## A YEAR OF CLINICAL, REGULATORY & COMMERCIAL PROGRESS

**FY2020 RESULTS & BUSINESS UPDATES** 

March 4, 2021

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.



## **Evolving our corporate identity**

Stock ticker to remain unchanged – Nasdaq/LSE AIM: HCM

Past group Identity:



**HUTCHISON CHINA MEDITECH** 

Past oncology/Immunology R&D operations identity:



HUTCHISON CHINA MEDITECH
Hutchison Medi Pharma

Group & all subsidiaries identity from now onwards:



## 1. OVERVIEW

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



Realizing the global potential of HUTCHMED's novel oncology assets



Building a fully integrated oncology business in China

## **Our Strengths**



Fully integrated 1,200 person R&D and commercialization platform built over 20 years

1

#### WORLD CLASS DISCOVERY & DEVELOPMENT CAPABILITY

**First** global-focused novel drug discovery company in China – established in the early 2000s

**600+** integrated R&D staff focused on oncology & immunological diseases >

2

## HIGHLY DIFFERENTIATED NME PORTFOLIO & GLOBAL PIPELINE

**10 innovative clinical NMEs** – all discovered in-house by HUTCHMED

**3 lead assets NDA filed/ approved** in China – all in late global development

3

#### DEEP PAN-CHINA MARKET ACCESS CAPABILITY

**420+** person oncology team – covering 2,100+ cancer centers in China

**Highly profitable** Other Ventures with 20 year commercial track-record in China

4

#### SEASONED MNC MGMT. TEAM – STRONG GOVERNANCE

**11 years** – median tenure of 14 person senior mgmt. team

**0 governance issues** during 14 years as a listed company

## **Differentiated portfolio**

## HUTCHMED

2 HIGHLY DIFFERENTIATED NME PORTFOLIO AND GLOBAL PIPELINE

#### All discovered in-house & designed for global differentiation

PRODUCT	MOA	DISCOVERY <sup>[1]</sup>	INDICATIONS	PARTNER	RIGHTS	CHINA <sup>[2]</sup>	GLOBAL <sup>[2]</sup>
Surufatinib (SULANDA®)	VEGFR 1/2/3, FGFR1 & CSF-1R	In-house (est. LOE ~2035)	Neuroendocrine tumors (NET), biliary tract, thyroid, solid tumors (multiple I/O combos)	None	HCM holds all WW rights	Marketed (non-pNET) NDA accepted (pNET)	<b>US NDA filing</b> started YE20 & <b>EU MAA planned</b> in 2021
Fruquintinib (ELUNATE®)	VEGFR 1/2/3	In-house (est. LOE ~2033)	Colorectal, gastric, NSCLC, solid tumors (multiple I/O & TKI combos)	Lilly	HCM has WW rights ex- China; 70%-80% of sales in China <sup>[4]</sup>	Marketed (Colorectal); Ph.III (Gastric)	<b>Ph.III US, EU, Japan</b> (Colorectal)
Savolitinib	c-MET	In-house (est. LOE ~2035)	NSCLC, kidney, gastric <sup>[3]</sup> , colorectal <sup>[3]</sup> (multiple I/O & TKI combos)	8	AZ has WW rights; China (30% royalty); ex-China (9- 18% tiered royalty)	NDA accepted (NSCLC mono) Ph.III (GC*, NSCLC combo*)	Ph.II/III global (multiple NSCLC) Ph.III global (PRCC*)
HMPL-689	РІЗКδ	In-house (est. LOE ~2040)	B-cell malignancies – indolent NHL	None	HCM holds all WW rights	Ph.lb/II (Treated >100 NHL pts.)	Ph.I US, EU, Aus (NHL)
HMPL-523	Syk	In-house (est. LOE ~2037)	ITP, B-cell malignancies – indolent non-Hodgkin's lymphoma (NHL)	None	HCM holds all WW rights	<b>Ph.Ib/II</b> (Treated >200 NHL pts.)	Ph.I US, EU, Aus (NHL)
HMPL-453	FGFR 1/2/3	In-house (est. LOE ~2039)	Cholangiocarcinoma	None	HCM holds all WW rights	Ph.II (IHCC)	-
Epitinib	EGFRm+	In-house (est. LOE ~2032)	Glioblastoma	None	HCM holds all WW rights	Ph.II (Glioblastoma)	-
HMPL-306	IDH 1/2	In-house (est. LOE ~2043)	Hematological malignancies, solid tumors	None	HCM holds all WW rights	Ph.I (Hem. malignancies)	Ph.I in planning (start H1 2021)
HMPL-295	ERK (MAPK pathway)	In-house	Solid tumors	None	HCM holds all WW rights	Ph.I planning to start in mid-2021	-
HMPL-653	Not Disc.	In-house	Solid tumors	None	HCM holds all WW rights	Target IND 2023	L (US/China)
HMPL-A83	Not Disc.	In-house	mAb – solid tumors, hematological malignancies	None	HCM holds all WW rights	Target IND 202	L (US/China)
HMPL-760	Not Disc.	In-house	Hematological malignancies	None	HCM holds all WW rights	Target IND 202	L (US/China)

<sup>\*</sup>In planning

<sup>[1]</sup> Approximate estimated Loss of Exclusivity (LOE) in key markets considering multiple patent families, extension, and regulatory protection; [2] Represents the most advanced clinical trial stage and indication; [3] Investigator initiated trials (IITs); [4] Subject to meeting pre-agreed sales targets, Lilly will pay HUTCHMED an estimated total of 70%-80% of ELUNATE® sales in the form of royalties, manufacturing costs and service payments.

## 6 assets in global development





**HIGHLY DIFFERENTIATED NME PORTFOLIO AND GLOBAL PIPELINE** 

#### Rapid expansion of our US/EU clinical & regulatory team

Program	Treatment	Indication	Target patient	Study name	Sites	Dose finding / safety run-in	Proof-of-concept	Registration
	Savolitinib + TAGRISSO®	NSCLC	2L/3L EGFRm; Tagrisso® ref.; MET+	SAVANNAH	Global	Oxnard/Ahn – DF/SMC		
	Savolitinib + IMFINZI® (PD-L1)	Papillary RCC	MET+			In planning		
Savolitinib	Savolitinib + IMFINZI® (PD-L1)	Papillary RCC *	All	CALYPSO	UK/Spain	Powles – Queen Mary's		
MET	Savolitinib + IMFINZI® (PD-L1)	Clear cell RCC *	VEGFR TKI refractory	CALYPSO	UK/Spain	Powles – Queen Mary's		
	Savolitinib	Gastric cancer *	MET+	VIKTORY	S Korea	Lee - Samsung Med. Ctr		
	Savolitinib	Colorectal cancer *	MET+		US	Strickler – Duke Uni		
	Surufatinib	NET	Refractory		US	Dasari/Yao – MD Anderson		
	Surufatinib	NET	Refractory		EU	Garcia-Carbonero – UCM		
Surufatinib VEGFR 1/2/3;	Surufatinib	Biliary tract cancer			US	Li – City of Hope		
FGFR1; CSF-1R	Surufatinib	Soft tissue sarcoma			US	Patel/Tapp – MD And/ MSKCC		
	<b>Suru.</b> + tislelizumab (PD-1)	Solid tumors			US/EU	In planning - IND cleared		
	Fruquintinib	Colorectal cancer	Refractory	FRESCO-2	US/EU/JP	Eng/Desari - MD And. [1]		
Fruguintinib	Fruquintinib	Breast cancer			US	Tripathy - MD And.		
VEGFR 1/2/3	Fruq. + tislelizumab (PD-1)	TN breast cancer			US	In planning - IND cleared		
	Fruq. + tislelizumab (PD-1)	Solid tumors			TBD	In planning - IND cleared		
HMPL-689	HMPL-689	Healthy volunteers			Australia			
РІЗКδ	HMPL-689	Indolent NHL			US/EU	Zinzani – U of Bologna		
HMPL-523	HMPL-523	Indolent NHL			Australia			
Syk	HMPL-523	Indolent NHL			US/EU	Strati/Abrisqueta – MD And. / Val´d	Hebron	
HMPL-306	HMPL-306	Solid tumors			US/EU	In planning - IND cleared		
IDH 1/2	HMPL-306	Hem. malignancies			US/EU	In planning - IND cleared		
	_							

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

## 8 assets in China development

...8-10 registration studies planned to start on 2021





2 HIGHLY DIFFERENTIATED NME PORTFOLIO AND GLOBAL PIPELINE

Program	Treatment	Indication	Target patient	Study name	Sites	Dose find / safety run-in	Proof-of-concept	Registration
	Savolitinib	NSCLC	MET Exon 14 skipping		China	Lu Shun - SH Chest Hosp.		
Savolitinib	Savolitinib + TAGRISSO®	NSCLC	2L EGFR TKI ref. NSCLC; MET+		China	In planning		
MET	Savolitinib + TAGRISSO®	NSCLC	Naïve MET+ & EGFRm NSCLC		China	In planning		
	Savolitinib	Gastric cancer	2L; MET+		China	In planning		
	Surufatinib	Pancreatic NET	All	SANET-p	China	Xu Jianming – #5 Med. Ctr.		
	Surufatinib	Non-Pancreatic NET	All	SANET-ep	China	Xu Jianming - #5 Med. Ctr.		
Surufatinib	Surufatinib	Biliary tract cancer	2L; chemotherapy refractory		China	Xu Jianming - #5 Med. Ctr.		
VEGFR 1/2/3;	Suru. + TUOYI® (PD-1)	NEN, ESCC, BTC			China	Shen Lin - BJ Univ. Tmr.		
FGFR1; CSF-1R	Suru. + TUOYI® (PD-1)	SCLC, GC, Sarcoma			China	Shen Lin - BJ Univ. Tmr.		
	Suru. + TUOYI® (PD-1)	TC, EMC, NSCLC			China	Shen Lin - BJ Univ. Tmr.		
	Suru. + TYVYT® (PD-1)	Solid tumors			China			
	Fruquintinib	Colorectal cancer	≥3L; chemotherapy refractory	FRESCO	China	Li Jin - Fudan Univ.		
	Fruq. + TAXOL®	Gastric cancer	2L	FRUTIGA	China	Xu Ruihua – Sun Yat Sen		
Fruquintinib	Fruq. + TYVYT® (PD-1)	CRC, EMC, RCC, HCC			China	Guanghai Dai - PLA Gen. (CRC)		
VEGFR 1/2/3	Fruq. + TYVYT® (PD-1)	GI tumors			China	Jin Li - SH East Hosp. (Others)		
	Fruq. + geptanolimab (PD-1)	CRC			China	Yuxian Bai - Harbin Med. Uni.		
	Fruq. + geptanolimab (PD-1)	NSCLC			China	Shun Lu – SH Chest Hosp.		
HMPL-689	HMPL-689	FL, MZL, MCL, DLBCL			China	Cao/Zhou – Fudan/ Tongji		
РІЗКδ	HMPL-689	CLL/SLL, HL			China	Cao/Zhou – Fudan/ Tongji		
HMPL-523	HMPL-523	B-cell malignancies	All		China	Multiple leads by sub-types		
Syk	HMPL-523	ITP	All		China	Yang - CN Hem. Hosp.		
HMPL-453	HMPL-453	IHCC			China	Jianming Xu – BJ 307 Hosp.		
FGFR 1/2/3								
HMPL-306	<b>HMPL-306</b> (IDH1/2)	Hem. malignancies			China			
HMPL-295	HMPL-295 (ERK, MAPK pathway)	Solid tumors			China	In planning - IND cleared		
Epitinib	Epitinib (EGFR)	Glioblastoma	EGFR gene amplified		China	Ying Mao - SH Huashan		

## 2. ONCOLOGY COMMERCIAL OPERATIONS

## 420+ person oncology commercial team



Expanding rapidly to support ELUNATE® and SULANDA® launches

## Broad drug marketing and distribution capabilities with long-standing operational track record



## 2,300+ oncology hospitals and 20,000+ oncology physicians covered

- Fully in-place mid-2020; in training until products launched
- Vast majority of new staff from successful China oncology companies
- Expansion planned for future product launches



## **China Oncology commercial team**

DEEP PAN-CHINA MARKET ACCESS



#### Blend of multinational and local oncology expertise



**Chief Commercial Officer** 



NOVARTIS



VP, Sales & Marketing







Director, Commercial





Director, Sales Force **Effectiveness & Training** 







Director, Marketing Research & New Business Development





Senior Marketing Director -Fruquintinib







**Associate Marketing** Director - Surufatinib





Associate Director, **Medical Marketing** 







**Regional Sales Director** North





Regional Sales Manager North I







Regional Sales Manager North II





East I

Regional Sales Manager







Regional Sales Manager East II





Regional Sales Manager Central





Regional Sales Manager South





Regional Sales Manager South West





## 3 novel drugs launched / in review



2021 Oncology consolidated revenues guidance \$110-\$130 million (vs. 2020 \$30.2m actual)



## Fruquintinib China commercial responsibility assumed Oct 2020

Receiving 70-80% of in-market sales as revenues in China [1]

#### Surufatinib launched in China Jan 2021

HUTCHMED owns all China rights

## Savolitinib potential approval as early as Q2 2021

First sale milestone in China \$25 million

Eligible for 30% royalty on China sales [2]









#### **Global registration study ongoing**

Potential NDA & MAA submissions in U.S., EU & Japan in 2022/2023

HUTCHMED owns all ex-China rights

#### US & EU filings to complete in 2021

Preparing for potential launch in 2022

HUTCHMED owns all ex-China rights

#### **AZ ex-China development**

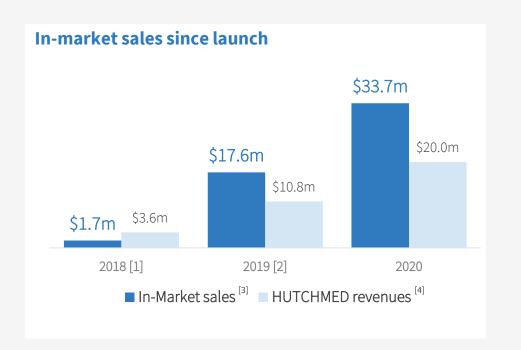
Phase III development in RCC & NSCLC targeted to start in 2021

Eligible for 9-18% royalty ex-China

## **ELUNATE®** commercial update



Sales growth accelerating since HUTCHMED assumed commercial role in Q4 2020



HUTCHMED Oncology team involved since Q4 2020					
	Lilly Sales Team	HUTCHMED	HUTCHMED Sales Team		
	Q1-Q3 2020	Q4 2020	Jan-Feb 2021*		
In-market sales [3]	\$23.5m	\$10.2m	\$14.3m		
YoY growth	+37%	+2,051% <sup>[2]</sup>	+116%		
HUTCHMED revenues [4]	\$12.8m	\$7.2m	\$10.2m		

#### Further activities to support continued acceleration

- KOL engagement plans in coordination with hospital listing expansion
- Life cycle management programs
- Synergy from surufatinib launch







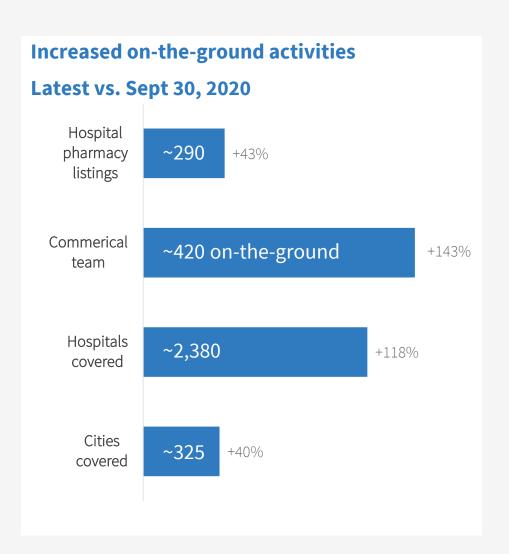
<sup>[1]</sup> ELUNATE® was launched in late November 2018. HUTCHMED revenues in 2018 primarily relate to manufacturing fees and royalties paid by Lilly.

<sup>[2]</sup> During Q4 2019, ELUNATE® in-market sales were affected by rebates and downward price adjustments required in the distribution channel in the lead up to NRDL inclusion effective Jan 1, 2020

## **ELUNATE®** commercial update



Sales benefitting from deeper coverage and increasing hospital listings



#### **Strong foundation**

✓ Clear clinical benefits continuously presented at medical conferences since 2018

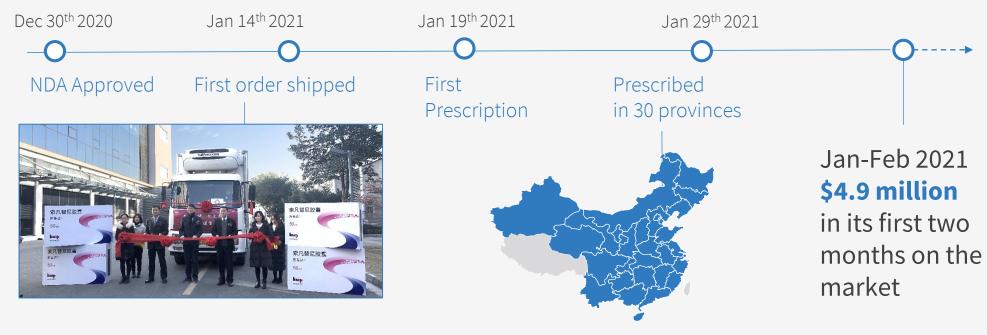


- ✓ Guideline inclusion<sup>[1]</sup>
  Class I recommendation (Level 1A evidence)
  for the treatment of 3L CRC regardless of RAS and
  BRAF gene status
- ✓ NRDL listing effective Jan 2020 enables broad patient access – Jan 2021 volume greater than full year 2019 [2]

## HUTCHMED

### **SULANDA®** launch

#### Executed within 3 weeks of NDA approval...just beginning



#### **Patient access**

 Eligible to negotiate for NRDL inclusion during 2021





## 3. 2020 REGULATORY ACHIEVEMENTS

## **Major regulatory achievements**

China



2020 was our most productive year in terms of regulatory progress



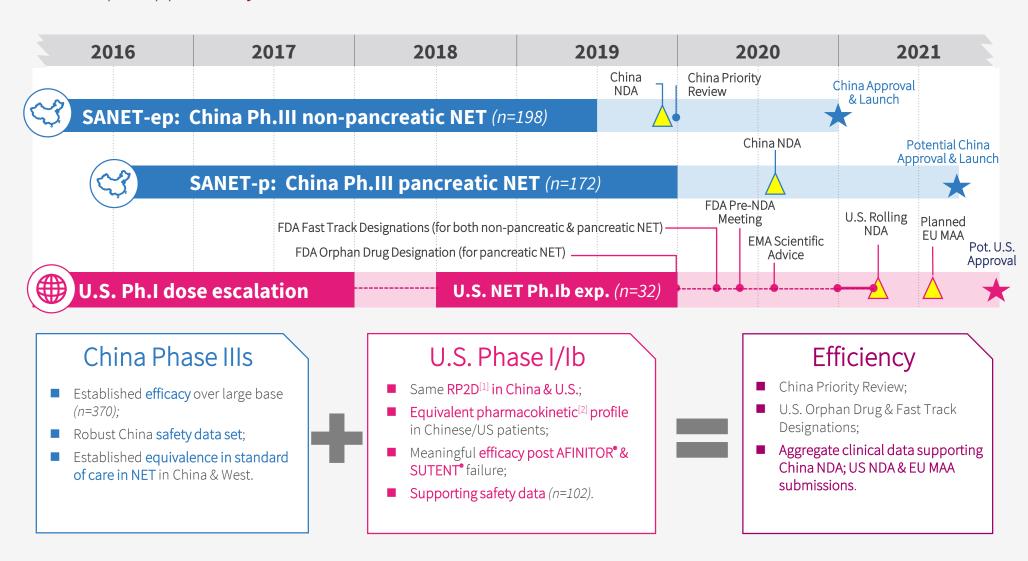
Q4 2020

Q4 2020

## China data support of US NDA & EU MAA



...unique opportunity for China-based innovators



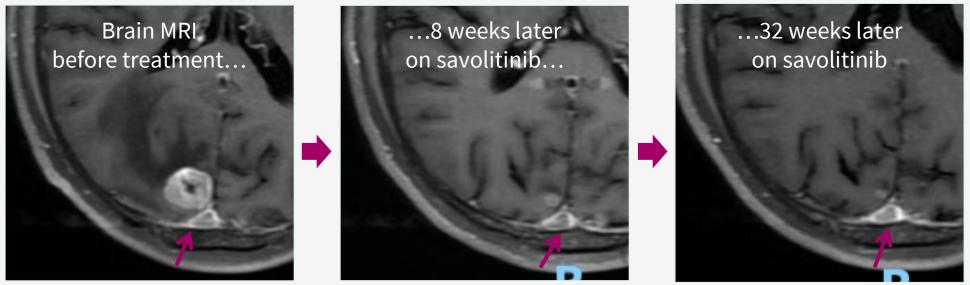
## 1st selective MET inhibitor NDA in China



Savolitinib for MET Exon 14 skipping NSCLC<sup>[1]</sup> NDA based on Phase II



#### 2. Anti-tumor activity observed in brain metastases<sup>[2]</sup>



## FRESCO-2 to support 3L+ mCRC US/EU/JP NDA



Regulatory alignment on fruquintinib across all major markets

#### Basis for US, EU, Japan filings



- FRESCO-2 + FRESCO + US CRC Ph Ib data, could support US NDA & EU MAA submissions in third-line and above metastatic colorectal cancer
- US Fast Track Designation → potential rolling submission
- Extensive list of supportive studies.

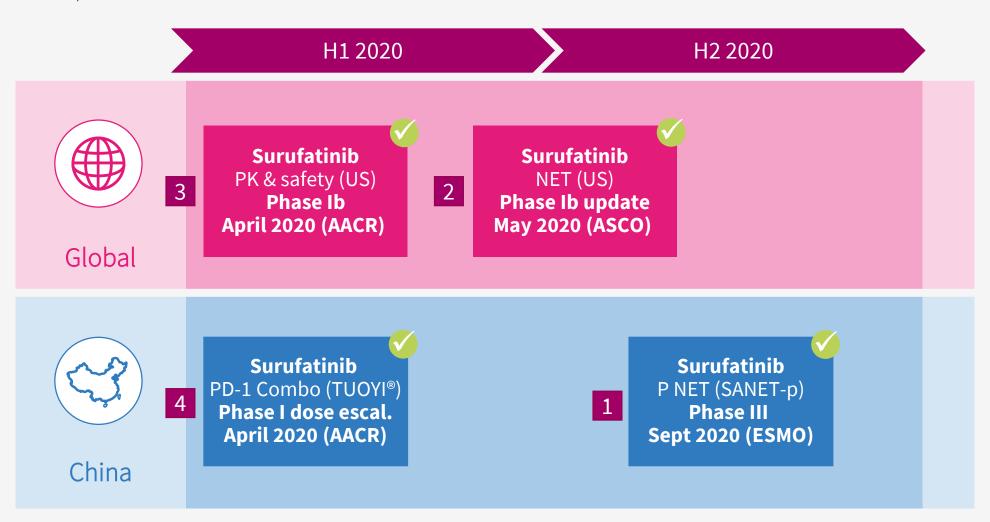
CONSISTENCY IN SAFETY	Phase Ib [1] United States		ase III Study d China <sup>[2]</sup>
Treatment arms	Fruq.	Fruq.	Placebo
Patients (n)	31	278	138
≥G3 AE (Safety population)	79.4%	61.1%	19.7%
VEGFR on-target related AEs ≥ G3:			
Hypertension	23.4%	21.2%	2.2%
Hand-Foot Syndrome	2.9%	10.8%	0.0%
Hepatic function (Liver function) AE	- s ≥ G3:		
ALT increased, ≥G3	<5%	0.7%	1.5%
AST increased, ≥G3	0%	0.4%	0.7%
Blood bilirubin increased, ≥G3	<5%	1.4%	1.5%
Tolerability: AE Leading to			
Dose reduction/interruption	41.2%	47.1%	13.1%
Treatment discontinuation	8.8%	15.1%	5.8%

## 4. CLINICAL DEVELOPMENT ACTIVITIES

## HUTCHMED

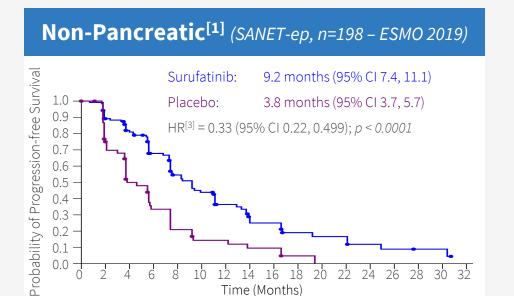
## Surufatinib clinical development activities

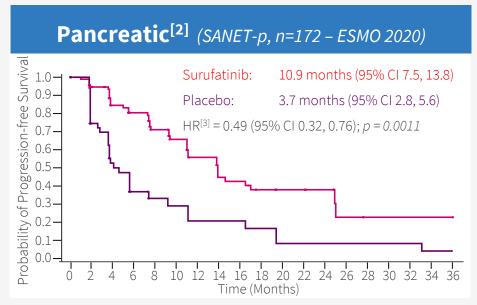
Data presentations in 2020



## 1 G1/2 Advanced NET

#### Pancreatic NET (SANET-p) data presentation





	China			US		E	EU5	
	Annual Incidence	Estimated Prevalence	mPFS	Annual Incidence <sup>[4]</sup>	Estimated Prevalence <sup>[4]</sup>	Annual Incidence <sup>[5]</sup>	Estimated Prevalence <sup>[5]</sup>	
Total NET	67,600	~300,000 (Est. China ratio <sup>[4]</sup> )		19,000	141,000	18,700	138,800	
Non- Pancreatic NET	~54,100	<b>~240,000</b> (Est. China ratio <sup>[4]</sup> )	9.2 mo. (SANET-ep Ph.III)	17,000	127,000	16,700	125,000	
Pancreatic NET	~13,600	~60,000 (Est. China ratio <sup>[4]</sup> )	10.9 mo. (SANET-p Ph.III)	2,000	14,000	2,000	13,800	

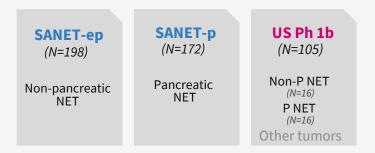
<sup>[1]</sup> Xu J, Shen L, Zhou Z, et al. Surufatinib in advanced extrapancreatic neuroendocrine tumours (SANET-ep): a randomised, double-blind, placebo-controlled, phase 3 study. Lancet Oncol. 2020;21(11):1500-1512. doi:10.1016/S1470-2045(20)30496-4; [2] Xu J, Shen L, Bai C, et al. Surufatinib in advanced pancreatic neuroendocrine tumours (SANET-p): a randomised, double-blind, placebo-controlled, phase 3 study. Lancet Oncol. 2020;21(11):1489-1499. doi:10.1016/S1470-2045(20)30493-9; ; [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] 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; [5] Estimated based on relative population versus the U.S.

## US NET bridging study



Encouraging surufatinib efficacy in everolimus & sunitinib refractory/intolerant patients

#### **Basis for NDA & MAA (US FDA / EMA)**

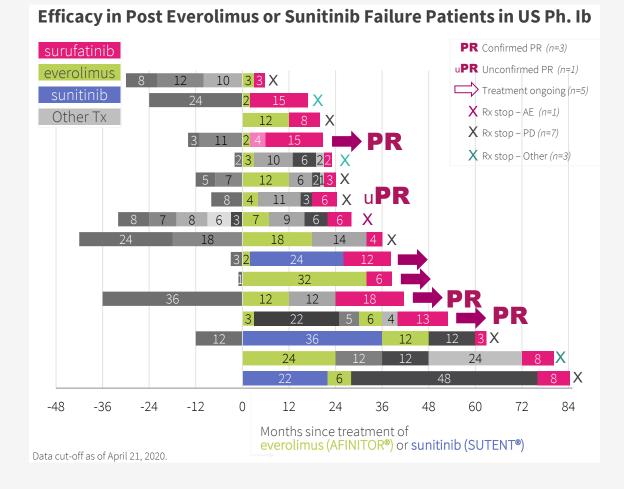


- SANET-ep + SANET-p + existing US NET patients data, could support US NDA & EU MAA submission;
- US Fast Track Designations → rolling sub;
- Extensive list of supportive studies.

## Similar PK and Toxicity Profile between China & US patients

- 300mg QD recommended in both populations;
- PK: C<sub>max</sub> & AUC<sub>tau</sub> <10% difference; no meaningful impact of race on exposure;
- Safety: similar dose intensities; US adverse events at or below China patients.

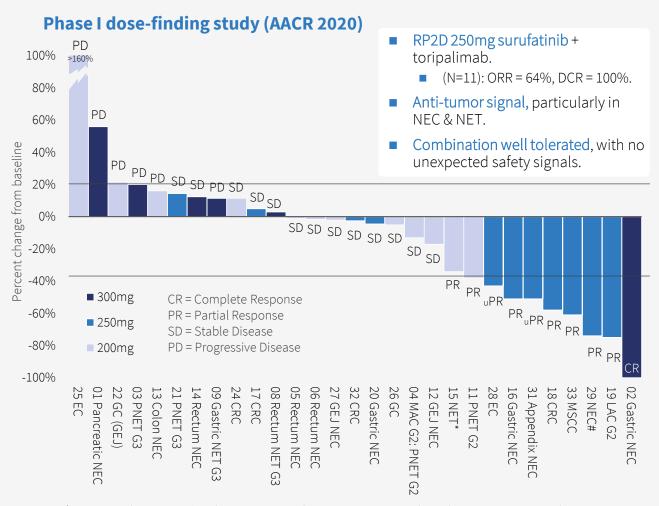
#### **Encouraging prelim. efficacy in heavily pre-treated US NET pts**

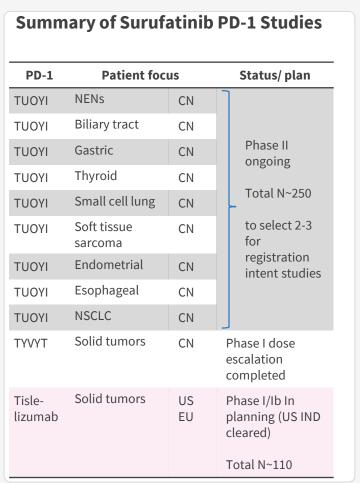






Encouraging anti-tumor efficacy for surufatinib plus TUOYI® (PD-1) combination



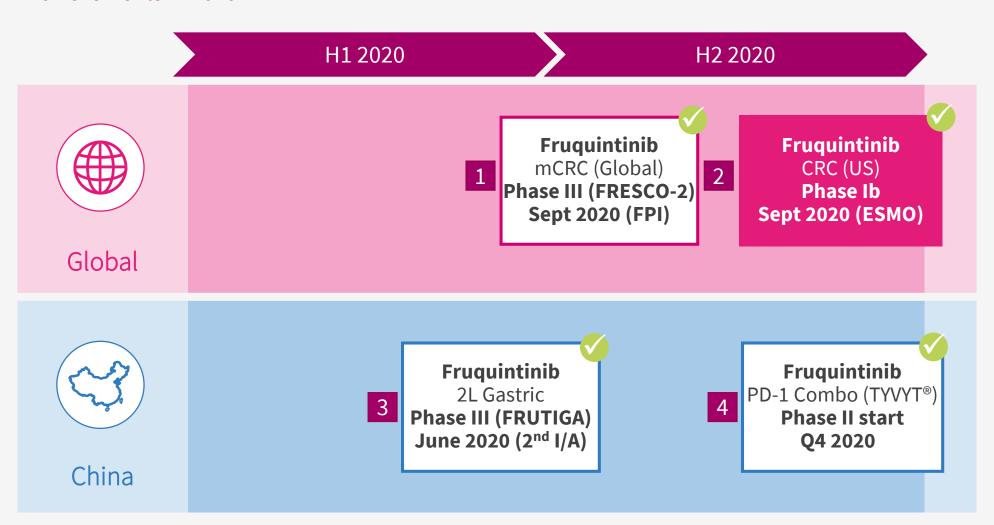


NET/NEN: neuroendocrine tumor/neoplasm; NEC: neuroendocrine carcinoma; CRC: colorectal carcinoma; GC: gastric adenocarcinoma; EC: esophageal squamous cell carcinoma; GE: gastroesophageal junction; MAC G2: mediastinal atypical carcinoid; PNET G2: Pancreas NET G2; MSCC: metastatic squamous cell carcinoma with unknown primary; NSCLC: non-small cell lung cancer; LAC: Lung atypical carcinoid; \*: Left supraclavicular lymph node neuroendocrine tumor; #: Merkel cell carcinoma.

## Fruquintinib clinical development activities



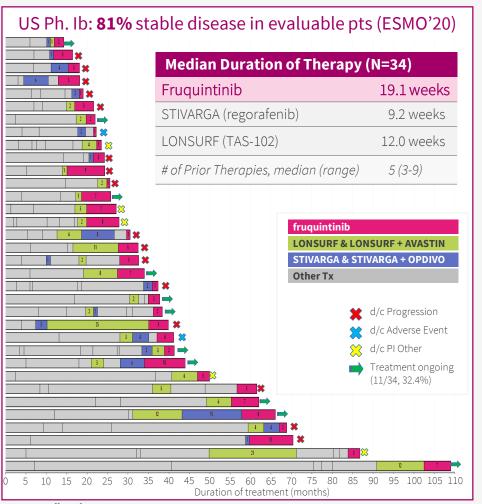
Achievements in 2020





## 2 FRESCO-2 initiation supported by US data

AACR, ASCO & ESMO presentations demonstrate compelling preliminary monotherapy efficacy and safety in heavily pre-treated US CRC patients





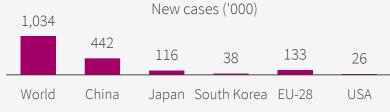
Data cut-off as of Aug 20, 2020



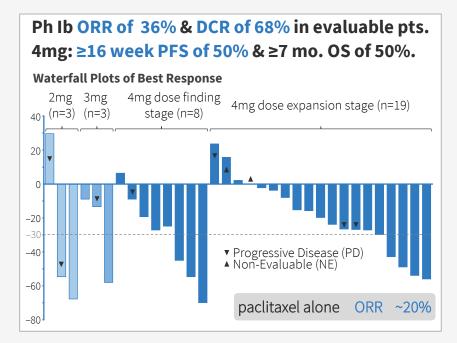
## FRUTIGA – 2L gastric combo with paclitaxel

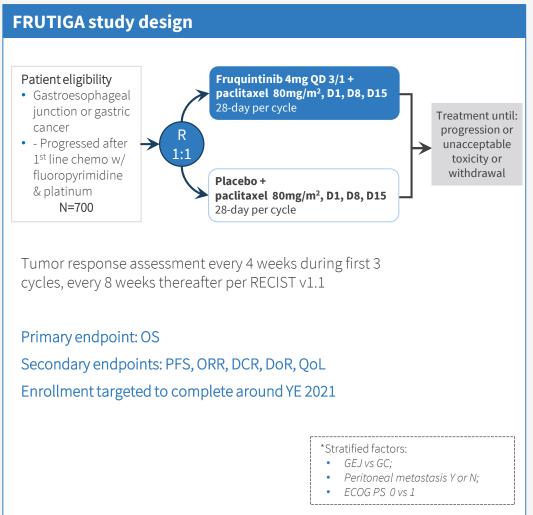
Ongoing – interim futility analysis Jun 2020 (~200 OS events)

Gastric (stomach) cancer is the 5<sup>th</sup> most common cancer globally –782,700 deaths/year



WHO, ACS, NCCR, Lancet, Frost & Sullivan Analysis.





## **Fruquintinib PD-1 combinations**



#### Fruquintinib selectivity highly suited for combinations

Tradaments selectivity ingitty surrect for combinations						
Inhibitors	Lenvatinib	Axitinib	Fruquintinib			
Selectivity for VEGFR	Relatively	selective	Highly selective			
VEGFR1 (nM)	22	3	33			
VEGFR2 (nM)	4	7	25			
VEGFR3 (nM)	5	1	0.5			
Phos-KDR (nM)	0.8	0.2	0.6			
Other kinases (IC50 < 100nM)	PDGFRα PDGFRβ FGFR1-4 Ret c-Kit	PDGFRα PDGFRβ c-Kit	none			

#### PD-1i/VEGFRi synergy in 1L Clear Cell RCC [2]



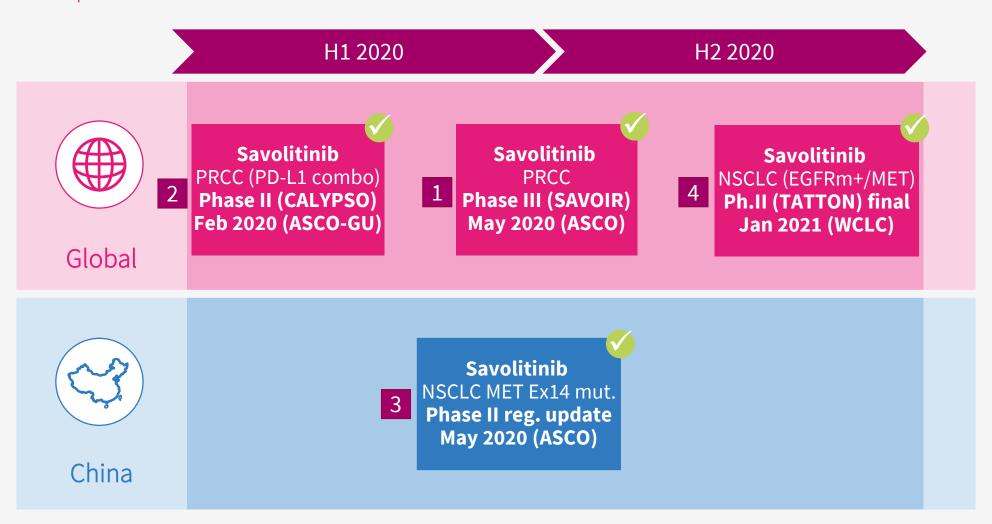
#### **Summary of fruguintinib PD-1 studies** PD-1 **Patient focus** Status/ plan Phase II ongoing TYVYT CRC CN Est. N~35 Hepatocellular TYVYT CN carcinoma Phase Ib/II ongoing; **TYVYT** Endometrial cancer CN Total est. N~120 TYVYT RCC CN to select 1-2 for registration intent studies **TYVYT** Other GI CN Tislelizumab US Phase I/Ib In planning **TNBC** Est. N~80 Tislelizumab Solid tumors Phase I/Ib In planning Est. N~60+ Geptanolimab Phase Ib ongoing CRC CN Est. N~15 Geptanolimab Phase Ib ongoing **NSCLC** CN Est. N~15

<sup>[1]</sup> Upadhaya S, Neftelino ST, Hodge JP, Oliva C, Campbell JR, Yu JX. Combinations take centre stage in PD1/PDL1 inhibitor clinical trials [published online ahead of print, 2020 Nov 11]. Nat Rev Drug Discov. 2020;10.1038/d41573-020-00204-y. doi:10.1038/d41573-020-00204-y. [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=1 pts. ORR=26.4% for CPS=1 pts.

## Savolitinib clinical development activities



Data presentations in 2020



### 1 Savolitinib in PRCC



#### SAVOIR 60 pt. data shows strong signal with monotherapy

2017		2018	2019	9
timolino	Study initiation Planned number of pts = 180 (90 savolitinib / 90 sunitinib)	60 pts e	nent stopped nrolled blitinib / 27 sunitinib)	Data cut-off (DCO)  23 pts remained on therapy (14 savolitinib / 9 sunitinib)

#### Anti-tumor activity

All 9 savo responders remained in response at DCO

[95% CI]	Savolitinib (N=33)	Sunitinib (N=27)
ORR*	9 (27) [13.3, 45.5]	2 (7) [0.9, 24.3]
PFS	7.0 [2.8, NC]	5.6 [4.1, 6.9]
	Hazard Ratio:	0.71 [0.37, 1.36]
DCR@6months	16 (48) [30.8, 66.5]	10 (37) [19.4, 57.6]
@ 12 months	10 (30) [15.6, 48.7]	6 (22) [8.6, 42.3]

<sup>\*</sup> One out of two sunitinib responders remained in response at DCO

#### Better tolerability

42% savo vs 81% sunitinib AE Gr. ≥3

	Savolitinib (N=33)	Sunitinib (N=27)
Treatment related AE Grade ≥3	8 (24)	17 (63)
Any AE Grade ≥3	14 (42)	22 (81)
Anemia	0	4 (15)
Hypertension	0	4 (15)
AST increased	5 (15)	2 (7)
ALT increased	4 (12)	2 (7)

#### Strong signal of potential overall survival benefit Savolitinib | Sunitinib Median. NC 13.2 [11.9, NC] [7.6, NC] mo. HR [95% CI]: **0.51** [0.21-1.17] *P=0.110* 0.8 Probability of OS Savolitinib (n=33) Sunitinib (n=27) Censored observations

Time From Randomization (Months)

### 2 Savolitinib + PD-L1 inhibitor



CALYPSO Savo/IMFINZI® combo tolerable, w/ durable efficacy

PD-1/PD-L1s important in non-ccRCC but need to see mature mPFS/mOS & further biomarker analysis [1]

#### MET+ **Papillary RCC** (~\$1.0b) ~8% of RCC

~ 28k new patients/yr.

#### MET-Papillary RCC (~\$1.0b)

~8% of RCC ~ 28k new patients/yr.

#### Other non-ccRCC (~\$0.6b)

~5% of RCC ~ 16k new patients/yr.

#### Savo mono.

All lines: (n=60) 27.3% mPFS 7.0 mo.

#### Tecentriq®+Cabometyx®

All lines: (n=15) ORR 40.0%

#### Keytruda® mono.

First line: (n=118) ORR 28.8% DCR 47.5%

#### Tecentrig®+Avastin®

All lines: (n=12) ORR 25.0%

#### Tecentrig®+Avastin®

All lines: (n=42) ORR 26.2%

#### Tecentrig®+Cabometyx®

All lines: (n=30) ORR 33.3%

DCR 93.3%

#### Savo + Imfinzi®

#### All lines: (n=41)

26.8% mPFS 4.9 mo. mOS 12.3 mo.

#### CALYPSO Interim Data First line: (n=27)

ORR 33.3%

#### Keytruda® mono. (all non-ccRCC)

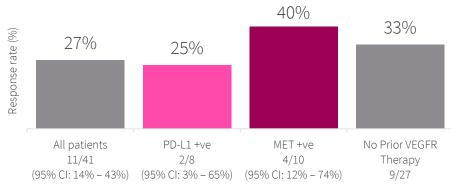
#### First line: (n=165)

26.7%

ORR

DCR 43.0% mPFS 4.2 mo. mOS 28.9 mo.

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



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



Further assessment of biomarkers (6 not assessable)

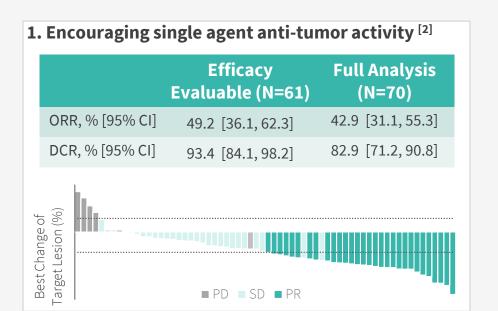
- Only MET+ overexpression assessed to date (10/41 positive, 25/41 negative);
- MET+ gene amp. / other MET aberrations to evaluate.

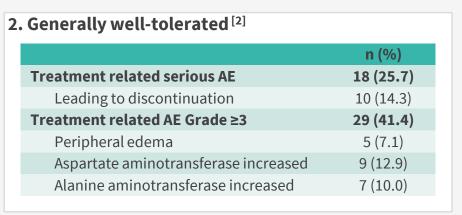
Phase III PRCC trial starting in mid-2021

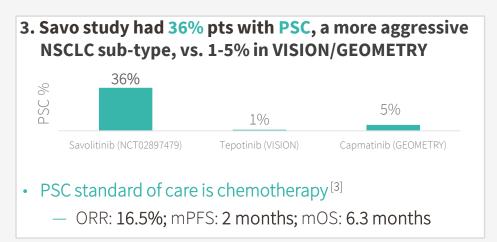
## **3 MET Exon 14 skipping NSCLC**[1]

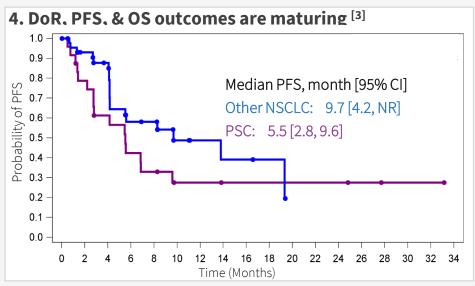


#### China NDA accepted in May 2020 based on data presented at ASCO 2020













#### TATTON B & D Data - efficacy

		TATTON Part D osimertinib 80 mg + savolitinib 300 mg		
	Part B1 (n=69) Prior third-generation EGFR-TKI	Part B2 (n=51)  No prior third- generation EGFR-TKI (T790M negative)	Part B3 (n=18)  No prior third- generation EGFR-TKI (T790M positive)	Part D (n=42)  No prior third- generation EGFR-TKI (T790M negative)
<b>Objective response rate</b> *, % [95% CI] Complete response, % Partial response, %	<b>33%</b> [22, 46] 0 33%	<b>65%</b> [50, 78] 0 65%	<b>67%</b> [41, 87] 0 67%	<b>62%</b> [46, 76] 0 62%
Non-response, % Stable disease (≥ 6 weeks) Progressive disease Not evaluable	42% 12% 13%	24% 6% 6%	33% 0 0	31% 2% 5%
Disease control rate <sup>#</sup> , % [95% CI]	<b>75%</b> [64, 85]	<b>88%</b> [76, 96]	<b>100%</b> [81, 100]	<b>93%</b> [81, 99]
Median DoR, months [95% CI]	<b>9.5</b> [4, 15]	<b>10.7</b> [6, 15]	<b>11.0</b> [2.8, NR]	<b>9.7</b> [5, 14]
Median PFS, months [95% CI]	<b>5.5</b> [4.1, 7.7]	<b>9.1</b> [5.5, 12.8]	<b>11.1</b> [4.1, 22.1]	<b>9.0</b> [5.6, 12.7]

<sup>[1]</sup> 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 <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 ≥4 weeks. # Disease control rate = confirmed complete response + confirmed partial response + stable disease at ≥5 weeks; CI, confidence interval; NR, not reached. Han JY, et al. Osimertinib + savolitinib in patients with EGFRm MET-amplified/overexpressed NSCLC: Phase Ib TATTON Parts B and D final analysis. WCLC January 2021 #FP14.03.





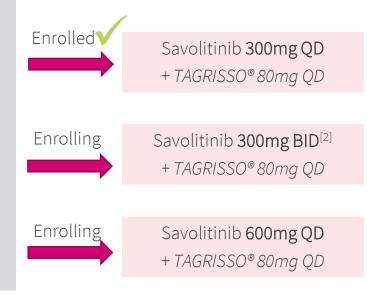


#### SAVANNAH designed for possible registration of 300mg QD [1]

#### In broadest TAGRISSO® (osimertinib) refractory population – FISH+ and/or IHC+ line agnostic population

## 2L+ LOCALLY ADV. / METASTATIC EGFRM+ NSCLC PATIENTS

- Progression on 1L or 2L TAGRISSO®;
- No prior chemo or immunotherapy;
- MET amplification / over-expression (central FISH/IHC or pre-existing local NGS);
- No prior MET inhibitor therapy;
- Stable/asymptomatic CNS mets. permitted;
- ECOG performance status 0-1.



#### PRIMARY ENDPOINT

■ 300mg QD ORR

#### SECONDARY ENDPOINTS

- 300mg QD
  - ORR by MET FISH+ / IHC+; PFS; DoR; OS
  - > Safety
- 300mg BID & 600mg QD
  - ➤ Efficacy (ORR; PFS; DoR; OS)
  - > Safety / tolerability

- SAVANNAH will also inform the design of planned global Phase III by mid-2021
  - optimal biomarker strategy (FISH/IHC);
  - optimal dose (300mg or 600mg);
  - optimal dose regimen (QD or BID); &
  - optimal dose line of treatment (post 1L or 2L TAGRISSO®)

[1] QD = Once daily dose; [2] BID = twice daily dose.

# HUTCHMED

# **Next wave of innovation**

### Current development status summary

#### HMPL-689 & HMPL-523

- China Ph.Ib dose expansions underway;
- U.S. & EU Ph.I multiple dose cohorts completed;
- To start multiple Ph.II/III reg. studies in 2021

#### **HMPL-453**

Ph.II initiated in intrahepatic cholangiocarcinoma in China.

#### **HMPL-306**

- 9<sup>th</sup> in-house discovered asset (IDH1/2) Ph.I;
- Addresses mutant IDH switching, from IDH1 to IDH2 or vice versa, a resistance mechanism.

#### **HMPL-295**

- 10<sup>th</sup> in-house discovered asset (ERK, MAPK pathway);
- Ph.I est. start mid-2021

Program	Treatment	Target Patient	Sites	Dose Finding / Safety Run-in	Proof-of-concept	Registration
	HMPL-689	Healthy volunteers	Australia			
HMPL-689	HMPL-689	Indolent NHL	US/EU			
ΡΙ3Κδ	HMPL-689	FL, MZL, MCL, DLBCL	China			*
	HMPL-689	Other iNHL subtypes	China			
HMPL-523	HMPL-523	Indolent NHL	US/EU/AU			
Syk	HMPL-523	B-cell malignancies	China			
Syn.	HMPL-523	ITP	China			
<b>HMPL-453</b> FGFR 1/2/3	HMPL-453	IHCC	China			
HMPL-306	HMPL-306	Hematological Malignancies	China			
IDH 1/2	HMPL-306	Hematological malignancies & solid tumors	US/EU	*		
HMPL-295 (ERK, MAPK pathway)	HMPL-295	Solid tumors	China	*		









# HMPL-689 a highly attractive PI3Kδ inhibitor

## Consistent efficacy profile across all cohorts

1. HMPL-689 – Phase I dose escalation [1]								
At Sept 15 cut-0 20mg / day: n=26 20mg / day: n=18 30mg / day: n=9, RP2D	ITT n=56	Evaluable n=52	CLL/SLL n=5	MZL n=7	FL n=23	MCL n=9	DLBCL n=9	HL n=3
Best response 40mg/day: n=3								
Complete Re Non evaluable: n=4	11	12	40	0	14	0	0	0
Partial Response, %	37	40	40	71	30	44	33	0
Stable Disease, %	34	37	0	29	39	56	11	67
Progressive Disease, %	11	11	20	0	4	0	33	33
Not Evaluable, %	7	na	0	0	9	0	22	0
Overall Response Rate (intent-to-treat)	48%		80%	71%	48%	44%	33%	0%
Overall Response Rate (efficacy evaluable)		52% 52	80% 5	71% 7	52% 21	44% 9	43% 7	0% 3
2. Competitive PI3Kδ inhibitors								
Overall Response Rate			CLL/SLL	MZL	FL	MCL	DLBCL	HL
Zydelig <sup>®</sup> (idelalisib) <sup>[2] [3]</sup>			58%	47%	54%	-	0%	-
n			26	15	72		9	
Aliqopa <sup>®</sup> (copanlisib) [2] [4]			-	<b>78%</b> 23	<b>59%</b> 104	-	-	-
Copiktra® (duvelisib) [2] [5] [6]			78%	39%	42%	50%	-	-
n .			95	18	83	10		
Ukoniq® (umbralisib) [2] [7] [8]			50%	49%	43%	17%	57%	-
Parsaclisib [9] [10] [11]			22 <b>33%</b>	69 <b>57%</b>	117 <b>70%</b>	6 70%/25%	7 <b>26%</b>	
n			33 <b>%</b> 6	100	10 <b>90</b> 108	1090/2390	<b>20%</b> 55	_
Zandelisib (intermittent dosing) [12]			100% ₃	-	<b>76%</b>	-	-	-

# **HMPL-689**



# Advantages in tolerability versus PI3K $\delta$ inhibitors

#### Incidence of select treatment emergent adverse events – all AEs / grade ≥3 AEs

	n	Neutropenia	Anemia	Thrombo- cytopenia	Diarrhea or colitis	Rash	ALT increased	AST increased	Pyrexia	Pneumonia	Hyper- tension	Hyper- glycemia
Zydelig® (idelalisib) <sup>[2]</sup>	146	53% / <b>25%</b> *	28% / 2%*	26% / 6%*	47% / 14%	21%/3%	50% / <b>19%</b>	41% / <b>12%</b>	28% / 2%	25% / 16%	na	na
Aliqopa® (copanlisib) [2]	168	32% / <b>25%</b>	na	22%/8%	36% / 5%	15% / 2%	na	na	na	21% / 14%**	35% / 27%	54%/39%
Copiktra® (duvelisib) [2]	442	34% / 30%	20% / 11%	17%/10%	50% / 23%	31%/9%	40% / <b>8%</b>	37% / <b>6%</b>	26% / 2%	21%/15%	na	na
Ukoniq <sup>®</sup> (umbralisib) [2]	221	33% / 16%*	27% / 3%*	26 % / 4%*	58% / 10%	18%/3%	33% / 8%	32% / 7%	na	PJP prophylaxis recommended	na	na
Parsaclisib (Dose escalation) [5]	72	44% / 20%*	31% / 8%*	35% / 10%*	36%/9%	31%/6%	28% / 1%	29% / 1%	18% / 1%	na	7% / 0%	10% / 1%
Parsaclisib (CITADEL-204/MZL) [6]	100	13% / 9%	14% / 5%	na	44%/11%	17% / 2%	26% / 4%	19% / 2%	13% / 1%	7% with PJP prophylaxis	na	na
Zandelisib (intermittent dosing) [7]	21	na / 14%	na / 0%	na / 0%	na / 4%	na / 2%	na / 0%	na / 0%	na	PJP prophylaxis	na	na
Zandelisib (Dose escalation) <sup>[8]</sup>	30	45% / 13%*	13% / 0%*	22% / 0%*	45% / 19%	42% / 13%	39% / <b>6%</b>	25% / <b>6%</b>	na	na	na	na
HMPL-689 <sup>[1]</sup>	56	43% / 11%	16% / 0%	11%/0%	<5% / <5%	11% / 5%	27% / 2%	21% / 2%	14% / 0%	25% / 16%	7% / 5%	11% / 2%

# Potential best-in-class IDH1/2 inhibitor



HMPL-306 – China Phase I underway, two US INDs cleared to start Phase I

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

TUMOR	% IDH MUTATION [1]					
	TOTAL	IDH1- R132	IDH2- R140	IDH2- R172		
Brain tumor						
Grade 2 and 3 glioma	60-80%	60-80%	0%	1%		
Secondary glioblastoma	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%		

### HMPL-306 is a potent IDH1/2 dual inhibitor

- IDH1 & 2 mutations are validated targets in R&R AML (IDH1i ivosidenib and IDH2i enasidenib)
- 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.

#### INDs cleared in China and US in 2020

#### China Phase I initiated July 2020

Aiming for recommended Phase 2 dose around YE2021

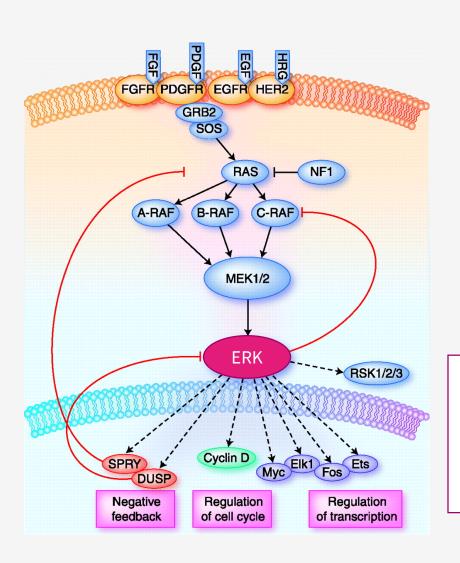
### US Phase I initiating after Oct 2020 IND clearance

First patient expected in H1 2021



# MAPK pathway represents major unmet need

HMPL-295 – the first of several HUTCHMED assets targeting MAPK pathway



#### The MAPK (RAS-RAF-MEK-ERK) signaling cascade

- ERK (extracellular signal-regulated kinases) a key component
- Pathway normal activation: ligand-dependent & tightly regulated by NF-1 and negative feedback
- *In tumors:* activating mutations in RAS, RAF and loss of the tumor suppressor NF1 leads to uncontrolled cell proliferation

#### ~50% of cancers have RAS or RAF mutation

- Increased mortality / poor OS
- Decreased the response to existing therapies including immunotherapy
- RAS: KRAS inhibitors in clinical trials
- BRAF/MEK: therapies approved induce initial rapid tumor regression, but acquire resistance developed due to MAPK pathway re-activation

ERK inhibition has the potential to overcome or avoid the intrinsic or acquired resistance from upstream mechanisms

HMPL-295, a highly selective ERK1/2 inhibitor, cleared for clinical trials in 2020 with Phase I to initiate in mid-2021

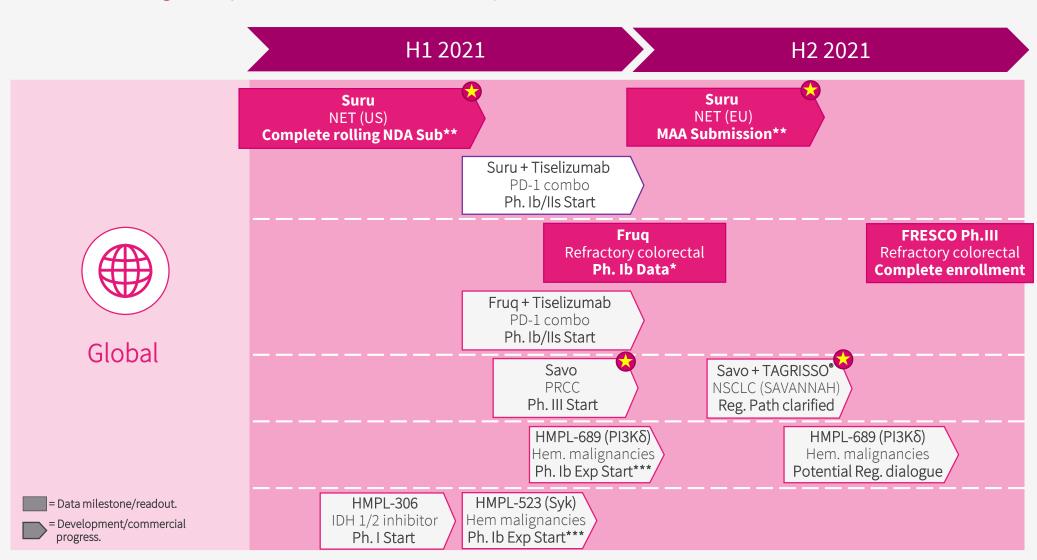
Source: Clin Cancer Res. 2010; 16: 3329-34.

# POTENTIAL UPCOMING CLINICAL & REGULATORY MILESTONES

# **Potential upcoming events**



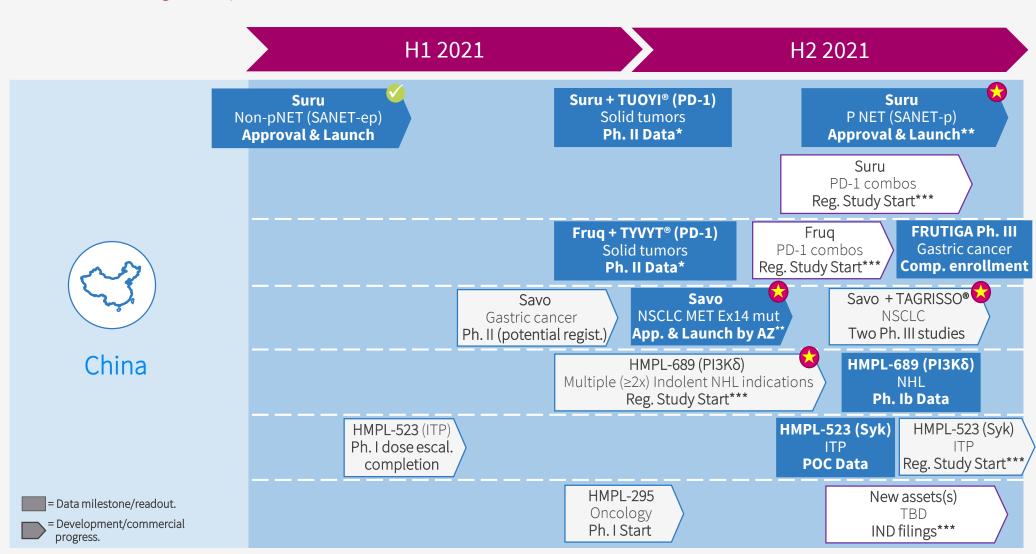
Clinical & regulatory milestones in US, EU & Japan



# **Potential upcoming events**



Clinical & regulatory milestones in China



# 5. OTHER OPERATIONAL DEVELOPMENTS

# HUTCHMED

# **Manufacturing Operations**

New Shanghai factory to support production post 2025

#### SUZHOU FACTORY – production up to 2025

- Built to produce ELUNATE®
- Manufacturing talent developed
- Suzhou is designed to U.S. GMP standards

#### SHANGHAI FACTORY

- Capex of \$130 million over 5 years
- Will fulfil additional global production requirements
- Additional capacity for expansion in large molecule production

Key Aspects	Suzhou Factory	New Shanghai Factory
Property Type	Leased	Owned
Land Size (sq.m.)	~1,800	~28,700 <b>(16x)</b>
Building Size (sq.m.)	~4,500 (Office: ~1,000)	~55,000 <b>(12x)</b> (Office: ~16,400)
Capacity (Cap & Tabs)	50 million	250 million <b>(5x)</b>
Growth Potential	No capacity for growth	Could expand to large molecules in long term











# **Immunology partnership**

Accelerating four HUTCHMED drug candidates

#### Overview

- 4 novel preclinical drug candidates discovered by HUTCHMED for the potential treatment of multiple immunological diseases
- Funded by Inmagene
- Companies working together to move candidates to IND
- Inmagene will pursue global clinical development

#### **Terms**

- HUTCHMED granted Inmagene four exclusive options (one per candidate) solely for the treatment of immunological diseases
- Option gives right to further develop, manufacture and commercialize that specific candidate worldwide
- HUTCHMED retains first right to co-commercialization in China
- Development milestones of up to US\$95 million
- Commercial milestones of up to US\$135 million
- Up to double-digit royalties



# FINANCIAL RESULTS, GUIDANCE AND SUMMARY





(in \$'000)

	As of De	ec 31,
	2020	2019
Assets		
Cash, cash equivalents & short term investments	435,176	217,168
Accounts receivable	47,870	43,254
Other current assets	47,694	56,600
Property, plant and equipment	24,170	20,855
Investments in equity investees	139,505	98,944
Other non-current assets	29,703	28,301
Total assets	724,118	465,122
Liabilities and shareholders' equity		
Accounts payable	31,612	23,961
Other payables, accruals and advance receipts	120,882	81,624
Long-term bank borrowings	26,861	26,818
Other liabilities	25,814	19,816
Total liabilities	205,169	152,219
Total Company's shareholders' equity	484,116	288,012
Non-controlling interests	34,833	24,891
Total liabilities and shareholders' equity	724,118	465,122

#### **Cash Position**

(at end December 2020)

- \$435m cash / cash eq. / ST inv. [1]
- \$69m additional unutilized banking facilities <sup>[2]</sup>
- \$27m in bank borrowings
- \$89m additional cash in JVs

#### 2020 Equity Financings:

- \$118m Nasdaq follow-on (Jan 2020) [3]
- \$100m PIPE with General Atlantic (Jul 2020)<sup>[4]</sup>
- \$100m PIPE with CPPIB (Nov 2020) [5]

# **Condensed Consolidated Statement of Operations**



(in \$'000, except share and per share data)

	Year Ended Dec 31,	
	2020	2019
Revenues:		
Oncology/Immunology – Marketed Products	19,953	10,766
Oncology/Immunology – R&D	10,262	16,026
Oncology/Immunology total revenues	30,215	26,792
Other Ventures	197,761	178,098
Total revenues	227,976	204,890
Expenses:		
Costs of revenues	(188,519)	(160,152)
R&D expenses	(174,776)	(138,190)
Selling and general administrative expenses	(61,349)	(52,934)
Total expenses	(424,644)	(351,276)
Loss from Operations	(196,668)	(146,386)
Other income	6,934	5,281
Loss before income taxes & equity in earnings of equity		
investees	(189,734)	(141,105)
Income tax expense	(4,829)	(3,274)
Equity in earnings of equity investees, net of tax	79,046	40,700
Net loss	(115,517)	(103,679)
Less: Net income attributable to non-controlling interests	(10,213)	(2,345)
Net loss attributable to HUTCHMED	(125,730)	(106,024)
Losses per share attrib. to HUTCHMED – basic & diluted	(0.18)	(0.16)
Losses per ADS attrib. to HUTCHMED – basic & diluted	(0.90)	(0.80)

### 2021 Guidance

- \$110-130m in consolidated
   Oncology/Immunology revenue
  - Accelerating growth on ELUNATE®
  - Full year sales on SULANDA®
  - Potential launch of savolitinib & first China sale milestone
- Rapid international expansion of organization & development on 6 oncology assets – U.S. & Europe R&D Expense grew to \$63.3 million (2019: 21.7m) while China stable at \$111.5 million (2019: \$116.5m)
- Continue evaluating non-core assets divestment opportunities
- Continue to monitor market conditions for listings on other stock exchanges such as Hong Kong & Shanghai

# **Summary**

# Oncology commercialization

**2021 Oncology consolidated revenues guidance \$110-130 million** from in-house commercial team (ELUNATE®) & product launches (SULANDA®, savolitinib planned)

# Savolitinib (MET) progress

Initiating 3+ Phase III combination studies in 2021, in parallel to potential 1st approval

# Hematology progress

HMPL-689 (PI3Kδ) entering potential registration studies supported by PoC data, while HMPL-523 (Syk) delivers promising PoC data, and HMPL-306 (IDH1/2) progress

#### **Combos**

**Exploring promising combinability of our assets in 2021 with anti-PD-1/PD-L1** (IMFINZI®, TUOYI®, TYVYT®) and other therapies (TAGRISSO®, TAXOL®)

# International organization ascending

**Filing 1st US FDA NDA and preparing team for potential US launch**, with global Phase III & many PoC studies enrolling

# **APPENDIX**



**Strategies** Realizing global potential of novel oncology assets **Building a fully integrated China oncology business Product Candidate Details Further Corporate Information** 

**A1** 

**HUTCHMED STRATEGY** 

# World class discovery engine





# Focus on Global Quality Innovation Proven & Validated at All Levels

- 15+ year track record in oncology, fully integrated 600+ person in-house scientific team
- 40+ oncology indications in development. 9 TKIs incl. VEGFR, c-MET, PI3K $\delta$ , Syk, FGFR & IDH
- 10+combo therapy trials with chemo, TKI & IO drugs. Superior selectivity enables combos
- 4 further in-house late pre-clinical molecules
- 2 validating collaborations





# **Superior Selectivity Profiles** Savolitinib ~1,000 times more selective to c-MET than next kinase (PAK3) [1] 1μM against **ELUNATE** ~250 times more selective to VEGFR3 than next non-VEGFR kinase (Ret) [2]

**HUTCHMED's Advanced Chemistry Approach Provides** 

# **Established global C&R infrastructure**

# HUTCHMED

1 WORLD-CLASS DISCOVERY & DEVELOPMENT CAPABILITY

HIGHLY DIFFERENTIATED NME PORTFOLIO AND GLOBAL PIPELINE

## Track record of breakthroughs

- Integrated development team of 120+ C&R & ~200 CMC staff located in Shanghai, Suzhou & Florham Park, New Jersey
- Broad bandwidth & capacity of R&D team enables smooth coordination of >25 trials globally & in China
- Important working relationships with China & global regulators potentially multiple new global registration studies in 2021
- At launch / filing stage on 3 lead assets major regulatory achievements



### Fruquintinib (ELUNATE® in China)

- 1st China-discovered & developed, unconditionally approved cancer therapy
- Global Ph.III started mid-2020, >150 sites in US, EU & JP
- Ideal combo candidate with limited off-target activity; favorable PoC results with chemo & TKIs

#### Savolitinib

- China NDA & Priority Review 1st NDA filing globally and first-inclass in China
- Global partnership with AZ China clinical by HUTCHMED
- Multiple global indications potentially 3 reg. studies 2021

#### Surufatinib (SULANDA® in China)

- 2 China NDAs (1 approved & 1 accepted) unpartnered
- US NDA submission using China Ph.IIIs & US Ph.Ib/II data (late 2020 through early 2021). EU to follow
- Dual-MoA anti-angiogenesis and immuno-oncology

# Seasoned executives - MNC veterans





**Selected Shareholders** 

## Global standards – Reputation & transparency

### **Management Team**



32/21

Christian Hogg Chief Executive Officer P&G



Weiguo Su Officer



Johnny Cheng Chief Financial Officer digital Myers Squibb Nestle



Junjie Zhou General Manager, SHPL



















MITSUI&CO.



















Chief Scientific **Pfizer** 



KPMG



30/20



Tom Held Head of Commercial. U.S.





International

Marek Kania

Managing Director &

Chief Medical Officer.



27/13

31/16

Zhenping Wu Pharmaceutical Sciences Roche **Pfizer** 



23/11

Bristol Myers Squibb **b** NOVARTIS

Chief Commercial

Hong Chen

Officer, China



NOVARTIS

30/1



May Wang Business Dev. & Strategic Alliances





Mark Lee Corporate Finance & Development





Charles Nixon General Counsel





Andrew Shih HR - Organization & Leadership Dev.





Yiling Cui





Enrico Magnanelli International Operations



#### 0 Issues

in governance in 14 years listed on AIM & 5 years on NASDAO





### **Track Record of Successful Partnerships**

Across functions verified by our long-term MNC partners







56

A1a

# REALIZING GLOBAL POTENTIAL OF NOVEL ONCOLOGY ASSETS

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





## 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

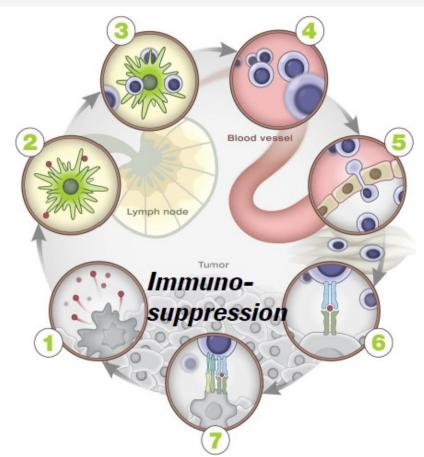
Insufficient T cell response

- 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

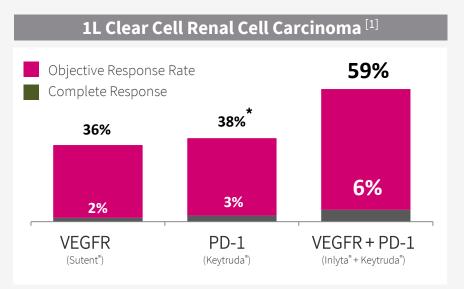
- Immunotherapies (address negative regulators)
- Vaccines

Need combinations of potent, yet tolerable drugs against specific targets

# **Immunotherapy combinations**



assets potentially ideal TKI combo partners for immunotherapy



	Inlyta®	Fruquintinib	Surufatinib
Selectivity	Relatively selective	Highly selective	Selective angio-immuno kinase inhibitor
Status	Launched	Launched	Launched
VEGFR1 (nM)	3	33	2
VEGFR2 (nM)	7	25	24
VEGFR3 (nM)	1	0.5	1
Phos-KDR (nM)	0.2	0.6	2
Other kinases (IC50 < 100nM)	PDGFRα PDGFRβ c-Kit	none	<b>CSF-1R</b> FGFR1 FLT3 TrkB
First Patent Expiration	2025/04/29 (US6534524B1)	2029 (without extension)	2030 (without extension)

Potent two-prong attack – BTD [2]:
Anti-angiogenesis + activated T-cell response

**Fruq. uniquely selective** – unlike other TKIs with off-target toxicity **Suru. inhibits TAM production** – amplifying PD-1 induced immune response

Multiple global immunotherapy combo deals...

# Managed by AstraZeneca



ccRCC/PRCC/ other solid tumors Innovent Biologics
fruquintinib/surufatinib

+ Tyvyt\* (PD-1)
Solid tumors

君实生物 Junshi Biosciences

Jointly managed by HUTCHMED & partners

surufatinib + Tuoyi® (PD-1)

Solid tumors



fruquintinib / surufatinib + tislelizumab (PD-1)

Solid tumors

Global PD-1 / PD-L1 combos – Development now underway / in planning on savo, fruq & suru

A1b

# BUILDING A FULLY INTEGRATED CHINA ONCOLOGY BUSINESS

# China: >25% of world cancer patients<sup>[1]</sup>





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

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



#### **HUTCHMED** is a first mover

- ELUNATE® launch in 3L mCRC; First ever in China [3]
- Deep pipeline 10 clinical drug candidates with 3 NDAs submitted in China



#### Major commercial opportunity

National Drug Reimbursement; Medical coverage



# **HUTCHMED** competence in China operations



A 17-year track record of 19% CAGR net income growth in our Other Ventures businesses



<sup>[1] 2003–2006</sup> incl. disco. operation; [2] Based on aggregate Non-GAAP net income / (loss) of consolidated subsidiaries and non-consolidated joint ventures of Other Ventures, please see appendix "Non-GAAP Financial Measures and Reconciliation"; [3] Includes the land compensation in SHPL of \$40.4 million from net income attributable to HUTCHMED in 2016, SHPL's R&D related subsidies of \$2.5 million from net income attributable to HUTCHMED in 2020.

**A2** 

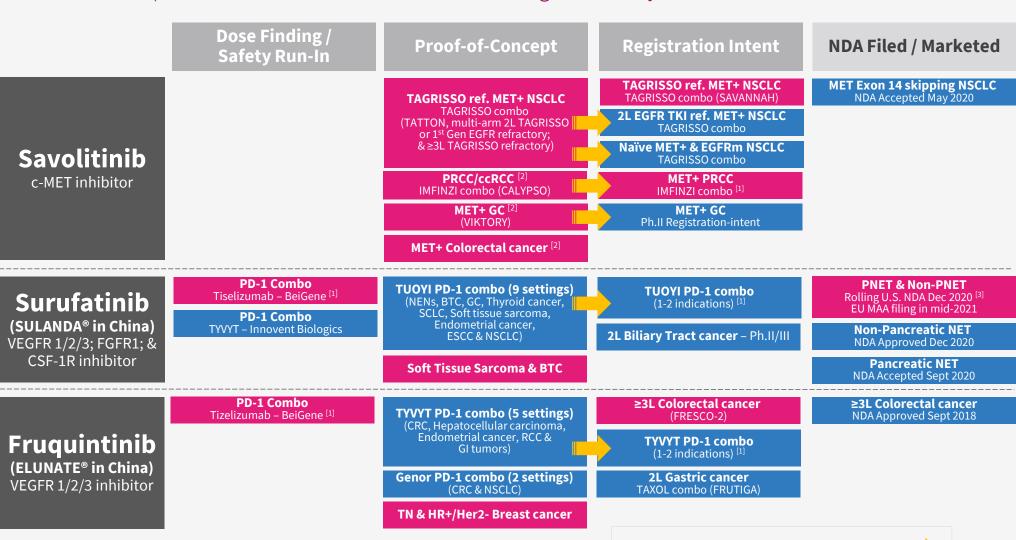
PRODUCT CANDIDATE DETAILS

# Maximizing the value of our lead assets

HUTCHMED

HIGHLY DIFFERENTIATED NME PORTFOLIO AND GLOBAL PIPELINE

2 marketed products, 3 NDAs under review & 8-10 reg. studies by mid-2021



[1] In planning; [2] Investigator initiated trials (IITs); [3] Initiated rolling U.S. NDA Dec 2020, target NDA completion H1 2021.

Note: TKI = Tyrosine kinase inhibitor; NDA = New drug application; NSCLC = Non-small cell lung cancer; GC = Gastric cancer; RCC = Renal cell carcinoma; NET = Neuroendocrine tumor; BTC = Biliary tract cancer; ESCC = Esophageal squamous cell carcinoma; SCLC = Small cell lung cancer; CRC = Colorectal cancer; GI = Gastrointestinal; TN = Triple negative.





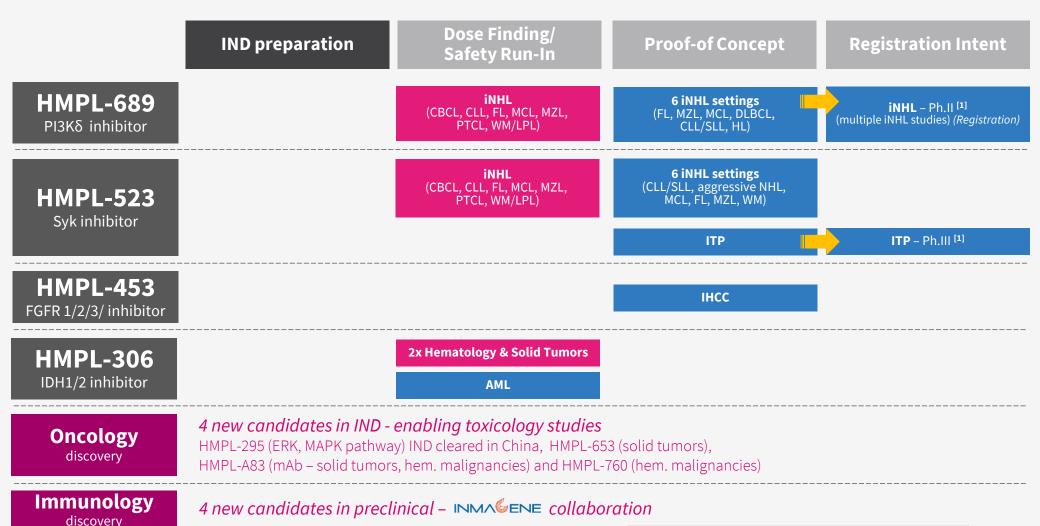




# **Deep NME early pipeline**

Multiple further waves of innovation progressing





[1] In planning, Note: iNHL = Indolent non-Hodgkin's lymphoma; CBCL = Cutaneous B-cell lymphoma; CLL/SLL = Chronic lymphocytic leukemia / Small lymphocytic lymphoma; FL = Follicular lymphoma; MCL = Mantle cell lymphoma; MZL = Marginal zone lymphoma; PTCL = Peripheral T-cell lymphoma; WM = Waldenström's macroglobulinemia; LPL = Lymphoplasmacytic lymphoma; DLBCL = Diffuse large B-cell lymphoma; ITP = Immune Thrombocytopenic Purpura; IHCC= Intrahepatic Cholangiocarcinoma; AML = Acute Myeloid Leukemia.











# **SAVOLITINIB**

A highly selective small molecule inhibitor of MET being developed broadly across MET-driven patient populations in lung cancer, gastric cancer and renal cell carcinoma

# Savolitinib development summary



#### Current status

#### **Strong position in NSCLC**

- MET Exon 14m NDA accepted in May 2020 & priority review;
- Savo/Tagrisso® SAVANNAH enrollment continues apace;
- Planning 2 China NSCLC Ph.IIIs.

### **Renewed RCC strategy**

- Savo monotherapy ~60 pt. SAVOIR data highly encouraging;
- Savo/Imfinzi® combo Planning global Ph.III.

#### Other exploratory studies

- Gastric monotherapy 50% ORR
   Planning China Ph.II registration study;
- Exploring colorectal.

Indication	Treatment	Target Patient	Study Name	Dose Finding / Safety Run-in	Proof-of-concept	Registration
	Savolitinib + Tagrisso®	2L+ EGFRm; Tagrisso ref.; MET+	SAVANNAH			
NECLE	Savolitinib	MET Exon 14 skipping				(NDA accepted) 🜟
NSCLC	Savolitinib + Tagrisso®	2L EGFR TKI ref.; MET+				*
	Savolitinib + Tagrisso®	Naïve MET+ & EGFRm				*
	Savolitinib + Imfinzi® (PD-L1)	MET+ Papillary RCC				*
Kidney	Savolitinib + Imfinzi® (PD-L1)	Papillary RCC **	CALYPSO			
	Savolitinib + Imfinzi® (PD-L1)	Clear cell RCC **	CALYPSO			
	Savolitinib	MET+ Gastric cancer **	VIKTORY			
Gastric & Colorectal	Savolitinib	2L; MET+ Gastric cancer				*
	Savolitinib	MET+ Colorectal cancer **				







# **Savolitinib**

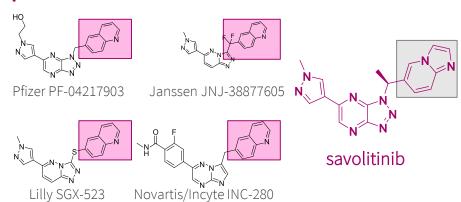


#### Potential first-in-class selective MET inhibitor

# 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.

# 3. Savolitinib design eliminates renal toxicity that first gen. selective MET inhibitors encountered – >1,100 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 [7]

		New Cases (2018)			
Indication	Amplification	Mutation	Over- Expression	Global	China
Gastric	10%	1%	41%	1,033,700	442,300
NSCLC	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 RCC	64%	70-100% [5]	55%	45,400	3,700
Clear Cell RCC	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

RCC = Renal Cell Carcinoma.

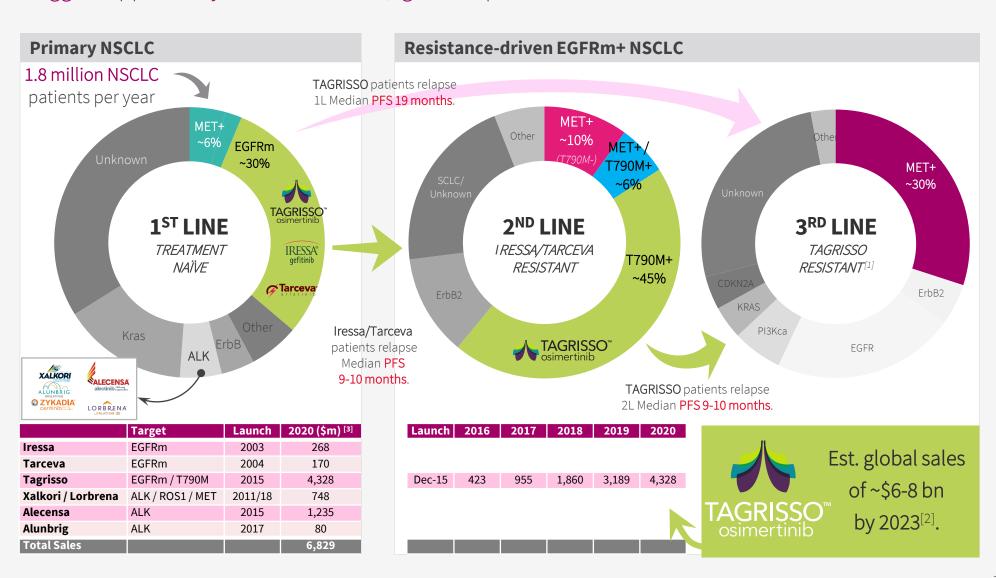
#### 4. AstraZeneca collaboration & 2016 amendment

- \$20m received upfront (Dec 2011);
- \$120m in development/approvals milestones (\$25m received as of Dec '20);
- Several hundred million in commercial milestones:
- Development: AZ pay 100% ex-China & 75% cost in China (HCM 25%);
  - Global PRCC Ph III: HCM contribute \$50m & equal share of additional
- From 9% up to 18% tiered royalty ex-China [8] & 30% flat rate China royalty on all product revenues.

# 4 NSCLC by driver aberration



Biggest opportunity is MET+ (mutant / gene amplified) NSCLC



# **Savolitinib - MET Exon 14 skipping NSCLC**



China's lead selective MET inhibitor

### Competitive landscape outside China:

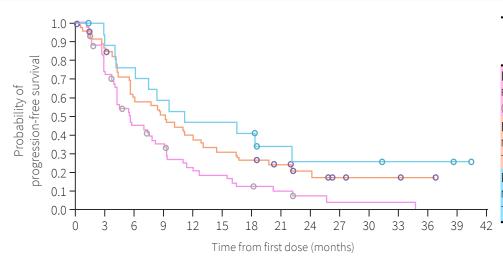
Treatment Line	MET aberration	N	BICR <sup>[1]</sup> ORR (%)	DCR (%)	mDoR (months)	mPFS (months)
Capmatinib [2]						
1L (cohort 5b)	Ex14 skipping	28	<b>68</b> [48, 84]	96 [82,100]	<b>12.6</b> [5.6,NE]	<b>12.4</b> [8.2,NE]
2/3L (cohort 4)	Ex14 skipping	69	<b>41</b> [29,53]	<b>78</b> [67,87]	9.7 [5.6,13.0]	<b>5.4</b> [4.2,7.0]
2L (cohort 6, group 2)	Ex14 skipping	31	<b>48.4</b> [30.2,66.9]	<b>90.3</b> [74.2,98.0]	6.93 [4.17, NE]	<b>8.11</b> [4.17, 9.86]
1L (cohort 5a)	Amp (GCN≥10)	15 <sup>[3]</sup>	40 [16,68]	<b>67</b> [38,88]	7.5 [2.6,14.3]	<b>4.2</b> [1.4,6.9]
2/3L (cohort 1a)	Amp (GCN≥10)	69	<b>29</b> [19,41]	71 [59,81]	<b>8.3</b> [4.2,15.4]	<b>4.1</b> [2.9,4.8]
Tepotinib [4]						
44% 1L, 56% ≥2L	Ex14 skipping	99 <sup>[5]</sup>	<b>46 .5</b> [36.4,56.8]	<b>65.7</b> [55.4,74.9]	<b>11.1</b> [7.2,NE]	8.5 [6.7,11.0]

## **TATTON B & D data - PFS**





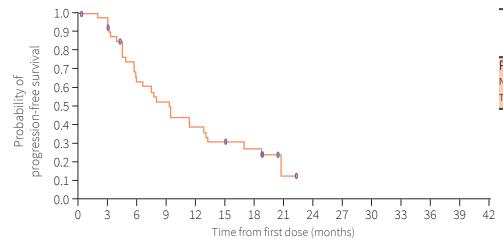
## TAGRISSO® + savolitinib in EGFR TKI refractory NSCLC



	Median PFS, months [95% CI]	Median (range) duration of response, months
Part B1 Prior third-generation EGFR-TKI; (600 mg [1]; n=69)	5.5 [4.1, 7.7]	9.5 [4.2, 14.7]
Part B2 No prior third-generation EGFR-TKI, T790M negative; (600 mg [1]; n=51)	<b>9.1</b> [5.5, 12.8]	<b>10.7</b> [6.1, 14.8]
Part B3  No prior third-generation EGFR-TKI,  T790M positive; (600 mg [1]; n=18)	<b>11.1</b> [4.1, 22.1]	<b>11.0</b> [2.8, NR]

#### Data-cut off date: March 4, 2020

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.



	Median PFS, months [95% CI]	Median (range) duration of response, months
Part D		
No prior third-generation EGFR-TKI,	9.0 [5.6, 12.7]	<b>9.7</b> [4.5,14.3]
T790M negative; (300 mg; n=42)		

#### Data-cut off date: March 4, 2020

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.



# TAGRISSO™ + savo in EGFR TKI refractory NSCLC



#### TATTON B & D data – AEs & tolerability

Event, n (%)	All Part B (n=138) osimertinib 80 mg + savolitinib 600 mg <sup>[1]</sup>	Part D (n=42) osimertinib 80 mg + savolitinib 300 mg [1]
Any AE	138 (100)	41 (98)
Any AE possibly related to savolitinib	124 (90)	32 (76)
AE grade ≥3	86 (62)	21 (50)
AE possibly causally related to study treatment leading to discontinuation of:		
Savolitinib	49 (36)	15 (36)
Osimertinib	24 (17)	8 (19)
Any AE leading to death	7 (5)	2 (5)
Any SAE	67 (49)	16 (38)

<sup>[1]</sup> 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 <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 savolitinib, for Parts B and D, respectively, Han JY, et al. Osimertinib + savolitinib in patients with EGFRm METamplified/overexpressed NSCLC: Phase Ib TATTON Parts B and D final analysis. WCLC January 2021 #FP14.03.

## HUTCHMED

### **TATTON B & D data - AEs & SAEs**

Most common AEs<sup>[1]</sup> independent of causality & SAEs (≥3%)<sup>[2]</sup>

	All Part B	(n=138)	Part D	(n=42)
AE*, n (%)	All	Grade	All	Grade
	grades	≥3	grades	≥3
Nausea	67 (49%)	4 (3%)	13 (31%)	0
Fatigue	48 (35)	6 (4)	4 (10)	0
Decreased appetite	47 (34)	5 (4)	6 (14)	1 (2)
Vomiting	46 (33)	6 (4)	5 (12)	0
Oedema peripheral	44 (32)	3 (2)	8 (19)	0
Diarrhoea	39 (28)	4 (3)	8 (19)	2 (5)
Paronychia	30 (22)	3 (2)	7 (17)	0
Pyrexia	29 (21)	1 (1)	6 (14)	0

AE*, n (%)	All Part B (n=138)		Part D (n=42)		
AL , II (90)	All grades	Grade ≥3	All grades	Grade ≥3	
Rash	26 (19%)	3 (2%)	8 (19%)	0	
Stomatitis	26 (19)	0	4 (10)	0	
Constipation	26 (19)	0	3 (7)	0	
Pruritus	24 (17)	1 (1)	5 (12)	0	
Headache	23 (17)	0	3 (7)	0	
Myalgia	22 (16)	3 (2)	6 (14)	1 (2)	
Cough	22 (16)	0	4 (10)	1 (2)	
AST increased	21 (15)	9 (7)	2 (5)	0	
Pneumonia	15 (11)	7 (5)	7 (17)	5 (12)	

SAE**, n (%)	All Part B (n=138)	Part D (n=42)
Pneumonia	7 (5%)	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

<sup>[1] ≥15%</sup> 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)

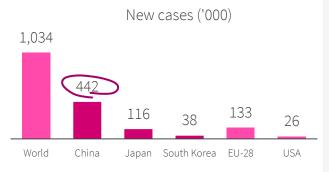
Sequist LV, Han JY, Ahn MJ, et al. Osimertinib plus savolitinib in patients with EGFR mutation-positive, MET-amplified, non-small-cell lung cancer after progression on EGFR tyrosine kinase inhibitors: interim results from a multicentre, open-label, phase 1b study. Lancet Oncol. 2020; S1470-2045(19)30785-5.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

## Savolitinib - MET+ gastric cancer



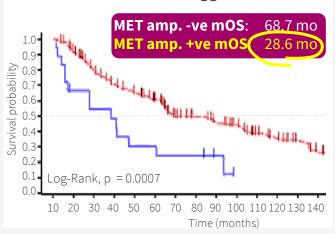
A major problem in east Asia – Japan, Korea & China

1. Gastric (stomach) cancer is the 5<sup>th</sup> most common cancer globally – 782,700 deaths/year

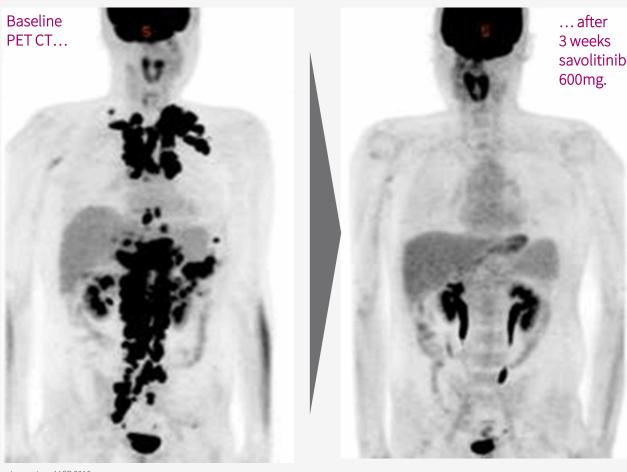


World Cancer Research Fund International, WHO, ACS, NCCR, Lancet, Frost & Sullivan Analysis.

2. MET+ disease is more aggressive [1]



## VIKTORY trial savolitinib arm – male, 34; surgery ruled-out; failed 4-cycles XELOX.



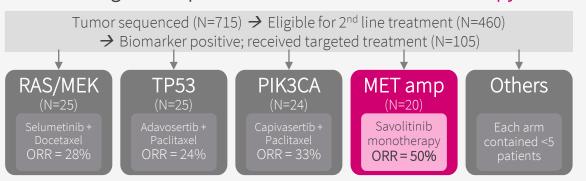
Jeeyun Lee, AACR 2016.

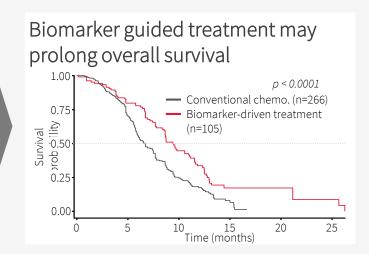
## Savolitinib potential in gastric cancer

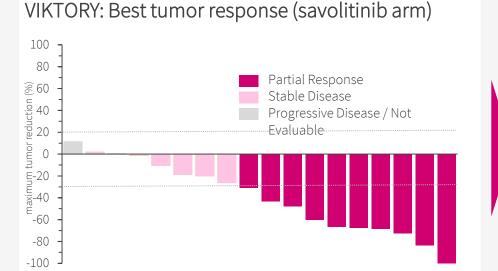


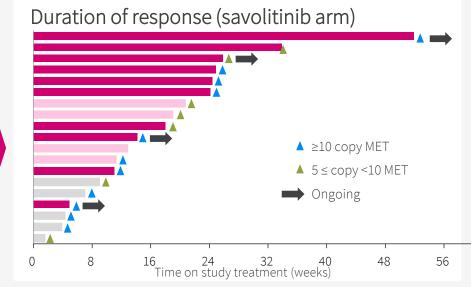
VIKTORY Phase II trial highly promising in MET+

#### VIKTORY: Highest response rate in savolitinib monotherapy arm











A2b

# SURUFATINIB (SULANDA® IN CHINA)

A small molecule inhibitor of VEGFR, FGFR & CSF-1R designed to inhibit tumor angiogenesis and promote the body's immune response against tumor cells via tumor associated macrophage regulation

## Surufatinib development summary



#### Current status

#### **China NET**

- Non-pancreatic NET NDA approved Dec 2020;
- Pancreatic NET NDA accepted.

#### **Global NET**

- U.S. NDA in late 2020 [1];
- Fast Track Designations for both pNET & non-pNET;
- EU MAA planned for mid-2021.

#### **PD-1 combos**

- TUOYI® (Junshi) Ph.II (in 9 solid tumor indications);
- TYVYT® (Innovent);
- Tislelizumab (BeiGene) [2].

#### **Biliary Tract Cancer**

Ph.II/III underway; interim analysis for futility in 2021

Indication	Treatment	Target Patient	Study Name	Dose Finding / Safety Run-in	Proof-of-concept	Registration
	Surufatinib	NET				(US NDA sub. started)
NET	Surufatinib	NET				(EU MAA planned)
NEI	Surufatinib	Pancreatic NET	SANET-p			(NDA accepted) 🜟
	Surufatinib	Non-Pancreatic NET	SANET-ep			(Approved) 🌟
втс	Surufatinib	Biliary tract cancer (BTC)				
ВІС	Surufatinib	2L; chemo ref. BTC				
STS	Surufatinib	Soft tissue sarcoma				
	Surufatinib + TUOYI® (PD-1)	NEN, ESCC, BTC				
22.4	Surufatinib + TUOYI® (PD-1)	SCLC, GC, Sarcoma				
PD-1 Combos	Surufatinib + TUOYI® (PD-1)	TC, EMC, NSCLC				
	Surufatinib + TYVYT® (PD-1)	Solid tumors				
	Surufatinib + tislelizumab (PD-1)	Solid tumors		*		







### **Surufatinib**



Overview of NET – ~141,000 patients in the U.S. [1][2][3]

#### What are neuroendocrine tumors ("NET")?

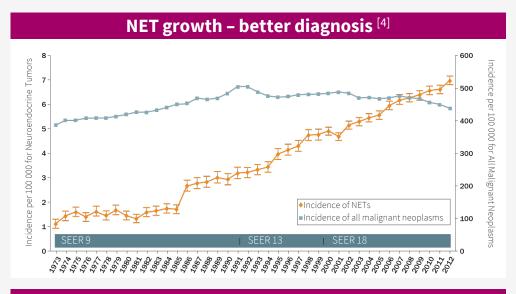
- ~2% of all malignancies.
- Tumor begins in the specialized cells of the body's neuroendocrine system. Cells have traits of both hormone-producing endocrine cells & nerve cells.
- Found throughout the body's organs. Most NETs take years to develop but some can grow fast.

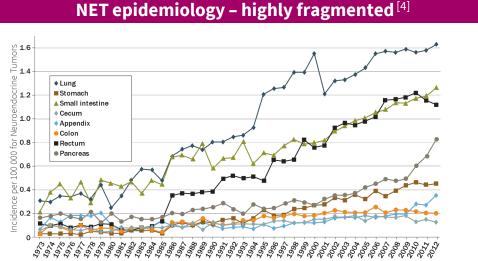
#### Hormone-related symptoms [1]

• 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.





## **High-level NET landscape**



Long-term disease – rapid deterioration in later stages [1][2][3]

#### Grade 1 (G1) NET Localized / Regional ~8-35% NET patients Somatostatin Analogue - Functional NET -**Treatment** – modulate/ Hormone related control symptoms related to symptoms: hormone overproduction & 94% flushing tumor growth: 78% diarrhea Octreotide: \$1.4b revenue (2020) 53% heart plaque Lanreotide: \$1.5b revenue (2020) 51% cramping Symptoms allow mOS: early diagnosis 16.2 yrs

Well Differentiated

Ki-67 Index ≤2; Mitotic Count <2





## G1/2 Advanced NET [1] (Ki-67 Index 0-20)









### Global opportunity in lung/other NETs & China wide-open

Site		est. %	Octreotide LAR	Lanreotide autogel	<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 [2]	Historical Ph. II  SSR over expression			RADIANT-4 [3]	SANET-ep
	Small bowel / appendix	9%	PROMID	CLARINET [2]	NETTER-1			RADIANT-4 [3]	SANET-ep
GI Tract	Colon & Rectum	31%		CLARINET [2]	Historical Ph. II SSR over expression			RADIANT-4 [3]	SANET-ep
Pancreas		6%		CLARINET [2]	Historical Ph. II  SSR over expression	Historical	PHASE III	RADIANT-3 [3]	SANET-p
Lung		20%						RADIANT-4 [3]	SANET-ep
Other	Other	~17%							SANET-ep
	Unknown Primary	~10%						RADIANT-4 [3]	SANET-ep

## ~141,000 NET patients in U.S. [1][2]

## HUTCHMED

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

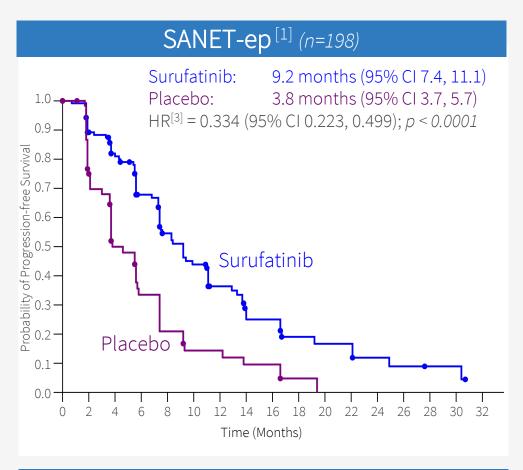
	Somatostatin Based Therapies			Kinase Inhibitor Therapies			
	Sandostatin° LAR (octreotide)	Somatuline Depot° (lanreotide)	Lutathera° ( <sup>177</sup> Lu-Dotatate)	Afinitor° (everolimus)	Sutent° (sunitinib)	Surufatinib (Approved in China)	
2020 Sales	\$1.4bn	\$1.5bn	\$0.4bn	\$1.1bn	\$0.8bn	-	
MOA [3]	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	Intravenous inj. (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>GEP-NETs: unresectable, well or moderately diff., (locally adv. or meta) GEP-NETs to improve PFS.</li> <li>Carcinoid Syndrome: 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., non-functional NET (unresectable, locally adv. or meta). Not for functional carcinoid tumors. <sup>[4]</sup></li> </ul>	differentiated pNET (unresectable locally adv. or meta).	<ul> <li>2 positive RCTs in pNET &amp; epNET in China</li> <li>epNET NDA approved in China; pNET under review</li> <li>US NDA filing started YE20.</li> </ul>	
Non-NET indication/s	Acromegaly; watery diarrhea from VIPomas.	Acromegaly.		<ul> <li>Adv. HR+ HER2-n breast cancer; adv. 2L RCC; renal angiomyolipoma and TSC.</li> </ul>	2L GIST; adv. RCC; high risk of recurrent RCC.		

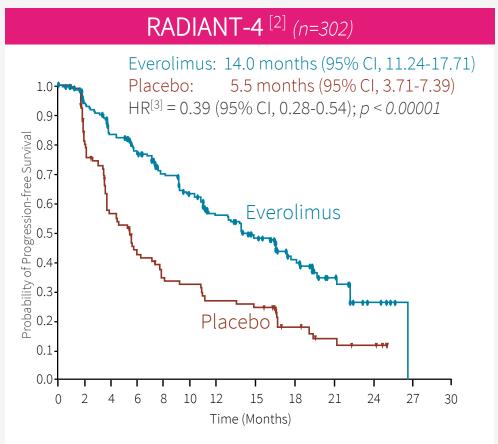
	Sandostatin°/ Placebo	Somatuline Depot° / Placebo	Lutathera° + Sando. LAR / Sando. LAR		tor <sup>®</sup> / :ebo	Sutent°/ Placebo		fatinib / acebo
mPFS (mo.) primary EP	14.3 / 6.0	NR / 18.0	NR / 8.5	pNET 11.0 / 4.6	Lung & GI NET 11.0 / 3.9	pNET: 11.4 / 5.5	Ph III pNET 10.9 / 3.7	Ph III non-pNET 9.2 / 3.8
HR	0.34	0.47	0.21	0.35	0.48	0.42	0.49	0.33
(p-value)	0.000072	<0.001	<0.0001	<0.001	<0.001	<0.001	0.0011	<0.0001
ORR	2% / 2%	NR	18%/3%	5% / 2%	2% / 1%	9% / 0%	19% / 2%	10%/0%
DCR	69% / 40%	NR	95% / 76%	73% / 51%	81% / 64%	72% / 60%	81%/66%	87% / 66%
Pivotal	PROMID	CLARINET	NETTER-1	RADIANT-3	RADIANT-4	A6181111	SANET-p	SANET-ep

## **G1/2 Advanced extra-pancreatic NET**



Investigator assessed median PFS





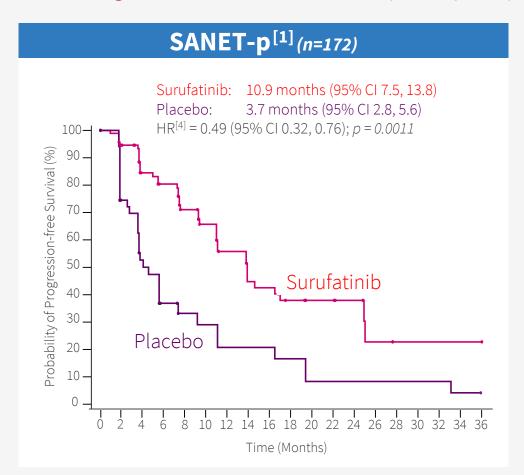
SANET-ep Primary (1°) endpoint was Investigator mPFS BIIRC [4] mPFS for supportive analysis not 1° or 2° endpoint RADIANT-4 Primary (1°) endpoint was BIIRC [4] mPFS Investigator mPFS not 1° or 2° endpoint

[1] Xu et al. "Surufatinib in advanced extrapancreatic neuroendocrine tumours (SANET-ep): a randomised, double-blind, placebo-controlled, phase 3 study." Lancet Oncol 2020. Published online September 20, 2020. https://doi.org/10.1016/S1470-2045(20)30496-4; [2] Yao et al. "Everolimus for the treatment of advanced, non-functional neuroendocrine tumors 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).

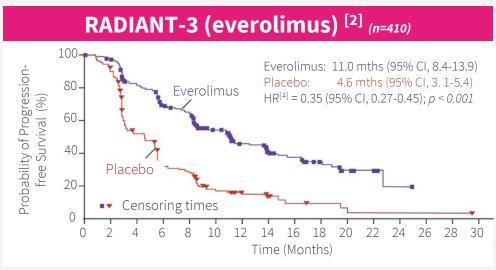
## HUTCHMED

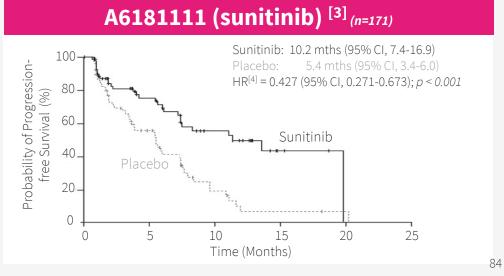
## **G1/2 Advanced pancreatic NET**

Investigator assessed median PFS (primary endpoints)



[1] Xu et al. "Surufatinib in advanced pancreatic neuroendocrine tumours (SANET-p): a randomised, double-blind, placebo-controlled, phase 3 study." Lancet Oncol 2020. Published Online September 20, 2020 https://doi.org/10.1016/S1470-2045(20)30493-9; [2] Yao et al. Everolimus for advanced pancreatic neuroendocrine tumors" N Engl J Med. 2011;364(6):514–23 DOI: 10.1056/NEJMoa1009290; [3] Raymond et al. Sunitinib malate for the treatment of pancreatic neuroendocrine tumors [published correction appears in N Engl J Med. 2011 Mar 17;364(11):1082]. N Engl J Med. 2011;364(6):501-513 DOI: 10.1056/NEJMoa1003825; [4] P-value from SANET-p is obtained from the stratified one-sided log-rank test; Hazard ratio is obtained from stratified Cox model; CI, confidence interval; HR, hazard ratio.





### Surufatinib vs. everolimus and sunitinib

6%

2%

8%

14%

RADIANT-4 [2

(n=302)

65%

35%

74% (73%: 75%)

26% (27%: 26%)

61%

25%

none

55%

n/a

n/a

HUTCHM

Broader range of tumor origins & later-stage patients

	Asia/China Extra- Pancreatic NET	SANET-ep [1] (n=198) (surufatinib vs placebo)
	Tsai et al. 2013	
Gastrointestinal Tract	58%	47%
Rectum Stomach Small Intestine Other GI	30% 7% 19% 3%	27% 10% 8% 3%
<b>Lung</b> Other Organ Site Thymus Liver	22%	12% 28% 7% 6%

**NON-PANCREATIC NET** 

SANET-ep [1]

(n=198)

16%

84%

60% (56%: 67%)

40% (44%: 33%)

67%

40%

10%

32%

34%

66%

Non-Pancreatic **Tumor Origin** 

Pathology grade

ECOG PS 0:1

Prior systemic

Number of organs ≤2

treatment

Mediastinum

**Unknown Origin** 

Other

Grade 1

Grade 2

control)

control)

PS 0 (treatment:

PS 1 (treatment:

Chemotherapy

Somatostatin

≥3 or unknown

Analogues

Targeted therapy

Any Prior Treatment

Adrenal Gland

	U.S. Extra- Pancreatic NET	RADIANT-4 <sup>[2]</sup> (n=302) (everolimus vs placebo)
	Yao et al. 2008	
Gastrointestinal Tract	50%	58%
Rectum Stomach Small Intestine Other GI	33% 8% 6% 4%	13% 4% <b>34%</b> 7%
Lung Thymus	21%	30% 1%
Unknown Origin		12%

PANCREATIC NET			
<b>SANET-p</b> <sup>[3]</sup> (n=172)	<b>RADIANT-3</b> <sup>[4]</sup> (n=410)	<b>A6181111</b> <sup>[5]</sup> (n=171)	
12% <b>88%</b>	<b>83%</b> 17%	n/a n/a	
67% (65% : 73%)	66% (67%: 66%)	55% (62% : 48%)	
33% (35% : 27%)	31% (30:32%)	44% (38% : 51%)	
<b>66%</b> 26% 9%	50% none	<b>69%</b> 66% none	

50%

64%

36%

36%

64%

36%

#### **SANET-ep** Enrolled more pts with poor prognosis.

		Survival
Primary Site	mOS	Rate @ 5-yr
Rectum	2.8y	28%
Stomach	2.4y	32%
Small Intestine	8.6y	69%

#### **RADIANT-4**

Did not enroll other extra-pancreatic **NET organ sites** incl. but not limited

Throat Thyroid Ovarv Kidnev Mediastinum Adrenal gland Retroperitoneal Ampulla vater Parathyroid Carotid body gland Liver

Broader pt. coverage.

SANET-ep

#### **Surufatinib**

Later-stage patients, more heavily pre-treated (incl. with targeted therapy) & weaker physical status. Likely due to later diagnosis in China & availability of everolimus.

involved Source: 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;

44%

49%

51%

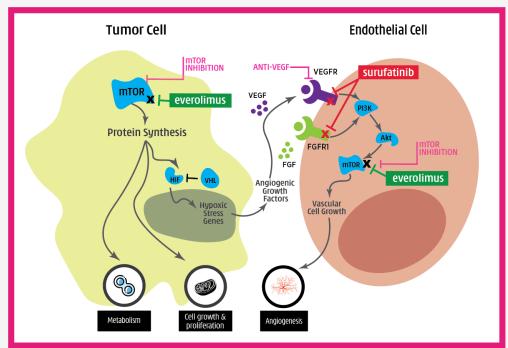
Colon-rectum in Tsai et al. (2013) report; Colon approximately 8% in Asian series (Shebani KO et al. (1999)); Colon-rectum in Yao et al. (2008) report; Colon approximately 4-7% in US/EU series (Niederle B et al. (2016)). [1] Xu et al. https://doi.org/10.1016/S1470-2045(20)30496-4; [2] Yao et al. doi: 10.1016/S0140-6736(15)00817-X; [3] Xu et al. https://doi.org/10.1016/S1470-2045(20)30493-9; [4] Yao et al. DOI: 10.1056/NEJMoa1009290; [5] Raymond et al. DOI: 10.1056/NEJMoa1003825

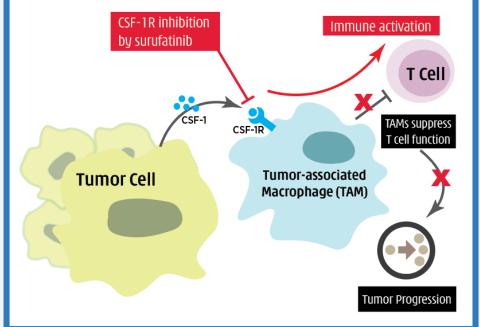
## HUTCHMED

## 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.







A2c

## FRUQUINTINIB (ELUNATE® IN CHINA)

A highly selective small molecule inhibitor of VEGFR 1/2/3 designed to improve kinase selectivity to minimize off-target toxicity and thereby improve tolerability

## Fruquintinib development summary



#### Current status

#### **ELUNATE® CRC China**

WRDL inclusion from January 1, 2020.

#### **CRC GLOBAL**

- U.S. Ph.Ib/II completed;
- FRESCO-2 Ph.III initiated in U.S., EU & Japan;
- US FDA Fast Track Designation.

#### **FRUTIGA Gastric Ph.III**

- 2<sup>nd</sup> Interim analysis in June 2020 complete;
- Increasing enrollment to 700-pts.

#### PD-1 combos

- TYVYT® (Innovent) Ph.II (in 5 solid tumor indications);
- Tislelizumab (BeiGene)\*;
- Geptanolimab (Genor).

Indication	Treatment	Target Patient	Study Name	Dose Finding / Safety Run-in	Proof-of-concept	Registration
CDC	Fruquintinib	Colorectal cancer (CRC)	FRESCO-2			
CRC	Fruquintinib	≥3L; chemotherapy ref. CRC	FRESCO			(Marketed) 🜟
Gastric	Fruquintinib + TAXOL®	2L gastric cancer	FRUTIGA			
Breast	Fruquintinib	Breast cancer				
	Fruquintinib + TYVYT® (PD-1)	CRC, EMC, RCC, HCC				
	Fruquintinib + TYVYT® (PD-1)	GI tumors				
PD-1	Fruquintinib + geptanolimab (PD-1)	CRC				
Combos	Fruquintinib + geptanolimab (PD-1)	NSCLC				
	Fruquintinib + tislelizumab (PD-1)	TN breast cancer		*		
	Fruquintinib + tislelizumab (PD-1)	Solid tumors		*		









### Fruquintinib & surufatinib both unique VEGFR TKIs

...potentially ideal VEGFR combos for immunotherapy

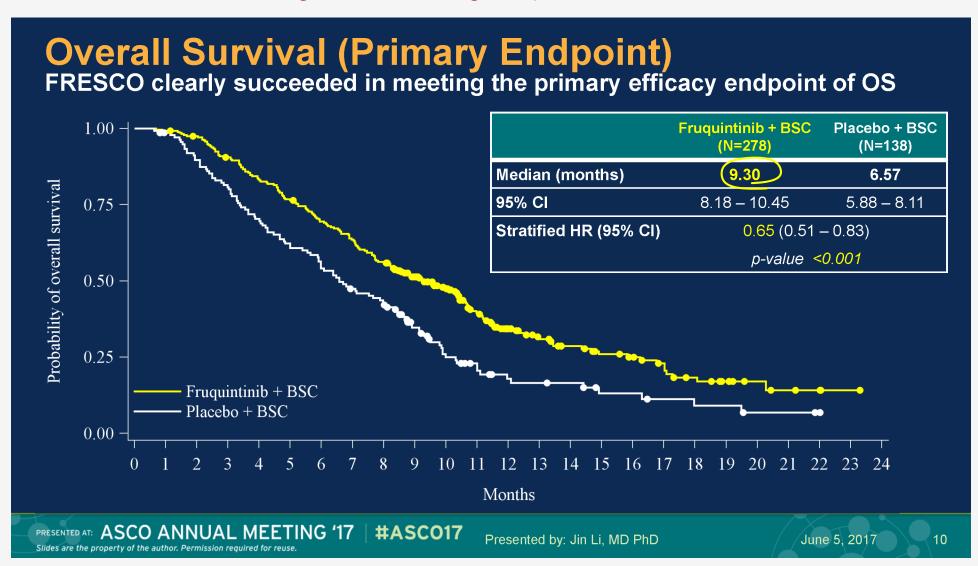
TKI	1st Generation 2		2no	d Generation		Next Generation		
Selectivity		Multiple targets		Relatively selective		re	Highly selective	Selective angio-immuno kinase inhibitor
Inhibitors	Sutent®	Nexavar®	Focus V®	Fotivda®	Lenvima®	Inlyta®	Fruquintinib	Surufatinib
Status	Launched	Launched	Launched	Launched	Launched	Launched	Launched	Approved
VEGFR1 (nM)	2	26	27	30	22	3	33	2
VEGFR2 (nM)	9	90	0.2	6.5	4	7	25	24
VEGFR3 (nM)	19	20	0.7	15	5	1	0.5	1
Phos-KDR (nM)	10	30	0.1-1	0.16	0.8	0.2	0.6	2
Other kinases (IC50 < 100nM)	PDGFRα PDGFRβ c-Kit Flt3 Ret CSF-1R	Raf-1 b-raf Flt3 P38 c-Kit Ret	PDGFRα PDGFRβ FGFR1-4 c-Kit	PDGFRα PDGFRβ EphB2 c-Kit Tie2	PDGFRα PDGFRβ FGFR1-4 Ret c-Kit	PDGFRα PDGFRβ c-Kit	none	CSF-1R FGFR1 FLT3 TrkB
First Patent Evniration				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

## HUTCHMED

## Fruquintinib – 3L+ colorectal cancer

Launched in China, initiated global Phase III reg. study



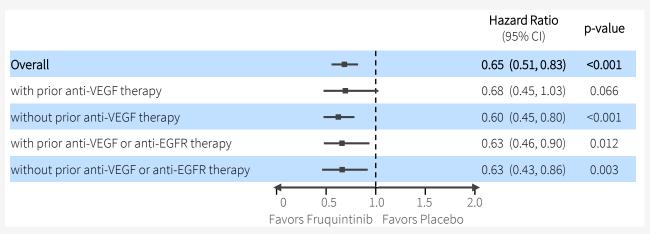
## **Efficacy advantage**





Third Line	FRESCO <sup>[1]</sup> Mainland China		CONCUR  Chinese Patients (Mainland China, Hong Kong, Taiwan) [2]		CONCUR  Mainland China, Hong Kong, Taiwan, Vietnam, South Korea		CORRECT	
Third-Line Metastatic Colorectal cancer								
Treatment arms	ELUNATE®	Placebo	STIVARGA®	Placebo	STIVARGA®	Placebo	STIVARGA®	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% +49	0.9 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	.9 1.8	2.0 +0.	3 1.7	3.2 +1	.5 1.7	1.9	+0.2 1.7
						10	0%AVASTIN®prior use	,
Median Overall Survival (mOS) (mo.)	9.3 +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® efficacy vs. Stivarga® in Chinese metastatic CRC pts;
- Advantage for ELUNATE® 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)



## **Stivarga® tox limitations**





Fruqu	uintinib	Capsul	es
-------	----------	--------	----

	ELUNATE® Fruquintinib Capsules	Stivarga® (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-RAFV600E	>10,000	19

#### Stivarga® liver toxicity black-box warning:

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

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

#### WARNING: HEPATOTOXICITY

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

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

#### **ELUNATE®** superior safety – advantage especially for liver mets patients

### **NRDL**



#### 2020 accessible pricing

## **Epidemiology** China Annual Incidence 380,000 patients [1] Surgery ~15% 1st-line treated 2<sup>nd</sup>-line treated 3<sup>rd</sup>-line treated >55,000 patients [2]

#### 2020 estimated penetration:

- ~39,500 cycles used (OOP & PAP);
- Average 4.7 months per patient;
- ~8,400 patients paid for ELUNATE®;
- Representing ~15% penetration.

#### **National Reimbursement Drug List (NRDL)**

#### Effective Jan 1, 2020:

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

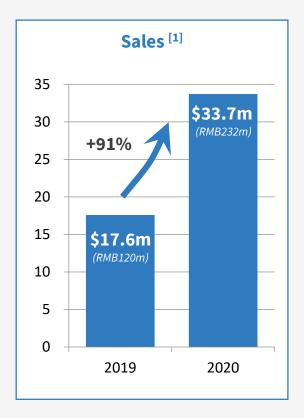
Costs p	er cycle <i>(all US\$)</i> <sup>[3]</sup>	With Medical Insurance	Without Medical Insurance
ELUNATE® (fruquintinib)	Pre-NRDL (without PAP) Post-NRDL	3,260 1,180	3,260 1,180
	3L CRC Pts Out-of- Pocket Cost	~350 [5]	~1,180
STIVARGA® (regorafenib)	3L CRC Pts Out-of- Pocket Cost	~670 [5]	~2,220

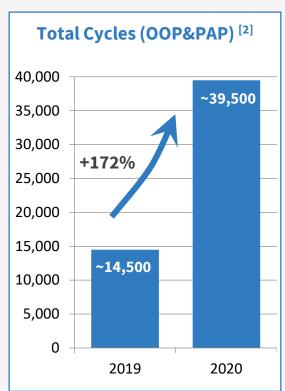
### FY 2020 performance





ELUNATE® early progress – set to expand rapidly





#### **2020 Lilly Amendment**

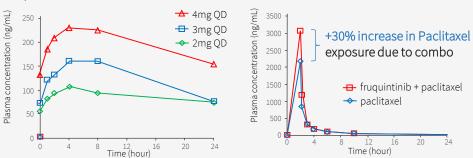
- Starting October 1, 2020, HUTCHMED took on all medical detailing, promotion & local/regional marketing activities across all of China:
- Lilly expected to pay HUTCHMED an est. 70%-80% of ELUNATE® sales in the form of royalties, mfg. costs & service payments [3];
- No upfront payment by HUTCHMED was made to secure these rights.

## Gastric combo with paclitaxel



#### Phase 2 results supports ongoing Phase III FRUTIGA

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



Waterfall Dints of Bost Desnonse

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

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

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

			wa	iterrali	ii Plots of Best Response
40	2mg (n=3)	3mg (n=3)	4mg dos	se findi e (n=8)	2 Ama doca avnancion crada in Figi
20	•				
-20 -30			<b>V</b>		
-40	•			ч	▼ Progressive Disease (PD) ▲ Non-Evaluable (NE)
-60					
90					paclitaxel alone ORR ~20%

4 AE profile in-line with expectations. Neutropenia – a paclitaxel AE – with 57.9% Grade >3 AEs. Similar to 60% seen ramcirumab (VEGF mAb) RAINBOW study paclitaxel combo in 2L gastric.

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%)

## **FALUCA – Third-line NSCLC Monotherapy**



#### Presented at WCLC 2019

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

- <u>Met all</u> secondary endpoints: mPFS; ORR; DCR; & DoR [1];
- 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.

#### Efficacy Endpoints (Intent-to-Treat) [2]

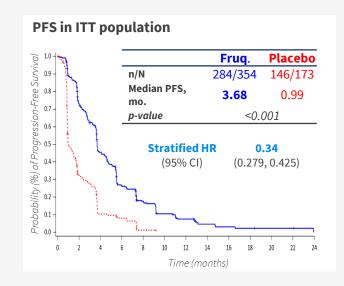
	Fruq. (N=354)	Placebo (N=173)	p-value
mOS (mths)	8.94	10.38	0.841
mPFS (mths)	3.68	0.99	<0.001
ORR	<b>13.8%</b> (49)	0.6% (1)	<0.001
DCR	<b>66.7%</b> (236)	24.9% (43)	<0.001

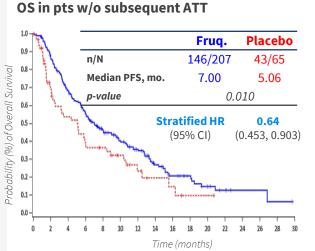
#### Good safety; most Grade ≥3 TEAEs targetrelated & clinically manageable.

Patient (%)	Fruq. (N=354)	Pbo (N=173)
TEAE ≥ 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

## 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









## **HMPL-689 & HMPL-523**

Targeting B-cell signaling for hematological cancers and immunology

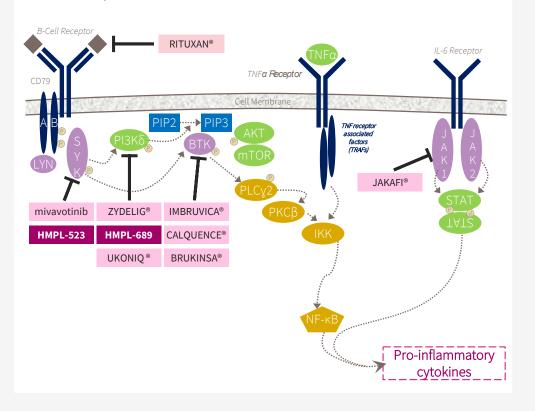
## HMPL-689 (PI3Kδ) & HMPL-523 (Syk)



Exciting targets emerging – our next wave of innovation

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

2020 sales: IMBRUVICA® \$6.6bn; ZYDELIG® \$0.1bn; JAKAFI® \$3.3bn; & RITUXAN® \$3.4bn [1][2].



## HMPL-689 (PI3Kδ inhibitor) Phase I/Ibs in China, US & EU ongoing

Designed to be a best-in-class inhibitor of PI3Kδ

- 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.

## HMPL-523 (Syk inhibitor) Large Phase Ib expansion in Australia & China

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

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

### **HMPL-689** – finding major room for improvement



Safety profiles of current PI3Kδ inhibitors are not good

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

Compound	Company	Indication	Status	Issue
Zydelig <sup>®</sup> idelalisib – PI3Kδ	Gilead	Relapsed CLL/SLL, FL	Approved	BOXED WARNING: FATAL AND SERIOUS TOXICITIES: HEPATIC, SEVERE DIARRHEA, COLITIS, PNEUMONITIS, INFECTIONS, and INTESTINAL PERFORATION
Aliqopa <sup>®</sup> copanlisib – PI3Κα/δ	Bayer	Relapsed FL	Approved [1]	BOXED WARNING: FATAL AND SERIOUS TOXICITIES: INFECTIONS, DIARRHEA OR COLITIS, CUTANEOUS REACTIONS, and PNEUMONITIS Need to spare PI3Kα
Copiktra <sup>®</sup> duvelisib – PI3Kγ/δ	Secura Bio/ CSPC <sup>[2]</sup>	Relapsed or refractory CLL/SLL Relapsed or refractory FL Peripheral T-cell lymphoma	Approved Approved [1] Phase II enrolling	BOXED WARNING: FATAL AND SERIOUS TOXICITIES: INFECTIONS, DIARRHEA OR COLITIS, CUTANEOUS REACTIONS, and PNEUMONITIS Need to spare PI3Kγ
Ukoniq® Umbralisib - PI3Kδ	TG Therapeutics	Previously treated MZL Previously treated FL Previously treated NHL, CLL	Approved [1] Approved [1] Phase IIb/III	Gastrointestinal & liver AEs
Parsaclisib	Incyte/ Innovent	FL, MZL, MCL Refractory myelofibrosis Autoimmune hemolytic anemia	NDA filing H2-2021 Phase III Phase II	Pending 12 months follow-up data from last responder [3]  Phase 2 studies required prophylaxis for pneumocystis  jirovecii pneumonia (PJP)
Zandelisib PI3Kδ	MEI/Kyowa Hakko Kirin	Relapsed or refractory FL  B-Cell Malignancies	Phase II (for pot. AA) Phase I/Ib	Progressing with intermittent dosing to mitigate immune related toxicities; all patients underwent prophylaxis for pneumocystis jirovecii pneumonia (PJP) [4]



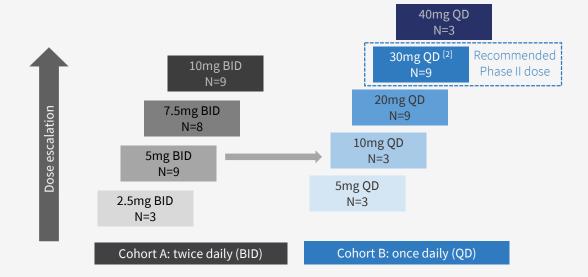
## HMPL-689 – designed to be better

Intent to improve safety...

#### **HMPL-689 – Advantages**

- Improved isoform selectivity sparing PI3K $\gamma$  & PI3K $\alpha$ .
- Improved potency at whole blood level over five-fold more potent than Zydelig® to cut compound related toxicity.
- Improved PK properties particularly efflux & drug/drug interaction due to CYP inhibition / induction, critical for combo therapy.

#### Dose escalation schema



#### Manageable toxicity profile [1]

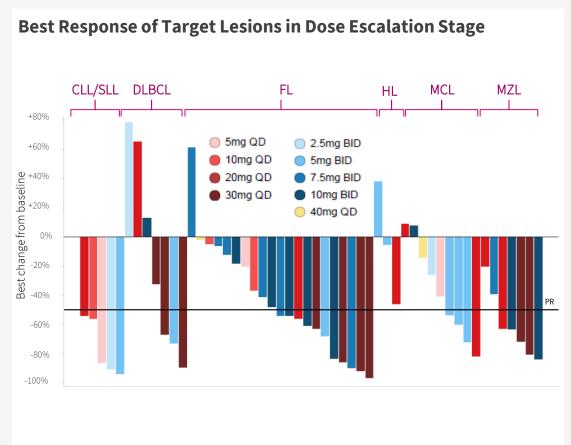
Treatment-emergent AEs	All dos	es (N=56) Grade
occurred in ≥ 5% of patients	grade	Siaue ≥3
Neutropenia	43%	11%
Leukopenia	29%	4%
ALT increased	27%	2%
Pneumonia	25%	16%
AST increased	21%	2%
Lipase increased	20%	5%
Cough	18%	-
Anemia	16%	-
Blood bilirubin increased	16%	2%
Mouth ulceration	14%	-
Pyrexia	14%	-
Upper respiratory tract infection	14%	-
Bilirubin unconjugated increased	13%	2%
Asthenia	11%	-
Blood creatinine increased	11%	-
Constipation	11%	-
Hyperglycemia	11%	-

[1] ASH 2020 Abstract #1135.

### **HMPL-689 – dose escalation**

## HUTCHMED

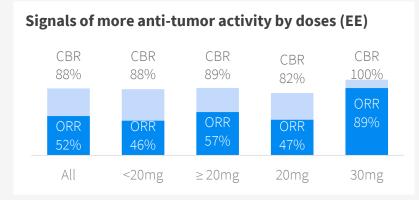
#### ...While maintaining efficacy



CLL/SLL: chronic lymphocytic leukemia/small lymphocytic lymphoma; FL: follicular lymphoma; MZL: marginal zone lymphoma; MCL: mantle cell lymphoma; DLBCL: diffuse large B cell lymphoma; HL: Hodgkin's lymphoma; NHL: non-Hodgkin's lymphoma.

NE: 2 DLBCL pts EOT due to AE (5mg BID) & voluntary withdraw (7.5 mg BID); 1 FL pt EOT due to AE (20 mg QD) before 1st tumor evaluation. 1 CLL arrive PR based on target lesion, as lymphocyte cell count increased assessed PD at C3D1.

11 (4-22)
37
34
11
48% (35-62)
82% (70-91)
5.6 months (0.7-23.2)
1.8 months (1.8-1.9)
9.2 months (3.9-NA)
10.1 months (5.5–15.7)
40% (27–57)

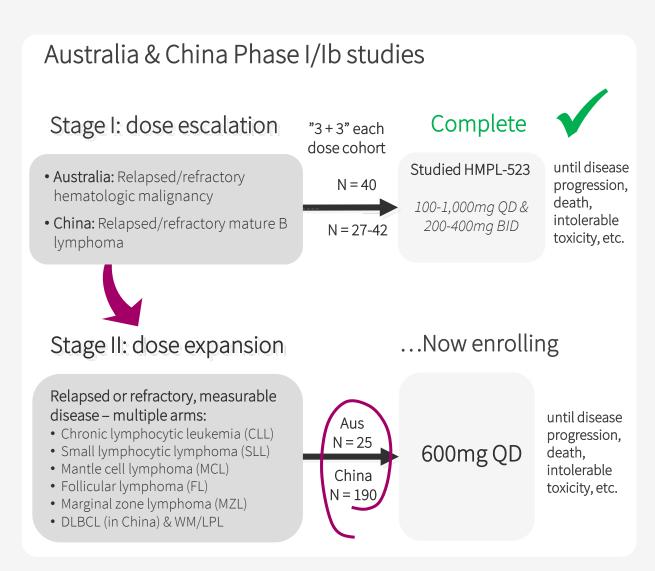


## HMPL-523 (Syk) in hematological cancer



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. Ib dose expansion study, total n>200, underway in ~30 active sites in Australia & China;
- U.S./E.U. Phase I/Ib enrollment underway, with 13 sites enrolling;
- These Phase I/Ib data will inform China registration study decisions.



[1] RP2D = Recommended Phase II doses.

A2e

**NEXT WAVE OF INNOVATIONS** 

## What is next from discovery?



Differentiated assets against multiple targets

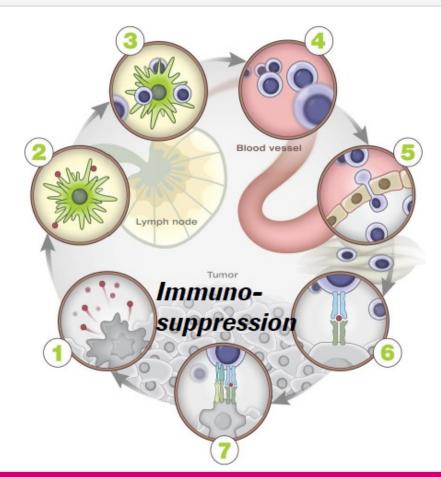
#### Priming & activations

Multiple mAb programs

#### Antigen release

- MET (savolitinib)
- EGFR (epitinib)
- Syk (HMPL-523)
- PI3Kδ (HMPL-689)
- FGFR (HMPL-453)
- IDH 1/2 (HMPL-306)
- ERK 1/2 (HMPL-295)

Multiple small molecule programs



#### Anti-angiogenesis

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

#### Negative regulators

- Treg (HMPL-689)
- CSF-1R (surufatinib)

Multiple small molecule & mAb programs

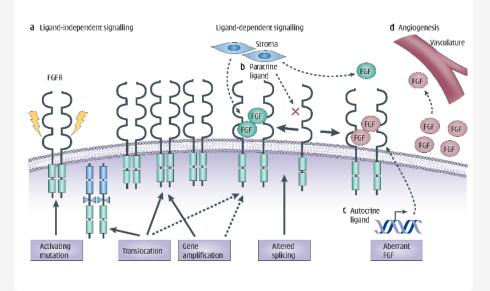
Creating highest-quality range of assets against novel targets for use in combos



### HMPL-453 - Phase II in China initiated

#### 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 w/multiple tumor types harboring low incidence.

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

**A3** 

FURTHER CORPORATE INFORMATION

### **Group Structure**

Main Entities / Offices



Hutchison China MediTech Group Level (Nasdaq/AIM: HCM)



Consolidated

Non-Consolidated

### Oncology/Immunology

Discovery, development, manufacturing & commercialization of novel oncology & immunology therapeutics (Ownership: 99.8%)

#### Shanghai

Discovery and development

Commercial

#### New Jersey

Clinical development & regulatory affairs

#### Suzhou

GMP-certified manufacturing

#### Beijing

Australia

E.U.

Others

#### Other Ventures

#### **Hutchison Sinopharm**

Rx Drug Commercialization

Partner: Sinopharm Group (HCM 51%)

#### Shanghai Hutchison Pharmaceuticals

Rx Drug Mfg & Commercialization

Partner: Shanghai Pharma (HCM: 50%)

#### Hutchison BYS<sup>[1]</sup>

Over-the-counter drugs

Partner: Guangzhou Pharma (HCM: 40%)

Consumer Healthcare<sup>[2]</sup>

[1] Hutchison Whampoa Guangzhou Baiyunshan Chinese Medicine Company Limited (HCM holds 50.0% through its 80.0% owned subsidiary Hutchison BYS (Guangzhou) Holding Limited), a JV with Guangzhou Baiyunshan Pharmaceutical Holdings Co. Limited which holds the other 50.0%.

[2] Mainly Hutchison Hain Organic Holdings Limited, a JV with The Hain Celestial Group, Inc.

## HUTCHMED

### Our Other Ventures have substantial value

- HUTCHMED's Other Ventures continue to perform well relative to our peer group.
- The market value, based on China Pharma median PE multiples, is approximately \$1.8 billion.<sup>[1]</sup>
- Given our share in the JVs, HUTCHMED's share of this value is approximately **\$0.9 billion**.

			NET SALES			NET IN	СОМЕ		VALUATION	[3]
(US\$ millions)	Code	<b>2019</b> Jan-Jun	<b>2020</b> Jan-Jun	<b>19-20</b> Growth	<b>2019</b> Jan-Jun	<b>2020</b> Jan-Jun	19-20 Growth	<b>2020</b> Margin	Market Cap.	P/E
HUTCHMED Other Ventures Subsidiaries/JVs <sup>[2]</sup>		367.1	365.2	-1%	57.0	62.4	9%	17%	n/a	n/a
Livzon Pharma	000513	705.6	727.9	3%	119.2	190.1	59%	26%	4,545	23
CR Double-Crane Pharma	600062	695.1	592.4	-15%	92.3	80.1	-13%	14%	1,726	12
Kunming Pharma	600422	536.6	489.2	-9%	34.4	32.4	-6%	7%	914	15
Zhejiang Pharma	600216	512.2	504.1	-2%	38.6	58.3	51%	12%	2,103	28
Tianjin Zhong Xin Pharma	600329	504.8	470.1	-7%	50.6	47.7	-6%	10%	1,624	21
Zhejiang Hua Hai Pharma	600521	379.0	472.2	25%	50.2	86.7	73%	18%	5,590	40
Shandong Xin Hua Pharma	000756	446.1	469.4	5%	23.4	26.9	15%	6%	666	17
Jiangsu Kang Yuan	600557	323.2	221.0	-32%	35.1	21.3	-39%	10%	855	19
Zhuzhou Qian Jin Pharma	600479	241.7	240.5	0%	14.8	13.6	-8%	6%	523	19
Jiu Zhi Tang	000989	241.2	261.9	9%	25.0	27.9	12%	11%	1,017	29
Peer Group Median (10 Comps. excl. HUTCHMED)		475.5	471.1	-1%	36.8	40.1	9%	9%	1,321	20

Peer Group: 10 companies (excl. HUTCHMED) selected are ALL listed and profitable mainland Chinese OTC/Rx pharma manufacturing companies, with a focus on similar product types, and 2020 Jan-Jun Net Sales in the ~\$200-750 million range.

Source: Company data, CICC

<sup>[1]</sup> Peer group/China Pharma multiple of 20x 2020 actual Net income after tax of \$90.2m, excluding one-time land compensation; [2] Total aggregate PRC domestic results of HUTCHMED's 6 Other Ventures companies (HBYS, SHPL, Hutchison Sinopharm, HHO, HHL & HCPL); [3] Market Capitalization and Price Earnings Ratios as at February 19, 2021: Trailing Twelve Month PE weighted averaged based on market capitalization.



### **Non-GAAP Financial Measures & Reconciliation**

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

- Consolidated Subsidiaries: includes Hutchison Sinopharm and others
- Non-consolidated joint venture: includes SHPL and HBYS

					IFI	RS							Ų	JS GAAP					19-20
(US\$ millions)	03	04	05	06	07	08	09	10	11	12	13	14	15	16	17	18	19	20	Growth
Revenues (Non-GAAP)	21.9	27.9	65.1	101.4	119.0	155.8	197.0	236.4	278.6	360.7	402.3	465.4	518.9	627.4	677.2	664.4	665.6	706.6	6%
Consolidated subsidaries	4.7	6.1	9.3	8.9	3.7	5.5	7.0	14.1	14.9	15.5	16.5	67.0	126.2	180.9	205.2	172.9	178.1	197.8	11%
Non-consolidated joint venture	17.2	21.8	55.8	92.5	115.3	150.3	190.0	222.3	263.7	345.2	385.8	398.4	392.7	446.5	472.0	491.5	487.5	508.8	4%
Total Revenues Growth	n/a	27%	133%	56%	17%	31%	26%	20%	18%	29%	n/a	16%	11%	21%	8%	-2%	0%	6%	
- GuanBao divested in Sept'2017	-	-	-	-	-	-	-	-	(11.4)	(50.5)	(51.6)	(49.7)	(40.7)	(45.0)	(38.6)	-	-	-	n/a
Adjusted Non-consolidated joint venture	17.2	21.8	55.8	92.5	115.3	150.3	190.0	222.3	252.3	294.7	334.2	348.7	352.0	401.5	433.4	491.5	487.5	508.8	4%
Adjusted Revenues (Non-GAAP)	21.9	27.9	65.1	101.4	119.0	155.8	197.0	236.4	267.2	310.2	350.7	415.7	478.2	582.4	638.6	664.4	665.6	706.6	6%
Total Adjusted Revenues Growth	n/a	27%	133%	56%	17%	31%	26%	20%	13%	16%	n/a	19%	15%	22%	10%	4%	0%	6%	
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>	<sup>4]</sup> 83.6	84.9	90.2	<sup>[5]</sup> 6%
Consolidated subsidaries	(10.3)	(4.9)	(2.9)	(2.4)	0.2	0.0	0.8	1.0	(0.4)	(1.1)	0.1	1.6	1.4	3.1	5.9	6.9	3.8	3.9	4%
Non-consolidated joint venture	(0.4)	1.3	5.1	9.1	11.0	14.7	20.7	26.9	30.5	34.2	39.6	47.2	52.7	60.2	71.4	76.7	81.1	86.3	6%
Net (loss)/income attrib. to HUTCHMED	(5.7)	(3.7)	(0.5)	1.2	4.5	5.9	9.3	<sup>2]</sup> 12.6	<sup>[2]</sup> 13.6 <sup>[</sup>	<sup>2]</sup> 14.6 <sup>[</sup>	<sup>2]</sup> <b>18.2</b> [	<sup>2]</sup> 22.8 <sup>[</sup>	<sup>2]</sup> 25.2 <sup>[</sup>	<sup>2]</sup> 29.9	<sup>[3]</sup> 37.5 <sup>[-</sup>	<sup>4]</sup> 41.4	41.5	44.0	<sup>[5]</sup> 6%
Consolidated subsidaries	(5.5)	(4.3)	(2.7)	(2.4)	0.2	0.0	0.8	1.0	0.0	(0.7)	0.2	1.3	1.0	1.8	3.9	4.8	2.9	2.8	-5%
Non-consolidated joint venture	(0.2)	0.6	2.2	3.6	4.3	5.9	8.5	11.6	13.6	15.3	18.0	21.5	24.2	28.1	33.6	36.6	38.6	41.2	7%
Net (loss)/income attrib. to HUTCHMED growth	n/a	35%	86%	340%	275%	31%	58%	35%	8%	7%	n/a	26%	10%	19%	25%	10%	0%	6%	

<sup>[1] 2003–2006</sup> 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 HUTCHMED 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 HUTCHMED for 2017; [5] Excluded the land compensation in HBYS of \$72.0 million from net income after tax and \$28.8 million from net income attributable to HUTCHMED for 2020.



July'17 – 15 new drugs in oncology<sup>[1]</sup> added to NRDL

		Ur	nit Pricing (U	S\$) <sup>[3]</sup>		Approximate Mor	nthly Pricing (	US\$) <sup>[3]</sup>	
Brand (generic)	Company	Dosage	Avg. Tender	Reimbursed	Δ%	Dosage	Avg. Tender	Reimbursed	Indication coverage
Herceptin® (trastuzumab)	Roche	440mg:20ml	\$3,298.81	\$1,125.93	-66%	Breast: 4mg/kg wk 1, 2mg/kg weekly <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 Q2W	\$11,590	\$4,440	Late-stage meta. CRC or advanced non-squamous NSCLC.
TheraCIM® [4] (nimotuzumab)	Biotech Pharma	50mg:10ml	\$435.26	\$251.85	-42%	100mg weekly	\$3,730	\$2,160	Combo with radiotherapy for EGFR+ Stage III/IV nasopharyngeal carcinoma.
Rituxan® (rituximab)	Roche	500mg:50ml <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)	J&J	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)	J&J	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.



Oct'18 – 17 new drugs in oncology added to NRDL

			Unit Pricin	g (US\$) <sup>[2]</sup>		Approximate Monthly P	ricing (US\$) [ <sup>[</sup>	2]	
Brand (generic)	Company	Dosage	Avg. Tender	Reimbursed	Δ%	Dosage [1]	Avg. Tender	Reimbursed	Indication coverage
Focus V® (anlotinib)	Sino Biopharm	12mg	\$127	\$70	-45%	12mg QD (2 wks-on/1-wk-off)	\$2,500	\$1,417	3L NSCLC
Oncaspar® (pegaspargase)	Hengrui	5ml: 3750 IU	\$560	\$429	-23%	≤2ml every 14 days	\$1,231	\$943	1L ALL
Vidaza® (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® (axitinib)	Pfizer	5mg	\$99	\$30	-70%	5mg BID	\$5,957	\$1,787	2L advanced renal cell carcinoma
Tagrisso® (osimertinib)	AstraZeneca	80mg	\$253	\$73	-71%	80mg QD	\$7,597	\$2,201	EGFR TKI refractory T790M+ NSCLC
Ninlaro® (ixazomib)	Takeda	4mg	\$3,234	\$710	-78%	4mg on Days 1, 8, 15 (28 day cycle)	\$12,934	\$2,839	2L multiple myeloma
Xalkori® (crizotinib)	Pfizer	250mg	\$123	\$37	-70%	250mg BID	\$7,407	\$2,245	Locally adv. or meta. ALK+ or ROS1+ NSCLC
Gilotrif® (afatinib)	Boehringer	40mg	\$116	\$29	-75%	40mg QD	\$3,483	\$863	NSCLC with EGFR
Tasigna® (nilotinib)	Novartis	200mg	\$39	\$14	-65%	400mg BID	\$4,645	\$1,635	CML
Votrient® (pazopanib)	Novartis	200mg	\$66	\$23	-65%	800mg QD	\$7,891	\$2,348	RCC
Sutent® (sunitinib)	Pfizer	12.5mg	\$49	\$22	-55%	GIST & RCC: 50mg QD pNET: 37.5mg QD	\$5,544 \$4,455	\$2,498 \$2,007	RCC, GIST, pNET
Stivarga® (regorafenib)	Bayer	40mg	\$52	\$28	-46%	160mg QD, 3-wks-on/1-wk-off *	\$4,368	\$2,352	Meta. CRC, GIST, HCC
Zykadia® (ceritinib)	Novartis	150mg	\$108	\$28	-74%	450mg QD	\$9,699	\$2,564	ALK+ adv. or meta. NSCLC
Zelboraf® (vemurafenib)	Roche	240mg	\$30	\$16	-47%	960mg BID	\$7,252	\$2,369	Melanoma
Erbitux® (cetuximab)	Merck	100mg	\$571	\$186	-67%	400mg/m2 initial dose, 250mg weekly	\$10,446	\$3,074	Colorectal cancer, head and neck cancer
Sandostatin LAR® (octreotide)	Novartis	20mg	\$1,169	\$835	-29%	20mg Q4W	\$1,169	\$835	GEP-NENs
Imbruvica® (ibrutinib)	JNJ	140mg	\$78	\$27	-65%	MCL: 560mg QD CLL & WM: 420mg QD	\$9,324 \$6,993	\$3,263 \$2,447	MCL, CLL/SLL

Source: Ministry of Human Resources and Social Security (MOHRSS); Yaozhi; China Merchants Securities Research; Citi Global Research; Frost & Sullivan. [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.



Nov'19 update – 8 new drugs in oncology<sup>[1]</sup>

			Unit Pricing	; (US\$) <sup>[2]</sup>		Approximate Month	nly Pricing (US	\$) <sup>[2]</sup>	
Brand (generic)	Company	Dosage	Avg. Tender	Reimbursed	Δ%	Dosage	Avg. Tender	Reimbursed	Indication coverage
Elunate® (fruquintinib)	HUTCHMED	5mg	\$161	\$58	-64%	5mg QD 3wks/1wk-off.	\$3,378	\$1,221	Metastatic colorectal cancer, 3L
Tyvyt® (sintilimab)	Innovent	10ml (100mg)	\$1,206	\$437	-64%	200mg Q3W	\$3,216	\$1,166	Classical Hodgkin's lymphoma, 3L
Saiweijian® (raltitrexed)	Sino Biopharm	2mg	\$232	\$103	-56%	3mg/m <sup>2</sup> Q3W	\$765	\$340	Colorectal cancer, 5-FU intolerable
Alecensa® (alectinib)	Roche	150mg	\$32	\$10	-70%	600mg, BID	\$7,689	\$2,343	NSCLC, ALK+
Lynparza® (olaparib)	AstraZeneca	150mg	\$68	\$26	-62%	300mg, BID	\$8,173	\$3,120	Epithelial ovarian, fallopian tube, or peritoneal cancer
Airuini® (pyrotinib)	Hengrui	80mg	\$39	\$13	-66%	400mg QD, 21 days	\$4,118	\$1,389	Breast cancer, HER2+, 2L
Perjeta® (pertuzumab)	Roche	420mg	\$2,892	\$762	-74%	840mg wk1, 420mg Q3W	\$8,676	\$2,286	Breast cancer, HER2+, neoadjuvant
Jakafi® (ruxolitinib)	Incyte / Novartis	5mg	\$20	\$9	-56%	Dose is based on patient's baseline platelet count:  • (a) >200 X 10 <sup>9</sup> /L: 20 mg BID  • (b) 100 X 10 <sup>9</sup> /L-200 X 10 <sup>9</sup> /L: 15 mg BID  • (c) 50 X 10 <sup>9</sup> /L to 100 X 10 <sup>9</sup> /L: 5 mg given BID	(a) \$4,800 (b) \$3,600 (c) \$1,200	(a) \$2,160 (b) \$1,620 (c) \$540	PMF, PPV-MF, PET-MF



Nov'19 update – 9 renewed drugs in oncology<sup>[1]</sup>

			Unit Pricing (	US\$) <sup>[2]</sup>		Approximate Montl	nly Pricing (US	\$) [2]	
Brand (generic)	Company	Dosage	'17 NRDL	'19 NRDL	Δ%	Dosage	'17 NRDL	'19 NRDL	Indication coverage
AiTan® (apatinib)	Hengrui	425mg <sup>[3]</sup>	\$30	\$27	-13%	850mg QD	\$1,823	\$1,594	3L gastric adenocarcinoma or GEJ with adenocarcinoma.
EnDu® (rh-endostatin)	Simcere	15mg	\$97	\$75	-22%	7.5mg/m² iv QD, 2wks/1wk-off	\$1,681	\$1,308	Late-stage NSCLC.
Epidaza® (chidamide)	Chipscreen	5mg	\$53	\$59	-11%	30mg QD, 2x per wk	\$2,843	\$2,533	2L+ Recurring or refractory peripheral T-cell lymph. (PTCL).
Herceptin® (trastuzumab)	Roche	440mg	\$1,169	\$846	-28%	3wks regimen: 8mg/kg wk1, 6mg/kg Q3W	\$1,276	\$923	Breast: Her2+; Her2+ meta. Her2+ late-stage meta. gastric.
Avastin® (bevacizumab)	Roche	100mg	\$307	\$231	-25%	3wks regimen: CRC: 7.5mg/kg Q3W NSCLC: 15mg/kg Q3W	CRC: \$1,844 NSCLC: \$3,689	CRC: \$1,385 NSCLC: \$2,769	Late-stage meta. CRC or advanced non-squamous NSCLC.
TheraCIM <sup>® [4]</sup> (nimotuzumab)	Biotech	50mg	\$262	\$221	-16%	100mg, QW	\$2,092	\$1,766	Combo with RT for EGFR+ III/IV nasopharyngeal carcinoma.
Tarceva® (erlotinib)	Roche	150mg	\$28	\$12	-56%	150mg, QD	\$841	\$374	Advanced NSCLC with limited EGFR gene mutation.
Nexavar® (sorafenib)	Bayer	200g	\$29	\$14	-53%	400g BID	\$3,519	\$1,662	RCC or HCC. meta. diff. thyroid after radio-iodine therapy.
Afinitor® (everolimus)	Novartis	5mg	\$23	\$20	-12%	RCC: 10mg, QD Pan-NETs: 10mg, QD	\$1,366	\$1,200	RCC after sunitinib or sorafenib. Pancreatic NETs. TSRA.



Dec'20 update – 13 new oncology drugs through negotiation<sup>[1]</sup>

			Unit Pricing	; (US\$) <sup>[2]</sup>		Approximate Moi	nthly Pricing (US\$)	) [2]	
Brand (generic)	Company	Dosage	Avg. Tender	Reimbursed	Δ%	Dosage	Avg. Tender	Reimbursed	Indication coverage
Lipusu® (paclitaxel liposome)	Luye Pharma	30mg	\$129	\$35	-73%	155mg/m <sup>2</sup> Q3W	\$1,470	\$399	1L+ metastatic ovarian cancer, breast cancer, 1L NSCLC
Ciptertin® (inetetamab)	3SBio	50mg	\$235	\$91	-61%	initial 4mg/kg, maintenance 2mg/kg	\$2,260	\$871	HER2+ metastatic breast cancer
Baizean® (tislelizumab)	BeiGene	100mg	\$1,644	\$335	-80%	200mg Q3W	\$4,385	\$894	3L relapsed or refractory classical Hodgkin's lymphoma, locally adv. or meta. urothelial cancer
Tuoyi® (toripalimab)	Junshi Biosciences	240mg	\$1,108	\$323	-71%	3mg/kg Q2W	\$1,662	\$485	Non-excisional or metastatic melanoma
AiRuiKa® (camrelizumab)	Hengrui	200mg	\$3,046	\$450	-85%	cHL&EC: 200mg Q2W NSCLC: 200mg Q3W HCC: 33mg/kg Q3W	\$6,092 \$4,062 \$40,209	\$601	3L relapsed or refractory classical Hodgkin's lymphoma, advanced HCC, 1L locally adv. or meta. non-squamous NSCLC, esophageal cancer
Xinfu® (flumatinib)	Hansoh Pharma	200g	\$27	\$10	-63%	600mg QD	\$2,430	\$900	Ph+ chronic myelogenous leukemia
Ameile® (almonertinib)	Hansoh Pharma	55mg	\$75	\$27	-64%	110mg QD	\$4,523	\$1,625	EGFR TKI refractory T790M+ locally advanced or metastatic NSCLC
Brukinsa® (zanubrutinib)	BeiGene	80mg	\$27	\$15	-44%	320mg QD	\$3,260	\$1,828	2L MCL, 2L CLL / SLL
Mekinist® (trametinib)	Novartis	2mg	\$142	\$57	-60%	2mg QD	\$4,254	\$1,705	BRAF V600M+ non-excisional or metastatic melanoma
Tafinlar® (dabrafenib)	Novartis	75mg	\$53	\$14	-74%	150mg BID	\$6,380	\$1,705	BRAF V600M+ non-excisional or metastatic melanoma
Lenvima® (lenvatinib)	Eisai	4mg	\$86	\$17	-81%	12mg QD	\$7,754	\$1,495	HCC
Xtandi® (enzalutamide)	Astellas Pharma	40mg	\$49	\$11	-78%	160mg QD	\$5,880	\$1,285	Castration-resistant prostate cancer (CRPC)
Zejula® (niraparib)	Zai Lab	100mg	\$128	\$31	-76%	300mg QD	\$11,534	\$2,769	Relapsed epithelial ovarian, fallopian tube or primary peritoneal carcinoma



Dec'20 update – 15 renewed drugs in oncology<sup>[1]</sup>

			Unit Pricir	g (US\$) [2]		Approximate Month	ly Pricing (US\$)		
Brand (generic)	Company	Dosage	Avg. Tender	Reimbursed	Δ%	Dosage	Avg. Tender	Reimbursed	Indication coverage
Focus V® (anlotinib)	Sino Biopharm	12mg	\$75	\$47	-37%	12mg QD (2 wks-on/1-wk-off)	\$1,515	\$952	3L NSCLC, 3L SCLC, STS
Oncaspar® (pegaspargase)	Hengrui	5ml: 3750 IU	\$584	\$458	-21%	≤2ml every 14 days	\$1,283	\$1,006	1L ALL
Inlyta® (axitinib)	Pfizer	5mg	\$32	Undisclosed	-	5mg BID	\$1,920	-	2L advanced renal cell carcinoma
Tagrisso® (osimertinib)	AstraZeneca	80mg	\$78	\$28	-64%	80mg QD	\$2,350	\$860	1L NSCLC harboring EGFR exon 19 deletions or exon 21 L858R mutations; EGFR TKI refractory T790M+ NSCLC
Ninlaro® (ixazomib)	Takeda	4mg	\$759	Undisclosed	-	4mg on Days 1, 8, 15 (28 day cycle)	\$2,277	-	2L multiple myeloma
Xalkori® (crizotinib)	Pfizer	250mg	\$40	\$35	-12%	250mg BID	\$2,400	\$2,112	Locally adv. or meta. ALK+ or ROS1+ NSCLC
Tasigna® (nilotinib)	Novartis	200mg	\$15	Undisclosed	-	400mg BID	\$1,800	-	CML
Votrient® (pazopanib)	Novartis	200mg	\$25	Undisclosed	-	800mg QD	\$2,510	-	RCC
Stivarga® (regorafenib)	Bayer	40mg	\$30	\$26	-12%	160mg QD, 3-wks-on/1-wk-off	\$2,520	\$2,217	Meta. CRC, GIST, HCC
Zykadia® (certinib)	Novartis	150mg	\$30	Undisclosed	-	450mg QD	\$2,700	-	ALK+ adv. or meta. NSCLC
Zelboraf® (vemurafenib)	Roche	240mg	\$17	Undisclosed	-	960mg BID	\$4,080	-	BRAF V600 Melanoma
Erbitux® (cetuximab)	Merck	100mg	\$199	Undisclosed	-	400mg/m² initial dose, 250mg QW	\$1,990	-	Colorectal cancer, head and neck cancer
Sandostatin LAR® (octreotide)	Novartis	20mg	\$892	Undisclosed	-	20mg Q4W	\$892	-	GEP-NENs
Imbruvica® (ibrutinib)	JNJ	140mg	\$29	Undisclosed	-	MCL: 560mg QD CLL & WM: 420mg QD	MCL: \$3,489 CLL&SLL: \$2,617	-	MCL, CLL/SLL, WM
Lynparza® (olaparib)	AstraZeneca	150mg	\$26	Undisclosed	-	300mg, BID	\$1,560	-	BRCAm epithelial ovarian, fallopian tube, or peritoneal cancer