	13 (31%)	0	Rash	26 (19%)	3 (2%)	8 (19%)	0
Fatigue	48 (35)	6 (4)	4 (10)	0	Stomatitis	26 (19)	0	4 (10)	0
Decreased	47 (34) 5	F (4)	6 (14) 1 (2)	Constipation	26 (19)	0	3 (7)	0	
appetite	47 (54)	5 (4)		Pruritus	24 (17)	1(1)	5 (12)	0	
Vomiting	46 (33)	6 (4)	5 (12)	0	Headache	23 (17)	0	3 (7)	0
Oedema	44 (22)	) 3 (2)	9 (10)	(10) 0	Myalgia	22 (16)	3 (2)	6 (14)	1 (2)
peripheral	44 (32)		0(19) 0	8 (19)	0	Cough	22 (16)	0	4 (10)
Diarrhoea	39 (28)	4 (3)	8 (19)	2 (5)	AST increased	21 (15)	9 (7)	2 (5)	0
Paronychia	30 (22)	3 (2)	7 (17)	0	Pneumonia	15 (11)	7 (5)	7 (17)	5 (12)
Pyrexia	29 (21)	1(1)	6 (14)	0					

SAE**, n (%)	All Part B (n=138)	Part D (n=42)
Pneumonia	5 (4%)	4 (10%)
Anaphylactic reaction	6 (4)	1 (2)
Pneumothorax	6 (4)	1 (2)
Pyrexia <sup>#</sup>	5 (4)	0
Dyspnoea	5 (4)	0
Drug hypersensitivity	4 (3)	1 (2)
Diarrhoea	4 (3)	1 (2)
Back pain	4 (3)	0
Pulmonary embolism	3 (2)	2 (5)

[1] ≥15% in either Part B or Part D for all grades; [2] ≥3% in either Part B or Part D for all grades. #The emergence of drug-related hypersensitivity AEs are characterised by events such as pyrexia; The emergence of hypersensitivity and anaphylaxis events led to a protocol amendment introducing a weight-based savolitinib dosing regimen (for the last group of patients enrolled in Part B) in parallel to the lower dose of savolitinib (300 mg) being tested (for all patients enrolled in Part D)

## Savolitinib – 2L NSCLC<sup>[1]</sup> combo w/ IRESSA<sup>®</sup> gefitinib<sup>®</sup> Encouraging in MET+ / T790M-, next step under discussion

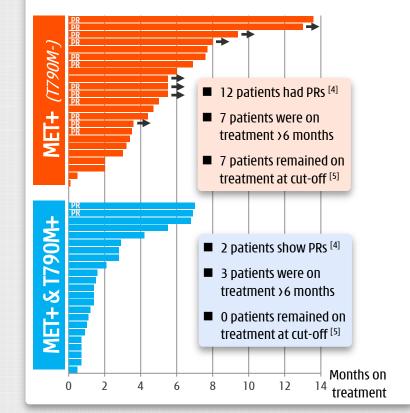
Savo / Iressa<sup>®</sup> combo in 1<sup>st</sup> gen. EGFRm-TKI refractory patients<sup>[2]</sup>...outstanding response in MET+ / T790M-

WCLC 2017	MET+ / T790M+ (n = 23)	MET+ <i>(T790M-)</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 / 1	Tagrisso <sup>®</sup> combo) <sup>[3]</sup>
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	MET+ / T790M+ (n = 18) Lancet Onc. 2020 <sup>[3]</sup>	MET+ <i>(T790M-)</i> (n = 51) Lancet Onc. 2020 <sup>[3]</sup>		
Confirmed response	12 (67%)	33 (65%)		
Stable disease≥ 6 weeks	6 (33%)	12 (24%)		
Progressive disease / death	0 (0%)	3 (6%)		
Not Evaluable	0 (0%)	3 (6%)		
MET status locally or centrally confirmed.				

### ...Iressa<sup>®</sup> combo - <u>6mo</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 plus gefitinib for patients with EGFR-mutant MET-amplified advanced NSCLC; [3] Sequist LV, Han JY, Ahn MJ, et al. Osimertinib plus savolitinib in patients with EGFR mutationpositive, MET-amplified, non-small-cell lung cancer after progression on EGFR tyrosine kinase inhibitors: interim results from a multicentre, open-label, phase 1b study. Lancet Oncol. 2020; S1470-2045(19)30785-5. doi:10.1016/S1470-2045(19)30785-5; [4] PR = Partial Response.

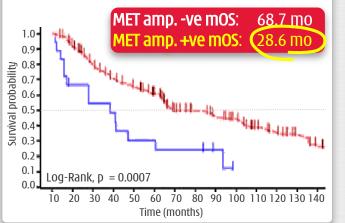
# Savolitinib – MET+ gastric cancer A major problem in east Asia – Japan, South Korea & China



1. Gastric (stomach) cancer is the 5<sup>th</sup> most common cancer globally -782,700 deaths/year 1,034 New cases ('000) 442 116 38 133 26



2. MET+ disease is more aggressive [1]

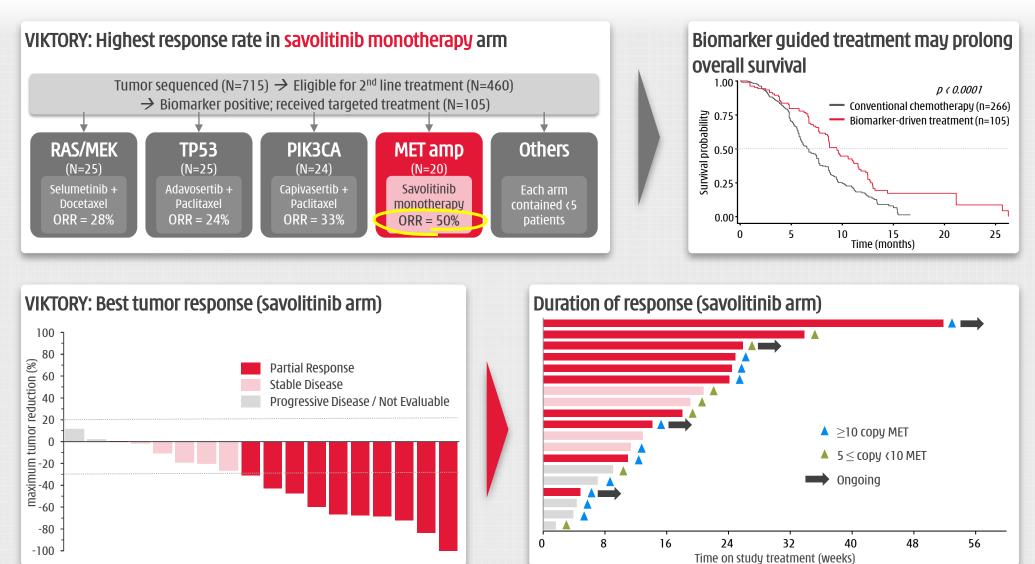


3. VIKTORY trial savolitinib arm – male, 34; surgery ruled-out; failed 4-cycles XELOX. **Baseline** ... after PET CT... 3 weeks savolitinib 600mg. Jeeyun Lee, AACR 2016.

[1] Catenacci, et al. "MET tyrosine kinase receptor expression and amplification as prognostic biomarkers of survival in gastroesophageal adenocarcinoma." Cancer. 2017 Mar 15; 123(6): 1061-1070. doi: 10.1002/cncr.30437. [2] Lee, et al. "Tumor genomic profiling guides metastatic gastric cancer patients to targeted treatment: The VIKTORY Umbrella Trial." Cancer Discov. 2019 Jul 17. pii: CD-19-0442. doi: 10.1158/2159-8290.CD-19-0442. <5 patients in all other arms.

## Savolitinib potential in gastric cancer VIKTORY Phase II trial highly promising in MET+ gastric cancer











Surufatinib Highly active TKI with unique angio-immuno activity

#### 100 000 for Neuroendocrine Tumors Tumor begins in the specialized cells of the body's neuroendocrine system. Cells have traits of both hormone-producing endocrine cells & nerve cells.

ncidence per

SEER 9

Overview of NET - ~170,000 patients in the U.S. [1][2][3]

Found throughout the **body's organs**. Most NETs take years to develop but some can grow fast.

What are neuroendocrine tumors ("NET")?

#### Hormone-related symptoms<sup>[1]</sup>

~2% of all malignancies.

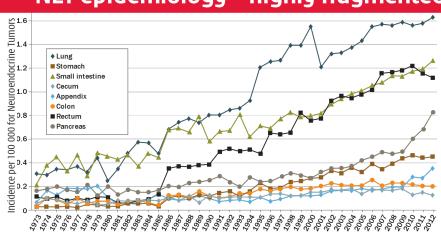
Functional NETs (~8-35% of patients) release hormones / peptides causing symptoms like diarrhea & flushing; Non-functional NETs have no symptoms.

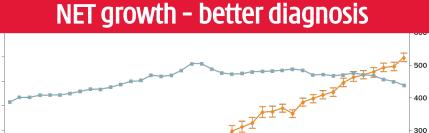
#### Differentiation & biomarkers for grading:

- Well differentiated: look like healthy cells grow slowly; **Poorly differentiated:** look less like healthy cells - grow quickly;
- Mitotic count Mitosis is process by which tumor cells grow & divide; Ki-67 index - Ki-67 a protein that increases as cells divide.

& Survival Outcomes in Patients With Neuroendocrine Tumors in the U.S. JAMA Oncol. 2017;3(10):1335-1342. [2] www.cancer.net (patient information from ASCO) - NET is a subtype of neuroendcrine neoplasms, NENs); [3] IQVIA 2019

NET epidemiology - highly fragmented





Incidence of NETs

Incidence of all malignant neoplasms



Incidence per 100 000 for All Malignant Neoplasms

76

200

# Surufatinib

# High-level NET landscape Long-term disease – rapid deterioration in later stages <sup>[1][2][3]</sup>



**mOS:** 

16.2 yrs.,

Well Differentiated

Ki-67 Index <2; Mitotic Count <2

#### ∽8-35% NET patients -Functional NET -

Hormone related symptoms:

> 94% flushing 78% diarrhea 53% heart plaque 51% cramping

Symptoms allow early diagnosis

#### Somatostatin Analogue

Treatment - modulate/ control symptoms related to hormone overproduction & tumor growth:

Octreotide: \$1.6b revenue (2019) Lanreotide: \$1.2b revenue (2019) G1/2 – Advanced NET Regional / Distant

∽60% NET patients - first diagnosis at advanced disease stage -Mostly non-Functional NET - TKIs<sup>[4]</sup>; chemo/ radiotherapy

> mos: 8.3 yrs.

**G3 – NET/NEC** Distant

No approved treatments - exploring *I/O*<sup>[5]</sup> + *TKI combos* 



**Poorly Differentiated** *Ki-67 Index >20; Mitotic Count >20* 

Moderately Differentiated Ki-67 Index 3-20; Mitotic Count 2-20

[1] Arvind Desari et. al. Trends in the Incidence, Prevalence, and Survival Outcomes in Patients With Neuroendocrine Tumors in the US, JAMA Oncol. 2017;3(10):1335-1342; [2] Van Cutsem et al. ESMO - Neuroendocrine Tumors Diagnostic & Therapeutic Challenges; [3] mOS = median overall survival; [4] TKIs = Tyrosine Kinase Inhibitors; [5] I/O = Immuno oncology/immunotherapy

## Surufatinib – China NET NET potential ~\$100-120m/yr.<sup>[1]</sup> – under treated/diagnosed



#### **Competitive landscape –** *China NET treatments*<sup>[1]</sup>

Brand	Indication/s	Launched		2017	2018	01-2019
SUTENT®	Pancr. NET	2007	Sales (US\$ million)	27	24	7
<i>(sunitinib - VEGFR)</i> Pfizer	(& GIST/RCC)		List Price (US\$/month)	4,455	NRDL Oct-18	2,007
<b>AFINITOR</b> <sup>®</sup>	Pancr. NET	2013	Sales (US\$ million)	9	13	3
<i>(everolimus – mTOR)</i> Novartis	(& 2L RCC)		List Price (US\$/month)	NRDL Jul-17	1,320	1,320
SANDOSTATIN LAR®	GEP-NENS <sup>[3]</sup>	2003	Sales (US\$ million)	14	15	5
<i>(octreotide – SSA</i> <sup>[2]</sup> ) Novartis			List Price (US\$/month)	1,169	NRDL Oct-18	835

#### Pancreatic-NET market est. ~\$10-15m/yr. - Non-Pancreatic NET market ~5-10X

# SANET-ep vs. RADIANT-4 - cannot compare SANET-ep broader range of tumor origins & later-stage patients



		Asia/China Extra- Pancreatic NET	SANET-ep (n=198) (surufatinib vs placebo)		U.S. Extra- Pancreatic NET	RADIANT-4 (n=302) (everolimus vs placebo)		<b>SANET-ep</b> Enrolled more pts with poor prognosis. Survival Rate
		Tsai et al. 2013	470/	Castrointectinal Tract	Yao et al. 2008	58%		Primary Site mOS @ 5-yr Rectum 2.8y 28%
	Gastrointestinal Tract Rectum Stomach Small Intestine	<b>58%</b> 30% 7% 19%	47% 27% 10% 8%	Gastrointestinal Tract Rectum Stomach Small Intestine	<b>50%</b> 33% 8% 6%	58% 13% 4% 34% 7%	4	Stomach 2.4y 32% Small Intestine 8.6y 69%
	Other GI Lung Other Organ Site	3% <b>22%</b>	3% 12% 28%	Lung Thymus	4% <b>21%</b>	30%		RADIANT-4
Tumor Origin	Thymus Liver Mediastinum Adrenal Gland		7% 6% 6% 2%					Did not enrol other extra-pancreatic NET organ sites incl. but not limited to
	Other <b>Unknown Origin</b>		8% <b>14%</b>	Unknown Origin		12%		Kidney Ovary SANET-ep
								Mediastinum Adrenal gland Retroperitoneal Ampulla vater Parathyroid gland Carotid body Liver COVErage.
Pathology grade	Grade 1 <b>Grade 2</b>		16% 84%			<b>65%</b>		
ECOG PS 0:1	PS 0 (treatment : control) PS 1		60% (56% : 67%) 40% (44% : 33%)			74% (73% : 75%) 26% (27% : 26%)		SANET-ep
Prior systemic treatment	(treatment : control) <b>Any Prior Treatment</b> Chemotherapy Targeted therapy Somatostatin Analogues		67% 40% 10% 32%			61% 25% none 55%	À	<b>Later-stage patients</b> , more heavily pre- treated (incl. with targeted therapy) & weaker physical status.
Multiple organ involvement		<b>66% with multiple organ</b> 76% had liver metastasi 47% had lymph nodes n 33% had bone metastas 26% had lung metastasi	<b>i involvement</b> s netastasis is		79% had liver metastasis 43% had lymph nodes me 19% had bone metastasis 22% had lung metastasis	etastasis		Likely due to later diagnosis in China & availability of everolimus.

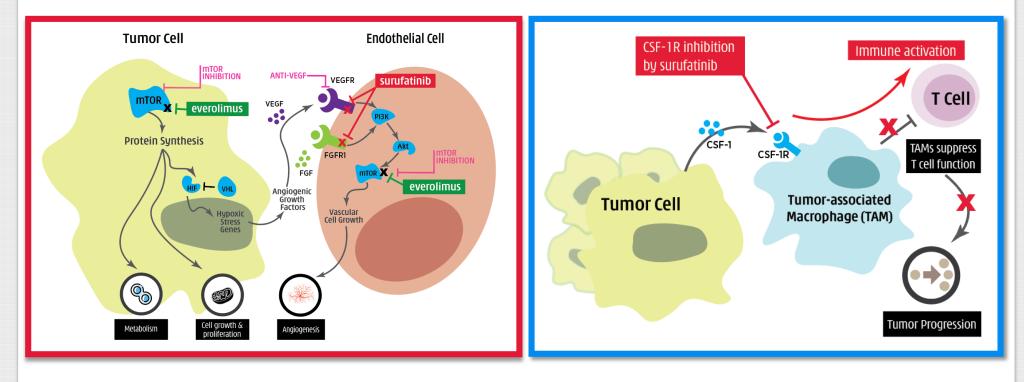
Source: Xu et al, ESMO 2019 #LBA76; Yao et al, Lancet 2016 387(10022) 968-77; Yao et al, JAMA Oncol 2017 3(10) 1335-42; Excludes 7% pancreatic NET in US series and 6% in Asia series; Colon-rectum in Tsai et al. (2013) report; Colon approximately 8% in Asian series (Shebani KO et al. (1999)); Colon-rectum in Yao et al. (2008) report; Colon approximately 4-7% in US/EU series (Niederle B et al. (2016)).

# Very different mechanism of action



**Everolimus** inhibits **mTOR** and blocks the effects caused by the loss of certain genes thereby reducing cell growth, proliferation, and angiogenesis.

Surufatinib inhibits VEGFR1/2/3 and FGFR1 blocking vascular cell growth and angiogenesis; as well as CSF-1R which limits the production of TAMs which cloak the cancer cell from T-Cell attack.



# ~170,000 NET patients in U.S.<sup>[1][2]</sup>

### U.S. NET treatment landscape - highly fragmented



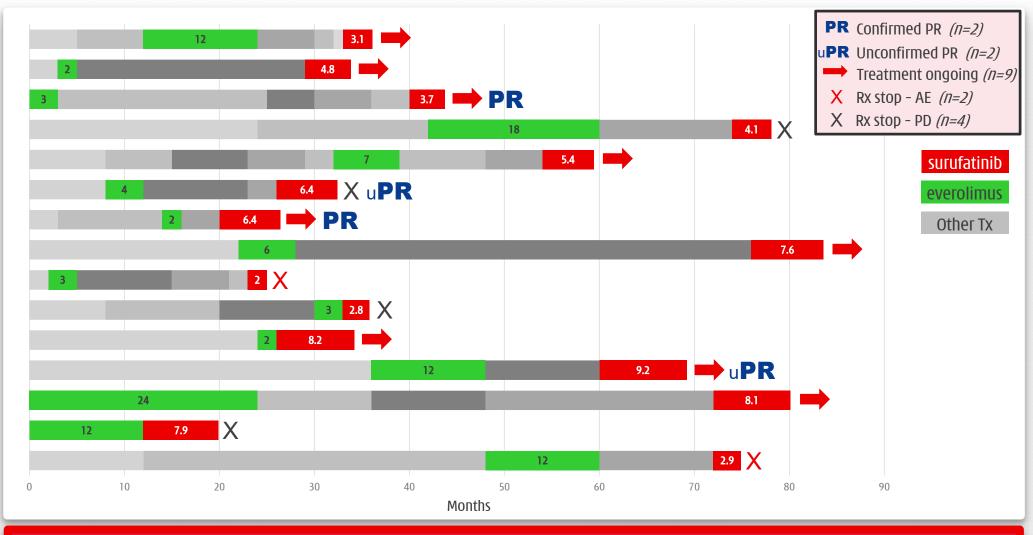
		Somatostatin Based Therapies	5	K	inase Inhibitor Therapies	
	Sandostatin <sup>®</sup> LAR (octreotide)	Somatuline Depot <sup>®</sup> (lanreotide)	Lutathera® ( <sup>177</sup> Lu-Dotatate)	Afinitor® (everolimus)	Sutent® (sunitinib)	Surufatinib (China NDA accepted)
2018 Sales	\$1.6bn	\$1.0bn	\$0.17bn	\$1.6bn	\$1.0bn	-
<b>MOA</b> <sup>[3]</sup>	Somatostatin analogue	Somatostatin analogue	Somatostatin receptor targeting radiotherapy	mTOR inhibition	Inhibits multiple receptor tyrosine kinases	VEGFR/FGFR1 & CSF-1R inhibition
Admin.	Subcutaneous or intramuscular inj. (LAR)	Subcutaneous injection	Subcutaneous injections (radio- qualified physicians).	Oral tablet	Oral capsules	Oral capsules
Shelf-life	3 years	2 years	72 hours	3 years	3 years	2+ years <sup>[5]</sup>
Dosage	2 wks: Sando. inj. 0.1-0.6mg per day; then 2 months Sando. LAR 20mg per 4 wks.	120mg inj. every 4 wks.	7.4GBq (one ~25ml vial) inj. every 8 wks - 4 doses total.	10mg orally once daily.	37.5mg taken orally once daily.	300mg orally once daily.
NET indication /s	LT treatment of severe diarrhea & flushing from meta. carcinoid tumors.	<ul> <li><u>GEP-NETs</u>: unresectable, well or moderately diff., (locally adv. or meta) GEP-NETs to improve PFS.</li> <li><u>Carcinoid Syndrome</u>: to reduce frequency of short-acting somatostatin rescue therapy.</li> </ul>	positive GEP-NETs.	<ul> <li><u>pNET</u>: progressive pNET (unresectable, locally adv. or meta).</li> <li><u>GI-NET or Lung NET</u>: progressive, well- diff., <i>non-functional</i> NET (unresectable, locally adv. or meta). Not for <i>functional</i> carcinoid tumors.<sup>[4]</sup></li> </ul>	<ul> <li><u>pNET</u>: Progressive, well- differentiated pNETs (unresectable locally adv. or meta).</li> </ul>	<ul> <li><u>Non-pNET</u>: SANET-ep study was in low- or intermediate- grade adv. non-pancreatic NET.</li> <li><u>pNET</u>: Phase III ongoing.</li> </ul>
Non-NET indication/s	• Acromegaly; watery diarrhea from VIPomas.	• Acromegaly.		• Adv. HR+ HER2-n breast cancer; adv. 2L RCC; renal angiomyolipoma and TSC.	• 2L GIST; adv. RCC; high risk of recurrent RCC.	

	Sandostatin® / Placebo	Somatuline Depot <sup>®</sup> / Placebo	Lutathera® + Sando. LAR / Sando. LAR	Afini Plac		Sutent® / Placebo		rufatinib / Placebo
mPFS (mo.) primary EP	14.3 / 6.0	NR / 18.0	NR / 8.5	pNET 11.0 / 4.6	Lung & GI NET 11.0 / 3.9	pNET: 11.4 / 5.5	Ph II pNET 19.4	Ph III non-pNET 9.2 / 3.8
HR	0.34	0.47	0.21	0.35	0.48	0.42	Ph III	0.33
(p-value)	0.000072	(0.001	(0.0001	<0.001	(0.001	(0.001	Ongoing	(0.0001
ORR	2% / 2%	NR	18% / 3%	5% / 2%	2% / 1%	9% / 0%	17% (Ph II)	10.3%
DCR	69% / 40%	NR	95% / 76%	73%/51%	81% / 64%	72% / 60%	90% (Ph II)	87%
Pivotal Trial	PROMID	CLARINET	NETTER-1	RADIANT-3	RADIANT-4	A6181111	SANET-p	SANET-ep

[1] Dasari A, et al.: Trends in the Incidence, Prevalence, & Survival Outcomes in Patients With Neuroendocrine Tumors in the U.S. JAMA Oncol. 2017;3(10):1335-1342; [2] www.cancer.net (patient information from ASCO) - NET is a subtype of neuroendocrine Tumors in the U.S. JAMA Oncol. 2017;3(10):1335-1342; [2] www.cancer.net (patient information from ASCO) - NET is a subtype of neuroendocrine neoplasms, NENs); [3] MOA = Mechanism of Action; [4] Afinitor is only approved for pancreatic neuroendocrine tumors in China; [5] 2-year stability studies completed so far; mPFS = median progression-free survival; HR = Hazard Ratio; ORR = objective response rate; DCR = Disease control rate.

## **Surufatinib efficacy post everolimus failure** U.S. Phase Ib (n=15) - pNET duration of treatment





#### Encouraging preliminary surufatinib efficacy post everolimus failure – different MOA<sup>[1]</sup>

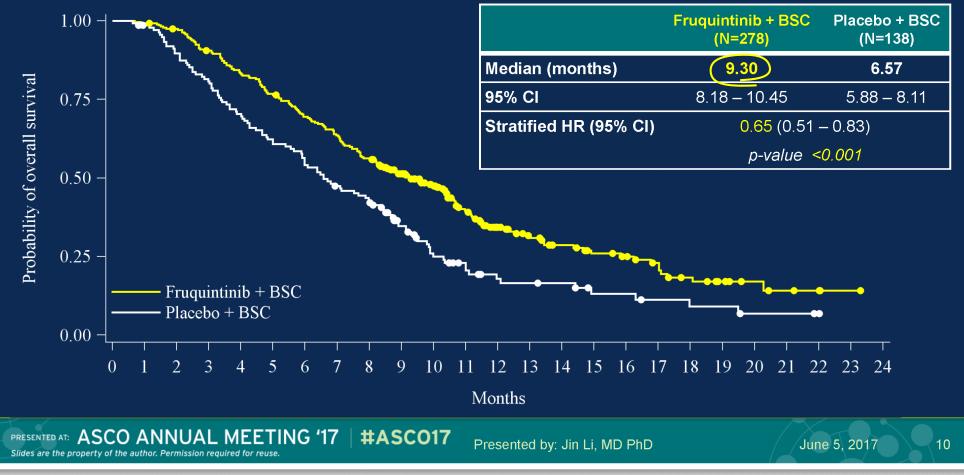




## **Fruquintinib – 3L/4L colorectal cancer** Develop in US/EU for rego/TAS-102 ref./intol. patients<sup>[1]</sup>



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



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

# ELUNATE<sup>®</sup> China VEGFR landscape



#### **Competitive landscape –** *small molecule VEGFR TKIs*

Brand	Indication/s	Launch		2011	2012	2013	2014	2015	2016	2017	2018	01-03 '19
STIVARGA®	<b>3L CRC</b> /2L GIST		Sales (US\$ million) [1]							5	21	62
<i>(regorafenib)</i> Bayer AG	2L HCC	Mar 2018	List Price (US\$/mo.)							4,368	NRDL Oct-18	2,352
NEXAVAR®	Unres. RCC & HCC		Sales (US\$ million) <sup>[1]</sup>	80	96	96	93	91	97	108	130	155
<i>(sorafenib)</i> Bayer AG	Diff. Thyroid can.		List Price (US\$/mo.)						7,250	NRDL Jul-17	3,610	3,610
SUTENT®	RCC, GIST, pNET	2007	Sales (US\$ million) [1]	9	33	41	21	26	29	27	24	29
<i>(sunitinib)</i> Pfizer			List Price (US\$/mo.) [4]							5,544	NRDL Oct-18	2,498
INLYTA®	2L adv. RCC	2015	Sales (US\$ million) <sup>[1]</sup>					3	12	16	13	20
<i>(axitinib)</i> Pfizer			List Price (US\$/mo.)							5,957	NRDL Oct-18	1,787
<b>VOTRIENT®</b>	RCC	2017	Sales (US\$ million) [1]							5	12	17
<i>(pazopanib)</i> Novartis			List Price (US\$/mo.)							7,891	NRDL Oct-18	2,348
AITAN®	3L Gastric can.	Dec 2014	Sales (US\$ million) <sup>[2]</sup>					∽45	∽126	219	258	<b>~180</b>
<i>(apatinib)</i> Hengrui			List Price (US\$/mo.)						2,870	NRDL Jul-17	1,810	1,810
FOCUSV®	<b>3L NSCLC</b>	June 2018	Sales (US\$ million) [3]								∽1 <b>9</b> 0	~268
<i>(anlotinib)</i> Sino Biopharn	ı		List Price (US\$/mo.)								NRDL Oct-18	981

#### Elunate<sup>®</sup> first 9 mo. sales progressing... relative to all MNC VEGFRi China launch sales<sup>[5]</sup>

[1] Frost & Sullivan; [2] Hengrui 2018 Annual report, dated on Feb 26, 2019; Goldman Sachs Research 2018 & GuoSen Securities research report July 20, 2017; [3] Sinobiopharm 2018 annual report/Citi Research May 2019. Note: Calculations assume at CER, using exchange rate of RMB6.74 per US\$1; [4] Sutent<sup>®</sup> price avg. pre-NRDL \$5,544 RCC/GIST, \$4,455 pNET; post-NRDL \$2,498/\$2,007; [5] MNC = multinationals, Nexavar<sup>®</sup> Yr.1 sales \$18.6m (2007) & Sutent<sup>®</sup> Yr.1 sales \$7.4m (2008).

# Fruquintinib & surufatinib both unique VEGFR TKIs ...potentially ideal VEGFR combo partners for immunotherapy



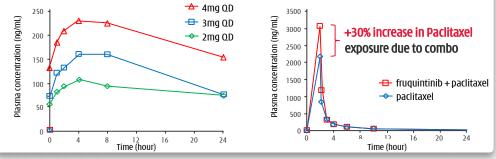
TKI	1 <sup>st</sup> Generation			2 <sup>nd</sup> Generation			Next Gei	neration
Selectivity	Multiple targets			Relatively selective			Highly selective	Selective anglo- immuno kinase inhibitor
Inhibitors	Sutent®	Nexavar®	Focus V <sup>®</sup>	Fotivda <sup>®</sup>	Lenvima®	Inlyta <sup>®</sup>	Fruquintinib	Surufatinib
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 (IC₅o < 100nM)	PDGFR <sub>α</sub> PDGFR <sub>β</sub> c-Kit Flt3 Ret CSF-1R	Raf-1 b-raf Flt3 P38 c-Kit Ret	PDGFR <sub>α</sub> PDGFRβ FGFR1-4 c-Kit	PDGFR <sub>α</sub> PDGFRβ EphB2 c-Kit Tie2	PDGFR <sub>α</sub> PDGFRβ FGFR1-4 Ret c-Kit	PDGFR <sub>α</sub> PDGFR <sub>β</sub> 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<sup>[1]</sup> production – amplifying PD-1 induced immune response

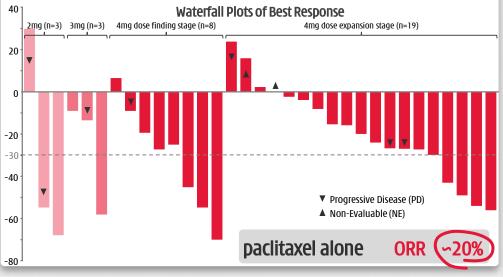
# **FRUTIGA – Gastric combo with paclitaxel** Phase III initiated Oct 2017 – 2<sup>nd</sup> interim analysis est. mid-2020



1. Dose proportional increase of fruquintinib AUC at steady state. Over 30% increase in paclitaxel drug exposure (mean AUC<sub>0-8</sub>) following multiple dose fruquintinib.



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



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).

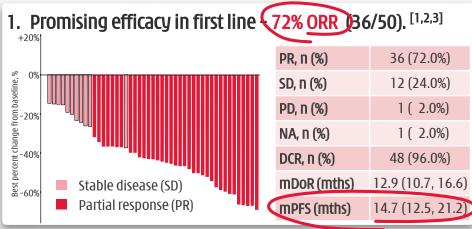
Characteristics (Unit)	Drug Expansion Stage (N=19) Fruquintinib 4 mg + paclitaxel 80 mg/m				
	Drug interruption	Drug reduction			
Dose modification with Fruquintinib N (%)	2 (10.5%)	2 (10.5%)			
Dose modification with Paclitaxel N (%)	5 (26.3%)	1 (5.3%)			

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%)			

### **Fruquintinib – 1L NSCLC combo w/** IRESSA<sup>®</sup> gefitinib Two small molecule TKIs allow for better management of tox.



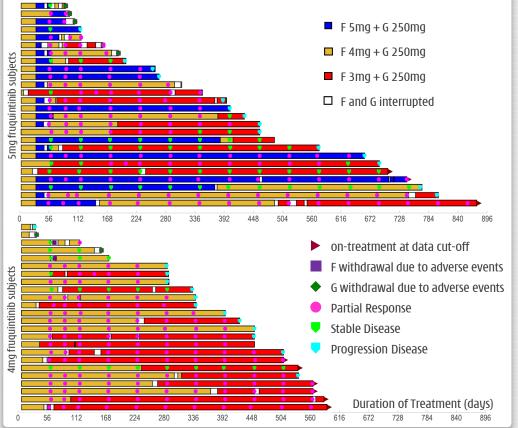


Data as of June 28, 2019.

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

Adverse Events ("AEs")	Iressa® or Tarceva® FLAURA <sup>[5]</sup> N = 277, n (%)	Avastin® + Tarceva®[6] N = 75, n (%)	5mg Fruq. + Iressa® N = 26, n (%) <sup>[3]</sup>	4mg Fruq. + Iressa® N = 24, n (%) <sup>[3]</sup>
All AEs, any grade	273 (98%)	≥74 (≥99%)	26 (100%)	24 (100%)
All AEs, Grade ≥3	124 (45%)	68 (91%)	17 (65%)	11 (46%)
AEs leading to death	6 (2%)	0 (0%)	3 (12%)	0 (0%)
AEs to VEGFRi disc.	NA	31 (41%)	6 (23%)	4 (16%)
Grade ≥3 AEs:				
Liver function	33 (12%)	6 (8%)	13 (50%)	3 (13%)
Hypertension	NA	45 (60%)	1 (4%)	1 (4%)
Proteinuria	NA	6 (8%)	3 (12%)	1 (4%)
Rash	13 (5%)	19 (25%)	0 (0%)	1 (4%)
Decreased appetite	22 (8%)	1 (1%)	NA	NA

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



[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, "Phase II Study of Fruquintinib plus Gefitinib in Stage IIIb/IV NSCLC Patients Harboring EGFR Activating Mutations", #4780 ESMO Asia, Singapore, November 23, 2019; [4] Drug discontinuation due to Grade 3 proteinuria and Grade 3 QTC prolonged; [5] Ramalingam S. et al, "LBA2\_PR Osimertinib vs standard of care (SoC) EGFR-TKI as first-line therapy in patients (pts) with EGFRm advanced NSCLC: FLAURA", ESMO 2017 Congress, Madrid, Spain, September 9, 2017; [6] Seto, T., et al, "erlotinib alone or with bevacizumab as first-line therapy in patients with advanced non-squamous non-small-cell lung cancer harbouring EGFR mutations (J025567); an open-label, randomised, multicenter, phase 2 study", The Lancet 2014, 15 (11) 1236-1244.

### FALUCA – Third-line NSCLC Monotherapy Presented at WCLC 2019



FALUCA Phase III (enrolled Dec. 2015 to Feb. 2018)

- Met all secondary endpoints: mPFS; ORR; DCR; & DOR <sup>[1]</sup>;
- Did not achieve primary endpoint of median OS, however:
  - Anti-tumor therapies after disease progression reduced OS diff.
  - Higher percentage of placebo pts received subsequent treatments.

Significant difference in subsequent anti-tumor treatments (ATT)

- **Chemotherapy:** Fruq. 29.7% vs. Placebo 53.8%
- Targeted therapies (VEGFi and/or EGFRi): Fruq. 20.9% vs. Placebo 31.2%
- Tagrisso® & anlotinib just approved in 2017

Efficacy Endpoints (Intent-to-Treat)<sup>[2]</sup>

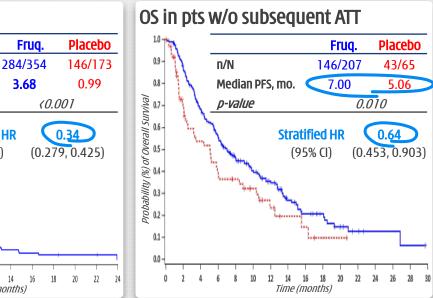
	Fruq. (N=354)	Placebo (N=173)	p-value
mOS (mths)	8.94	10.38	0.841
mPFS (mths)	3.68	0.99	(0.001
ORR	13.8% (49)	0.6% (1)	(0.001
DCR	<b>66.7%</b> (236)	24.9% (43)	(0.001

Stratified HR

(95% CI)

### Good safety; most Grade $\geq$ 3 TEAEs target-related & clinically manageable.

laryet-related &	LIIIICally III	allayeavie.
Patient (%)	Fruq. (N=354)	Pbo (N=173)
TEAE $\geq$ Grade 3	216 (61.2%)	47 (27.6%)
Leading to discontinuation	37 (10.5%)	9 (5.3%)
Leading to interruption	61 (17.3%)	7 (4.1%)
Leading to dose reduction	85 (24.1%)	2 (1.2%)
Hypertension	74 (21.0%)	5 (2.9%)
Hand-foot syndrome	39 (11.0%)	0



[1] mOS = median Overall Survival; mPFS = median Progression-Free Survival; ORR = Objective Response Rate; DCR = Disease Control Rate; DOR = Duration of Response; HR = hazard ratio; 95% CI = 95% Confidence Interval; [2] Lu, et al. "A Randomized Phase III trial of Fruquintinib versus Placebo in Patients with Advanced Non-Small Cell Lung Cancer (FALUCA)." WCLC 2019 Abstract #MA14.05; [3] Lu, et al. Randomized, Double-Blind, Placebo-Controlled, Multicenter Phase II Study of Fruquintinib After Two Prior Chemotherapy Regimens in Chinese Patients With Advanced Non-Squamous Non–Small-Cell Lung Cancer. Journal of Clinical Oncology 36, no. 12 (April 20 2018) 1207-1217. DOI: 10.1200/JCO.2017.76.7145; [4] Li, et al. Effect of Fruquintinib vs Placebo on Overall Survival in Patients With Previously Treated Metastatic Colorectal Cancer: The FRESCO Randomized Clinical Trial. JAMA. 2018 Jun 26;319(24):2486-2496. doi: 10.1001/jama.2018.7855. \* Post-hoc analysis.

**PFS in ITT population** 

n/N

**D-Value** 

Median PFS, mo.

0.9 -

0.8

0.7 -

0.5 -

0.4 -

0.3 -

0.2 -

Probability (%) of Progression-Free Survival



## 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:
- C Freedom to operate in selecting & pursuing any future indications in China;
- Solution A series a series of the series of
- Section 2017 Freedom to collaborate with any third-party in clinical development; and
- Source Possible promotion rights in 30-40% of China for Elunate<sup>®</sup>.<sup>[2]</sup> 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 <sup>[1]</sup> Development Costs – Paid by Lilly LCI Development Costs – Paid by Chi-Med	70% 30%	0%
LCI Regulatory Approval Milestones – Paid to Chi-Med <sup>[3]</sup>	12.5	20.0
Royalty Payments – Paid to Chi-Med [4]	15 - 20%	15 - 29%
<b>Co-Promotion Rights in China</b> (% of provinces) <b>Co-Promotion Service Fees</b> – paid to Chi-Med (% Net Sales)	0% 0%	<b>30 - 40%</b> Not disclosed

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

[1] LCI = Life Cycle Indication; [2] upon achievement of a non-fruquintinib related Eli Lilly commercial action; [3] Lifecycle Indication - China - per LCI, up to 3 LCIs; [4] On Total Molecule Sales in China triggered upon launch of 1st LCI







HMPL-523 (Syk) in hematological cancer Phase I/Ib ongoing in Australia, China, US & EU



- Extensive Ph.I dose escalation study now complete in Australia & China (total n>60);
- RP2D<sup>[1]</sup> determined & large Ph. lb dose expansion study, total n>200, underway in ~30 active sites in Australia & China;
- U.S./E.U. Phase I/Ib enrollment underway, with 12 sites enrolling;
- These Phase I/Ib data will inform China registration study decisions in 2020.

#### **Complete** Stage I: dose escalation "3 + 3" each dose cohort until disease • Australia: Relapsed/refractory Studied HMPL-523 N = 38 progression. hematologic malignancy 100-1,000mq QD & death. • China: Relapsed/refractory mature B 200-400mq BID intolerable N = 27 lymphoma toxicity, etc. Stage II: dose expansion ...Now enrolling Relapsed or refractory, measurable disease – multiple arms: until disease Aus Chronic lymphocytic leukemia (CLL) progression, N = 40 Small lymphocytic lymphoma (SLL) 600mg QD death, Mantle cell lymphoma (MCL) China intolerable Follicular lymphoma (FL) N = 190toxicity, etc. • Marginal zone lymphoma (MZL) DLBCL (in China) & WM/LPL

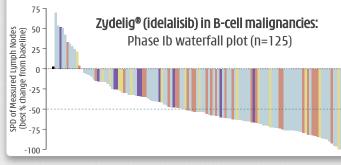
### Australia & China Phase I/Ib studies

# HMPL-689 – Phase I/Ib ongoing in China, US & EU Designed to be a best-in-class inhibitor of PI3K $\delta$



#### 1. PI3K $\delta$ now a proven target.

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



#### 2. PI3K $\delta$ inhibitors being developed in a very broad range of indications.

Compound		Indication	Status	Issue
Zydelig <sup>®</sup> (idelalisib) PI3K&	Gilead	Chronic lymphocytic leukaemia, non-Hodgkin's lymphoma	Marketed	<b>High incidence of liver toxicity</b> seen with idelalisib (150mg bid)
AMG-319 PI3Kδ	Amgen	B-cell lymphoma, non-Hodgkin's lymphoma, T-cell lymphoma, chronic lymphocytic leukaemia	Phase I Trial	
Copiktra®	Verastem/	Relapsed or refractory chronic lymphocytic leukaemia / small lymphocytic lymphoma	Approved	<b>Need to spare PI3Ky</b> serious infection seen &
(duvelisib) PI3Kγ/δ		Relapsed or refractory follicular lymphoma	Approved <sup>[2]</sup>	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 <sup>[2]</sup>	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<sup>®</sup>, Copiktra<sup>®</sup> & Aliqopa<sup>®</sup>.

Enzyme IC <sub>50</sub> (nM)	HMPL-689	Zydelig®	<b>Copiktra<sup>®</sup></b>	Aliqopa®
ΡΙ3Κδ	0.8 (n = 3)	2	1	0.7
PI3K $\gamma$ (fold vs. PI3K $\delta$ )	114 <b>(142x)</b>	104 <mark>(52x)</mark>	2 (2X)	6.4 <b>(9x)</b>
PI3K <sub>α</sub> (fold vs. PI3Kδ)	>1,000 <b>(&gt;1,250x)</b>	866 <mark>(433x)</mark>	143 (143x)	0.5 (1X)
PI3Kδ human <u>whole blood</u> CD63+	3	14	15	n/a
ΡΙ3Κβ (fold vs. ΡΙ3Κδ)	87 <mark>(109x)</mark>	293 <b>(147x)</b>	8 (8X)	3.7 <b>(5x)</b>

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







### HMPL-453 (FGFR)

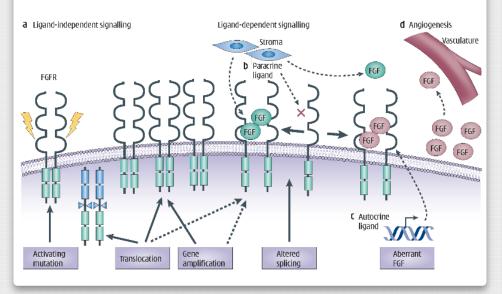
Aim to establish proof-of-concept

## HMPL-453 – Phase II in China set to initiate 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.

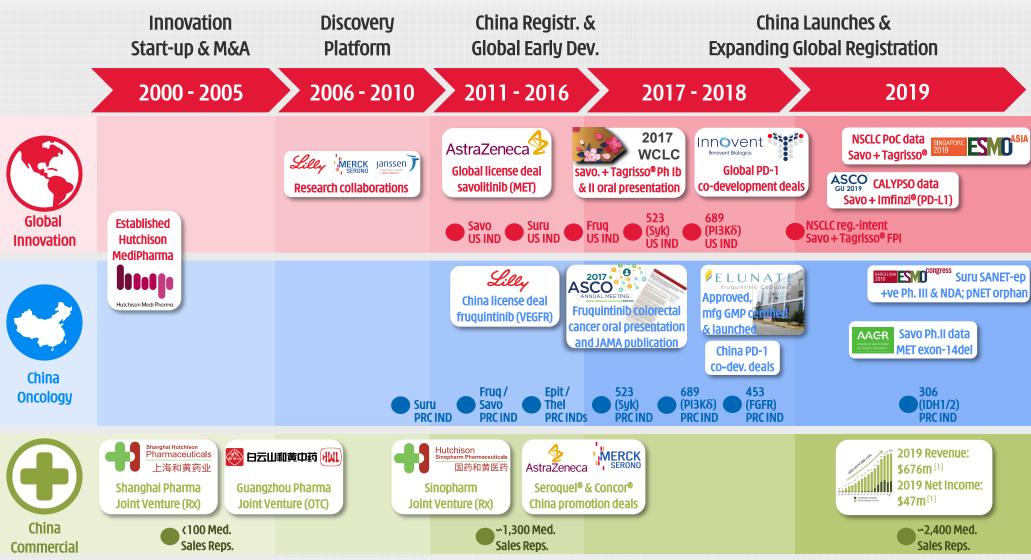
	Gene amplification	Gene translocation	Gene mutation
FGFR1	Lung squamous (7~15%) H&N squamous (10~17%) Esophageal squamous (9%) Breast (10~15%)	Lung squamous (n/a) Glioblastoma (n/a) Myeloproliferative syndrome (n/a) Breast (n/a)	Gastric (4%) Pilocytic astrocytoma (5~8%)
FGFR2	Gastric (5∽10%) Breast (4%)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



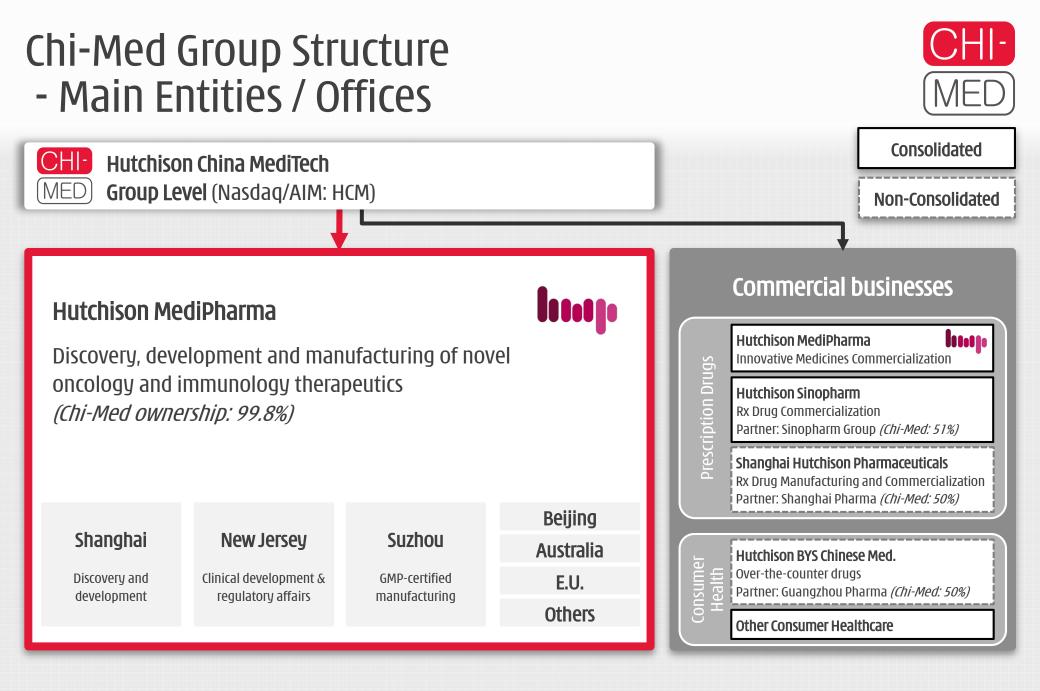




# Important milestones in Chi-Med's evolution



[1] Based on aggregate Non-GAAP revenues and net income attributable to Chi-Med of consolidated subsidiaries and non-consolidated joint ventures of Commercial Platform.





# 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 \$1.9-2.0 billion.<sup>[1]</sup>
- Given our share in the JVs, Chi-Med's share of this value is approximately \$0.9-1.0 billion.

			NET SALES			NET IN	NCOME		VALUATION	3]
	Code	2018 Jan-Jun	2019 Jan-Jun	18-19 1H Growth	2018 Jan-Jun	2019 Jan-Jun	18-19 1H Growth	2019 1H Margin	Market Cap.	P/E
CHI-MED Commercial Platform Subsidiaries/JVs <sup>[2]</sup>		360.3	367.1	2%	55.1	57.0	3%	16%	n/a	n/a
Li Zhu Pharma	000513	652.1	705.6	8%	96.5	119.2	24%	17%	4,017	23
Shandong Dong E E Jiao Kunming Pharma	000423 600422	426.6 483.5	270.0 536.6	-37% 11%	123.4 26.2	27.5 34.4	-78% 32%	10% 6%	3,157 1,110	21 18
Zhejiang Kang En Bai Pharma	600572	510.9	521.4	2%	78.6	60.0	-24%	12%	2,382	32
Tianjin Zhong Xin Pharma Zhangzhou Pien Tze Huang	600329 600436	444.6 343.4	504.8 413.5	14% 20%	45.0 86.7	50.6 108.1	12% 25%	10% 26%	1,463 8.602	17 45
Jiangsu Kang Yuan	600557	263.5	323.2	23%	29.2	35.1	20%	11%	1,205	17
Zhuzhou Qian Jin Pharma Jiu Zhi Tang	600479 000989	212.7 257.2	241.7 241.2	14% -6%	11.5 46.5	14.8 25.0	29% -46%	6% 10%	509 995	13 37
Wuhan Jian Min Pharma	600989	153.5	158.9	3%	8.3	8.1	-2%	5%	368	32
Peer Group Median (10 Comps. excl. Chi-Med)		385.0	368.4	-4%	45.8	34.8	24%	9%	1,334	22
All 61 Listed thina Pharma. Companies Median		263.5	264.7	0%	29.9	27.5	8%	10%	1,065	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 2019E Net Sales in the ~\$400-1,400 million range.

Source: Company data, Deutsche Bank, FactSet

(US\$ millions)

[1] Peer group/China Pharma multiple of 21x-22X 2019 actual Net income after tax of \$90.8 million; [2] Total aggregate PRC domestic results of Chi-Med's 6 Commercial Platform companies (HBYS, SHPL, Hutchison Sinopharm, HHO, HHL & HCPL); [3] Market Capitalization and Price Earnings Ratios as at **December 4, 2019**: Trailing Twelve Month PE weighted averaged based on market capitalization.

# Non-GAAP Financial Measures and Reconciliation (1/3)



Reconciliation of Adjusted Group net cash flow activities:	rs excluding f	Reconciliation of Adjusted Innovation Platf operating loss:	orm segm	ent	
	2019 Guidance	2019 Actual		2018	2019
Cash and cash equivalents and short-term investments at end of year	180-210	217.2	Segment operating loss – Innovation Platform	(104.6)	(133.3)
Less: Cash and cash equivalents and short-term investments at beginning of year	(300)	(301.0)	Less: Segment revenue from external customers – Innovation Platform	(37.6)	(16.0)
Add: Net cash used in financing activities for the year		1.5			
Adjusted Group net cash flows excluding financing activities	(90) - (120)	(82.3)	Adjusted Innovation Platform segment operating loss	(142.2)	(149.3)

(US\$ millions unless otherwise stated)

# Non-GAAP Financial Measures and Reconciliation (2/3)



#### Reconciliation of GAAP sales and net income/(loss) attributable to Chi-Med to CER:

	Year E	nded	Cha	nge Amo	ount	Cl	hange %		
\$'Million (except %)	December 31, 2019	December 31, 2018	Actual	CER	Exchange effect	Actual	CER	Exchange effect	
Consolidated sales - Group	204.9	214.1	(9.2)	(2.0)	(7.2)	-4%	-1%	-3%	
Consolidated sales - Commercial Platform	188.9	176.5	12.4	19.2	(6.8)	7%	11%	-4%	
— Prescription Drugs	154.5	136.4	18.1	24.4	(6.3)	13%	18%	-5%	
— Consumer Health	34.4	40.1	(5.7)	(5.2)	(0.5)	-14%	-13%	-1%	
Non-consolidated joint venture sales	487.5	491.5	(4.0)	17.2	(21.2)	-1%	3%	-4%	
- SHPL	272.1	275.7	(3.6)	7.9	(11.5)	-1%	3%	-4%	
– HBYS	215.4	215.8	(0.4)	9.3	(9.7)	0%	4%	-4%	
Total Sales - Commercial Platform (Non-GAAP)	676.4	668.0	8.4	36.4	(28.0)	1%	5%	-4%	
Consolidated net loss attributable to Chi-Med	(106.0)	(74.8)	(31.2)	(34.7)	3.5	-42%	-46%	4%	
Innovation Platform	(133.2)	(104.4)	(28.8)	(34.2)	5.4	-28%	-33%	5%	
Commercial Platform	47.4	43.4	4.0	5.9	(1.9)	9%	13%	-4%	
— Prescription Drugs	37.5	34.1	3.4	4.7	(1.3)	10%	14%	-4%	
– Consumer Health	9.9	9.3	0.6	1.2	(0.6)	7%	12%	-5%	

# Non-GAAP Financial Measures and Reconciliation (3/3)



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

Prescription Drugs: includes our Consolidated subsidiaries (Hutchison Sinopharm and HMPL) 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				18-19
(US\$ millions)	03	04	05	06	07	08	09	10	11	12	13	14	15	16	17	18	19	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	668.0 [5]	676.4 [5]	] 1%
Prescription Drugs	17.2	21.8	23.3	23.2	28.1	<i>39.5</i>	54.4	71.2	92.4	116.5	138.2	204.9	286.6	372.3	411.0	412.1	426.6	4%
- Consolidated subsidiary	-	-	-	-	-	-	-	-	-	-	-	50.2	105.5	149.9	166.4	136.4	154.5	13%
- 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	272.1	-1%
Consumer Health	4.7	6.1	41.8	<i>78.2</i>	90.9	116.3	142.6	165.2	186.2	244.2	264.1	260.5	232.3	255.1	266.2	255.9	249.8	-2%
- Consolidated s ubsidiaries	4.7	6.1	<i>9.3</i>	<i>8.9</i>	3.7	5.5	7.0	14.1	14.9	15.5	16.5	16.8	20.7	31.0	38.8	40.1	34.4	-14%
- Non-consolidated joint venture	-	-	32.5	<i>69.3</i>	87.2	110.8	135.6	151.1	171.3	228.7	247.6	243.7	211.6	224.1	227.4	215.8	215.4	0%
Total Sales Growth	n/a	27%	133%	56%	17%	31%	26%	20%	18%	29%	n/a	16%	11%	21%	8%	-1%	1%	
- GuanBao divested in Sept'2017	-	-	-	-	-	-	-	-	(11.4)	(50.5)	(51.6)	(49.7)	(40.7)	(45.0)	(38.6)	-	-	n/a
Adjusted Consumer Health	4.7	6.1	41.8	<i>78.2</i>	90.9	116.3	142.6	165.2	174.8	193.7	212.5	210.8	191.6	210.1	227.6	255.9	249.8	-2%
- Adjusted Non-consolidated joint venture	0.0	-	32.5	<i>69.3</i>	87.2	110.8	135.6	151.1	159.9	178.2	196.0	1 <i>94.0</i>	170.9	179.1	188.8	215.8	215.4	0%
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	<i>668.0</i> <sup>[5]</sup>	676.4 <sup>[5]</sup>	<sup>1</sup> 1%
Total Adjusted Sales Growth	n/a	27%	133%	56%	17%	31%	26%	20%	13%	16%	13%	19%	15%	22%	10%	5%	1%	
Net (loss)/income after tax (Non-GAAP)	(10.7)	(3.6)	2.2	6.7	11.2	14.7	21.5	27.9	30.1	33.1	39.7	48.8	54.1	63.3 <sup>[3]</sup>	77.3 <sup>[4]</sup>	85.6 <sup>[5]</sup>	90.8 [5]	] 6%
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	65.9	<i>69.3</i>	5%
- Consolidated subsidiary	-	-	-	-	-	-	-	-	-	-	-	0.1	0.6	1.6	2.4	6.1	8.0	30%
- 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>	61.3	3%
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	19.7	21.5	9%
- Consolidated subsidiaries	(10.3)	(4.9)	<i>(2.9)</i>	(2.4)	0.2	-	0.8	1.0	(0.4)	(1.1)	0.1	1.5	0.8	1.5	3.5	2.8	1.7	-38%
- 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	1 <i>9.8</i>	17%
% 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%	12.8%	1 <i>3.4%</i>	
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 <sup>[2]</sup>	22.8 [2]	25.2 <sup>[2]</sup>	<b>29.9</b> <sup>[3]</sup>	37.5 [4]	<b>43.4</b> <sup>[5]</sup>	47.4 [5]	9% I
Prescription Drugs	(0.2)	0.6	1.0	0.7	0.9	1.4	3.0	5.9	7.1	8.8	11.2	13.2	<i>15.9</i>	20.7	26.5	34.1	37.5	10%
Consumer Health	(5.5)	(4.3)	(1.5)	0.5	3.6	4.5	6.3	6.7	6.5	5.8	7.0	9.6	<i>9.3</i>	<i>9.2</i>	11.0	<i>9.3</i>	<i>9.9</i>	7%
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%	16%	9%	

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 \$2.5 million from net income attributable to Chi-Med for 2017;
 5 In 2019, the results of innovative medicines developed by the Innovation Platform to Commercial Platform- Prescription Drugs business. 2018 information has been revised for comparison purpose.

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



		U	Init Pricing (US\$	) [3]		Approximate Mor	thly Pricing (U	<b> \$\$)</b> <sup>[3]</sup>	
Brand (generic)	Company	Dosage	Avg. Tender	Reimbursed	$\Delta$ %	Dosage	Avg. Tender	Reimbursed	Indication coverage
Herceptin® (trastuzumab)	Roche	440mg:20ml	\$3,298.81	\$1,125.93	-66%	Breast: 4mg/kg wk 1, 2mg/kg weekly. <sup>[2]</sup>	\$4,500	\$1,540	Breast: Her2+; Her2+ meta. Her2+ late-stage meta. gastric.
Avastin® (bevacizumab)	Roche	100mg:4ml	\$772.74	\$296.00	-62%	10mg/kg 0.2W.	\$11,590	\$4,440	Late-stage meta. CRC or advanced non-squamous NSCLC.
TheraCIM <sup>®[4]</sup> (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 <sup>[2]</sup>	\$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 <sup>[2]</sup>	\$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 <sup>[2]</sup>	\$47.85	\$30.22	-37%	850mg QD.	\$2,870	\$1,810	3L gastric adenocarcinoma or esophageal junction with adenocarcinoma.
Velcade® (bortezomib)	L&L	3.5mg <sup>[2]</sup>	\$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)	L&L	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 <sup>[2]</sup>	\$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 <sup>[2]</sup>	\$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			Approximate Monthly F	Pricing (US\$) <sup>[2]</sup>		
Brand (generic)	Company	Dosage		Reimbursed	$\Delta$ %	Dosage <sup>[1]</sup>	Avg. Tender	Reimbursed	Indication coverage
Focus V <sup>®</sup> (anlotinib)	Sino Biopharr	n 12mg	\$127	\$70	-45%	12mg QD (2 wks-on/1-wk-off)	\$1,783	\$981	3L NSCLC
Oncaspar <sup>®</sup> (pegaspargase)	Hengrui	5ml:3750 IU	\$560	\$429	-23%	$\leq$ 2ml every 14 days	\$1,231	\$943	1L ALL
Vidaza <sup>®</sup> (azacitidine)	Celgene	100mg	\$378	\$152	-60%	1 <sup>st</sup> 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 <sup>®</sup> (axitinib)	Pfizer	5mg	\$99	\$30	-70%	5mg BID	\$5,957	\$1,787	2L Advanced renal cell carcinoma
Tagrisso <sup>®</sup> (osimertinib)	AstraZeneca	80mg	\$253	\$73	-71%	80mg QD	\$7,597	\$2,201	EGFR TKI refractory T790M+ NSCLC
Ninlaro <sup>®</sup> (ixazomib)	Takeda	4mg	\$3,234	\$710	-78%	4mg on Days 1, 8, 15 (28 day cycle)	\$12,934	\$2,839	2L Multiple myeloma
Xalkori <sup>®</sup> (crizotinib)	Pfizer	250mg	\$123	\$37	-70%	250mg BID	\$7,407	\$2,245	Locally adv. or meta. ALK+ or ROS1+ NSCLC
Gilotrif <sup>®</sup> (afatinib)	Boehringer	40mg	\$116	\$29	-75%	40mg QD	\$3,483	\$863	NSCLC with EGFR
Tasigna <sup>®</sup> (nilotinib)	Novartis	200mg	\$39	\$14	-65%	400mg BID	\$4,645	\$1,635	CML
Votrient <sup>®</sup> (pazopanib)	Novartis	200mg	\$66	\$23	-65%	800mg QD	\$7,891	\$2,348	RCC
Sutent <sup>®</sup> (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 <sup>®</sup> (regorafenib)	Bayer	40mg	\$52	\$28	-46%	160mg QD, 3-wks-on/1-wk-off *	\$4,368	\$2,352	Meta. CRC, GIST, HCC
Zykadia <sup>®</sup> (certinib)	Novartis	150mg	\$108	\$28	-74%	450mg QD	\$9,699	\$2,564	NSCLC
Zelboraf <sup>®</sup> (vemurafenib)	Roche	240mg	\$30	\$16	-47%	960mg BID	\$7,252	\$2,369	Melanoma
Erbitux <sup>®</sup> (cetuximab)	Merck	100mg	\$571	\$186	-67%	400mg/m2 initial dose, 250mg weekly	\$10,446	\$3,074	Colorectal cancer, head and neck cancer
Sandostatin LAR <sup>®</sup> (octreotide)	Novartis	20mg	\$1,169	\$835	-29%	20mg Q4W	\$1,169	\$835	GEP-NENs
Imbruvica <sup>®</sup> (ibrutinib)	INI	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.

\* Price amended to account for 3-weeks on, 1 week off regimen.

## National Reimbursement Drug List Pricing ("NRDL") Nov'19 update - 8 new & 9 renewed drugs in oncology<sup>[1]</sup>



			Unit Pricing (	US\$) <sup>[3]</sup>		Approximate Mo	nthly Pricing (l	JS\$) <sup>[3]</sup>	
Brand (generic)	Company	Dosage	Avg. Tender	Reimbursed	$\Delta$ %	Dosage	Avg. Tender	Reimbursed	Indication coverage
Elunate <sup>®</sup> (fruquintinib)	Chi-Med	5mg	\$149	\$53.77 -	64%	5mg QD 3wks/1wk-off.	\$3,350	\$1,210	Metastatic colorectal cancer, 3L
Tyvyt <sup>®</sup> (sintilimab)	Innovent	10ml	\$1,114	\$404.41 -	64%				Classical Hodgkin's lymphoma, 3L
Saiweijian <sup>®</sup> (raltitrexed)	Sino Biopharm	2mg	\$234	\$95.16 -	59%				colorectal cancer, 5-FU intolerable
Alecensa <sup>®</sup> (alectinib)	Roche			Undisclosed					NSCLC, ALK+
Lynparza <sup>®</sup> (olaparib)	AstraZeneca			Undisclosed					Epithelial ovarian, fallopian tube, or peritoneal cancer
Airuini <sup>®</sup> (pyrotinib)	Hengrui			Undisclosed					Breast cancer, HER2+, 2L
Perjeta <sup>®</sup> (pertuzumab)	Roche			Undisclosed					Breast cancer, HER2+, neoadjuvant
Jakafi <sup>®</sup> (ruxolitinib)	Incyte / Novartis			Undisclosed					PMF, PPV-MF, PET-MF

			Unit Pricing	(US\$) <sup>[3]</sup>		Approximate Monthly Pr	ricing (US\$)	[3]	
Brand (generic)	Company	Dosage	'17 NRDL	'19 NRDL	$\Delta\%$	Dosage	'17 NRDL	'19 NRDL	Indication coverage
AiTan® (apatinib)	Hengrui	425mg <sup>[2]</sup>	\$29.03	\$24.56	-15%	850mg QD.	\$1,740	\$1,470	3L gastric adenocarcinoma or GEJ with adenocarcinoma.
EnDu® (rh-endostatin)	Simcere	15mg	\$89.62	\$69.70	-22%	7.5mg/m² iv QD 2wks/1wk-off.	\$1,430	\$1,120	Late-stage NSCLC.
Epidaza® (chidamide)	Chipscreen	5mg	\$54.77	\$48.79	-11%	30mg QD, 2x per wk.	\$2,820	\$2,510	2L+ Recurring or refractory peripheral T-cell lymph. (PTCL).
Herceptin® (trastuzumab)	Roche			Undisclosed					Breast: Her2+; Her2+ meta. Her2+ late-stage meta. gastric.
Avastin® (bevacizumab)	Roche			Undisclosed					Late-stage meta. CRC or advanced non-squamous NSCLC.
TheraCIM <sup>®[4]</sup> (nimotuzumab)	Biotech			Undisclosed					Combo with RT for EGFR+ III/IV nasopharyngeal carcinoma.
Tarceva® (erlotinib)	Roche			Undisclosed					Advanced NSCLC with limited EGFR gene mutation.
Nexavar® (sorafenib)	Bayer			Undisclosed					RCC or HCC. meta. diff. thyroid after radio-iodine therapy.
Afinitor <sup>®</sup> (everolimus)	Novartis			Undisclosed					RCC after sunitinib or sorafenib. Pancreatic NETs. TSRA.

Source: National Healthcare Security Administration (NHSA); Goldman Sachs equity research.

[1] Excluding botanical oncology drugs; [2] Reference SKU or reference recommended dosage for monthly pricing calculation; [3] Calculation assumes an exchange rate of CN¥7.03 per US\$1; [4] Marketed as Tai Xin Sheng® in China.





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Thank you