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# R&D UPDATE

August 25, 2021

# DISCLOSURES

Certain statements contained in this presentation and in the accompanying oral presentation, other than statements of fact that are independently verifiable at the date hereof, may constitute forward-looking statements. Examples of such forward-looking statements include statements regarding BeiGene's research, discovery, and pre-clinical and early-stage clinical programs and plans; recent clinical data for BeiGene's product candidates and approvals of its medicines; the conduct of late-stage clinical trials and expected data readouts; additional planned commercial product launches; and the advancement of and anticipated clinical development, regulatory milestones and commercialization of BeiGene's medicines and drug candidates. Actual results may differ materially from those indicated in the forward-looking statements as a result of various important factors, including BeiGene's ability to demonstrate the efficacy and safety of its drug candidates; the clinical results for its drug candidates, which may not support further development or marketing approval; actions of regulatory agencies, which may affect the initiation, timing and progress of clinical trials and marketing approval; BeiGene's ability to achieve commercial success for its marketed medicines and drug candidates, if approved; BeiGene's ability to obtain and maintain protection of intellectual property for its technology and medicines; BeiGene's reliance on third parties to conduct drug development, manufacturing and other services; BeiGene's limited experience in obtaining regulatory approvals and commercializing pharmaceutical products and its ability to obtain additional funding for operations and to complete the development and commercialization of its drug candidates and achieve and maintain profitability; the impact of the COVID-19 pandemic on BeiGene's clinical development, commercial, regulatory and other operations, as well as those risks more fully discussed in the section entitled "Risk Factors" in BeiGene's most recent periodic report filed with the SEC, as well as discussions of potential risks, uncertainties, and other important factors in BeiGene's subsequent filings with the SEC. All information in this presentation is as of the date of this presentation, and BeiGene undertakes no duty to update such information unless required by law.

Some of the clinical data in this presentation relating to BeiGene's investigational drug candidates is from pre-clinical studies or early phase, single-arm clinical trials. When such data or data from later stage trials are presented in relation to other investigational or marketed drug products, the presentation and discussion are not based on head-to-head trials between BeiGene's investigational drug candidates and other products unless specified in the trial protocol. BeiGene is still conducting pre-clinical studies and clinical trials and, as additional patients are enrolled and evaluated, data on BeiGene's investigational drug candidates may change.

This presentation and the accompanying oral presentation contain data and information obtained from third-party studies and internal company analysis of such data and information. BeiGene has not independently verified the data and information obtained from these sources. Forward-looking information obtained from these sources is subject to the same qualifications noted above.

# TODAY'S AGENDA

- Welcome and Introduction:
   Creating an Ecosystem of Innovation
   John V. Oyler
- Our Scientific Ecosystem of Excellence Xiaodong Wang
- R&D Capability and Portfolio Overview Lai Wang, Ph.D.
- Hematology Oncology Clinical
   Program Update
   William Novotny, M.D.

• Solid Tumor Clinical Program Update Yong (Ben) Ben, M.D.

- New Modalities and Future Pipeline Lai Wang, Ph.D.
- The Road Ahead John V. Oyler

Questions and Answers

# CREATING AN ECOSYSTEM OF INNOVATION

John V. Oyler Co-Founder, Chairman and CEO

# OUR MISSION: PATIENTS FIRST THROUGH SCIENCE AND AFFORDABILITY

Help transform the biotechnology industry, creating impactful medicines that will be affordable and accessible to far more cancer patients around the world through science and business innovation **BEICENE** 

# WE SAW THE ONCE-IN-A-LIFETIME OPPORTUNITY TO ACCELERATE GLOBAL DEVELOPMENT 10+ YEARS AGO...

Internal, global, and modernized efficiency of clinical science to create time and cost advantages and broaden patient access

# BUILDING A GLOBAL ECOSYSTEM OF COST AND SPEED COMPETITIVE ADVANTAGES

Innovation with speed and lower cost to better serve patients around the world

### FULLY INTEGRATED END-TO-END ~7,000 COLLEAGUES - 5 CONTINENTS - 17 GEOGRAPHIES

#### Research

- One of world's largest oncology teams (650+) with 50+ preclinical programs
- Passionate, entrepreneurial science-based culture
- Internal team (not CROs)

#### Development

 1,700+ internal clinical development colleagues, predominately CRO-free

 More inclusive development (e.g., Australia, China, Poland) enabling ~ 1/3 cost & time savings

• ~50 assets in clinical & commercialization stage

#### Manufacturing

- In-house capabilities bring cost, agility, and flexibility to internal and external programs
- Built to state-of-the-art standards & technologies (e.g., 1st paperless facility in China)
- Building toward 200,000 L of biologics capacity in Guangzhou

#### Commercial

- 2,900+ in China, competitively positioned, science-based leadership
- 150+ competitive footprint in U.S.
- Expanding presence in multiple countries/regions, including underserved areas

# OUR SCIENTIFIC ECOSYSTEM OF EXCELLENCE

Xiaodong Wang, Ph.D. Co-Founder & Chairman of Scientific Advisory Board

# **RESPONDING TO GREATEST PATIENT NEED**

Estimated 19.3 million new cancer cases and almost 10.0 million cancer deaths in 2020 worldwide



Our broad pipeline covers 80% of the world's cancers by incidence



# TO REACH **BILLIONS** OF PATIENTS WORLDWIDE



# R&D CAPABILITY AND PORTFOLIO OVERVIEW

Lai Wang, Ph.D. Global Head of R&D



# **R&D HIGHLIGHTS**

# **R&D Engine Built at Scale to Fuel Innovation with Efficiency and Quality**

- Highly productive preclinical research team with proven track record experienced major expansion recently
- Clinical development is largely CRO-free, with unique global operating model

# Robust Innovative Pipeline Focusing on Internal Combinations

- Close to 100 programs, including 50+ at preclinical stage, 30+ at clinical stage and 10+ at commercial stage
- Strong heme and solid tumor franchise anchored on BTK and PD-1 programs

# SCIENCE-DRIVEN PRECLINICAL CAPABILITIES EXPERIENCED MAJOR EXPANSION RECENTLY

### **Evolution Through Two Stages**

#### Stage I: Establishing Core Capabilities and Talent for Early Success (2011-2018)

- I 50-200 scientists
- Proven track-record with 3 internally discovered, commercially available medicines
- Small scale, only 1-2 clinical candidates a year

#### Stage 2: Expanding New Capabilities and Talent for Future Success (2019 – Present)

- 650+ scientists now growing to 800 by end of 2021
- Investing in new platforms and capabilities and driving efficiencies through portfolio management
- ▶ 50+ preclinical programs, ~50% with first-in-class potential
- A burst of new clinical molecules expected in the next few years

### **New Platforms/Capabilities**



CDAC, Chimeric Degradation Activating Compound; ADC, Antibody Drug Conjugate.

# AN INTERNAL CLINICAL DEVELOPMENT TEAM EQUIPPED TO RUN GLOBAL DEVELOPMENT AT SCALE

### Strong Internal Global Development Capabilities

- Through the extensive internalization effort in the last few years, we are now largely CROfree, allowing better control over quality, speed, cost and site / PI engagement
- Currently 1,700+ global clinical development colleagues, close to 700 outside of China, growing to 2,000+ by end of 2021
- Internal training academy allows for continuous growth of the team and promotes quality and efficiency

# BeiGene's Global Clinical Trial Footprint



- ▶ 95+ clinical trials initiated since 2013 in 40+ geographies
- I3,000+ patients and healthy volunteers enrolled, close to half outside of China

# UNIQUE OPERATING MODEL TO DRIVE SPEED & COST ADVANTAGE

#### **Investigator Fee Per Patient**



Source: Grants Manager median benchmark data for those countries in which BeiGene has enrolled at least one patient. Each region/country is weighted for the number patients enrolled to date.

#### Clinical Trial Enrollment Rate (Pts/Site/Month)



Source: Internal data on file; Benchmark from Citeline, filtered for NSCLC 1L studies. Includes data only from the same countries that BeiGene has run clinical studies. Data exported on 9 Aug 2021 ANZ/ SK, Australia and New Zealand/ Slovakia



**BEICENE** 



# TRANSLATING SCIENCE INTO BETTER AND NOVEL MEDICINES



SM, Small Molecule; mAb, Monoclonal Antibody; ADC, Antibody Drug Conjugate; TAA, Tumor Associated Antigen; CDAC, Chimeric Degradation Activating Compound; BsAb, Bispecific Antibody; TsAb, Trispecific Antibody; CAR-NK, Chimeric Antigen Receptor-Natural Killer Cell

# ENHANCING INNOVATION AND ACCESS THROUGH COLLABORATION

#### We follow the science virtually anywhere that leads to innovation – independently and via collaboration

NOVEL COMBINATIONS	Extend our existing franchise in large indications tisle combinations: +sitra; +zanidatamab; +Leap asset
NEXT-GEN TECHNOLOGIES	ADC CAR-NK mRNA Bispecifics
Set EUSAPharma EIG 男 泰 BIO-THERA	Ambrx Oassemblybio OSeagen leaptherapeutics
Singlomics 丹序生物 SHORELINE	
TREATE Zymeworks	Celgene   Ulli Bristol Myers Squibb" AMCEN° U NOVARTIS Company

"BeiGene is evaluating sitravatinib in patients with a variety of tumor types with the goal of accelerating development of treatment options for more patients. Our collaboration with BeiGene has supported the expansion of our world-wide development capabilities by leveraging their clinical expertise in China."

**CHARLES M. BAUM, M.D., Ph.D.** President and CEO of Mirati Therapeutics

"Partnering with BeiGene was a key component of our development and commercialization strategy for ZW25 and ZW49. This collaboration allows Zymeworks to leverage BeiGene's resources and expertise to accelerate the development of our most advance product candidates and broaden our reach globally including in a key region of the world."

ALI TEHRANI President and CEO of Zymeworks



PARTNER

# DEEP CLINICAL AND COMMERCIAL PIPELINE

COMPOUND	DOSE ESC.	DOSE ESC. DOSE EXPANSION PIVOTAL FILED /	FILED /	COMMERCIA	DADTHER		DOSE ESC.	DOSE ESC. DOSE EXPANSION		PIVOTAL		FILED / C	COMMERCIAL				
COMPOUND	PHIa	PHIb	PH2*	PH2**	PH3	MARKETED	RIGHTS	PARTNER	COMPOUND PHIa	PHIa	РНІБ	PH2*	PH2**	PH3	MARKETED	RIGHTS	PARTNE
Zanubrutinib (BTK)	MCL, WM, CLL/SLL, MZL, FL (P	hase 2), Lupus (Ph	ase 2), DLBCL (Phas	se I)			Global		BLINCYTO <sup>®</sup>	ALL			· <b></b> •				
Tislelizumab (PD-1)	MSI-H or dMMR solid tumors, E (combo; Phase 1)	SCC, HCC, NSCLO	, cHL, UC, NK/T-cel	l lymphoma (Phase	e 2), solid turnors,	B-cell malignancies	Global ex-NVS territory <sup>†</sup>	Novartis	KYPROLIS <sup>®</sup>	мм							
Pamiparib (PARP)	gBRCA+ OC, PSOC (Phase 3), H	IER2- BRCA+ BC (	Phase 2), Glioblastor	na (Phase 2), Solio	d tumors (Phase I	)	Global		XGEVA®	MM, Solid tumors							
BGB-A1217 (TIGIT)	NSCLC, Cervical Cancer (Phase	2), ESCC (Phase 2	), Solid tumors (Phas	se I)			Global		LUMAKRAS (KRAS G12C)	Solid tumors, NSCLC, CRC							
BGB-A445 (OX40)	Solid tumors						Global		Pavurutamab^^ (BCMA x CD3)	мм							
BGB-A425 (TIM-3)	Solid tumors						Global		AMG 176 (MCL-1)	Myeloid malignancies							
BGB-A333 (PD-L1)	Solid tumors						Global		AMG 330^ (CD33 x CD3)	Myeloid malignancies							
BGB-11417 (BCL-2)	B-cell malignancies						Global		AMG 673^^ (CD33 x CD3)	AML							
BGB-15025 (HPKI)	Advanced solid tumors						Global		AMG 427^^ (FLT3 x CD3)	AML							
BGB-10188 (ΡΙ3Κδ)	B-cell malignancies, Solid tumors	5					Global		Tarlatamab^^ (DLL3 x CD3)	SCLC, Neuroendocrine							
Lifirafenib (RAF dimer)	B-RAF/K-RAS/N-RAS mut. Solid t	tumors					Global		Acapatamab^^ (PSMA x (D3)	Prostate Cancer						China	Amgen
REVLIMID	мм								AMG 509 (STEAP / XmAb <sup>®</sup> antibady)	mCRPC							
ABRAXANE	BC, NSCLC, Pancreas adenocarc	inoma					China	Bristol Myers Squibb	AMG 506 (FAP x 4-1 BB DARPin)	Solid tumors							
VIDAZA	Myelodysplastic Syndromes								AMG 256 (PD I x II 12 mutein)	Solid tumors							
SYLVANT	Multicentric Castleman's disease	2					China	ELISA Pharma	AMG 650 (KIE/84)	Solid tumors							
QARZIBA	Neuroblastoma			PH2** PH3 PH3 MARKET  rase 1) rase 1) rase 1) rase 1) race 1) race 1)		Chind	2004 Mania	AMG 91000 (CLDN/8.2 × CD3)									
BAT1706	Metastatic colorectal cancer				PH3 PH2EU/ Ph3 PH3 PARKETED	China	BIO-THERA	AMG 19900 (MUCL7 x CD3)									
Sitravatinib <sup>111</sup> (multi-kinase inhibitor)	NSCLC, RCC, OC, MEL, HCC, G	CIGEJC					Asia ex-Japan, AU, NZ	Mirati Therapeutics	AMG 199 (MOC17 x CD3)								
Zanidatamab <sup>1111</sup> (HER2, bispecific antibody)	BC, GEA, Biliary track (Phase 3)						Asia ex-Japan, AU, NZ	Zymeworks	AMG 774	Solid turnors							
Z₩49 (HER2, bispecific ADC)	HER2+ cancers						Asia ex-Japan, AU, NZ	Zymeworks	AMG 397 (MCL-1)	Myeloid malignancies							
BGB-3245 (B-RAF)	Solid tumors						Asia ex-Japan	SpringWorks <sup>1</sup>	*Selected assets and indicatio	ons are presented. **So	me indicatio	ons will not	require a n	on-pivotal	Ph2 clinica	l trial prior to l	peginning
SEA-CD70 (CD70)	MDS, AML						Asia ex-Japan, AU, NZ	Seattle Genetics	pivotal Ph2 or Ph3 clinical tria ^ BiTE, ^^ HLE BiTE, † Nova	als. ***Confirmatory cli rtis owns commercial r	nical trials ights in Un	post approvited States.	val are requ Canada, Me	ired for ac exico, the l	celerated o European U	r conditional ap nion. United K	provals.
DKN-01 (DKK1)	GCIGEJC						Asia ex-Japan, AU, NZ	Leap Therapeutics	Norway, Switzerland, Iceland	I, Liechtenstein, Russia,	and Japan.	tt ABX is s	suspended in	n China. †	† Mirati is	also conducting	g its own
ABI-H0731 (HBV core inhibitor)	Chronic Hepatitis B Virus								leukemia, HLE BiTE: Half-life	extended Bi-specific T-	-cell engage	ers, BC: bre	ast cancer,	GC/GEJ: ga	astric cance	r/gastroesopha	geal
ABI-H2158 (HBV core inhibitor)	Chronic Hepatitis B Virus		-				China	Assembly Bio	junction, HCC: hepatocellula Hodgkin's lymphoma, N/SCL	r carcinoma, IND: Inve C: non-/small cell lung (	stigational l cancer, OC	New Drug, 2: ovarian ca	MEL: melan incer, RCC:	oma, MM: renal cell	multiple m carcinoma,	yeloma, NHL: ı SM: small mole	non- ecule; I.
ABI-H3733 (HBV core inhibitor)	Chronic Hepatitis B Virus								By MapKure, a JV with SpringWorks.								

# HEMATOLOGY ONCOLOGY CLINICAL PROGRAM UPDATE

William Novotny, M.D. Head of Clinical Development, Hematology



# SUMMARY OF HEMATOLOGY CLINICAL AND COMMERCIAL PORTFOLIO



# () AML/MDS/ALL BGB-11417 (BCL-2) BGB-A425 (TIM-3) VIDAZA Vidaza AMG 330/673 (CD33 x CD3) AMG 427 (FLT3 x CD3) AMG 176/397 (MCL-1) SEA-CD70 (CD70)

3	Myeloma
	BGB-11417 (BCL-2)
	AMG 701 (BCMA x CD3)
	AMG 176/397 (MCL-1)



# **BEST-IN-CLASS** BTK INHIBITOR



Designed to Overcome Shortcomings of Other BTK Inhibitors	Superiority Over Ibrutinib in Clinic	Broad Indications Approved/In Development	
<ul> <li>Reduced off-target toxicity</li> </ul>	<ul> <li>Better efficacy</li> </ul>	► CLL ► MZL	
<ul> <li>Achieved sustained target inhibition in disease originating tissues</li> </ul>	<ul><li>Better safety</li><li>More convenient</li></ul>	► MCL ► FL ► WM	



# BETTER KINASE SELECTIVITY THAN IBRUTINIB, ACALABRUTINIB AND ITS METABOLITE M27



- Kinome analysis (370 kinases panel) at concentrations of 100 times of each molecule's IC50 for BTK (not at 1 μM which is not adjusted for BTK potency); acalabrutinib and its metabolite M27 are much weaker BTK inhibitors
- Zanubrutinib demonstrated better kinase selectivity (7 off-target kinases with >50% inhibition) compared to ibrutinib (17), acalabrutinib (15), and M27 (23) at 100X IC50



# FAVORABLE PHARMACOKINETIC PROFILE COMPARED TO ACALABRUTINIB AND IBRUTINIB

### FREE DRUG CONCENTRATION TIME PROFILES RELATIVE TO IC50



Note: These data are from separate analyses. Limitations of cross-trial comparisons apply. Source: I. Kaptein, et al. *Blood.* 2018;132:1871. 2. Ou, et al. *Leuk Lymphoma*. In press. 3. Marostica, et al. *Cancer Chemother Pharmacol.* 2015;75:111-121.0 QD, Once a Day; BID, Twice a Day



Vs. Ibrutinib

# BROAD CLINICAL DEVELOPMENT PROGRAMS ACROSS MULTIPLE HEMATOLOGIC MALIGNANCIES AND BEYOND ONCOLOGY



PHASE

PHASE

PHASE





# ZANUBRUTINIB VS. BENDAMUSTINE + RITUXIMAB: STATISTICALLY SIGNIFICANT IMPROVEMENT IN PFS DEMONSTRATED IN TREATMENT-NAÏVE CLL/SLL PATIENTS

**Study Identifier:** 

Primary Endpoints: PFS by IRC in Cohort I

BGB-3111-304, NCT03336333

Key Secondary Endpoints: Cohort I – ORR, OS, DOR, safety







## ZANUBRUTINIB VS. IBRUTINIB: SUPERIORITY IN OVERALL RESPONSE RATE BY INVESTIGATOR ASSESSMENT DEMONSTRATED IN RELAPSED OR REFRACTORY CLL/SLL PATIENTS

**Study Identifier:** 

Primary Endpoints: ORR by Investigator Assessment

BGB-3111-305, NCT03734016

Key Secondary Endpoints: PFS, OS, safety

#### **ORR** by Investigator Assessment

	Zanubrutinib (n=207)	lbrutinib (n=208)			
Primary endpoint:	<b>78.3</b> %	62.5%			
ORR (PR+CR)	P<0.001				
CR/CRi	1.9%	1.4%			
PR	75.8%	61.1%			
ORR (PR-L+PR+CR)	88.4%	81.3%			

#### **PFS** by Investigator Assessment



#### **Overall Survival**



The PFS and OS analyses were not pre-specified.

More results including safety data were presented at *EHA 2021*. Hillmen P et al. Abstract LB1900. ORR, Overall Response Rate; CR, Complete Response; PR, Partial Response; PFS, Progression-Free Survival; OS, Overall Survival



BGB-3111-AU003



# ZANUBRUTINIB LONG-TERM FOLLOW-UP: DEEP AND DURABLE RESPONSES IN PATIENTS WITH RELAPSED OR REFRACTORY CLL/SLL

**Study Identifier:** 

Primary Endpoints: safety, RP2D

BGB-3111-AU-003, NCT02343120

Key Secondary Endpoints: ORR, CR rate, PR rate, PFS, OS, DOR



Investigator-Assessed Response	N = 103
ORR (PR + CR)	90.3%
ORR (PR-L or better)	94.2%





BGB-3111-206



# ZANUBRUTINIB LONG-TERM FOLLOW-UP: DEEP AND DURABLE RESPONSES IN PATIENTS WITH RELAPSED OR REFRACTORY MCL

**Study Identifier:** 

BGB-3111-206, NCT03206970

Primary endpoint: ORR by IRC Secondary endpoints: PFS, DOR, TTR

#### **Best Overall Response by IRC**

Response by IRC	N=86
ORR (CR+PR)	83.7%
Best Response	
CR	77.9%
PR	5.8%

#### More results including safety data were presented at EHA 2021. Song Y, et al. Abstract EP789.

#### Median PFS = 33 month (95% CI: 19.4, NE)







### ZANUBRUTINIB VS. IBRUTINIB: DIRECTIONALLY FAVORABLE VGRP+CR RATE AND PFS/OS OBSERVED IN MYD88<sup>MUT</sup> WALDENSTRÖM MACROGLOBULINEMIA PATIENTS

**Study Identifier:** 

Primary Endpoints: CR and VGPR by IRC

BGB-3111-302, NCT03053440

Key Secondary Endpoints: MRR(CR+VGPR+PR), DOR, PFS, Safety

#### **CR+VGPR**





#### OS



+Adjusted for stratification factors and age group. ++ Adjusted for stratification factors and age group. p-value is for descriptive purpose only. More results including safety data were presented at: 1) ASCO 2020. Tam CS et. al. Abstract S225. 2) EHA 2020. Dimopoulos M et al. Abstract EPI 180. VGPR, Very Good Partial Response





# ZANUBRUTINIB VS. IBRUTINIB: IMPROVED TOLERABILITY OF ZANUBRUTINIB COMPARED TO IBRUTINIB

ALPINE Res Travel Travel Travel Travel			ASPEN Preus 3 Madematrian Vacandolulirenia		
Category, n(%)	Zanubrutinib (n=204)	lbrutinib (n=207)	Category, n(%)	Zanubrutinib (n=101)	lbrutinib (n=98)
Any AE	195 (95.6)	205 (99.0)	Patients with $\geq 1$ AE	98 (97.0)	97 (99.0)
Grade ≥3 AE	114 (55.9)	106 (51.2)	Grade ≥3 AE	59 (58.4)	62 (63.3)
Serious AE	56 (27.5)	67 (32.4)	Serious AE	40 (39.6)	40 (40.8)
AE leading to death	8 (3.9)	12 (5.8)	AE leading to death	1 (1.0)	4 (4.1)
AE leading to dose reduction	23 (11.3)	25 (12.1)	AE leading to dose reduction	14 (13.9)	23 (23.5)
AE leading to dose interruption	81 (39.7)	84 (40.6)	AE leading to dose interruption	47 (46.5)	55 (56.1)
AE leading to treatment discontinuation	16 (7.8)	27 (13.0)	AE leading to treatment discontinuation	4 (4.0)	9 (9.2)

I) EHA 2021. Hillmen P et al. Abstract LB1900. 2) EHA 2020. Dimopoulos M et al. Abstract S225.



ent Rate

0 Cumulativ

No. of Subjects at Risk

Zanubrutinib

2.5%

12

Months

15

- Zanubrutinib

Ibrutinib

Censored



**Safety Outcomes** 

# **ZANUBRUTINIB VS. IBRUTINIB:** FAVORABLE AFIB/FLUTTER OUTCOMES





# FAVORABLE CLINICAL PHARMACOLOGY PROFILE PROVIDES CLINICAL CONVENIENCE FOR PATIENTS

Flexible dosing regimen

- Can be administered as QD or BID (1, 2)
- Can be administered with or without food (1)

Less drug-drug interactions (DDI); ability to be safely and effectively co-administered with a broader variety of concomitant medicines

- Can be co-administered with strong or moderate CYP3A inhibitors at a reduced dose (1, 3, 4)
- Can be co-administered with acidreducing agents (including PPIs) (3, 5)

Improved safety for organ -dysfunctional patients (1, 5, 7)

- No dose modification needed for patients with renal impairment or mild-moderate hepatic impairment.
- Can be used for patients with severe hepatic impairment at a reduced dose

I. Brukinsa USPI 2019.

- 2. Ying C Ou et al. Rationale for once-daily or twice-daily dosing of zanubrutinib in patients with mantle cell lymphoma, Leuk Lymphoma. 2021
- 3. Kun Wang et al. Comprehensive PBPK model to predict drug interaction potential of Zanubrutinib as a victim or perpetrator, CPT Pharmacometrics Syst Pharmacol. 2021
- 4. Song Mu et al. Effect of rifampin and itraconazole on the pharmacokinetics of zanubrutinib (a Bruton's tyrosine kinase inhibitor) in Asian and non-Asian healthy subjects, Cancer Chemother Pharmacol. 2019
- 5. Ying C Ou et al. Population Pharmacokinetic Analysis of the BTK Inhibitor Zanubrutinib in Healthy Volunteers and Patients With B-Cell Malignancies, Clin Transl Sci. 2021
- 6. Ying C Ou et al. Evaluation of drug interaction potential of zanubrutinib with cocktail probes representative of CYP3A4, CYP2C9, CYP2C19, P-gp and BCRP, Br J Clin Pharmacol. 2020
- 7. Ying C Ou et al. A phase I, open-label, single-dose study of the pharmacokinetics of zanubrutinib in subjects with varying degrees of hepatic impairment, Leuk Lymphoma. 2020



# GLOBAL FOOTPRINT CONTINUES TO EXPAND TO **70+ COUNTRIES**\*

#### 2019

#### 3 submissions

- MCL and CLL in CN
- MCL US approval

### 2020

#### 22 submissions

- I5 for MCL CN approval
- 7 for WM
- ► I for CLL CN approval

### 202 I

#### 40+ submissions planned

- 9-12 expected approvals
- ► 22 for MCL 4 approvals (CA, AE, IL, CL)
- I3 for WM 2 approvals (CA, CN)
- 8 for MZL
- 3 for CLL



### BE1CENE

# BGB-II4I7 (BCL-2) AN INVESTIGATIONAL BCL-2 INHIBITOR WITH POTENTIAL BEST-IN-CLASS PROPERTIES

- Better selectivity and more potent than venetoclax, potentially translating to deeper target inhibition and better safety
- Potential to overcome resistance to venetoclax (GI0IV mutant)
- A key molecule in heme portfolio
  - Monotherapy or in combination with zanubrutinib for CLL/NHL
  - Potential new entry to AML/MDS/MM space

		-
Protein	BGB-11417 IC <sub>50</sub> (nM)	Venetoclax IC <sub>50</sub> (nM)
Bcl-2	<b>0.014</b> ± 0.0021	<b>0.20</b> ± 0.015
Bcl-2- G101V	<b>0.59</b> ± 0.08	<b>34</b> ± 3.8

**Highly Potent** 







# BCL-2 PROGRAM CLINICAL PROGRESS UPDATE

- Dose escalation ongoing for monotherapy, combination with zanubrutinib, and combination with azacitidine
- Clinical activity observed at the first dose level 40 mg
- Currently at 320 mg dose level, PK exposure at 80 mg equivalent to that of venetoclax at 400 mg, its approved dose for CLL
- Pivotal studies expected to be initiated in 2022



### ZANUBRUTINIB, OUR BEST-IN-CLASS BTK INHIBITOR, IS THE CORNERSTONE MOLECULE FOR TREATING B CELL MALIGNANCIES

- More efficacious in ALPINE, safer in ALPINE and ASPEN, and more convenient than ibrutinib in clinic
- **•** Great combination opportunities with Bcl-2, PI3-Kδ, Revlimid, etc.
- Additional new molecules in the pipeline
  - BTK-CDAC to overcome resistance
  - CAR-NK, bispecifics, ADC, etc.

### Growing AML/MDS portfolio

- Our investigational Bcl-2 inhibitor, BGB-11417, has better pharmacological properties than venetoclax, similar story as ZANUBRUTINIB
- Interesting new molecules targeting CD70, TIM3, CD33, FLT3, & MCL-1
# SOLID TUMOR CLINICAL PROGRAM UPDATE

Yong (Ben) Ben, M.D. CMO, I/O

# SUMMARY OF SOLID-TUMOR CLINICAL PORTFOLIO



**BEICENE** 



# DIFFERENTIATED PD-I ANTIBODY WITH ENGINEERED FC TO AVOID ADCP

#### Nivo/Pembro with FcyRI-binding



#### Tislelizumab without FcyRI-binding



#### Tislelizumab for cHL Patients



#### **Competitor Data for Trials with Similar Patient Population**

	ORR	CR	mPFS
Camrelizumab	<b>76.0%</b> (95%Cl: 64.7-85.1)	26.7%	<b>22.5m</b> (95%Cl: 14.7-NR)
Sintilimab	<b>85.4%</b> (95%Cl: 76.7-91.8)	<b>29.2</b> %	<b>18.6m</b> (95%Cl: 14.4-22.3)

Long-term results including safety data were presented at EHA 2021: 1. Song Y et al. Abstract S207. Cross trial comparison; competitor data from EHA Library. Wu J. 06/09/21; 324614; S206 / Journal of Clinical Oncology 37, no. 15\_suppl (May 20, 2019) 7533-7533 / Journal of Clinical Oncology 38, no. 15\_suppl (May 20, 2020) 8034-8034.



# CLINICAL DEVELOPMENT PROGRAM



#### 5 Approvals and 4 sBLAs under Review in China

- Approved: IL non-sq NSCLC, IL sq-NSCLC, 2/3L HCC, 2/3L UBC and R/R cHL
- Under review: IL NPC, 2/3L NSCLC, 2L ESCC and 2/3L MSI-High

#### II Additional Registrational Trials On-going/Planned

- 7 as monotherapy or in combo with SOC expected to readout in next 36 months
- 4 combination registration trials ongoing/planned to be initiated with ociperlimab, zanidatamab or sitravatinib

#### Collaboration with Novartis

- For N. America, Europe and Japan filing and marketing opportunities
- Explore therapeutic potential of tislelizumab in combination strategies

#### **Global Program Positioned for Global Access**

- 5,300+ pts treated with tislelizumab on BeiGene clinical trials, 1,700+ pts from exmainland China
- Collaboration with Boehringer Ingelheim, one of the world's leading biologics manufacturers

STRONG SOLID TUMOR PORTFOLIO CENTERED AROUND PD-I

Overcome PD-1 Ab resistance via TME re-modeling

Sitravatinib, TIM3, PI3Kô, DKK1



Enhance PD-1 response via releasing other immune checkpoint inhibition TIGIT, HPKI





Activate T cell via stimulation agonists

**OX40**, bispecific T cell engager

Sensitize tumors to PD-1 Ab via immunogenic cell death Zanidatamab, ZW49





# OCIPERLIMAB (TIGIT)

AUGMENTING PD-I'S ANTI-TUMOR ACTIVITY IN PD-I SENSITIVE TUMORS

- Positioned to be combined with tislelizumab in PD-1 sensitive tumors
  - Mechanistically synergize with PD-I
- Highly potent, with intact Fc function
  - Fc function shown to be critical to activity preclinically
- One of the most advanced anti-TIGIT antibodies
  - 200+ pts have been dosed with oci/tisle combination
    - Early efficacy observed
    - Generally well-tolerated, no additional safety signals detected
  - Two Phase 3s initiated earlier this year, broad clinical programs ongoing



Phase 3 Molecules	Company	# of On-going Ph3 Trials	Fc Function	Phase 3 Indications
Tiragolumab	Roche	5	Intact	NSCLC; ES- SCLC; ESCC
Ociperlimab	BeiGene	2	Intact	NSCLC
Vibostolimab	Merck	I.	Intact	NSCLC
Domvanalimab	Arcus/ (Gilead)	L	Null	NSCLC

## OCIPERLIMAB (TIGIT) CURRENT CLINICAL DEVELOPMENT PROGRAM

	Phase Ia	Phase Ib	Phase 2	Phase 3
Untreated, locally advanced, unresectable NSCLC	AdvanTIG-301, Ociperlimab + Tislelizur	mab + CRT		Global
IL PD-LI+ NSCLC	AdvanTIG-302, Ociperlimab + Tislelizur	mab		Global
IL All-Comer NSCLC	AdvanTIG-205, Ociperlimab + Tislelizur	mab + Chemo (planned 2H 2021)	Global	
IL LS-SCLC	AdvanTIG-204, Ociperlimab + Tislelizur	mab + CRT	Global	
2/3L CC	AdvanTIG-202, Ociperlimab + Tislelizur	mab	Global	
2/3L ESCC	AdvanTIG-203, Ociperlimab + Tislelizur	mab	Global	
IL HCC	AdvanTIG-206, Ociperlimab + Tislelizur	mab + bevacizumab biosimilar	China	
NSCLC (sq, non-sq, PD-L1+, CPI), ES-SCLC, ESCC, EAC, HNC, GC	BGB-900-105, Ociperlimab +Tislelizum	ab <u>+</u> chemo Global		



## SITRAVATINIB REVERSE PD-1 RESISTANCE BY CHANGING TUMOR MICROENVIRONMENT

- A potent inhibitor of VEGFR and tumor-associated macrophage (TAM) family kinases (AxI, Tyro3, & Mer) to reverse immune resistance
  - Inhibit TAM activities (polarization and efferocytosis) that appear to be critical in establishment of an immunotolerant state
  - Block angiogenesis
- Clinical activity in combination with PD-I
  - OS in PD-1 failed NSCLC: 15.6 months
  - ORR in PD-1 failed bladder cancer: 27.3%
  - ORR in PD-1 failed melanoma: 24.0%
  - ORR in immune-cold PROC: 26.4%
- Pivotal studies in NSCLC initiated

#### **Clinical Development Plan**

In collaboration with Mirati for Asia (ex-Japan), Australia and New Zealand

	Phase Ia	Phase Ib	Phase 2	Phase 3
Sq. & Non-sq. NSCLC, RCC, OC, Melanoma	BGB-900-103, -	Tislelizumab		BeiGene
HCC, GC	BGB-900-104, -	-Tislelizumab		BeiGene
Sq. & Non-sq. NSCLC	BGB-A317-sitra	watinib-301, +Tislel	izumab	BeiGene

Mirati conducting other studies, including the Phase 3 SAPPHIRE study in NSCLC



# BROAD REGISTRATION PROGRAMS IN NSCLC

On-going Approval Filing

NSCLC	Tislelizumab		Ociperlimab	Sitravatinib
Neoadjuvant/ Adjuvant	<b>Tisle (PD-1) + Chemo</b> RATIONALE-315			
Untreated, Locally Advanced, Unresectable			<b>Oci (TIGIT) + Tisle + CRT</b> AdvanTIG-301	
I <sup>st</sup> Line	<b>Tisle (PD-1) + Chemo</b> (SQ) RATIONALE-307	<b>Tisle (PD-1) + Chemo</b> (Non-SQ) RATIONALE-304	<b>Oci (TIGIT) + Tisle (PD- LI+)</b> AdvanTIG-302	
2 <sup>nd</sup> or 3 <sup>rd</sup> Line	<b>Tisle (PD-1 naïve)</b> RATIONALE-303			<b>Sitra + Tisle (PD-1 R/R)</b> BGB-A317-sitravatinib-301

## ZANIDATAMAB (ZW25) HER2XHER2 BISPECIFIC HER2 ANTIBODY TARGETING HER2 EXPRESSING TUMORS

- Target two distinct HER2 epitopes, ECD2 (trastuzumab binding domain) and ECD4 (pertuzumab binding domain)
  - Increased tumor cell binding and enhanced HER2 internalization compared with trastuzumab
  - Superior to trastuzumab and pertuzumab in preclinical studies
- Encouraging anti-tumor activity as monotherapy across multiple HER2-expressing tumor types
  - 33% ORR in heavily-pretreated HER2 breast cancer
  - 33% ORR in late-line HER2 gastric cancer
  - 40% ORR in refractory HER2 biliary tract cancer
- Pivotal studies ongoing/planned
  - Phase 2 single arm study in 2L+ BTC initiated Oct 2020
  - Phase 3 in IL GC planned to be initiated in H2 2021

#### **Clinical Development Plan**

In collaboration with Zymeworks for Asia (ex-Japan), Australia and New Zealand



**BEICENE** 

# BGB-445 (OX-40)

## UNIQUE OX-40 AGONIST ANTIBODY THAT DOES NOT BLOCK LIGAND-BINDING

- Distinguished MOA vs. other OX-40 antibodies in the clinic
  - Non-ligand blocking to retain OX-40 ligand signaling in APC cells
  - Maximize OX-40 signaling in T cells
- Clinical progress
  - Dose escalation on-going as monotherapy and in combination with tislelizumab
  - No DLT to date
  - Clinical activity observed in both monotherapy and combination-therapy cohorts
  - Plan to start dose expansion/Phase 2 later this year or early next year

Other OX40 antibodies in clinic are ligand-blocking, which lose OX-40L signal in APC cells



BGB-445 is a non-ligand-blocking OX40 Ab which retains OX-40L signal on APC cells and maximizes T cell activation



## BGB-15025 (HPKI) POTENTIAL FIRST-IN-CLASS HPKI INHIBITOR

- Positioned to be combined with tislelizumab in PD-1 sensitive tumors
  - HPK1, an intracellular immune checkpoint to negatively regulate T cell activity
  - Based on MOA, HPK1 inhibitor will synergize with anti-PD-1 antibody
- HPK1 is a hard-to-hit target
  - Challenging to achieve good potency and selectivity due to very low ATP Km
- BGB-15025 has good pharmacological properties
  - Good kinome selectivity
  - Robust anti-tumor efficacy
  - Well-tolerated in animal studies
- Clinical progress
  - Phase I dose escalation on-going







## **DEEP SOLID-TUMOR PORTFOLIO**

Broad global clinical development programs built around tislelizumab, a differentiated PD-1 antibody Multiple combination strategies to enhance response and overcome resistance New assets entering late stage & new wave of molecules planned in the next section

# NEW MODALITIES AND PLANNED PIPELINE

Lai Wang, Ph.D. Global Head of R&D

## BEICENE ENHANCE THE NEXT WAVE OF DRUG DISCOVERY BY INVESTING IN NEW MODALITIES

### 2011-2020





## BEICENE ENHANCE THE NEXT WAVE OF DRUG DISCOVERY BY INVESTING IN NEW MODALITIES





## **Power of New Platforms**

- Drugging the undruggable
- Precise tumor targeting



CDAC, Chimeric Degradation Activating Compound; ADC, Antibody Drug Conjugate; BsAb, Bispecific Antibody; TsAb, Trispecific Antibody





## **Power of New Platforms**

- Drugging the undruggable
- Precise tumor targeting



# DRUGGING THE UNDRUGGABLE



## CHIMERIC DEGRADATION ACTIVATING COMPOUND (CDAC) PLATFORM



## Advantage

Differentiated from traditional small molecules by inducing target degradation

- Ability to target proteins with or without enzymatic activity
- Does not require strong and sustained target binding
- Eliminate the scaffolding function of the target

# OUR DIFFERENTIATION

Build strong capability to leverage a wider spectrum of E3 ligases: ubiquitous and tissue specific

- Reduce dose-limiting toxicities
- Overcome E3-relevant drug resistance
- Broaden substrate spectrum



**BEICENE** 

CDAC

## **BROAD CDAC PROGRAMS**





**BCR Signaling** BTK CDAC to target **BTK** scaffolding function



Receptor **Tyrosine Kinase** CDAC A to treat lung cancer



**Cell Cycle** CDAC B applicable in heme



**Apoptosis** CDAC C applicable in heme and solid tumors



**TLR** signaling CDAC D applicable in heme and I-O therapies



Cytokine signaling CDAC E applicable in heme and I-O therapies



**Transcription**/ epi-genetics CDAC F applicable in heme

# BTK CDAC PROGRAM FOR B CELL MALIGNANCES

- **Next generation** BTK inhibitor to enhance BTK franchise
  - ► To overcome zanubrutinib and other BTKi resistance
  - To destroy non-kinase (scaffolding) function
- BGB-16673, BeiGene's first CDAC molecule planned for clinic
  - Good pharmacological properties
    - Highly potent and selective
    - Good oral bioavailability
    - Long t<sub>1/2</sub>
  - Well-tolerated in animal studies
  - First patient expected to be dosed in 2H202I

#### BGB-16673 Can Overcome C481S Resistance

C481S Tumor Model











## **Power of New Platforms**

- Drugging the undruggable
- Precise tumor targeting





# TUMOR SELECTIVE TARGETING APPROACH

## Tumor Associated Antigen (TAA) Toolbox **10+** TAAs are being developed, many associated with multiple tumor types Lung Cancer Breast Cancer Gastric Cancer (11 TAAs) Esophageal Cancer (5 TAAs) Colon Cancer (8 TAAs) Gio Kidney Cancer (2 TAAs) D Liver Cancer (I TAAs) Lymphoma (2 TAAs) 0<sub>6</sub>⊖ Leukemia ⊙0 (I TAAs) **Inactive Pro-Cytokine**

# Tumor Selective Immune Cell Activation

- Bi/Tri-specific TAA-Immune cell engager
- CAR-NK
- Immune stimulating-ADC

## **Tumor Targeted Cell Killing**

Toxin delivering-ADC

## **Tumor Selective Cytokine Release**

**BEICENE** 

# ACTIVATING INTRA-TUMOR IMMUNE CELLS



**BEICENE** 

### Tumor Selective Immune Cell Activation via TAA-oriented Engagement of T/NK Receptors

BsAb/TsAb

#### **TAA-targeting**

Mono-specific or dual targeting





### Comprehensive BsAb/TsAb Discovery Platforms to Fulfil Diverse Formats and MoAs



### Llama SdAb platform Small Ab unit to be easily used in diverse formats



#### Human SdAb phage library Bypass humanization



**Common light chain platform** IgG-like w/o additional engineering on Fab



**ScFv engineering platform** Convert Fab to good-behave ScFv

# OFF-THE-SHELF TUMOR TARGETING NK CELLS

## BE<mark>IC</mark>ENE



### MoA of CAR-NK



#### Collaboration



iPSC differentiation NK function enhancement Unique CAR platforms GMP capability readiness ScFv generation IND-enabling studies Global clinical development & commercialization

BeiGene

#### Positioning

- **Shoreline Biosciences collaboration:** fast-track first wave products to development
- Internal capability build-up: enable Shoreline Biosciences collaboration and pipeline expansion

#### **Planned Portfolio**

- Start with iPSC-derived, allogeneic CAR-NK cell therapies
- Expand therapeutic cell types and utilities as appropriate

# TAA DIRECTED MISSILE



ADC

#### Toxin-based ADC as Targeted Bomb



#### Immune-Stimulating Antibody Conjugate (ISAC)

 Tumor-targeted delivery of immune agonists to turn COLD tumors HOT



# **NEXT GENERATION ADC** designed to have superior efficacy and safety

- Site specific conjugation via Ambrx technology to achieve homogeneous drug antibody ratio (DAR)
- Innovative linker design to improve ADC biophysical property and stability, and to enhance tumorselective cleavage



# TUMOR SELECTIVE CYTOKINE ACTIVATION



### **Scientific rationale**

- Cytokine therapy to activate the immune system has been an attractive treatment modality in clinical cancer research
- Severe systemic toxicity hinders the broad application of immune-stimulating cytokines

**BeiGene's pro-cytokine design** for tumor-specific cytokine-mediated immune cell activation with minimized peripheral adverse events

- **Complete** masking in circulation
- Full recovery of cytokine activity once cleaved
- Prolonged half life of the pro-cytokine form
- Plug-and-play format applicable to broad cytokine spectrum

MoA of pro-cytokine



## PRECLINICAL PIPELINE



- 50+ programs ongoing, close to half with first-in-class potential
- IO+ planned to clinic in the next 24 months

## Going beyond oncology

- First immunology/inflammation asset planned to enter clinic later this year
- BeiGene institute to focus on early technology and translational science

# CANCER-ATTACKING IMMUNE MODULATING STRATEGIES

## TAA DRIVEN THERAPEUTICS NOT INCLUDED



**BEICENE** 

CDAC

**Pro-cytokine** 

SM, mAb

0



# ONLY THE BEGINNING

Productive first decade establishes scientific track record

- Zanubrutinib and tislelizumab set stage for leading franchises
- Deep portfolio built around these cornerstone assets many moving into pivotal stage

BeiGene's R&D model stands to deliver significant innovation in the next decade

- Discovery engine has flourished and diversified
- Clinical pipeline expected to accelerate
- Poised to develop novel, science-based combinations
- Unique clinical development model and truly global, inhouse capabilities at scale, speed and cost advantages

# CLOSING REMARKS



# OUR TRUE NORTH

Continued innovation and expansive growth planned in next 10 years

Amazing research across number of platforms with scale and talent

Future innovation requires deep portfolio with breadth of modalities (combinations)



4

2

One of the largest research organizations with executional and cost focus

Speed, cost advantage to accelerate combination strategy



## **Q&A PARTICIPANTS**



Xiaodong Wang, Ph.D. Chairman of Scientific Advisory Board & Co-Founder



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Josh Neiman Chief Commercial Officer, North America and Europe

# THANKYOU

## ABBREVIATIONS

Ab	Antibody
Acala	Acalabrutinib
ADC	Antibody drug conjugate
ADCP	Antibody-dependent cellular phagocytosis
AE	Adverse Event
AE/UAE	United Arab Emirates
AFIB	Atrial fibrillation
ALL	Acute lymphocytic leukemia
ALT	Alanine aminotransferase
AML	Acute myeloid leukemia
ANZ	Australia and New Zealand
APAC	Asia Pacific
APC	Antigen-presenting cells
APC	Acute pancreatic cancer
AST	Aspartate aminotransferase
BC	Breast cancer
BCL-2	B-cell lymphoma-2
BCRP	Breast cancer resistance protein
BI	Boehringer Ingelheim
BIC	Bayesian Information Criterion
BIC	Best-in-Class
BID	Twice a Day
BLA	Biologics License Application
BMS	Bristol Myers Squibb
BR	Bendamustine and Rituximab
BsAb	Bispecific Antibody
BTC	Biliary tract cancer
втк	Bruton's Tyrosine Kinase
BTK-CDAC	Bruton tyrosine kinase-Chimeric Degradation Activating Compound
CA	California

CA	Canada
CAR	Chimeric antigen receptor
CAR-NK	Chimeric antigen receptor-Natural Killer cell
CC/CRC	Colorectal Cancer
CCRCC	Clear cell renal cell carcinoma
CDAC	Chimeric Degradation Activating Compound
Chemo	Chemotherapy
cHL	Classical Hodgkin's Lymphoma
СНО	Chinese hamster ovary
CI	Confidence interval
CL	Chile
CLD	Chronic liver disease
CLL	Chronic lymphocytic leukemia
СМС	Chemistry, Manufacturing and Controls
CN	China
COVID-19	Coronavirus disease 2019
CR	Complete Response
CRBN	Cereblon
CRO	Contract Research Organization
CRT	Conformal Radiation Therapy
CRT	Concurrent Radiochemotherapy
СТ	Computed tomography
стс	Circulating tumor cell
CTL	Cutaneous T-cell lymphoma
DAR	Drug to antibody ratio
DC	Discontinuation from study treatment
DCP	Decision Check Point
DDI	Drug-drug interaction
del(17p)	Deletion of 17p

# ABBREVIATIONS (CONT'D)

DKKI	Dickkopf-1
DLBCL	Diffuse large B-cell lymphoma
DLT	Dose limiting toxicity
dMMR	Deficient Mismatch Repair
DOR	Duration of Response
EAC	Esophageal adenocarcinoma
ECD2	Extracellular domain 2
ECD4	Extracellular domain 4
EHA	European Hematology Association
EMEA	European Medicines Evaluation Agency
EMEA	Europe Middle East Africa
Er+	Oestrogen receptor positive
ESCC	Esophageal Squamous-Cell Carcinoma
ES-SCLC	Extensive-stage small cell lung cancer
EU	European Union
Fab	Fragment of antigen binding
FCR treatment	Fludarabine, cyclophosphamide and rituximab treatment
FcγR	Fcy receptor
FDA	US Food and Drug Administration
FIC	Fractional inhibitory concentration
FIC	First-in-Class
FIH	First-in-human
FL	Follicular lymphoma
GBM	Glioblastoma
gBRCAm	Germline Breast Cancer mutation
GC	Gastric cancer
GC/GRJC	Gastric cancer/Gastroesophageal Junction Cancer
GCTB	Giant cell tumor of bone
GEA	Gastroesophageal Adenocarcinoma
GMP	Good Manufacturing Practice

нс	Headcount
HCC	Hepatocellular Carcinoma
Heme	Hematological malignancies
HER2	Human Epidermal Growth Factor Receptor 2
HL	Hepatic lipase
HL	Hodgkin lymphoma
HLE BITE	Half-life extended Bi-specific T-cell engagers
HNC	Head and neck cancer
HNSCC	Head and Neck Squamous Cell Carcinoma
HPK-I	Hematopoietic progenitor kinase I
HR	Hazard ratio
HRD	Homologous recombination deficiencies
HTH	Head to Head
HTS	High-throughput sequencing
1/1	Immunology/Inflammation
IgG-like BsAb	IgG-like bispecific antibodies
IGHV	Immunoglobulin heavy chain variable region
IL	Israel
IL	Interleukin
IND	Investigational new drug
I-O	Immuno-oncology
IPD	Integrated Product Development
iPSC	Induced pluripotent stem cells
IRC	Independent Review Committee
ISAC	Immune Stimulating Antibody Conjugate
ISR	Incurred sample reanalysis
JP	Japan
KOL	Key Opinion Leader
KRAS	Kirsten Rat Sarcoma Viral Oncogene Homolog
LN	Ludus Nedhritis
## **BEICENE**

## ABBREVIATIONS (CONT'D)

MA	Massachusetts
mAb	Monoclonal Antibody
MCL	Mantel Cell Lymphoma
McI-I	Myeloid cell leukemia I
mCRPC	Metastatic Castration-Resistant Prostate Cancer
MDM2 gene	Murine double minute 2 gene
MDS	Myelodysplastic syndromes
MDSC	Myeloid-derived suppressor cell
MEL	Melanoma
MM	Multiple Myeloma
MoA	Mechanism of Action
mPFS	Median Progression-free survival
Mpfs	Progression-free survival duration in months
MPN	Myeloproliferative neoplasm
MRD	Minimal Residual Disease
MRI	Magnetic resonance imaging
mRNA	Messenger RNA
MRR (CR+VGPR+PR)	Major response rate
MSI-H	Microsatellite Instability High
MZL	Marginal zone lymphoma
Mφ/DC	Macrophages (MΦ) /dendritic cells (DC)
N/SCLC	Non-/small cell lung cancer
NA	Not applicable
NALA	North America Latin America
NCT	National Clinical Trial Network
NDA	New Drug Application
NE	Not Evaluable
NHL	Non-Hodgkin's lymphoma

NK cell	Natural killer cells
NMPA	National Medical Products Administration
Non-sq. NSCLC	Non-squamous non-small cell lung cancer
NPC	Nasopharyngeal carcinoma
NR	Not reached
N-RAS	Neuroblastoma RAS Viral Oncogene Homolog
NRDL	National Reimbursement Drug List
NSCLC	Non-small-cell lung cancer
Obin	Obinutuzumab
oc	Ovarian Cancer
oci	Ociperlimab
ORR	Overall Response Rate
OS	Overall Survival
OX-40 Ab	OX-40 Antibody
OX-40 L	OX-40 Ligand
PARP	Poly ADP-Ribose Polymerase
Pca	Prostate Cancer
PD	Disease Progression
PD-I	Programmed Cell Death protein I
PD-LI	Programmed Cell Death Ligand I
Pembro	Pembrolizumab
PFS	Progression-Free Survival
P-Gp	P-glycoprotein
ΡΙ3Κδ	Phosphoinositide 3-kinase delta
PO	Oral
PPI	Proton Pump Inhibitor
PR	Partial Response
PROC	Platinum-resistant Ovarian Cancer
PSOC	Platinum-sensitive Ovarian Cancer
pt	Patient

## **BEICENE**

## ABBREVIATIONS (CONT'D)

QD	Once a Day
R&D	Research and Development
R/R	Relapsed / Refractory
R/R cHL	Relapsed/refractory Classical Hodgkin Lymphoma
RCC	Renal Cell Carcinoma
RP2D	Recommended Phase 2 Dose
sBLA	Supplemental Biologics License Application
ScFv	Single-Chain Fragment Variable
SCLC	Small Cell Lung Cancer
SD	Stable Disease
SEC	Securities and Exchange Commission
Sitra	Sitravatinib
SK	South Korea
SLL	Small Lymphocytic Lymphoma
SM	Small Molecule
SOC	Standard of Care
SPD	Solution Program Development
Sq. NSCLC	Squamous Non-small Cell Lung Cancer
SVP	Subvirus Particle
TAA	Tumor associated antigen
ТАМ	Tamoxifen
TCR	T cell receptor
TIGIT	T cell immunoreceptor with Ig and ITIM domain
TIM-3	T-cell immunoglobulin and mucin-domain containing-3
Tisle	Tislelizumab
ткі	Tyrosine kinase inhibitor
TLR	Toll-like receptor
TME	Tumor Microenvironment
TN	Treatment-naïve
TPD	Technology Platform Development

TR	Technical Review
TsAb	Trispecific Antibody
TTR	Time to response
Ub	Ubiquitin
UBC	Urinary bladder cancer
UC	Urothelial carcinoma
US	United States
VEGF	Antivascular endothelial growth factor
VEGFR	Vascular endothelial growth factor receptor
VGPR	Very Good Partial Response
w/o	without
WM	Waldenström's Macroglobulinemia
YTD	Year to Date