NASDAQ: IDYA



IDEAYA Biosciences

Improving Lives
Through Transformative
Precision Medicines



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Other

This presentation concerns anticipated products that are under clinical investigation and which have not yet been approved for marketing by the FDA or any other country regulatory authority. These anticipated products are currently limited by Federal law to investigational use, and no representation is made as to their safety or effectiveness for the purposes for which they are being investigated.

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IDEAYA Biosciences Highlights

Leading Precision Medicine Oncology Biotechnology Company Advancing Potential First-in-Class Therapies

Target Milestone Guidance on Broad Pipeline of 6 Clinical & 3 Preclinical (IND-enabling) Programs:

PHASE 2/3 PHASE 1/2 PRECLINICAL IDE397 (MAT2A) IDE275 / GSK959 (WERNER) NEXT GEN PROGRAMS DAROVASERTIB (PKC) Phase 1/2 mono expansion ongoing Daro + Crizo 1L HLA-A2(-) MUM Medical conference update – H1 2025 • IDE892 DC (MTA-cooperative PRMT5) potential registrational Ph2/3 IDE397 + Trodelvy® (Trop2-ADC) IND submission – Mid-2025 IDE161 (PARG) Targeting Expansion in MTAP UC – Q1'25 median PFS readout – by YE 2025 IDE034 DC (B7H3/PTK7 Bi-Specific Phase 1 mono expansion ongoing Clinical data update in MTAP UC – 2025 Daro + Crizo Ph2 1L MUM median ADC) IND submission - H2 2025 Expansion into NSCLC OS readout – 2025 IDE161 + Merck's anti-PD-1, KEYTRUDA® Daro Ph2 Neoadjuvant UM clinical *IDE397 + PRMT5* (pembrolizumab) IDE251 DC (KAT6/7) IND submission – data and regulatory update - H1 Wholly-owned clinical combo with • Phase 1 expansion in EC – 2025 H2 2025 2025 IDE892 (IDEAYA PRMT5) – H2 2025 IDE161 + Topo1i-ADC Daro Ph3 Neoadjuvant UM Enable clinical combo(s) – 2025 IDE849 / SHR-4849 (DLL3 ADC) registrational trial initiation – H1 Clinical data update and combo 2025 IDE705 / GSK101 (POL THETA) initiation with IDE161 - 2025 Phase 2 expansion (\$10M Milestone)

Pharma Collaborations











~\$2B in potential milestones

Financials and Investor Relations

~\$1.1B to fund operations at least into 2028 1, 2

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- (1) Includes aggregate of \$1.1 billion of cash, cash equivalents and marketable securities as of December 31, 2024
- (2) IDEAYA's Form 10-K dated February 18, 2025, as filed with the U.S. Securities and Exchange Commission
- KEYTRUDA® is a registered trademark of Merck Sharp & Dohme LLC, a subsidiary of Merck & Co, Inc, Rahway NJ, USA



IDEAYA Precision Medicine Oncology Platform

Fully-Integrated Target, Biomarker, Drug Discovery and Translational Capabilities

Drug Discovery and Pharmacological Validation

Structure Based Drug Design

Small Molecule Chemistry Protein Degrader Capabilities

Dual CRISPR, CRISPR, Chemogenomics **Genetically Engineered Models**

Bioinformatics, including AI Algorithms

Target & Biomarker

Discovery and Validation

- Key emerging novel targets identified, such as Werner Helicase, PARG and Pol Theta Helicase
- DECIPHER™ Dual CRISPR SL Library in DDR Cell Lines in collaboration with UCSD
- PAGEO™ Paralogous Gene Evaluation in Ovarian in collaboration with Broad Institute
- Machine Learning and Multi-Omics platform

Translational Research and Opportunity Expansion



Genomics – DNA and RNA Analysis Proteomics – Protein Expression Profiling Tissue (IHC, IF) and Liquid Biopsies Analysis

- transformative combinations
 - Opportunity expansion through broad cell panel screening

clinical biomarkers and

Translational research to define

- Pharmacodynamic biomarker analysis to confirm target modulation and correlation with clinical activity
- Crystal structures for SL discovery programs obtained to enable structure-based design
- INQUIRE™ Chemical Library proprietary, expert-curated small-molecule library
- HARMONY™ Machine-Learning engine empowers drug discovery platform
- Differentiated clinical / candidate compounds discovered, including IDE397, IDE275 (GSK959), IDE161, and IDE705 (GSK101)



IDEAYA Precision Oncology Drug Discovery Platform & IND Engine

AI/ML & Structure Based Drug Design to Deliver Potential First-in-Class Development Candidates

Structural Biology & Structure Based Drug Design

Full suite of capabilities in structural biology, biophysics, & computational chemistry

Ligand bound co-crystal structures resolved to enable Structure Based Drug Design for each of the Synthetic Lethality drug-discovery programs

Multiple potential "first-in-world" co-crystal structures resolved, including for Werner Helicase, PARG and Pol Theta Helicase

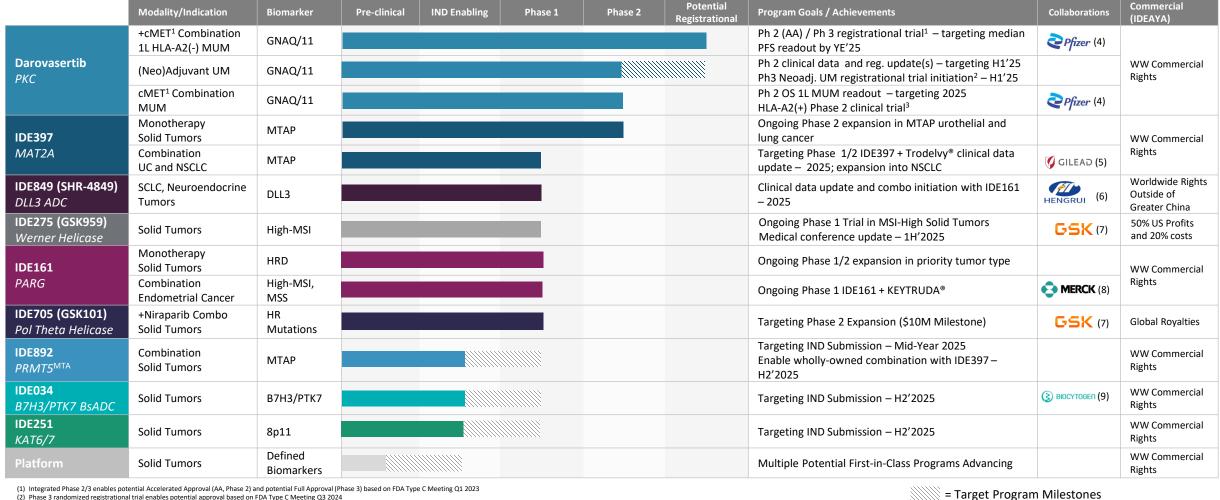
INQUIRE™ Proprietary Chemical Library

Expert-curated HTS library to enhance hit discovery capabilities against novel SL targets classes

Enhances IDEAYA's SL Drug Discovery Platform and competitive differentiation

AI/ML Enabled Computational Drug Discovery¹ Make Fewer Compounds Make Better Compounds **HARMONYTM ML and** FEP discovery engine In-synthesis Accurate properties prediction **Hypothesis** Creative design solutions Compoun AI/ML to Accelerate Time to IND for Potential First-in-Class DCs

IDEAYA's Potential First-in-Class Precision Medicine Oncology Pipeline



⁽¹⁾ Integrated Phase 2/3 enables potential Accelerated Approval (AA, Phase 2) and potential Full Approval (Phase 3) based on FDA Type C Meeting O1 2023

MAT2A = Methionine Adenosyltransferase 2a, MTAP = Methylthioadenosine Phosphorylase, MTA = Methylthioadenosine Phosphor PKC = Protein Kinase C, MUM = Metastatic Uveal Melanoma, UM = Uveal Melanoma, Crizo = Crizotinib, UC = Urothelial Cancer, NSCLC = Non-Small Cell Lung Cancer, WW = Worldwide, HLA-A2(-) = HLA-A2(+) = HLA-A2(-) = PLA-A2(-) =



⁽²⁾ Phase 3 randomized registrational trial enables potential approval based on FDA Type C Meeting Q3 2024

⁽³⁾ Targeting enrollment of additional HLA-A2(+) patients in ongoing IDE196-001 Phase 2 clinical trial

⁽⁴⁾ Pursuant to Pfizer Clinical Trial Collaboration and Supply Agreements for Darovasertib/Crizotinib Combination; IDEAYA retains all Darovasertib Commercial Rights

⁽⁵⁾ Pursuant to Gilead Clinical Study Collaboration and Supply Agreement for IDE397 + Trodelvy*, a Trop-2 directed antibody-drug conjugate (ADC); the Company will sponsor the study and Gilead will provide Trodelvy at no cost. Gilead retains all commercial rights to Trodelvy.

⁽⁶⁾ Pursuant to exclusive license agreement with Jiangsu Hengrui Pharmaceuticals Co., Ltd for worldwide rights outside of Greater China

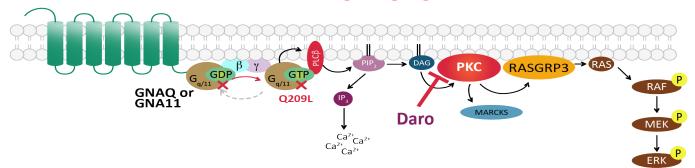
⁽⁷⁾ Pursuant to GSK Collaboration, Option and License Agreement: Polθ: Global Royalties; WRN: 50/50 US Profits + ex-US Royalties

⁽⁸⁾ Pursuant to Merck Clinical Trial Collaboration and Supply Agreement for IDE161 + Keytruda*, an anti-PD-1 therapy; the Company will sponsor the study and Merck will provide Keytruda at no cost

⁽⁹⁾ Pursuant to exclusive worldwide licensing and option agreement with Biocytogen

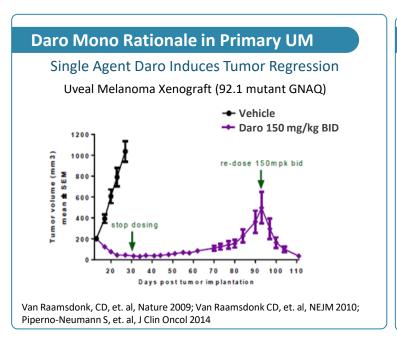
Darovasertib: Potential to Broadly Impact Uveal Melanoma (UM) Potential First-in-Class and Best-in-Class in (Neo)adjuvant UM and Metastatic UM (MUM)

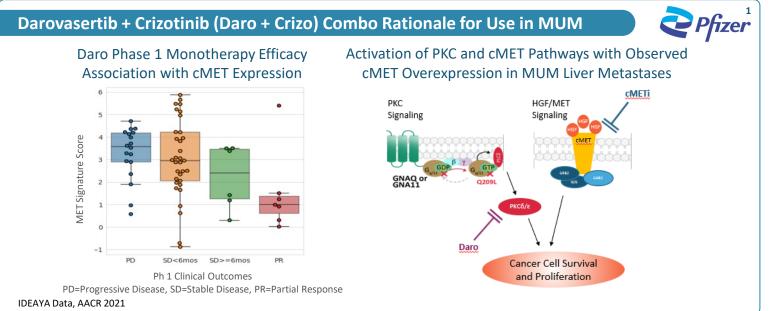
Mutations in GNAQ / GNA11 activate PKC Signaling, a genetic driver of Uveal Melanoma



Darovasertib is an oral, potent and selective PKC inhibitor GNAQ or GNA11 (~95%) and other upstream mutations activate PKC signaling in UM and MUM patients

UM is typically treated with radiation and/or enucleation, with no approved systemic therapies for Neoadjuvant UM MUM occurs in approximately 50% of UM patients and predominantly as liver metastasis in ~90% of MUM patients, with no approved therapies for HLA-A*02:01 negative MUM

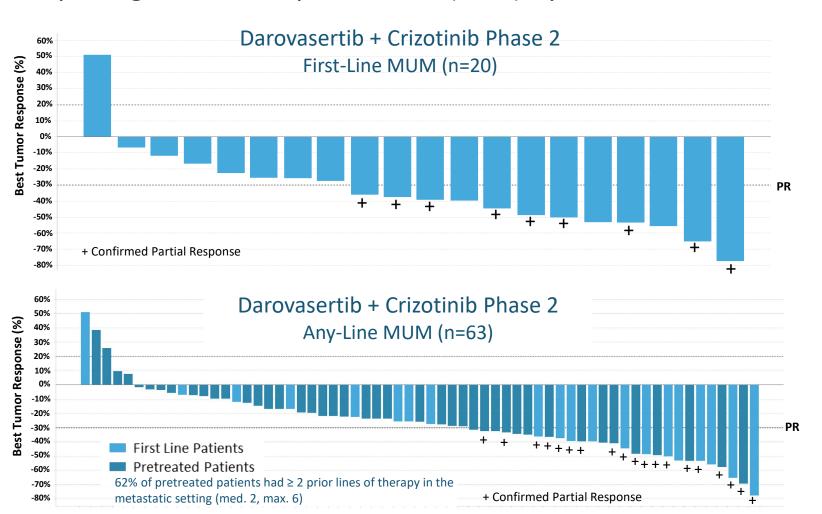






Daro + Crizo Phase 2 Efficacy: First-Line MUM and Any-Line MUM

Compelling Overall Response Rate (ORR) by RECIST 1.1 Observed



Confirmed 45% ORR and 90% DCR

Response by RECIST 1.1 First-Line MUM	Evaluable (N=20)
Confirmed ORR (9/20)	45%
Tumor Shrinkage (19/20)	95%
>30% Tumor Shrinkage (12/20)	60%
Best Overall Response	
cPR (9/20)	45%
SD (9/20)	45%
DCR (18/20)	90%

Confirmed 30% ORR and 89% DCR

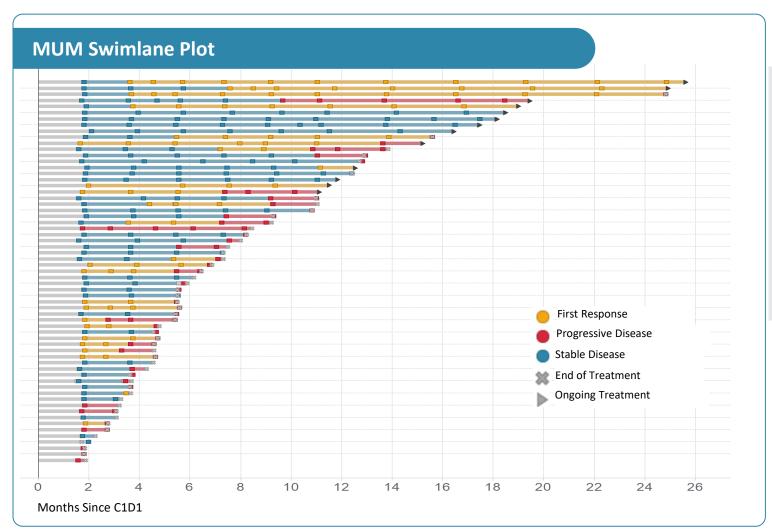
Response by RECIST 1.1 Any-Line MUM	Evaluable (N=63)
Confirmed ORR (19/63)	30%
Tumor Shrinkage (58/63)	92%
>30% Tumor Shrinkage (27/63)	43%
Best Overall Response	
cPR (19/63)	30%
SD (37/63)	59%
DCR (56/63)	89%





Median PFS in First-Line, Any-Line and Hepatic-Only MUM

Observed Compelling Median Progression Free Survival with Encouraging Trend



Darovasertib + Crizotinib Phase 2

Median Progression Free Survival

- First-Line (n=20): 7.1 months
- Any-Line (n=63): 6.8 months
- Hepatic-Only (n=19): 11.0 months

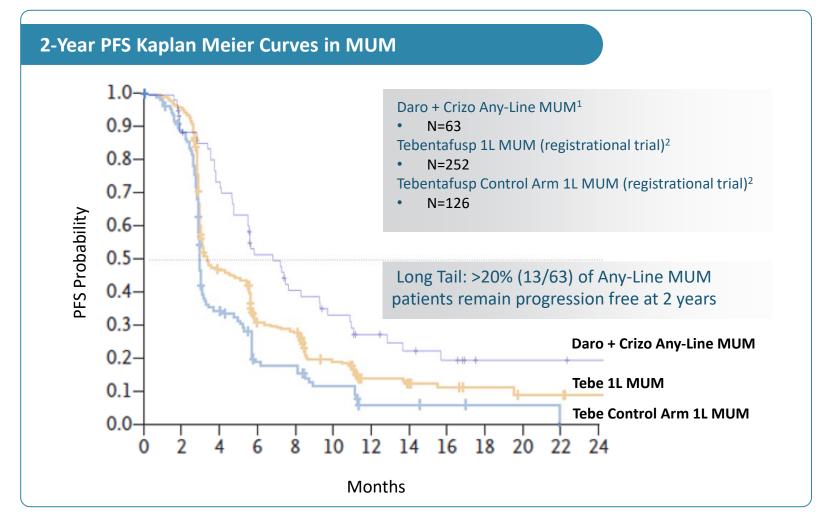
Treatment Duration – Observations

- ~50% of patients treated > 6 months
- ~30% of patients treated > 1 year



2-Year PFS Kaplan Meier (KM) Curve: Daro + Crizo in Any-Line MUM¹

Daro + Crizo Combo Observes a Promising 2-Year PFS KM Curve and a "Long Tail" Effect





Darovasertib + Crizotinib Combination Clinical Summary in MUM

Highly Differentiated Clinical Efficacy & AE Profile Observed^{1, 2}

	Darovasertib + Crizotinib	Cabozantinib Mono / Crizotinib Mono	Selumetinib + DTIC	lpi + Nivo	Tebentafusp
Target / Mechanism	PKC + cMET	сМЕТ	MEK + Chemotherapy	CTLA4 + PD-1	HLA-A2-0201 Bi-Specific
Study Name(s)	NCT03947385	A091201 ³ / NCT05063058 ⁴	NCT01974752 ⁵	NCT02626962 ⁶	IMCgp100-102 ⁷
Population	1L/2L/3L+ MUM (n=63)	1L+ MUM (n=31) / 1L (n=6) 2L (n=1) MUM	1L+ MUM (n=97)	1L MUM (n=52)	2L+ MUM (n=127)
Patient Selection	NA	NA / MET Overexpression	NA	NA	HLA-A2-positive
Drug Form	Oral Tablets	Oral Capsules	Oral Capsules + chemo	IV infusion	IV Infusion (Weekly)
Tolerability (Grade ≥3 Drug-Related AE)	31%	51.6% / NA	63% (All Cause)	58%	46.5%
% of Patients with Tumor Shrinkage	First-Line = 95% / Any-Line = 92% / Hepatic Only = 100% ⁸	23% ⁹ / NA	35% ⁹	27% ⁹	44% ⁹
Confirmed ORR% (by RECIST 1.1)	First-Line = 45% / Any-Line = 30% / Hepatic Only = 37% ⁸	0% / 0%	3%	11.5% (not confirmed ORR)	4.7%
Median PFS	First-Line: 7.1 months / Any-Line: 6.8 months / Hepatic-Only: 11.0 months ⁸	2 months / NA	2.8 months	3 months	2.8 months

⁽¹⁾ Cross-trial comparisons are not based on head-to-head studies and are presented for informational purposes; no direct comparisons are being made



⁽²⁾ ESMO 2022: Dimitriou, F, et. al: IPI + Nivo Combo in HLA-A2-0201 MUM reports ~6% ORR (2 PRs out of 33 patients)

⁽³⁾ Randomized Phase II Trial and Tumor Mutational Spectrum Analysis from Cabozantinib versus Chemotherapy in Metastatic Uveal Melanoma (Alliance A091201); Clin Cancer Res 2020;26:804–11

⁽⁴⁾ European Journal of Cancer, Leyraz, et. al, 2022; 146-155

⁽⁵⁾ Journal of Clinical Oncology, Carjaval, et. al, 2018; 1232-1239

⁽⁶⁾ ASCO 2021, Piulats, J, et. al, Ipi = Ipilimumab, Nivo = Nivolumab, ORR% did not require PR/CR confirmation

⁽⁷⁾ Based on Immunocore reported 2L+ study data (to reflect comparative patient population) and by independent review and ORR% was with confirmed PRs

⁽⁸⁾ ESMO 2023 Proffered Presentation McKean, M, et al: Preliminary analysis of unlocked database as of 08/22/2023 by investigator review; data cutoff based on treatment Day 1 of Cycle 1 (C1D1) as of 9/22/2022

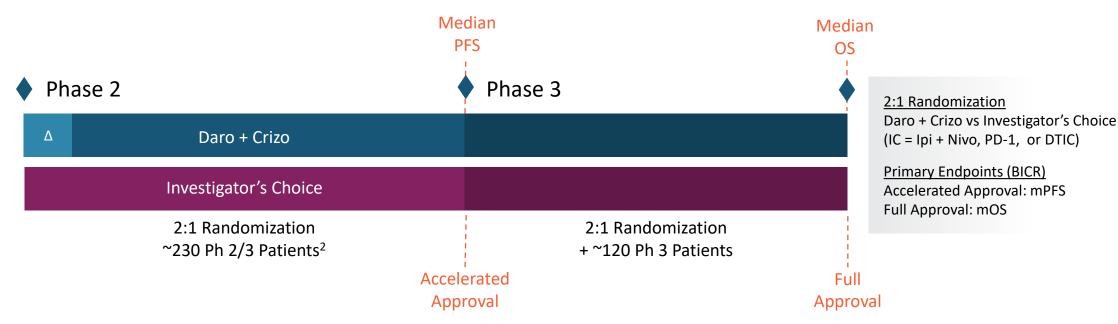
⁽⁹⁾ Estimated from Waterfall plot

Accelerated Approval Trial Design in First-Line HLA-A2 Negative MUM

FDA Guided Design: Open Label Ph2/3 Evaluation of Daro + Crizo vs. Investigator's Choice¹

FDA Project FrontRunner: Target First-Line approval strategy to enhance patient benefit in MUM **FDA Accelerated Approval:** Address unmet need in MUM, which has limited effective treatment options

Integrated Phase 2/3 Study within Study Enables Potential Accelerated Approval & Full Approval



FDA Fast Track and EMA SME Status Designation for Daro + Crizo in MUM



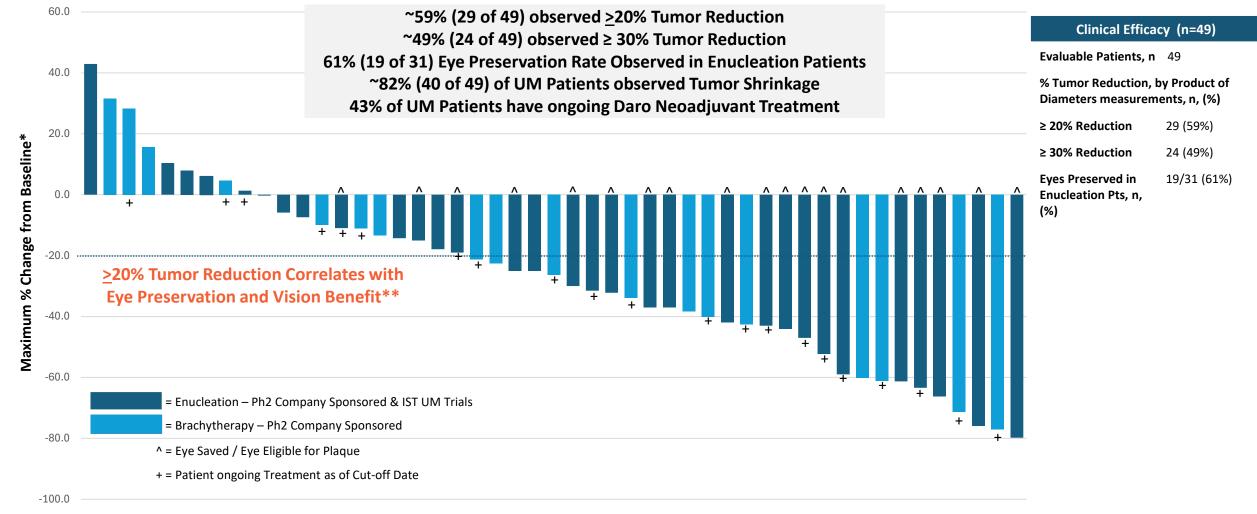
Clinicaltrials.gov: NCT05987332

⁽²⁾ Phase 2 study contemplates data set of n=200 patients randomized 2:1 with treatment arm at move forward dose in support of potential accelerated approval based on mPFS

^a Nested study to confirm move forward dose: (i) Daro 300 mg BID + Crizo 200 mg BID or (ii) Daro 200 mg BID + Crizo 200 mg BID

Daro = Darovasertib, Crizo = Crizotinib, MUM = Metastatic Uveal Melanoma, HLA-A2 = HLA-A2*02:01 Serotype, IC = Investigator's Choice, Ipi = Ipilimumab, Nivo = Nivolumab, DTIC = Dacarbazine

Darovasertib Neoadjuvant Therapy: Ph2 Company Sponsored & Ph2 IST UM Trials 61% (19 of 31) Observed Eye Preservation and 49% (24 of 49) with >30% Tumor Reduction*

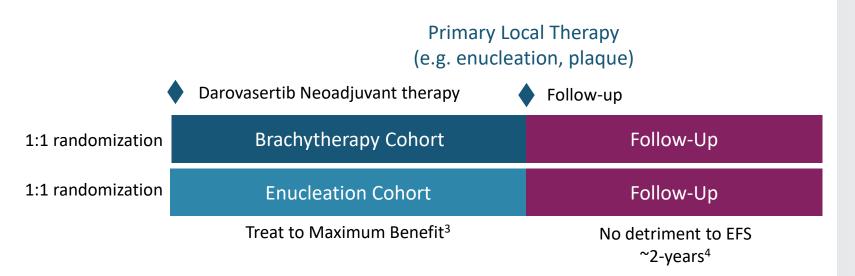




IST = Investigator Sponsored Trial

Preliminary Darovasertib Neoadjuvant UM Phase 3 Trial Design¹

Paradigm Shifting Opportunity to Save the Eye and Protect Vision



Primary Endpoints²

- Cohort 1: Time to Vision Loss
- Cohort 2: Eye Preservation

Secondary Endpoints

 Cohort 1 and 2: No detriment to Event Free Survival (EFS). Initial EFS readout anticipated in ~2-years

FDA discussion ongoing for use of ORR as potential surrogate and composite endpoint for earlier approval scenarios

Three Independent Approaches for Demonstrating Clinical Benefit With Approval Pathway

Enucleation Cohort \rightarrow Save the Eye

Brachytherapy Cohort → Protect Vision

Follow-up → No detriment to EFS



⁽¹⁾ Protocol finalization pending FDA Type B meeting

⁽²⁾ FDA briefing book notes clinical endpoint target to exceed a lower bound of 10% for eye preservation rate with a 95% confidence interval

⁽³⁾ Treatment to maximum benefit: continued observation of ocular tumor shrinkage

⁽⁴⁾ Estimate of initial no detriment EFS readout of UM patients with high risk of metastatic disease

Darovasertib and Uveal Melanoma Patient Journey

High Unmet Need and Multiple First-Line Opportunities in UM and MUM¹

+95% of UM patients harbor GNAQ/GNA11 mutation

	Uveal Melanoma Patient Journey			
	Neoadjuvant UM	Adjuvant UM	MUM	
HLA-A2-Negative (~70% of UM / MUM) ²	Daro Phase 2/3 Enucleation Define Approval Path Path Daro Paro Paro Paro Paro Paro Paro Paro P	Daro Phase 2	Daro + Crizo Registrational Trial Accelerated Approval Full Approval	
HLA-A2-Positive (~30% of UM / MUM) ²	Approval Appro Path Path	No FDA A	Daro + Crizo Target NCCN / Compendia Listing	
Target Treatment Duration	≥6 months	≥6 months	mPFS + ~3 months	
Target Clinical Endpoints	Eye Preservation, Time to Vision Loss No detriment to EFS	Relapse Free Survival	ORR, mPFS, mOS	
Annual Incidence ³	~12K	~12K	~4-5k	

FDA Orphan Drug Designation in Uveal Melanoma⁴; FDA Fast Track Designation in Metastatic Uveal Melanoma Phase 2/3 Registrational Trial Ongoing in HLA-A2 negative 1L MUM for both Accelerated and Full Approval

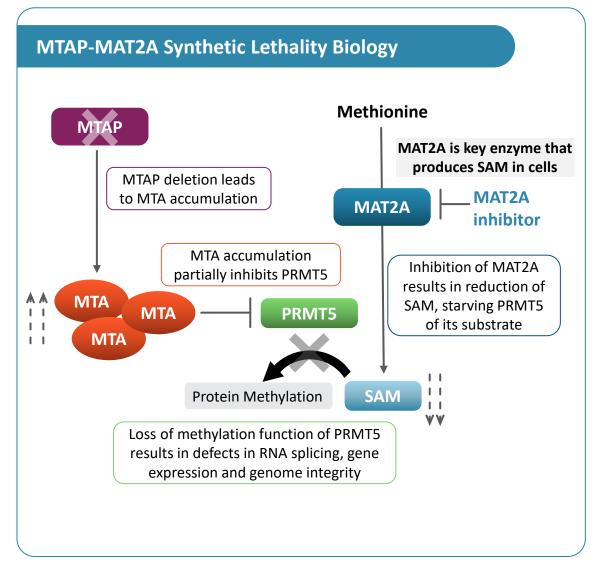
⁽¹⁾ No FDA approved systemic therapies in multiple UM and MUM indications across the patient journey

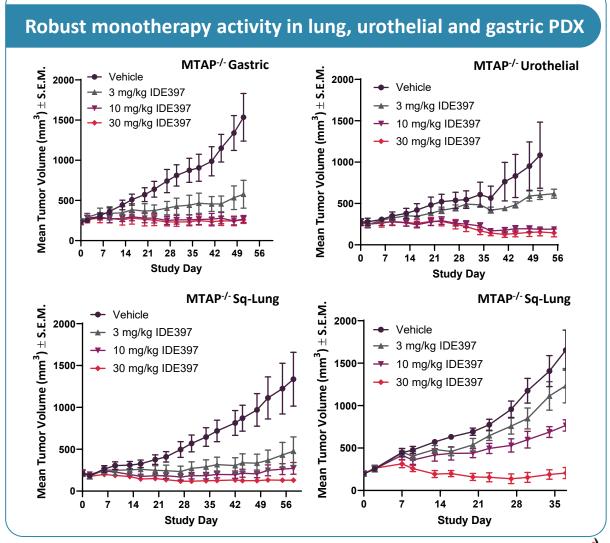
⁽²⁾ IDEAYA data: ~70% of MUM patients were HLA-A2-negative based on 149 patients tested for HLA-A2 status as presented at ESMO 2023

⁽³⁾ Annual incidence for North America, Europe and Australia (as applicable), based on market research analysis

MAT2A Inhibition is Synthetic Lethal with MTAP-Deletion

Strategies to address MTAP-/- Prevalence in ~15% of all Solid Tumors



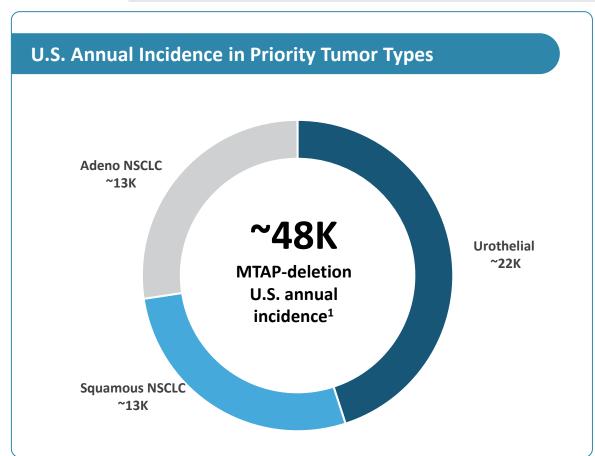


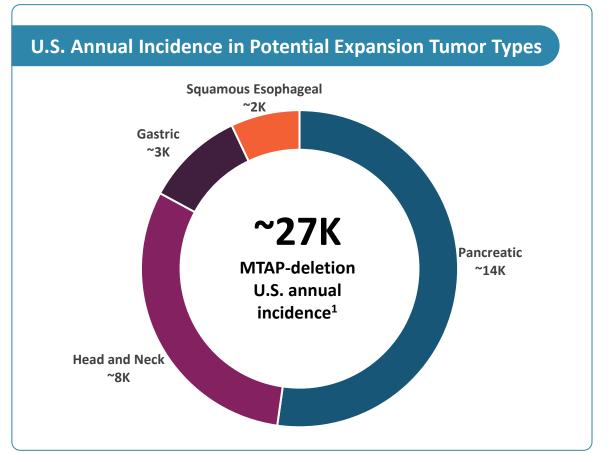


IDE397: Phase 2 Potential First-in-Class MAT2A Inhibitor

~48k U.S. Annual Incidence in MTAP-Deletion NSCLC and Urothelial Cancer

High Unmet Need: No FDA-Approved Therapies for MTAP-Deletion Solid Tumors



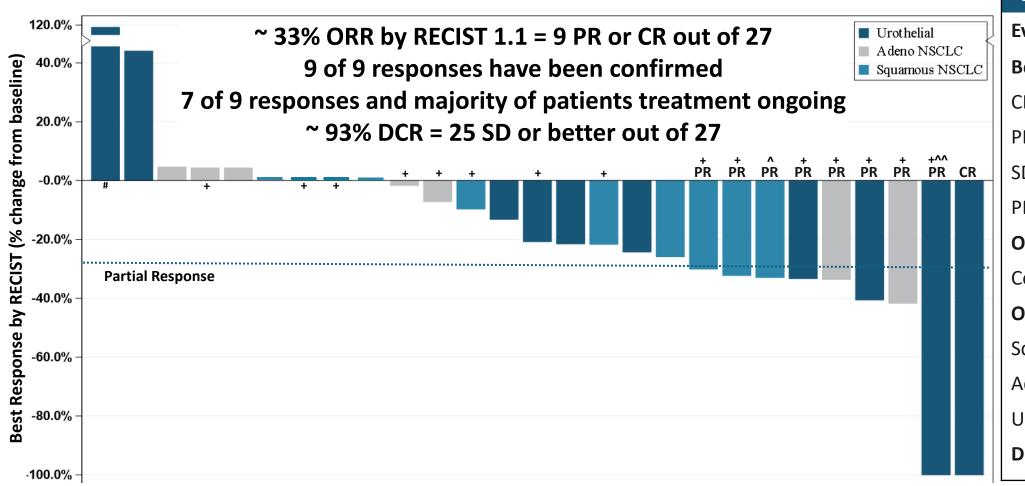






IDE397 Phase 1 Clinical Efficacy in MTAP-Deletion NSCLC & UC

Best Response by RECIST 1.1 at 30mg QD Phase 2 expansion dose¹



Efficacy by RECIST 1.1 ¹	
Evaluable Pts	27
Best Response, n (9	%)
CR	1 (4)
PR	8 (30)
SD	16 (59)
PD	2 (7)
ORR, n (%)	9 (33)
Confirmed, n^^	9
ORR, n (%), by Tumor (n)	
Squam NSCLC (8)	3 (38)
Adeno NSCLC (9)	2 (22)
Urothelial (10)	4 (40)
DCR, n (%)	25 (93)

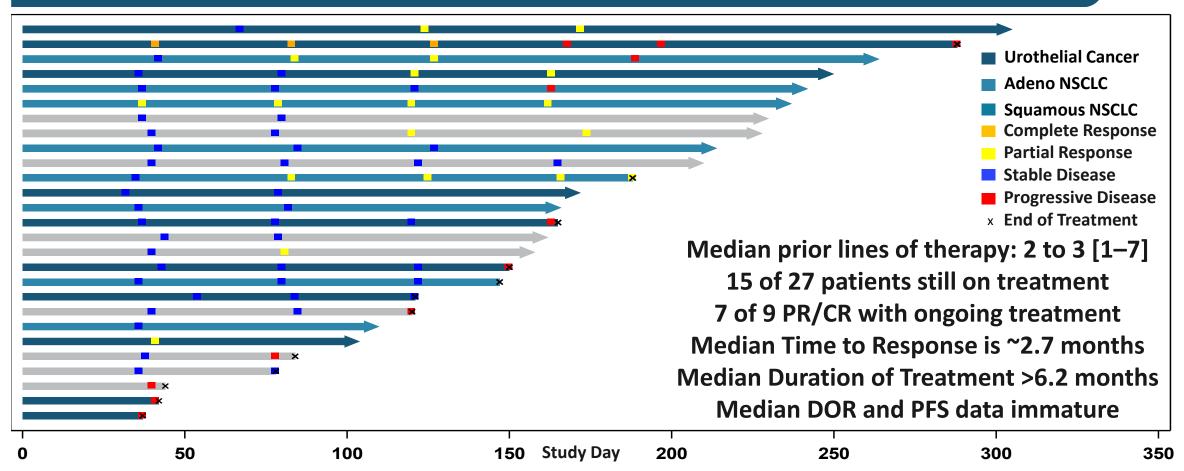




IDE397 Phase 1 Clinical Efficacy in MTAP-Deletion NSCLC & UC

Time on treatment at 30mg QD Ph2 Expansion Dose

NSCLC & Urothelial Cancer Efficacy Evaluable Patients Treated at 30 mg (n=27)¹



⁽¹⁾ Evaluable Patients: Treated with ≥1 cycle (21 days) of IDE397 at 30 mg expansion dose and with ≥1 post-baseline scan(s) Data from an unlocked, unverified database as of 22AUG2024 data cut off. The confirmed complete response urothelial patient progressed after the week 18 scan due to a drug-unrelated AE dose holiday and then restarted treatment. Two patients confirmed response after the data cut.

PFS = Progression Free Survival, DOR = Duration of Response





IDE397 Confirmed CR by RECIST 1.1 in UC Patient With MTAP-Deletion

Case Report and CT-Scan Images

Baseline Characteristics:

60+ years old male urothelial carcinoma

Treatment History:

- Neo-adjuvant cisplatin/gemcitabine
- Left nephro-ureterectomy
- Adjuvant Nivolumab

Recurrent disease while on adjuvant immunotherapy

RECIST 1.1 Evaluation:

CR by RECIST 1.1 at week 6 and confirmed at week 12

Urothelial Carcinoma with MTAP-Deletion: Maintained CR at Week 18 **Baseline** Week 18 **Enlarged Retrocaval Lymph Node, Maintained Complete Response at** 1.5 cm short axis week 18 scan



ENA 2024 EORTC NCI AACR 36th Symposium

IDE397 + Sacituzumab Govitecan Confirmed PR by RECIST 1.1 in Urothelial with MTAP-Deletion and FGFR3-TACC3 Fusion

Case Report and CT-Scan Images

Baseline Characteristics:

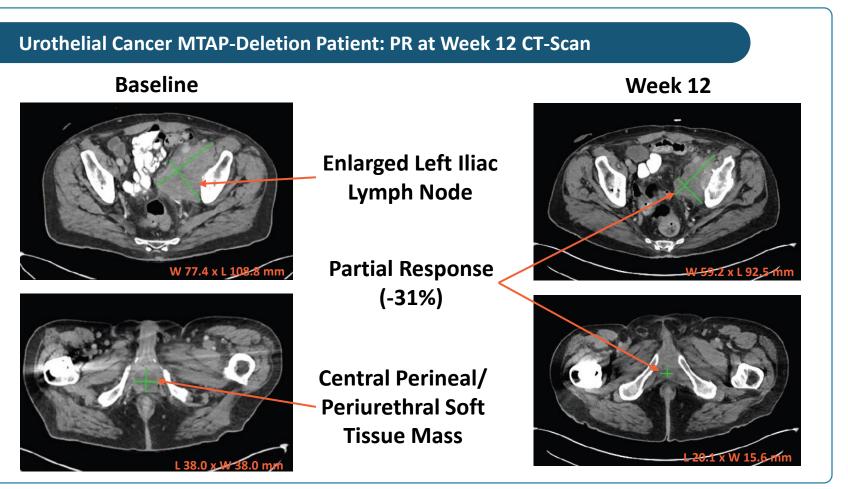
60+ years old male with Urothelial Cancer and MTAPdeletion and FGFR-TACC3 fusion

Treatment History:

- Transurethral resection
- Best response of PD to Enfortumab Vedotin (EV) + Pembrolizumab, and Erdafitinib

Clinical Evaluation:

PR by RECIST 1.1 at week 12, and confirmation at next scan with treatment ongoing

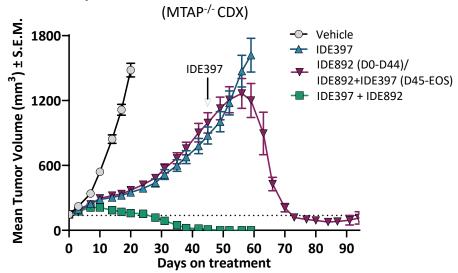




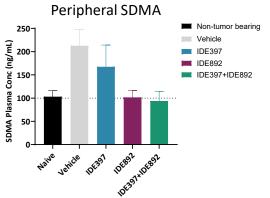
IDE892 DC: Potential Best-in-Class MTA-Cooperative PRMT5 Inhibitor Target to Enable Wholly-Owned Clinical Combo with IDE397/MAT2A in H2 2025¹

MTA-templated target binding Potent biochemical inhibition PRMT5 activity (%) $IC_{50} < 10 \text{ nM}$ -10 IDE892 Log[M] **Persistent MTA-dependent Robust pathway modulation** target occupancy (Surface plasmon resonance) Total SDMA H4R3-SDMA ■ SmB-SDMA $C_{50}s < 10 \text{ nM}$ IDE892 Log[M] Time (h) MTAP-/--specific cell killing Correlation of Cell Features and IDE892 AUC across >800 cancer cell lines Copy number WDR77 -log10(P-value) -log10(P-value) variation PRMT5 CLNS1A DMRTA1 -0.2 0.0 0.0 Spearman Correlation (r) Spearman Correlation (r)

Exceptional IDE397 combination benefit



Pathway sparing in normal tissue

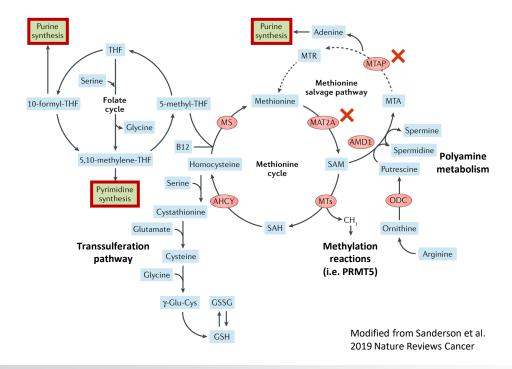




TRODELVY® + IDE397 Combination in MTAP-Deletion Urothelial Cancer

Disordered methionine metabolism underpins IDE397 combination opportunity with TOP1i-based ADC

IDE397 perturbs both nucleotide pools and mRNA splicing in MTAP-/- cells



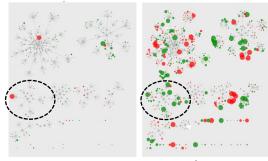
Key clinical correlates underscore combination opportunity

- MTAP-/- UC: shorter survival, lower RR to CPI, shorter PFS and OS with Enfortumab-Vedotin (EV)
- MTAP-/- status is associated with pemetrexed antitumor activity in UC
- A small study evaluating Trodelvy activity in UC post-EV suggested preferential activity in MTAP^{-/-} tumors (RR 50% vs. 19% post EV)
- IDE397 demonstrated monotherapy efficacy in MTAP-/- UC (CR observed in a patient with short DFI post platinum and CPI, other tumor reductions and molecular responses seen)

Metabolic perturbation by IDE397 selectively interacts with MTAP

Metabolite Cytoscape

Global (untargeted) metabolic profiling of MTAP^{wt} vs MTAP^{-/-}+/- IDE397

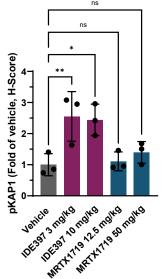


MTAP WT +/- IDE397 MTAP-/- +/- IDE397

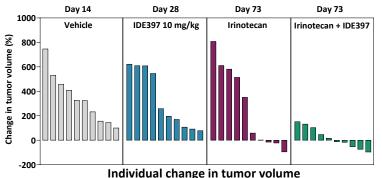
Ovals indicate nucleotide subcluster (purine/pyrimidine); green-decrease, red-increase FDR< 0.05

IDE397 provokes DDR response in vivo

HCT116 MTAP-/- CDX QD 6 days



TOP1i combination delivers durable benefit in challenging RT112/84 UC CDX model



Individual Change in tumor volume
Individual tumors measured on day of study group termination as indicated;
termination timing was based on endpoint criteria for tumor volume



IDE397 Phase 1/2 Clinical Development Plan in MTAP-Deletion Solid Tumors

Clinical Strategic Focus on High Conviction Rational Combinations

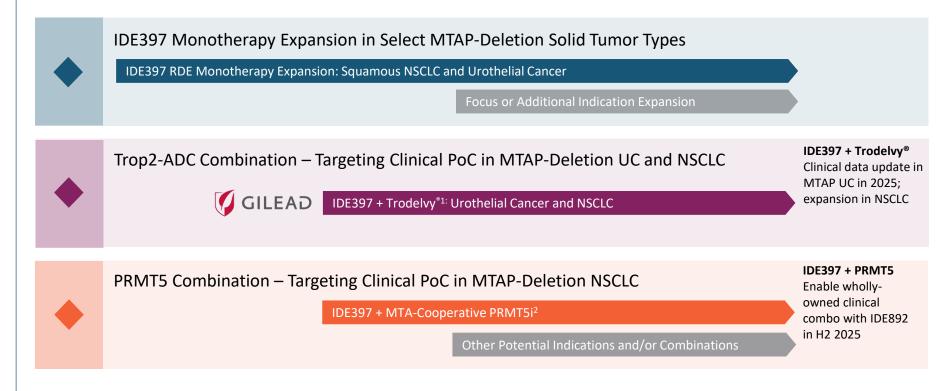
IDF397 - Clinical Profile

Exposure-Dependent Pharmacokinetic (PK) Profile with low C_{max}:C_{min}

Robust Pharmacodynamic (PD) Response observed

Monotherapy Expansion demonstrated clinical efficacy with Responses in Multiple High-**Priority Tumor Types in Dose** Expansion, including a Complete Response

IDE397 is strategically well positioned to evaluate both monotherapy and clinical combinations in MTAP-deletion solid tumors



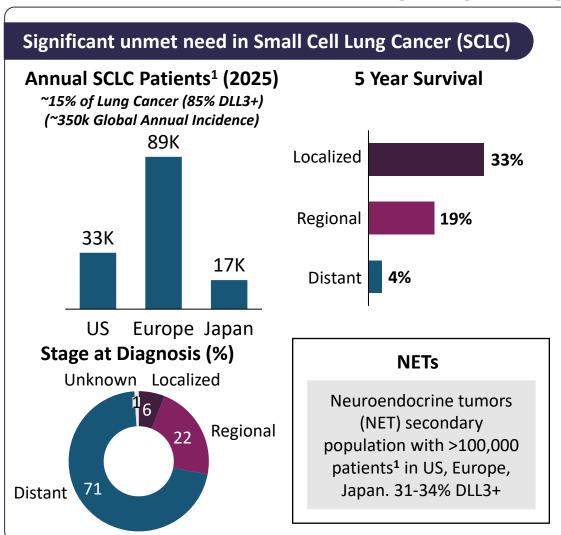


Trodelvy® = Gilead's Trop-2 directed ADC

IDE892, IDEAYA PRMT5 inhibitor in IND-enabling studies

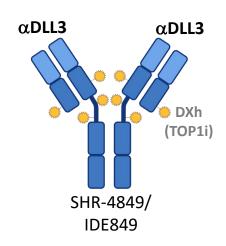
IDE849 (SHR-4849): Phase 1 DLL3 TOP1i ADC

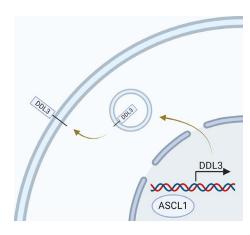
First-in-Class Potential and Targeting Lineage Survival Oncogene Activity



IDE849 (SHR-4849) potential first-in-class/best-in-class

The SCLC lineage survival oncogene, ASCL1, directly promotes DLL3 expression



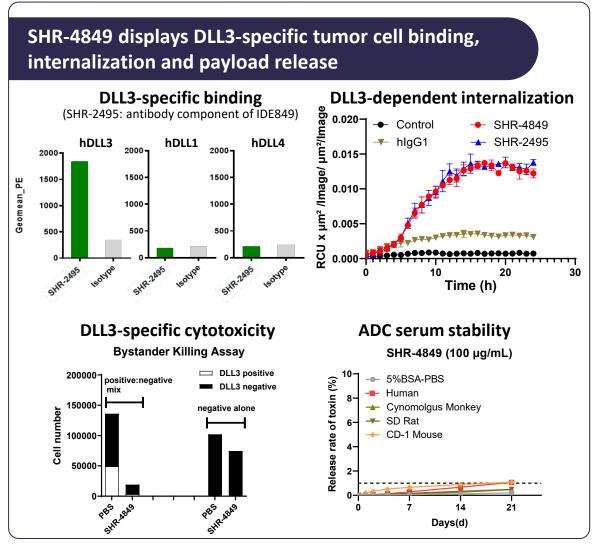


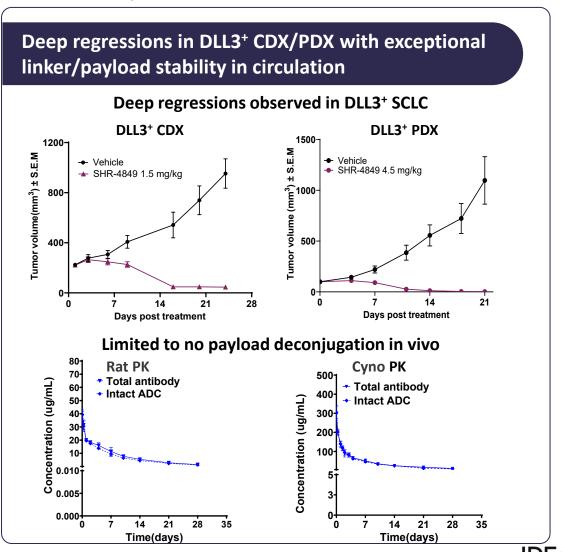
- DLL3 expression driven by the tumor-essential ASCL1 TF
- Humanized antibody with strong affinity and high selectivity
- Proprietary TOP1i payload (~4,000 patients treated)
- Internalization-dependent cleavable linker
- Optimized DAR value of 8
- High plasma stability
- 120X estimated therapeutic index



IDE849 (SHR-4849): Well-tolerated Robust Antitumor Activity in DLL3+ SCLC

DLL3-Specific Tumor Cell binding, Internalization and Payload Release



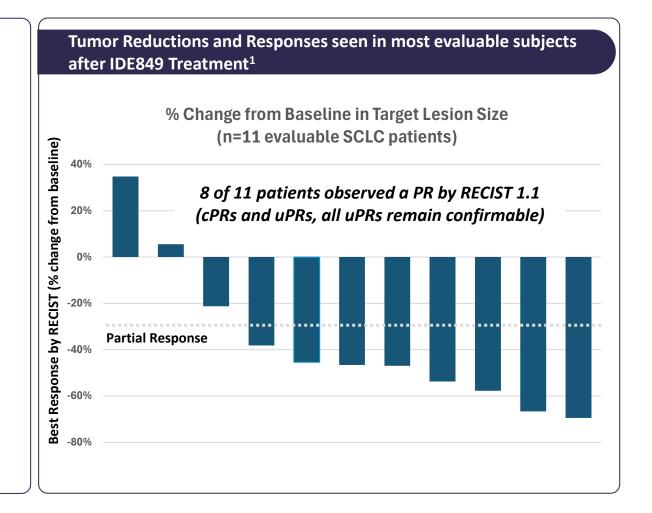


IDE849 (SHR-4849): Potential First-in-Class with Preliminary Ph1 Clinical PoC

Phase 1 FIH Study of DLL3 Topo-1-Payload ADC in Pre-Treated SCLC Patients

Phase 1 Dose Escalation in China in Pre-Treated SCLC Patients¹

- Preliminary Clinical PK Summary
 - Dose dependent increase in exposure
 - Promising T-Ab to ADC ratio
- Preliminary Clinical Efficacy Summary²
 - 8 of 11 evaluable SCLC patients observed a partial response by RECIST 1.1, resulting in a ~73% ORR (confirmed and unconfirmed, all unconfirmed PRs remain confirmable)
- Preliminary Clinical Safety Summary
 - TRAEs were largely Grade 1 or 2
 - No AE leading to discontinuation (related or unrelated)
 - Maximum tolerated dose has not yet been reached
 - Most commonly observed TRAEs: white blood cell count decreased, anemia, neutrophil count decreased, nausea and platelet count decreased



⁽¹⁾ All unconfirmed responses pending further evaluation

⁽²⁾ Clinical efficacy summary at therapeutic dose levels

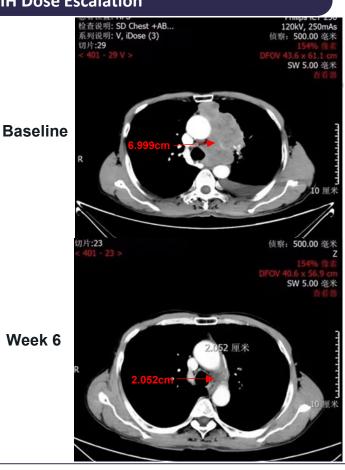
IDE849 (SHR-4849): Potential First-in-Class with Preliminary Ph1 Clinical PoC

Pre-Treated SCLC Patient Case Study and Preliminary IDEAYA Clinical Development Plan

Case Example in Phase 1 FIH Dose Escalation

A 70-year-old male with extensive stage SCLC who had failed prior PD-L1 and platinum doublet treatment

The subject was treated with IDE849 and achieved PR at Week 6 with a 70.6% reduction in the large mediastinal tumor mass



IDE849 Phase 1/2 Clinical Development Plan

IDE849 Monotherapy Dose Escalation and Expansion



IDE849 Combination with IDE161/PARG



Preliminary Clinical Strategy:

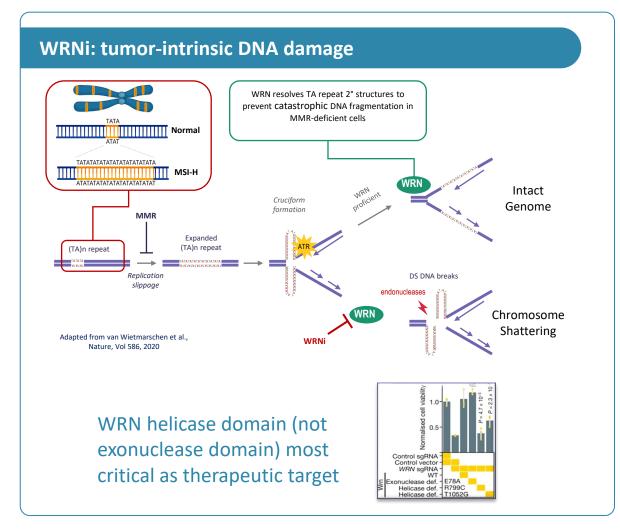
- Potential monotherapy path in 2L plus SCLC
- Evaluate clinical combinations, including with SOC, in 1L SCLC
- Evaluate NETs as monotherapy, including potential basket trial
- Target to enhance durability with IDE849 + IDE161/PARG combo

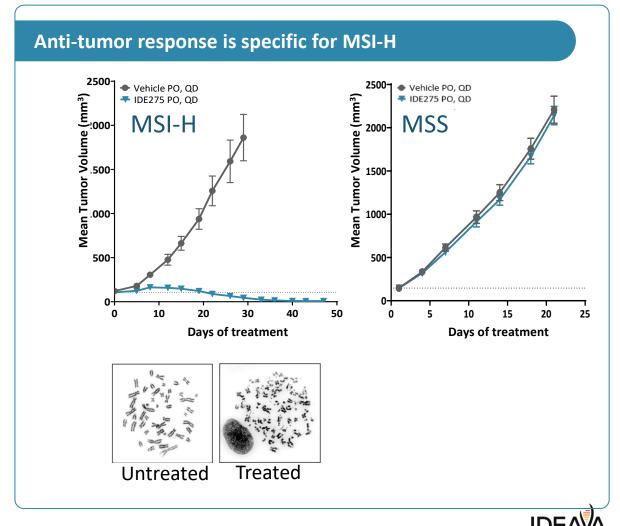




IDE275 (GSK959): Potential First-in-Class Ph1 Werner Helicase Inhibitor 55K

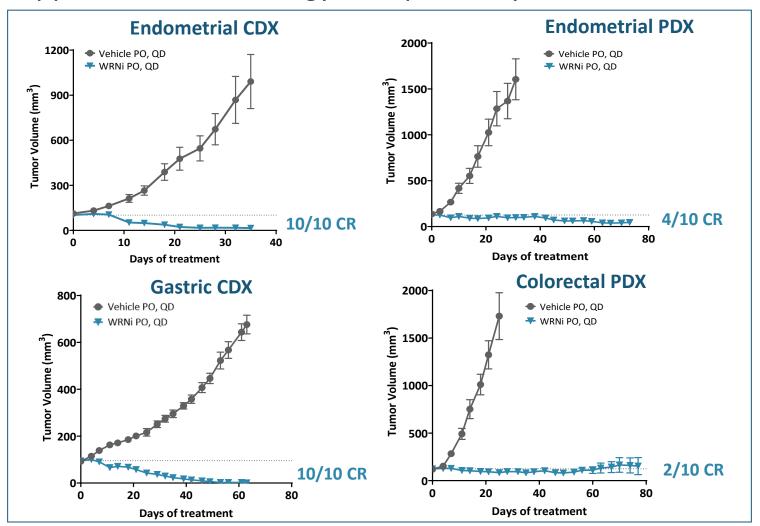
WRN Helicase Activity is Specifically Essential for Survival of MSI-high/dMMR Cancer Cells

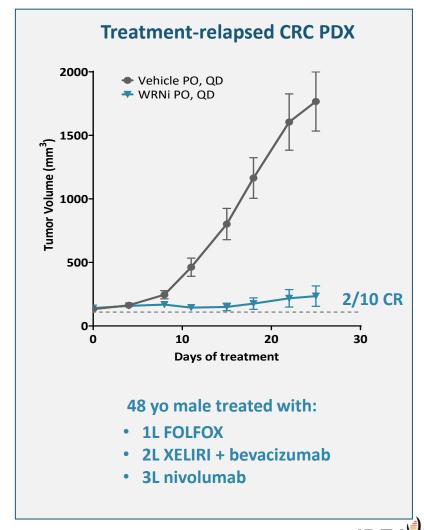




IDE275 (GSK959) Werner Helicase Inhibitor Demonstrates Robust Anti-Tumor Activity in MSI-H and Heavily Pre-Treated Tumors

Supports Clinical Strategy to Expand beyond MSI-H Colorectal Cancers





IDE275 (GSK959): Phase 1 Werner Helicase Inhibitor



Clinical Development Plan

IDE275 (GSK959) Werner Helicase Inhibitor

- IDE275 (GSK959) has demonstrated robust and selective synthetic lethality preclinically in the high microsatellite instability (MSI-High) biomarker setting
- Phase 1 clinical trial enrolling patients having tumors characterized by MSI-High (NCT06710847)

Werner Clinical Development Plan

PART 1: Monotherapy Dose Escalation

Monotherapy IDE275 (GSK959)

- ≥18 years old
- >3 months life expectancy
- dMMR/MSI-H tumor
- Advanced (unresectable/metastatic or recurrent)
- Must have exhausted SOC.

PART 3: Combination Dose Escalation

Combination IDE275 (GSK959) + anti PD-1

- ≥18 years old
- >3 months life expectancy
- dMMR/MSI-H tumor
- Advanced (unresectable/metastatic or recurrent)
- Must have exhausted SOC



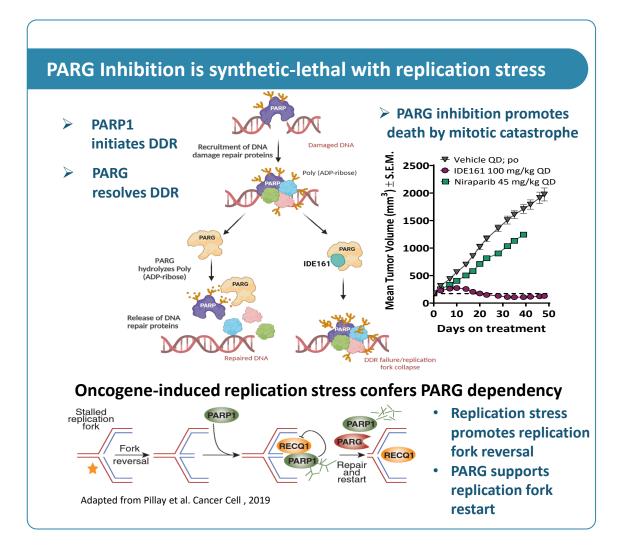
PART 2: Monotherapy Dose Expansion

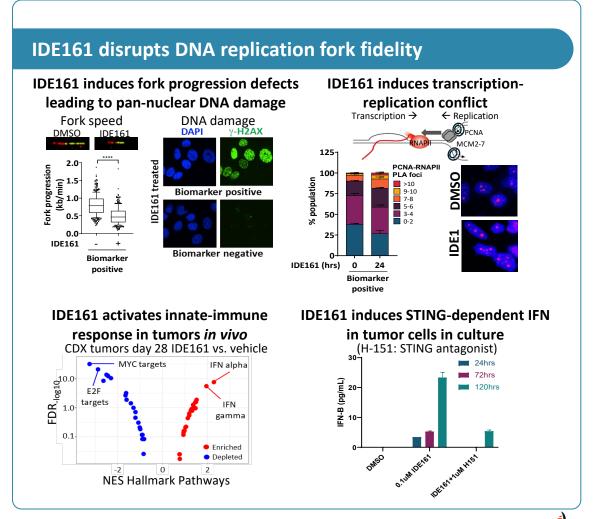
• Histological diagnosis of CRC or EC

GSK Strategic Partnership: 50/50% US Profit Share and ex-US Royalties, ~\$1B Milestones, incl. up to \$20M Preclinical / Ph1 Clinical; Cost Share 80% (GSK) / 20% (IDEAYA); Potential Combination with GSK's Jemperli™, a PD-1 IO Agent



IDE161: Potential First-in-Class Phase 1 PARG Inhibitor

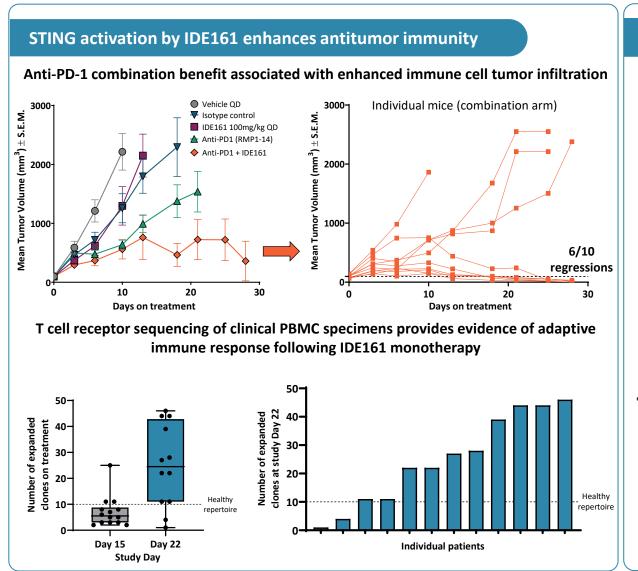






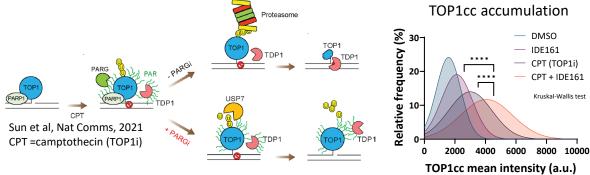
IDE161 Combination Strategies with PD-1 and TOP1i-ADCs

High Conviction Mechanistic Rationale with Potentially Broad Development Opportunity



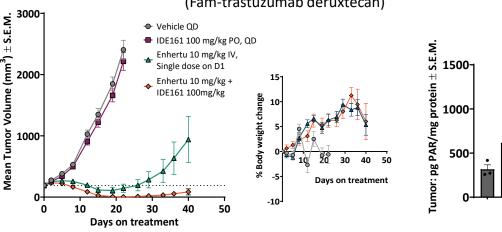
IDE161/TOP1i DDR interaction enhances ADC efficacy

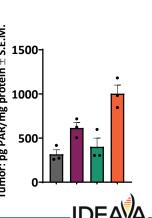
Dual inhibition of TOP1 & PARG induces unresolvable DNA-protein crosslinks



Potential for IDE161 as platform / backbone combo partner for TOP1i-ADCs

HER2⁺ lung; 60% complete responses in combination (Fam-trastuzumab deruxtecan)





IDE161 Phase 1/2 Clinical Development Plan in Solid Tumors

Clinical Strategic Focus on Rational Combinations with TOP1i-ADCs and PD-1

IDE161 Phase 1/2 – Monotherapy and Combination Clinical Development Plan

IDE161 Monotherapy Dose Escalation and Expansion in HRD Solid Tumors¹



Dose Escalation



Expansion Cohort: Priority Tumor Type

IDE161 + KEYTRUDA® (pembrolizumab) in Endometrial Cancer





IDE161+ KEYTRUDA in Endometrial Cancer – Targeting 2025 clinical expansion

IDE161 Topo ADC Combination Opportunities Validated Preclinically





IDE161+ IDE849 (DLL3 ADC)





IDE161+ IDE034 (B7H3/PTK7 Bispecific Ab-TOP1i ADC DC

Activity in PARPi- and Platinum-Resistant Settings **Differentiated Sensitivity** relative to PARPi's

Targeting Improved Safety Profile relative to PARPi's

IDE161 monotherapy expansion initiated in priority tumor type

IDE161 + Keytruda clinical combo FPI achieved

Targeting to enable IDE161 + TOP1i-ADC clinical combinations in 2025

Facile peripheral PD Biomarker for PARGi based on measurement of PAR in blood samples (PBMC's)

Clinicaltrials.gov: NCT05787587

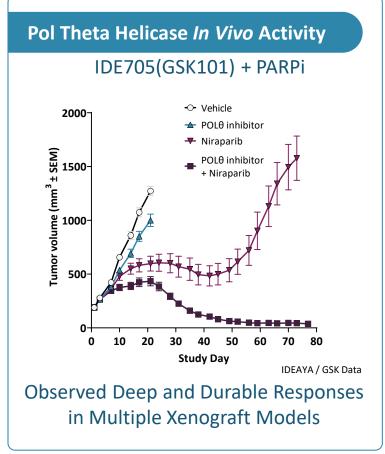
⁽²⁾ Pursuant to Merck Clinical Trial Collaboration and Supply Agreement for IDE161 + Keytruda®, Merck's anti-PD-1 therapy; the Company will sponsor the study and Merck will provide Keytruda at no cost (3) Pursuant to exclusive license agreement with Jiangsu Hengrui Pharmaceuticals Co., Ltd for worldwide rights outside of Greater China

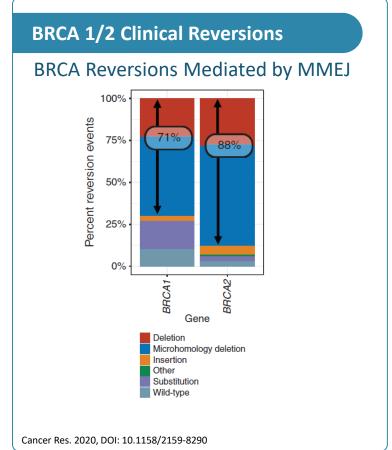
⁽⁴⁾ Pursuant to exclusive worldwide licensing and option agreement with Biocytogen

GSK

IDE705 (GSK101): Potential First-in-Class Ph1 Pol Theta Helicase Inhibitor

Phase 1 in Combination with Niraparib (PARPi)







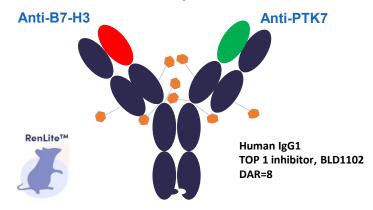
GSK Strategic Partnership: Global Royalties with GSK covering all Costs, ~\$1B Milestones, incl. up to \$20M Preclinical / Ph1 Clinical Potential Combination with GSK's Zejula™, a PARP Inhibitor



IDE034 DC: Potential First-in-Class B7H3/PTK7 TOP1i Bi-Specific ADC¹

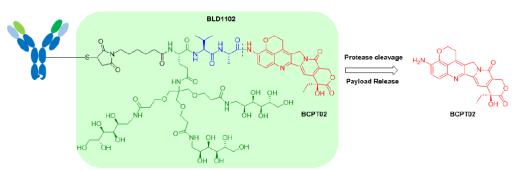
Dual Tumor-Associated Antigen Targeting for Potential Enhanced Therapeutic Window

IDE034: B7H3/PTK7 Bispecific Ab-TOP1i ADC1

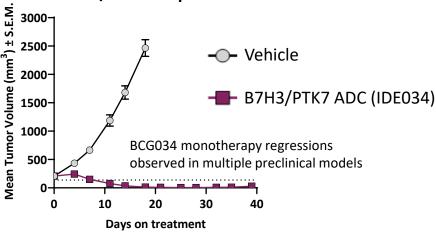


Knobs-into-holes

Proprietary Topoisomerase I Linker-Payload



B7H3+/PTK7+ Expression PDX Model



- Enhanced tumor versus normal cell binding
- Enhanced internalization efficiency
- Meaningful double-positive disease population²

Indication	B7H3/PTK7 Double Positive %
Lung	29.8%
Colorectal ³	45.9%
HNSCC	27.1%
Ovarian	23.1%

Substantial addressable B7H3/PTK7 patient population

⁽¹⁾ IDE034: B7H3/PTK7 Top1i Bispecific ADC development candidate (DC). Exclusive worldwide licensing and option agreement with Biocytogen; IND-enabling studies ongoing with IND-filing targeted in H2 2025

⁽²⁾ IDEAYA analysis of Human Protein Atlas

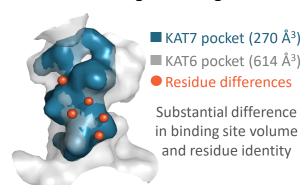
⁽³⁾ Human Protein Atlas annotates colorectal cancer as bowel cancer DAR = Drug Antibody Ratio, IND = Investigational New Drug

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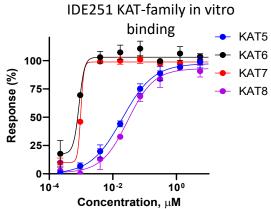
IDE251 DC: Dual KAT6/7 Inhibitor with High Selectivity vs KAT Family¹

Potent Pathway Modulation Delivers Robust Biomarker-Specific Single-Agent Activity

IDE251 solves considerable design challenge

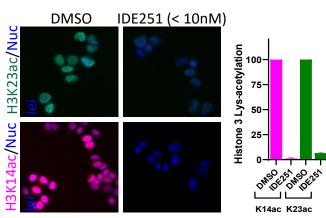


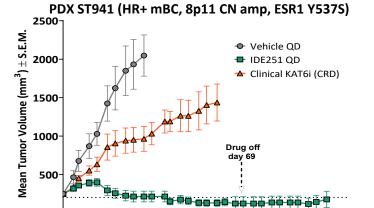
Wide biochemical selectivity window



Concentration, μM

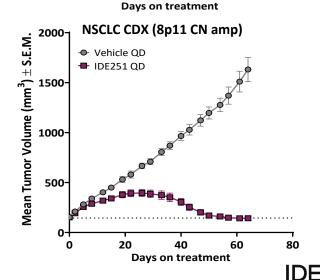
On-target Kac modulation ER+ mBC model



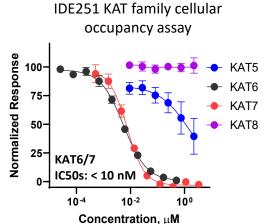


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Durable anti-tumor activity



Strong and selective cellular target binding



Building a Fully-Integrated Biotech in Precision Medicine Oncology

Foundational Potential First-in-Class Clinical Pipeline and Drug Discovery Platform

CLINICAL PROGRAMS

Ph 2/3 – Darovasertib (PKC) ¹
Ph 2 – IDE397 (MAT2A) ¹
Ph 1 – IDE849 (DLL3 ADC) ²
Ph 1 – IDE275 (Werner Helicase) ³
Ph 1 – IDE161 (PARG) ¹
Ph 1 – IDE705 (Pol Theta Helicase) ³

DEVELOPMENT CANDIDATES

IDE892 (PRMT5) – Targeting IND Mid-Year 2025 IDE034 (B7H3/PTK7 Bi-Specific ADC 4) – Targeting IND H2 2025 IDE251 (KAT6/7) – Targeting IND H2 2025

PRECLINICAL PROGRAMS

Multiple Potential First-in-Class

Programs Advancing

6 Clinical Programs

Targeting 3 IND Filings

Darovasertib Registration-Enabling Trial with Potential Accelerated Approval in HLA-A2(-) MUM and Ph3 registrational trial targeted in Neoadjuvant UM is tractable for commercial execution and provides path to potential product revenue to reinvest in broad *first-in-class* pipeline

Potential First-in-Class Precision Medicine Oncology Pipeline, including Darovasertib (Ph2/3), IDE397 (Ph 2), IDE849 (Ph1), IDE275 / GSK959 (Ph 1), IDE161 (Ph 1), IDE705 / GSK101 (Ph 1), IDE892 (IND-enabling), IDE034 (IND-enabling), and IDE251 (IND-enabling)

Strong Balance Sheet with ~\$1.1B⁵ and opportunity for milestone payments with cash runway into at least 2028

Pharma Collaborations including Pfizer, Gilead, Merck, Hengrui, and GSK partnership with ~\$2 billion³ in potential milestones



⁽¹⁾ Clinical Trial Collaboration and Supply Agreements, independently with Pfizer (Darovasertib + Crizotinib), Gilead (IDE397 + Trodelvy®), and Merck (IDE161 + KEYTRUDA®); IDEAYA retains all commercial rights to its products

⁽²⁾ IDE849 (SHR-4849): DLL3 Top1i Antibody Drug Conjugate. Exclusive license agreement with Jiangsu Hengrui Pharmaceuticals Co., Ltd for worldwide rights outside of Greater China

³⁾ IDE705 (GSK101) Pol Theta Program Cost Share = 100% GSK with ~\$1B Milestones and WW Royalties; IDE275 (GSK959) Werner Helicase Program Cost Share = 80% GSK / 20% IDEAYA with ~\$1B Milestones, 50/50 US Profit Share and Ex-US Royalties

⁴⁾ IDE034: B7H3/PTK7 Top1i Bispecific ADC development candidate. Exclusive worldwide licensing and option agreement with Biocytogen

⁵⁾ Includes aggregate of \$1.1 billion of cash, cash equivalents and marketable securities as of December 31, 2024