

# DEVELOPING PRECISION MEDICINES FOR THE TREATMENT OF CANCER

Corporate Presentation – May 2024



# FORWARD-LOOKING STATEMENTS

This presentation contains forward-looking statements. Such statements include, but are not limited to, statements regarding our research, preclinical and clinical development activities, plans and projected timelines for ziftomenib, tipifarnib and KO-2806, plans regarding regulatory filings, our expectations regarding the relative benefits of our product candidates versus competitive therapies, and our expectations regarding the therapeutic and commercial potential of our product candidates. The words “believe,” “may,” “should,” “will,” “estimate,” “promise,” “plan”, “continue,” “anticipate,” “intend,” “expect,” “potential” and similar expressions (including the negative thereof) are intended to identify forward-looking statements. Because such statements are subject to risks and uncertainties, actual results may differ materially from those expressed or implied by such forward-looking statements. Risks that contribute to the uncertain nature of the forward-looking statements include: our preclinical studies and clinical trials may not be successful; the U.S. Food and Drug Administration (FDA) may not agree with our interpretation of the data from clinical trials of our product candidates; we may decide, or the FDA may require us, to conduct additional clinical trials or to modify our ongoing clinical trials; we may experience delays in the commencement, enrollment, completion or analysis of clinical testing for our product candidates, or in the reporting of data from such clinical testing, or significant issues regarding the adequacy of our clinical trial designs or the execution of our clinical trials may arise, which could result in increased costs and delays, or limit our ability to obtain regulatory approval; our product candidates may not receive regulatory approval or be successfully commercialized; unexpected adverse side effects or inadequate therapeutic efficacy of our product candidates could delay or prevent regulatory approval or commercialization; and we may not be able to obtain additional financing. Additional risks and uncertainties may emerge from time to time, and it is not possible for Kura’s management to predict all risk factors and uncertainties.

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# Investment Highlights

<b>Targeted Oncology</b>	Advancing a pipeline of novel investigational therapies, forging new scientific and clinical paths to give patients a better chance for long-term, durable remissions
<b>Proprietary Pipeline</b>	<b>Menin Inhibitor Program (Ziftomenib)</b> <ul style="list-style-type: none"><li>• Potential to address up to 50% of acute leukemias through monotherapy and combinations</li><li>• 35% CR rate among 20 patients with NPM1-mutant AML treated at recommended Phase 2 dose</li><li>• Positive preliminary combination data in NPM1-m and KMT2A-r AML, including 100% CR rate with 7+3 in 1L, 56% CR/CRh rate with ven/aza in R/R menin inhibitor naïve patients and mitigation of differentiation syndrome</li><li>• Breakthrough Therapy Designation granted by FDA for the treatment of patients with R/R NPM1-mutant AML</li><li>• Completion of enrollment in Phase 2 registration-directed trial in NPM1-mutant AML expected by mid-2024</li></ul>
	<b>Farnesyl Transferase Inhibitor Programs (Tipifarnib &amp; KO-2806)</b> <ul style="list-style-type: none"><li>• Durable responses observed with tipifarnib as a monotherapy in R/M HRAS-mutant HNSCC patients</li><li>• Compelling safety profile and activity observed with tipifarnib plus alpelisib in PIK3CA-dependent HNSCC</li><li>• Preclinical data support clinical combinations of next-gen FTI KO-2806 with adagrasib and cabozantinib</li><li>• Clinical collaboration with BMS to evaluate KO-2806 and adagrasib in KRAS<sup>G12C</sup>-mutated NSCLC</li><li>• Now dosing patients in dose-escalation trial of KO-2806 as monotherapy and in combination with cabozantinib in ccRCC; combo with adagrasib expected to start by mid-2024</li></ul>
<b>Strong Financials</b>	<ul style="list-style-type: none"><li>• \$25 million strategic equity investment from Bristol Myers Squibb</li><li>• \$527 million in cash as of March 31, 2024* provides runway into 2027</li></ul>

\* Cash, cash equivalents and short-term investments



# Experienced Leadership Team and Board of Directors

## Leadership Team



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Chief Executive Officer



**Teresa Bair, J.D.**  
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**Stephen Dale, M.D.**  
Chief Medical Officer



**Kathy Ford**  
Chief Operating Officer



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**Pete De Spain**  
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**Tom Doyle**  
Senior Vice President,  
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**Roger Bakale, Ph.D.**  
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**Carol Schafer**

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**Faheem Hasnain**  
Lead Independent Director

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**Diane Parks**

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**Helen Collins, M.D.**

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**Mary Szela**

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**Thomas Malley**

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# Drug Candidate Pipeline

PROGRAM	CLINICAL TRIAL	STUDY STARTUP	DOSE-ESCALATION	DOSE-VALIDATION	REGISTRATION DIRECTED	ANTICIPATED MILESTONE
ZIFTOMENIB Menin Inhibitor	KOMET-001 Monotherapy (Relapsed/refractory)	NPM1-mutant acute myeloid leukemia (AML)				Complete enrollment of 85 patients by mid-2024
		KMT2A-rearranged acute lymphoblastic leukemia				Now dosing patients
		Non-NPM1-mutant / Non-KMT2A-rearranged AML				Now dosing patients
	KOMET-007 Combination with venetoclax + azacitidine (Relapsed/refractory)	NPM1-mutant AML				Identify RP2D by mid-2024 / Advance to frontline AML in 2H2024
		KMT2A-rearranged AML				
	KOMET-007 Combination with cytarabine + daunorubicin (Frontline)	NPM1-mutant AML				Identify RP2D by mid-2024
		KMT2A-rearranged AML				
	KOMET-008 Combinations with gilteritinib, FLAG-IDA, LDAC (Relapsed/refractory)	NPM1-mutant AML				Now dosing patients
KMT2A-rearranged AML						
TIPIFARNIB Farnesyl Transferase Inhibitor (FTI)	KURRENT-HN Combination with alpelisib	PIK3CA-dependent head and neck squamous cell carcinoma (HNSCC)				Identify optimal biologically active dose by end of 2024 / Present preliminary data in 1H2025
KO-2806 Next-Generation FTI	FIT-001 Monotherapy, combinations with cabozantinib and adagrasib	Solid tumors				Now in dose escalation as monotherapy
		Clear cell renal cell carcinoma (ccRCC)				Now dosing patients in combo with cabozantinib
		KRAS <sup>G12C</sup> -mutant non-small cell lung cancer (NSCLC)				Dose first patients in combo with adagrasib by mid-2024

# ZIFTOMENIB: MENIN-KMT2A/MLL INHIBITOR IN ACUTE LEUKEMIAS



# Ziftomenib Demonstrates Potential to Become a Cornerstone of AML Therapy

**Targets foundational mutations at the core of up to 50% of AML cases**

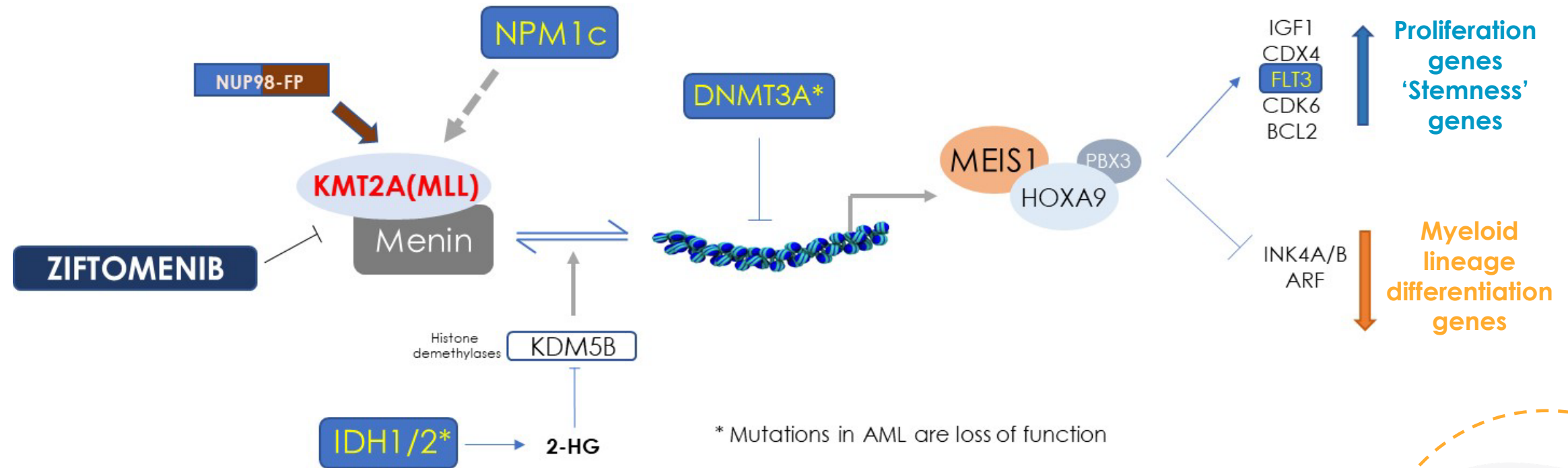
- **Compelling clinical data support frontline opportunity**
  - Good tolerability profile, enabling continuous administration in combination with SOC
  - Combinations appear to mitigate the risk of differentiation syndrome
  - No observed or predicted drug-drug interactions
  - Encouraging preliminary evidence of clinical activity
- **Strong investigator enthusiasm as evidenced by rapid enrollment across studies**
  - First 20 patients enrolled in KOMET-007 combination trial in less than four months
  - Now dosing patients in KOMET-008 combination trial with SOC, including FLT3 inhibitor
  - KOMET-001 monotherapy registrational trial expected to complete enrollment by mid-2024



# Ziftomenib Targets the Menin-KMT2A Pathway, a Foundational Target in AML

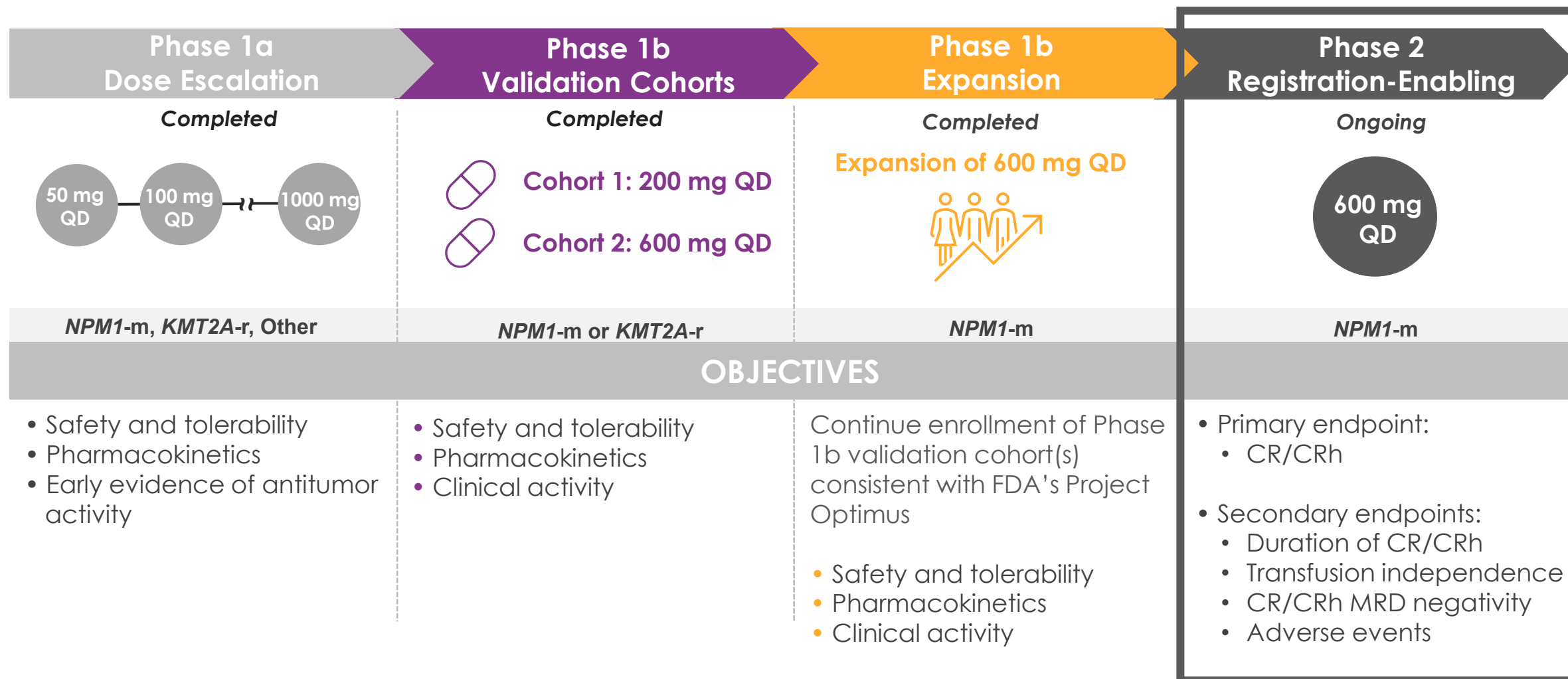


- *NPM1*-m and *KMT2A*-r drive overexpression of *HOXA9/MEIS1* genes, critical for transformation to AML
- *KMT2A*(MLL) sits upstream from major AML targets (*i.e.*, *FLT3*, *IDH1/2*, *DNMT3A*)
- *KMT2A*(MLL)-dependent genes contribute to therapeutic resistance and relapse to current therapies
- Menin inhibition downregulates *HOXA9/MEIS1*, leading to differentiation of leukemic blasts





# KOMET-001 Phase 1/2 Study of Ziftomenib in Relapsed / Refractory AML

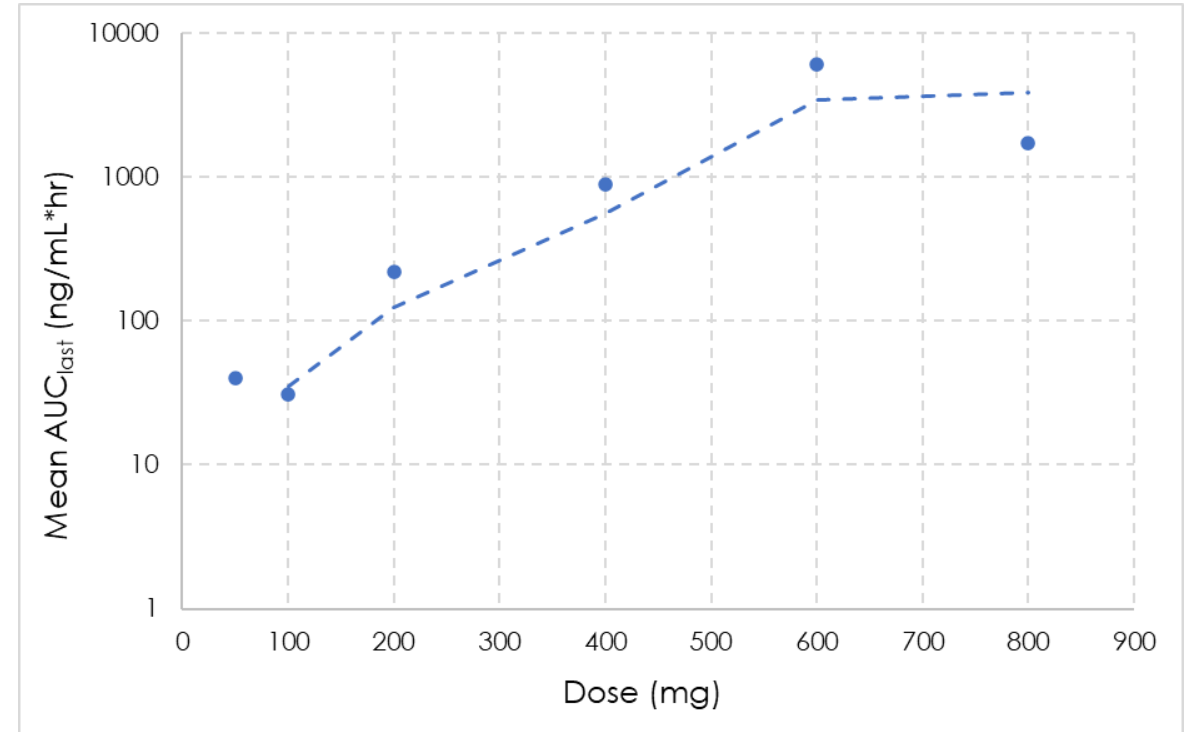




# Ziftomenib Demonstrates Optimal Pharmaceutical Properties

Clinical data from KOMET-001 demonstrate:

- Ziftomenib demonstrates a dose-dependent increase in exposure up to RP2D at 600 mg
- Ziftomenib is not a clinically meaningful CYP3A4 substrate
  - No dose adjustment of ziftomenib needed when administered with a CYP3A4 inhibitor (e.g., azoles)
- Ziftomenib is not a clinically meaningful CYP3A4 inhibitor
  - No dose adjustment needed for CYP3A4 substrates (e.g., venetoclax)
- No drug-induced QTc prolongation observed at any dose





## Ziftomenib Demonstrates Encouraging Safety Profile in Phase 1b

- Differentiation syndrome (DS) appears manageable in NPM1-m monotherapy patients with mitigation strategy
  - 20% rate of mild to moderate DS
- Rates of DS in KMT2A-r monotherapy patients were 38.5% at 200 mg and 37.5% at 600 mg; potential to mitigate in combination
- DS is an on-target adverse event and represents evidence of clinical activity
- No reports of drug-induced QTc prolongation
- Maintained count recovery suggests no drug-induced myelosuppression



# Ziftomenib has Highly Differentiated Monotherapy Activity

40% of NPM1 patients achieved a CR during course of study

Best Overall Response	600 mg
<b>NPM1-m Phase 1a + 1b</b>	<b>(n=20)</b>
<b>CR</b>	<b>7 (35.0)</b>
CR/CRh	7 (35.0)
CRC	8 (40.0)
MRD negativity	4 (50.0) <sup>1</sup>
ORR	9 (45.0)
<b>KMT2A-r Phase 1a + 1b</b>	<b>(n=18)</b>
CR/CRh	2 (11.1)
CRC	3 (16.7)
MRD negativity	3 (100.0)
ORR	3 (16.7)

(preliminary data as of April 12, 2023)

## Differentiated CR Rates vs. SOC in Heavily Pretreated Patients

	MUTATION	CR %	mDOR	MEDIAN PRIORS
<b>Ziftomenib</b> 600mg QD	<b>NPM1m</b>	<b>35%</b>	<b>8.2 mo*</b>	<b>3</b>
	<b>FLT3m</b>	<b>33%</b>	-	
	<b>IDH 1/2</b>	<b>50%</b>	-	
<b>Gilteritinib</b>	<b>FLT3m</b>	14.2%	14.8 mo	1
<b>Enasidenib</b>	<b>IDH2</b>	19%	8.2 mo	2
<b>Ivosidenib</b>	<b>IDH1</b>	25%	10.1 mo	2

\*Median DoR for CRc without censoring at HSCT  
Source: USPI's

- **High activity, durable responses and favorable profile suggest potential for ziftomenib to become a backbone therapy across the continuum of AML care**

<sup>1</sup> MRD was assessed for 6/8 CRC patients; 4 of those 6 patients (67%) tested were MRD negative  
CRC includes CR, CRh, CRi, CRp; ORR includes CR, CRh, CRi, CRp, MLFS

# Case Studies Highlight Meaningful Durability and Favorable Tolerability



Durable CR for 36 cycles on ziftomenib  
in 8<sup>th</sup> line including 2 HSCTs

## 44 yo female with *NPM1*-m, *DNMT3A* and *IKZF1* AML 7 Prior Tx

Baseline bone marrow blasts: 14%

ziftomenib at 200 mg

Response

- CRmrd- after Cycle 1
- CRmrd- through Cycle 36

Enthusiasm among investigators and  
patients to utilize ziftomenib earlier  
and initiate maintenance

## 22 yo male with *NPM1*-m AML 1 Prior Tx (refractory to 7+3)

Baseline bone marrow blasts: 90%

ziftomenib at 600 mg

Response

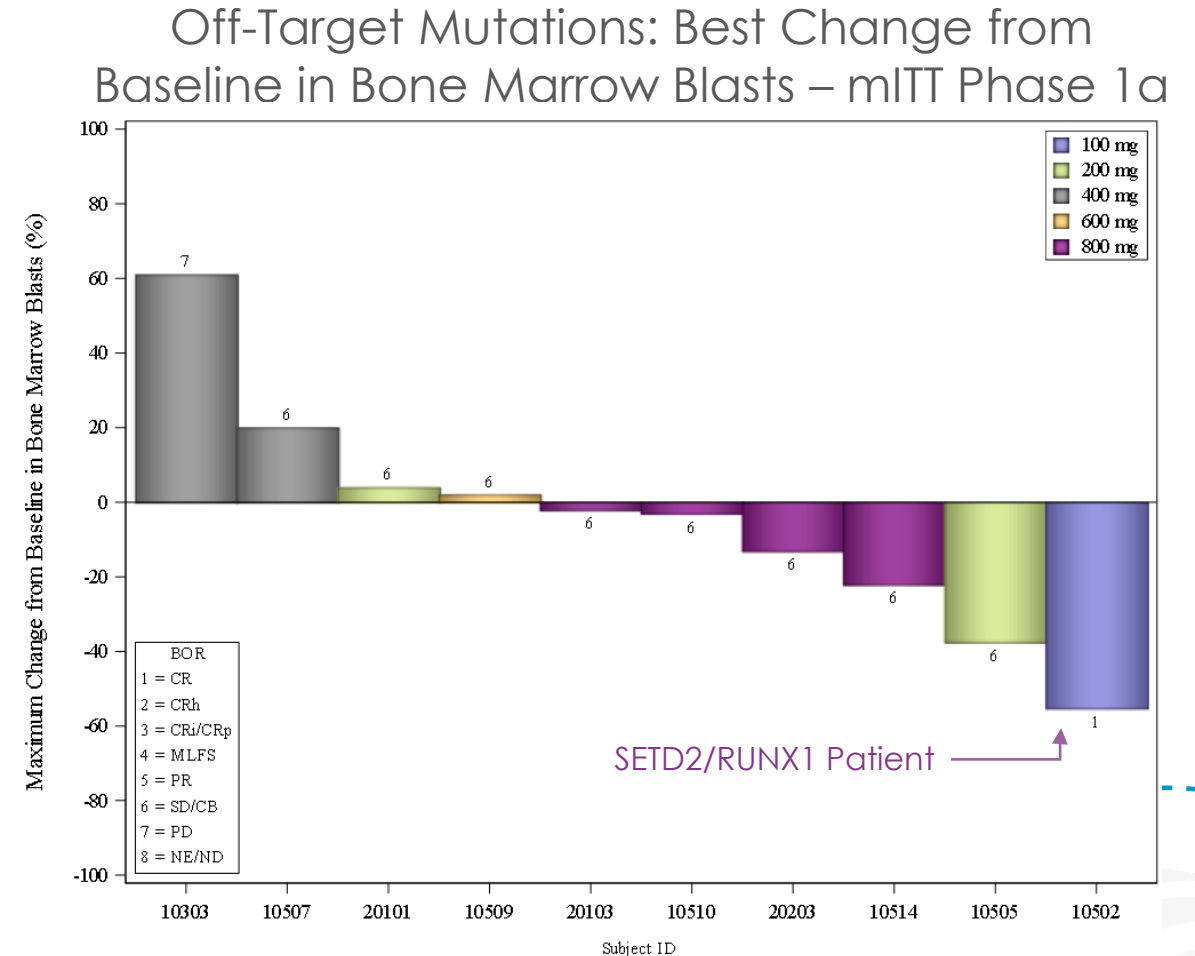
- CRmrd- after Cycle 1
- HSCT
- CRmrd- maintained on Cycle 2 post-HSCT

# Targeting the Menin-KMT2A Pathway has Potential to Benefit a Broader Subset of AML Patients



## Ziftomenib has Demonstrated Evidence of Activity in Non-*NPM1*-m/*KMT2A*-r Patients

- *SETD2*/*RUNX1* patient achieved a CR at 100mg dose in Phase 1a
- Notable evidence of blast reduction in range of off-target patients
- KOMET-001 study will continue to evaluate additional AML populations
- Potential to be incorporated into KOMET-007/008 combination studies





# Summary: KOMET-001 Phase 1 Clinical Trial of Ziftomenib

## **Ziftomenib demonstrates an encouraging safety profile and tolerability**

- Reported events most often consistent with features and manifestations of underlying disease
  - No evidence of drug-induced QTc prolongation
  - Differentiation syndrome, an on-target effect, manageable with mitigation strategy

## **Clinical activity of ziftomenib monotherapy is optimal at the 600 mg daily dose**

- Positive *NPM1*-m benefit/risk balance with pronounced activity and 35% CR rate (n=20)
- High levels of ziftomenib tissue penetration likely drive clearance of extramedullary disease
- Emergence of resistance mutations has been observed at a much lower rate relative to certain competition

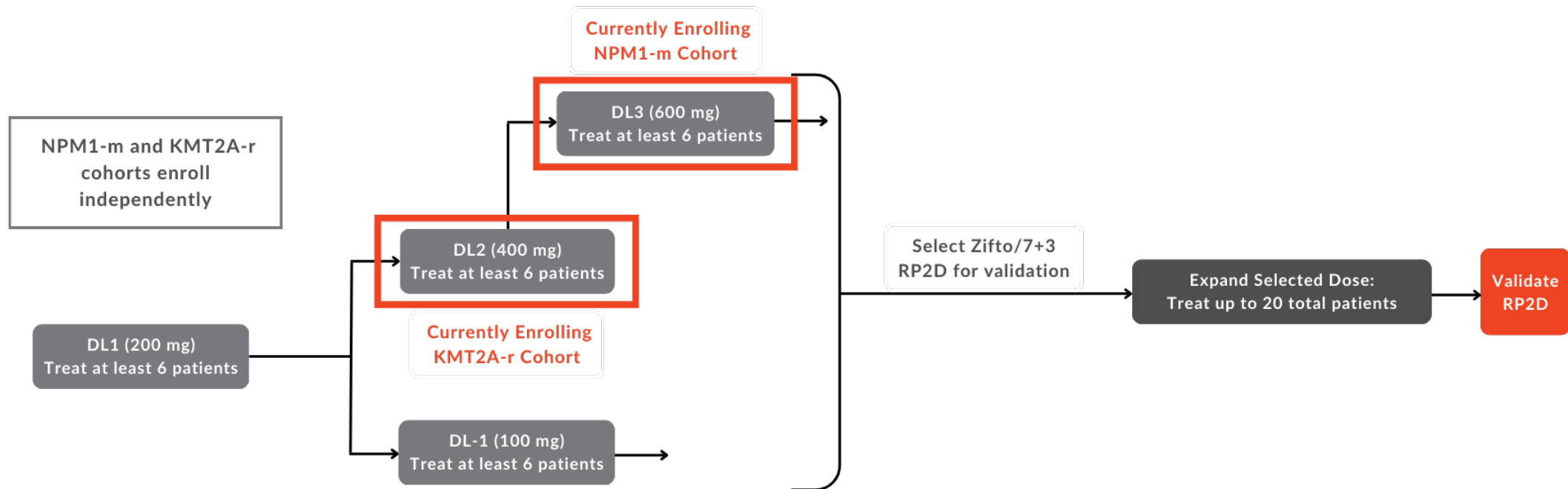
## **Monotherapy data supportive of combination strategies**

- No predicted adverse drug-drug interactions
- Optimization of *KMT2A*-r benefit/risk planned via combination strategies to maximize time on treatment
- Oral, QD dosing allows for convenient administration and combination with standards of care



# KOMET-007: Phase 1 Combination Trial of Ziftomenib in Patients with Newly Diagnosed or R/R AML

## Ziftomenib/cytarabine/daunorubicin (7+3) combination



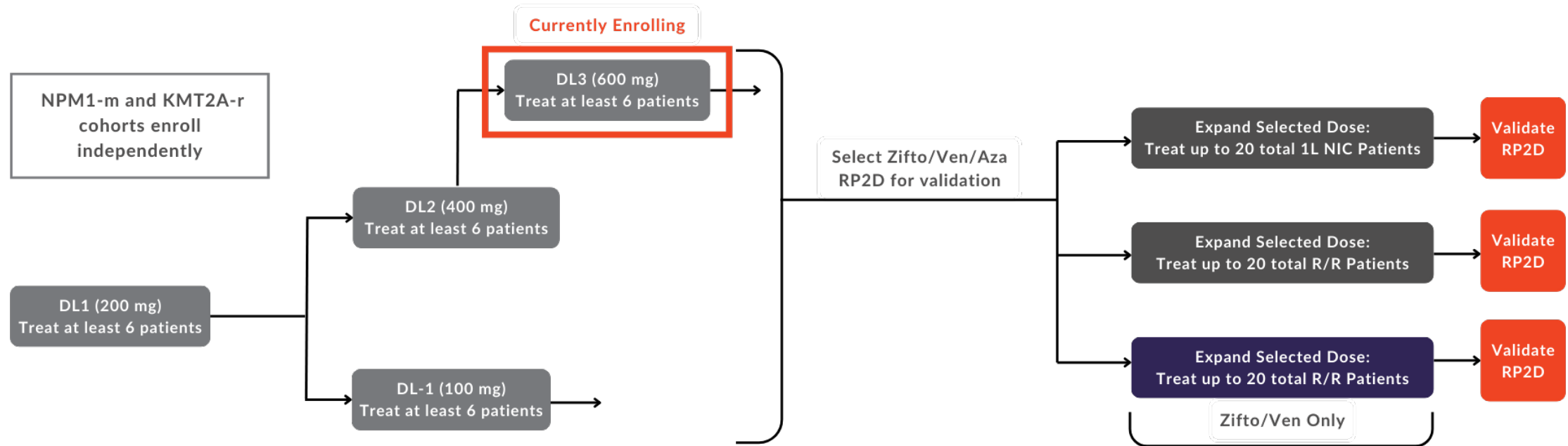
- Ziftomenib dosing begins on Cycle 1 Day 8 and be administered continuously thereafter
- Cytarabine administered on Cycle 1 Day 1-7; administration of an additional cycle based on C1 bone marrow biopsy results
- Daunorubicin administered on Cycle 1 Day 1-3; administration of an additional cycle based on C1 bone marrow biopsy results
- Dose escalation conducted in patients with adverse risk\*

\*Age ≥ 60 years and/or treatment-related AML and/or adverse risk cytogenetics per ELN

DL = ziftomenib dose level; zifto = ziftomenib; 7+3 = cytarabine/daunorubicin; RP2D = recommended Phase 2 dose; 1L = first-line; IC = intensive chemotherapy

# KOMET-007: Phase 1 Combination Trial of Ziftomenib in Patients with Newly Diagnosed or R/R AML

## Ziftomenib/venetoclax/azacitidine combination



- Ziftomenib dosing begins on Cycle 1 Day 8 and be administered continuously thereafter
- Venetoclax administered per label in 28-day cycles with adjustments to cycle length based on Cycle 1 bone marrow biopsy results
- Azacitidine administered per label on Cycle 1 Day 1-7 of each cycle with additional cycles based on bone marrow biopsy results



# KOMET-007: Promising Safety and Tolerability Profile in Combination

## Combinations mitigate risk of differentiation syndrome (DS)

Grade $\geq$ 3 TEAEs ( $\geq$ 10%)	n (%)
<b>Patients with Grade <math>\geq</math> 3 TEAEs</b>	<b>18 (90)</b>
Platelet count decreased	6 (30)
Febrile neutropenia	5 (25)
White blood cell count decreased	4 (20)
Pneumonia	3 (15)
Hypoxia	2 (10)
Neutrophil count decreased	2 (10)
Sepsis	2 (10)
Thrombocytopenia	2 (10)

Grade $\geq$ 3 Ziftomenib-Related AEs (All)	n (%)
<b>Patients with Grade <math>\geq</math> 3 Ziftomenib-Related AEs</b>	<b>6 (30)</b>
Platelet count decreased	3 (15)
Anemia	1 (5)
Febrile neutropenia	1 (5)
Leukopenia	1 (5)
Neutrophil count	1 (5)
Thrombocytopenia	1 (5)

- No DS events reported
- No dose-limiting toxicities (DLTs) observed to date, including delayed hematologic count recovery
- No QTc prolongation observed
- TEAEs consistent with underlying disease and backbone therapies



# 100% CR rate with Ziftomenib and 7+3 in 1L Patients with Adverse-Risk AML\*

- Anticipated CR/CRi rate with 7+3 in all-comer 1L adverse risk patients: 32-33%<sup>1,2</sup>

1L Adverse-Risk Group n=5	CR Rate (n)
Overall (NPM1-m + KMT2A-r)	100% (5)
NPM1-m only (n=4)	100% (4)
KMT2A-r only (n=1)	100% (1)

- All patients treated in initial dose cohort (200 mg) in combination with 7+3

Preliminary data as of January 11, 2024

<sup>1</sup> Lancet et al. *Blood*. 2014 May 22;123(21):3239-46.

<sup>2</sup> Lin et al. *Blood Adv*. 2021 Mar 23;5(6):1719-1728.

\*Age ≥ 60 years and/or treatment-related AML and/or adverse risk cytogenetics per ELN



## Ziftomenib + Ven/Aza with Pronounced Activity in Menin Inhibitor Naïve Patients

- ~35-45% CR/CRi rate is expected in ven-naïve relapsed/refractory patients<sup>1</sup>
- Anticipated CR/CRi rate in KMT2A-r AML following two prior therapies <10%<sup>2</sup>
- 53% ORR in mITT population (n=15, including six menin experienced patients)
- 40% (6/15) of patients treated with ven/aza received prior treatment with a menin inhibitor

Menin Inhibitor Naïve Group n=9	ORR (n)	CR/CRi Rate (n)	CR/CRh Rate (n)
Overall (NPM1-m + KMT2A-r)	78% (7)	67% (6)	56% (5)
NPM1-m (n=5)	100% (5)	80% (4)	60% (3)
KMT2A-r (n=4)	50% (2)	50% (2)	50% (2)

- All patients treated in initial dose cohort (200 mg) in combination with Ven/Aza
- Enrollment in 600 mg dose cohort ongoing

Preliminary data as of January 11, 2024

<sup>1</sup> Stahl, M. et al., *Blood Advances* 5(5), 1552-1564 (2021)

<sup>2</sup> Issa, G. et al. *Blood Cancer J.* 11, 162 (2021)

ORR includes CR, CRh, CRi, MLFS



# Ziftomenib + Ven/Aza Able to Drive Responses in Venetoclax Failures

- Expected response rates following ven/aza ~ 0-20%<sup>1-4</sup>
- Anticipated CR/CRi rate in KMT2A-r AML following two prior therapies < 10%<sup>5</sup>

Venetoclax Experienced Group n=10	ORR (n)	CR/CRi Rate (n)	CR/CRh Rate (n)
Overall (NPM1-m + KMT2A-r)	40% (4)	30% (3)	30% (3)
NPM1-m (n=5)	60% (3)	40% (2)	40% (2)
KMT2A-r (n=5)	20% (1)	20% (1)	20% (1)

- All patients treated in initial dose cohort (200 mg) in combination with Ven/Aza
- Enrollment in 600 mg dose cohort ongoing

Preliminary data as of January 11, 2024

<sup>1</sup> Zainaldin, C. et al., Lymphoma 63(13):3245-3248 (2022);

<sup>2</sup> Chan, O. and Walker, A., Hematology 702-708 (2023);

<sup>3</sup> Maiti A, et al., Haematologica. 2021; 106(3):894-898;

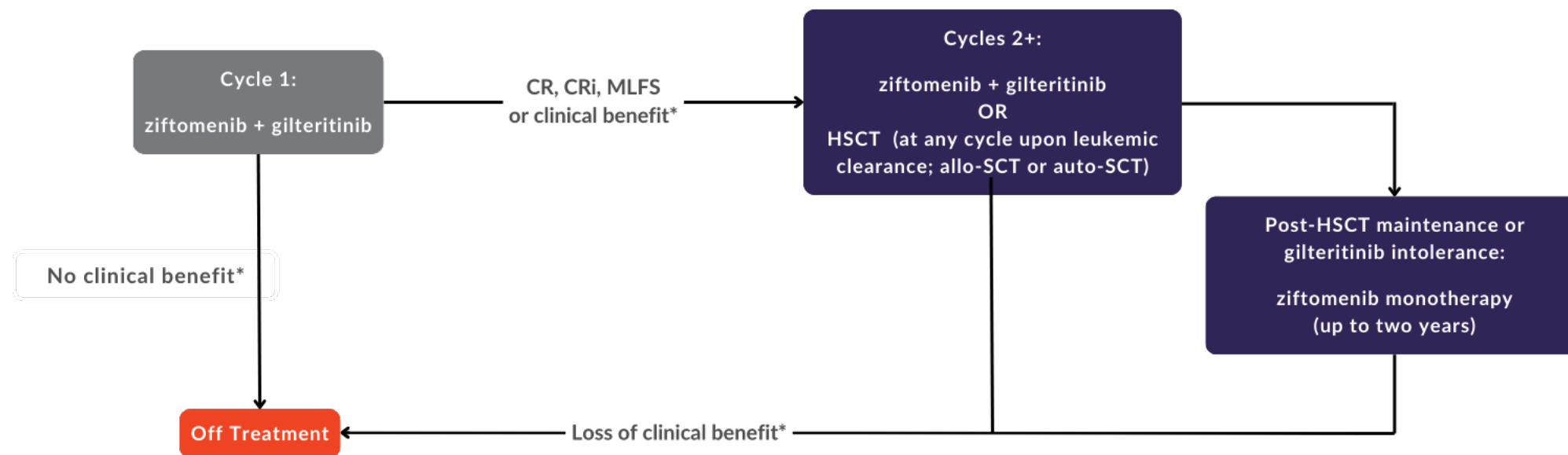
<sup>4</sup> Issa, Syndax ASH Investor Event (Dec. 2023)

<sup>5</sup> Issa, G. et al. Blood Cancer J. 11, 162 (2021)

ORR includes CR, CRh, CRi, MLFS

# KOMET-008: Phase 1 Combination Trial of Ziftomenib in Patients with R/R AML

## Ziftomenib + gilteritinib combination



- Phase 1a Dose Escalation
  - Arm A: Ziftomenib in combination with FLAG-IDA or LDAC or gilteritinib (illustrated above) in relapsed or refractory (R/R) NPM1-mutant AML
  - Arm B: Ziftomenib in combination with FLAG-IDA or LDAC in R/R KMT2A-rearranged AML
- Patients must also have documented FLT3 mutation if receiving gilteritinib

\*Per investigator discretion

CR, complete remission; CRi, CR with incomplete count recovery; MLFS, morphologically leukemia-free state; HSCT, hematopoietic stem-cell transplantation; allo-SCT, allogeneic stem-cell transplantation; auto-SCT, autologous stem-cell transplantation

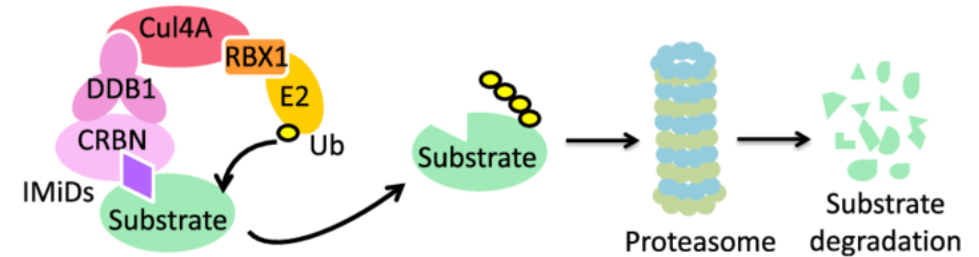




# Targeting Foundational Mutations has Transformed Deadly Hematologic Cancers into Chronic Diseases

## Multiple Myeloma

- Until the 2000's, there were few treatment options for multiple myeloma, and the median survival was 2–3 years.
- With the advent of immunomodulatory drugs (IMiDs) and proteasome inhibitors (PIs) in the 2000's, the outcomes of patients are now significantly improving.
- Many patients can now live with their disease > 10 years.
- **IMiDs have become a cornerstone of treatment for patients with multiple myeloma and are used in combinations at all stages of disease.**

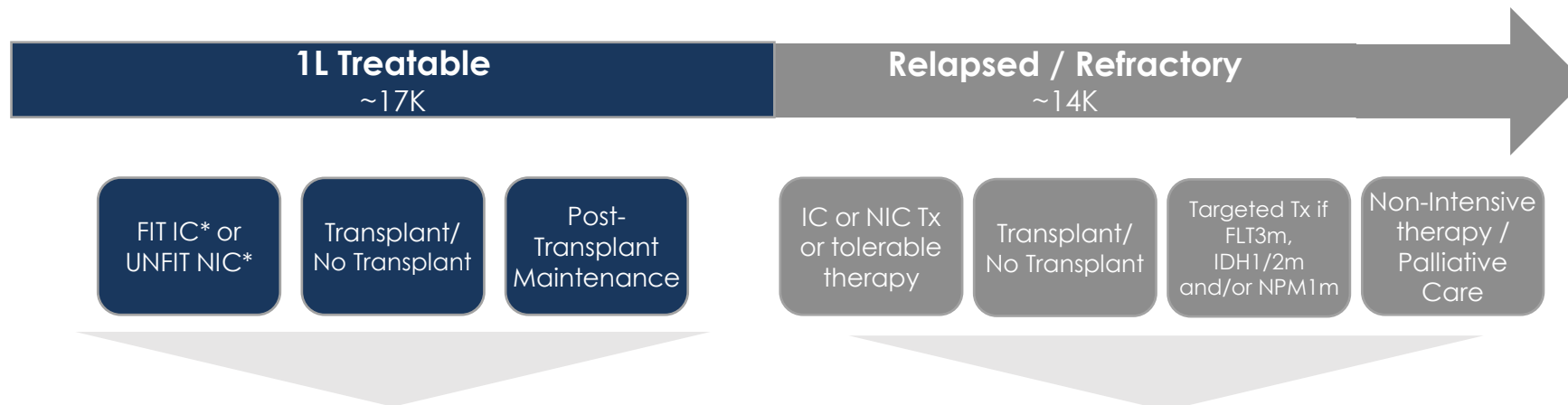
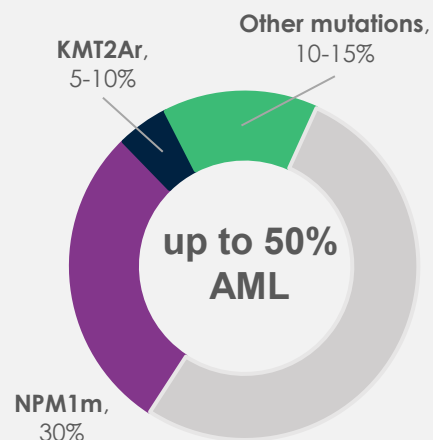


**IMiD combinations increased 5yr OS from 35% to > 65%; class generated ~\$15B in revenues at peak**



# We are Investigating Ziftomenib Across the AML Continuum in up to 50% of Patients for Whom Menin-KMT2A Pathway is a Disease Driver

## Prevalence of Ziftomenib Eligible Patients



### KOMET-007

- 1L Zifto + Ven / Aza
- 1L Zifto + 7+3

### Investigator/ Company Sponsored Studies

- Post-HSCT Maintenance

### KOMET-001

- R/R NPM1m AML

### KOMET-007

- R/R Zifto + Ven/Aza

### KOMET-008

- R/R Zifto + FLAG-IDA
- R/R Zifto + LDAC
- R/R Zifto + gilteritinib

\*FIT IC = patients eligible for induction chemotherapy; UNFIT NIC = patients eligible for non-intensive chemotherapy



# Ziftomenib Offers a Multi-Billion-Dollar Opportunity in AML and Beyond

## Potential to Transform Outcomes Across the Continuum of Care

### Relapsed / Refractory

- Initial approval represents **30% of potential patients**
- KOMET-001 registration-directed study for FDA full approval
- **BTD granted in R/R NPM1-mutant AML** indicating potential for substantial improvement over available therapies

### Frontline / Maintenance

- Significant opportunity in 1L AML and Maintenance
- Safety, tolerability and clinical activity anticipated to be ideal for combinations with SOC and with maintenance indication

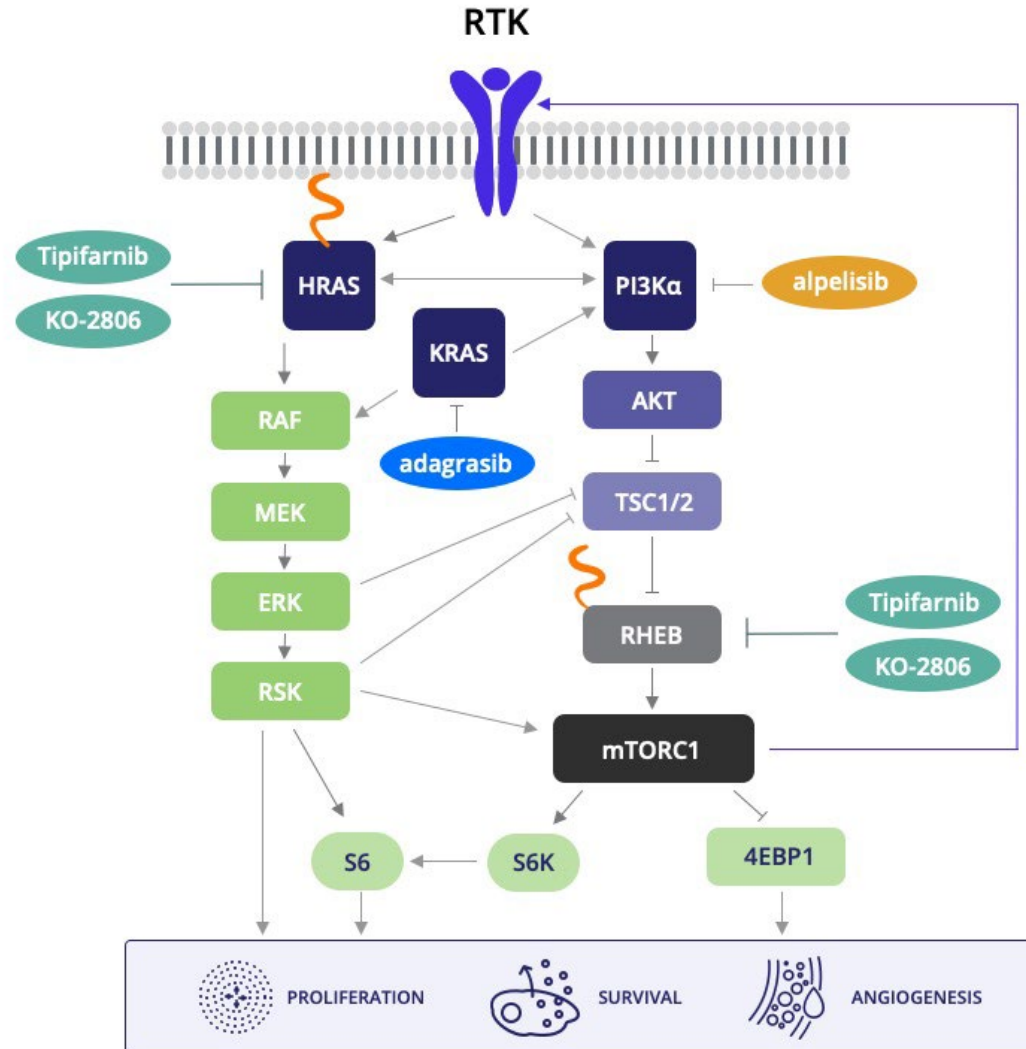
### Other Indications

- Compelling **additional opportunities beyond AML** offer multi-billion-dollar potential
- Translational data support potential in **solid tumor and non-oncology indications**
- **Next-generation menin inhibitor** currently under development

# TIPIFARNIB: FARNESYL TRANSFERASE INHIBITOR (FTI)



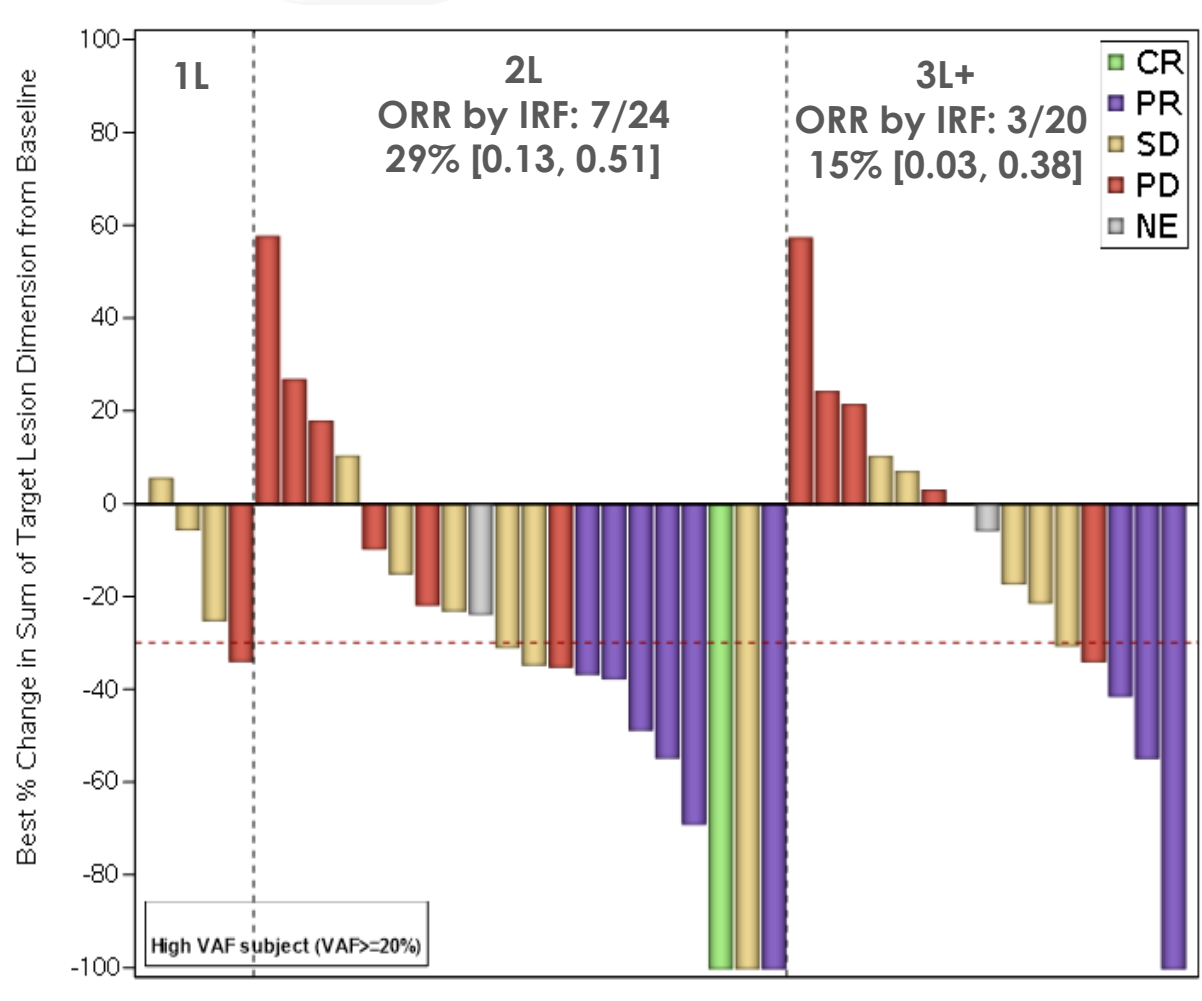
# Therapeutic Applications of Farnesyl Transferase Inhibitors



- Dysregulated RAS-MAPK and PI3Kα/AKT/mTOR signaling are key drivers of various cancers. Targeted cancer therapies such as alpelisib and adagrasib slow tumor progression by inhibiting individual elements in this complex signaling pathway
- However, resistance to these treatments develops through compensatory activation of complementary proteins, including receptor tyrosine kinases and mTOR
- Farnesyl Transferase Inhibitors (FTIs) can blunt the compensatory reactivation process by inhibiting farnesylation-mediated activation of additional proteins in the pathway – HRAS and RHEB
- By combining targeted therapies with FTIs, we believe we can reshape treatment options for many cancer patients



# Tipifarnib Shows Clinical Benefit in HRAS mutant HNSCC



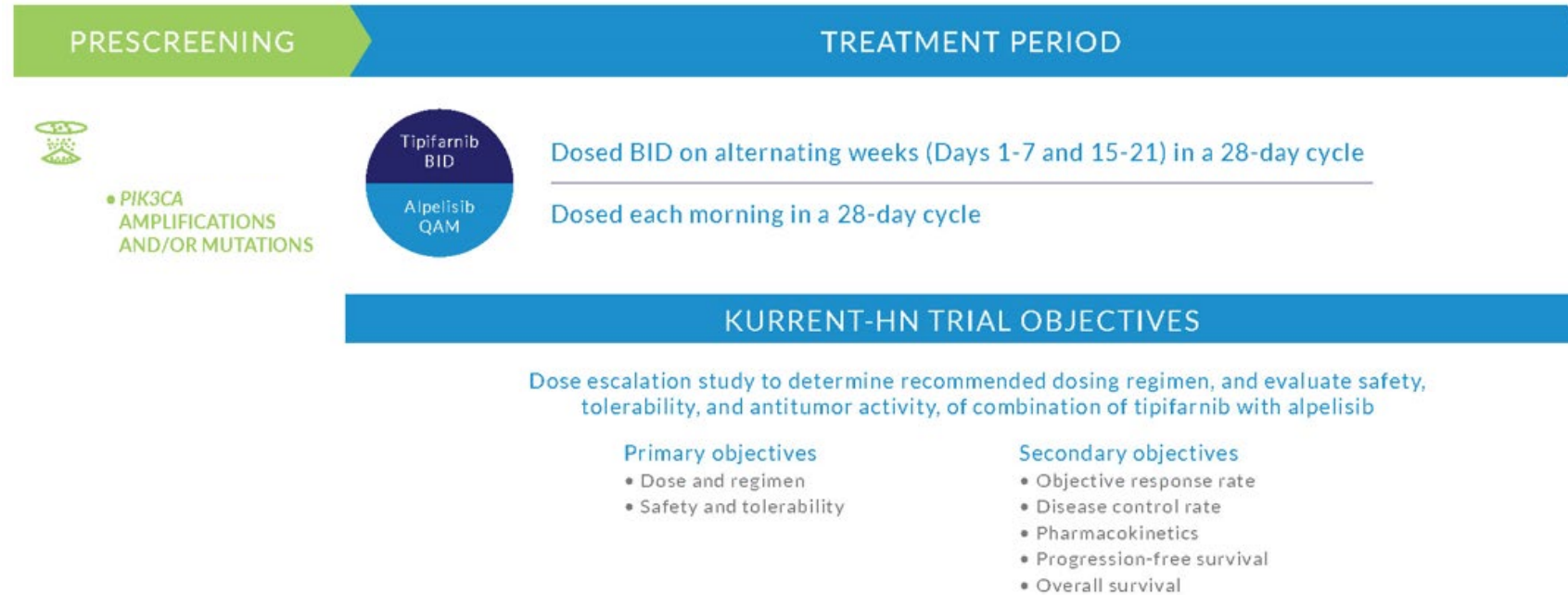
**6/10 responders had BOR of PD in the last prior line with IO-based therapies**  
**PFS in these ranged from 1-5 months vs. 6 –27 months on tipifarnib**

## Patients with **High VAF** in mITT (N=50)

	Investigator Assessment	Independent Review Facility
Best Overall Response, n (%)		
Confirmed CR	1 (2)	1 (2)
Confirmed PR	14 (28)	9 (18)
SD	17 (34)	14 (28)
PD	6 (12)	14 (28)
NE	12 (24)	12 (24)
DCR, n (%) [95% CI]	32 (64) [0.49, 0.77]	24 (48) [0.34, 0.63]
ORR, n (%) [95% CI]	15 (30) [0.18, 0.45]	10 (20) [0.10, 0.34]
mDoR, months [95% CI]	5.6 [3.88, 9.23]	6.5 [3.88, -]
mPFS, months [95% CI]	3.7 [2.60, 5.55]	2.6 [1.87, 4.40]

mITT: Patients treated with at least one dose of Tipifarnib. CR, complete response; PR, partial response; BOR, best overall response; IO, immuno-oncology; SD, stable disease; PD, progressive disease; NE, not evaluable; -, not calculable; ORR: objective response rate; DCR, disease control rate; mDoR, median duration of response; mPFS, median progression free survival.

# KURRENT-HN: PHASE 1/2 Combination Trial of Tipifarnib and Alpelisib in Patients with HNSCC

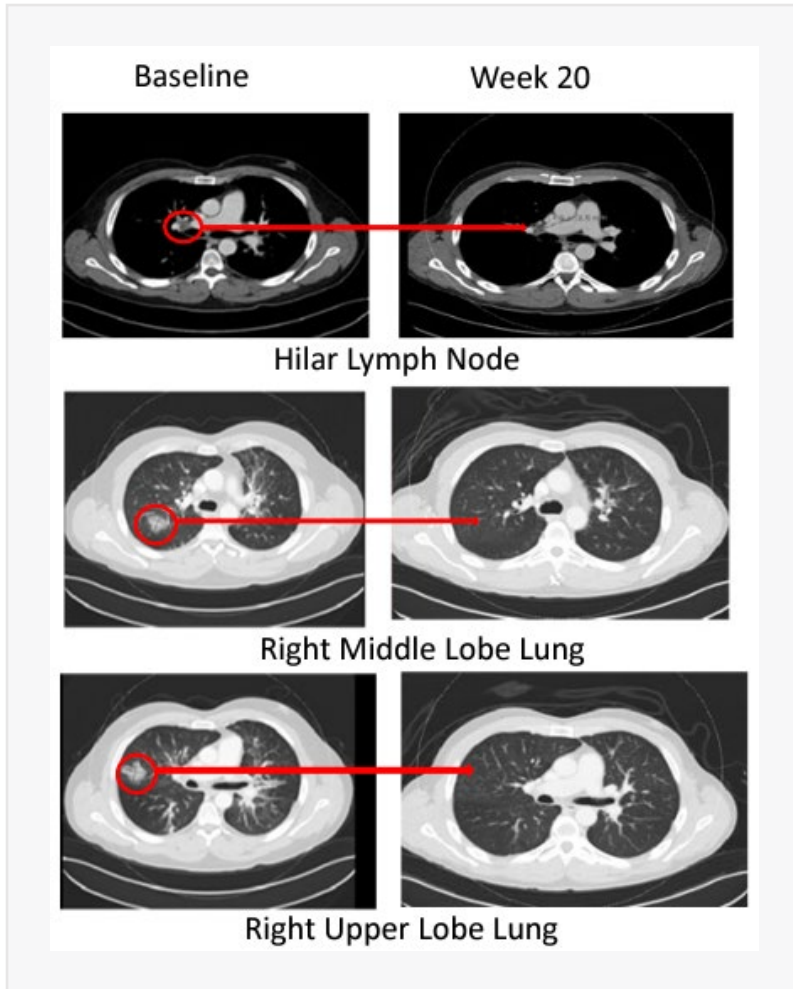


**Phase 1 clinical trial of tipifarnib and alpelisib in patients with recurrent/metastatic PIK3CA-amplified and/or PIK3CA-mutated HNSCC**

- Clinical collaboration to evaluate tipifarnib in combination with alpelisib for the treatment of patients with HNSCC whose tumors have PIK3CA mutation and/or amplification
- Under the collaboration, Kura sponsors the trial and supplies tipifarnib and Novartis supplies alpelisib



# Durable Clinical Response Observed in Patient with PIK3CA-dependent HNSCC



- 35yo, male, nonsmoker, HPV16 positive
- SCC of tonsil Stage III cT4N2M0; PD-L1 CPS = 60
- Prior Treatments
  - CDDP/rad for 1 mo (Nov-Dec2019), BOR UNK
  - Cemiplimab/ISA101b (Jun-Nov2021), BOR PD
- PIK3CA R88Q mutation (44%) and HRAS OE (3+ staining in 100% of tumor cells) by IHC from May 2021 biopsy
- DL1 tipifarnib, DL2 alpelisib; completed 6 cycles
- G1/2 TRAE, G3 lipase elevation; presented clinical benefit and improvement in respiratory symptoms
- 81% reduction in target lesions after 1 cycle of treatment
- 84% reduction in target lesions after 3 cycles (BOR)
- Continued on-study for >27 weeks maintaining QoL

**Additional data from KURRENT-HN clinical trial expected to be presented in 1H2025**

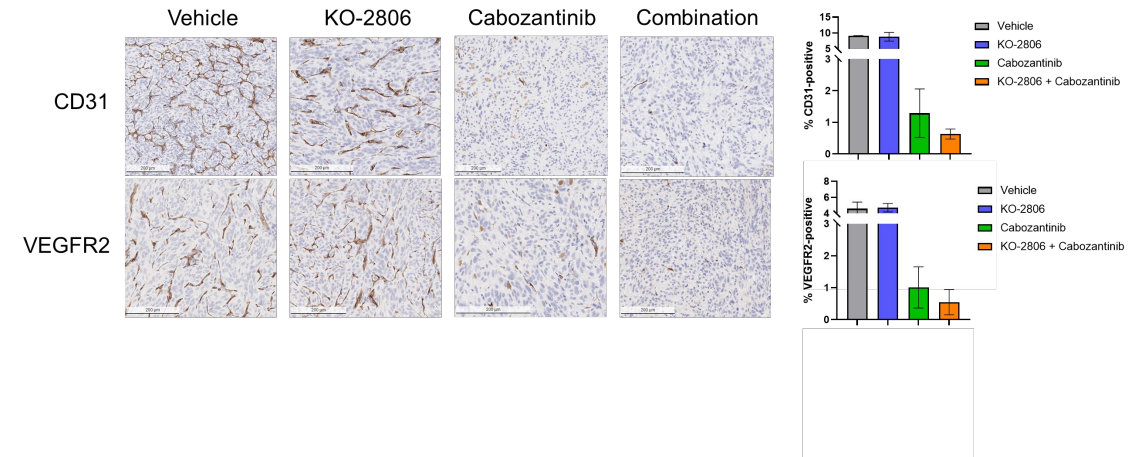
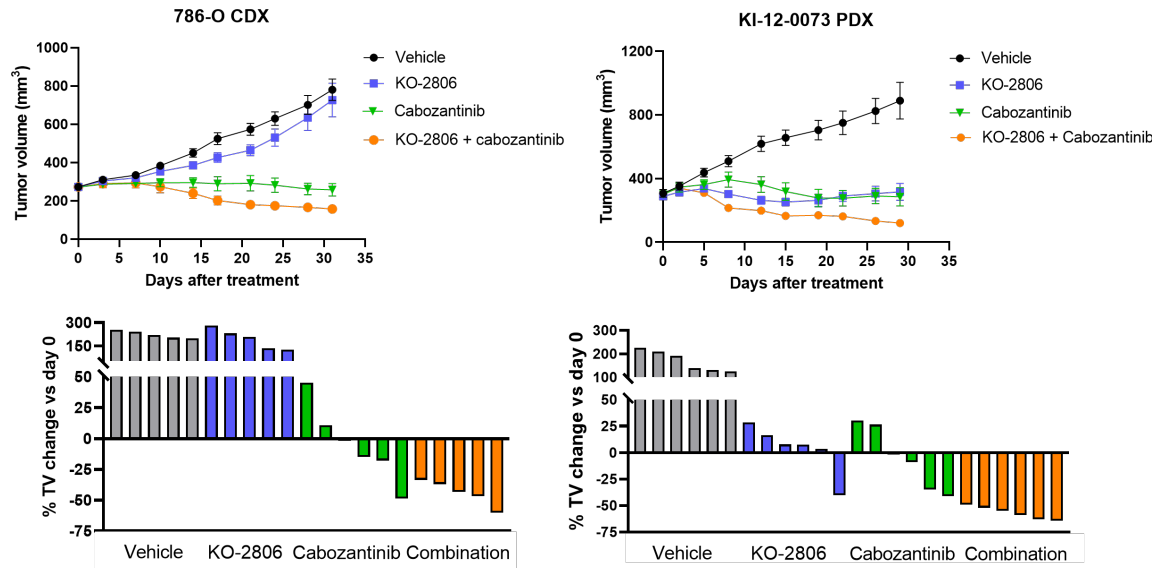
# KO-2806: NEXT- GENERATION FTI

# Combination of KO-2806 and Cabozantinib Demonstrates Synergistic Activity in ccRCC CDX & PDX Models



KO-2806 potentiates antitumor activity of cabozantinib in ccRCC models

KO-2806 enhances the anti-angiogenic activity of cabozantinib *in vivo*



IHC of 786-O tumors and quantitation

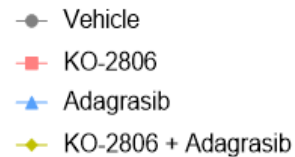
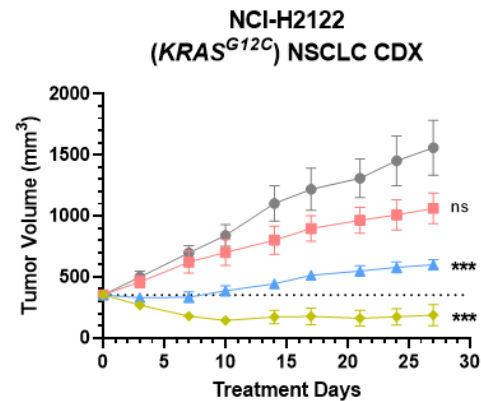
- KO-2806 enhances the anti-angiogenic activity of cabozantinib *in vivo*, as observed by decreased expression of vascular markers in 786-O tumors
- Studies are ongoing to further define the basis of the synergy of the combination

# Combination of KO-2806 to Enhance Antitumor Efficacy of KRAS<sup>G12C</sup> Inhibitor in NSCLC

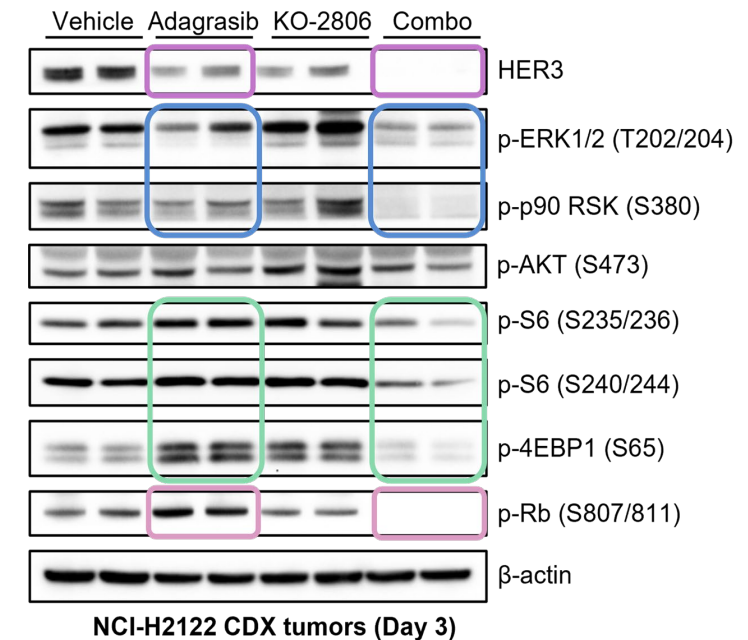
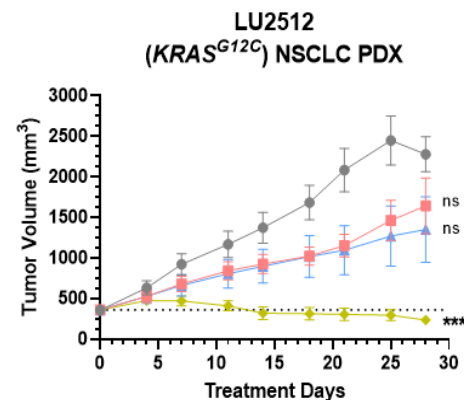


Combination of KO-2806 with a KRAS<sup>G12C</sup> inhibitor causes tumor regressions in patient-derived and cell-derived NSCLC xenografts

Combination of KO-2806 with a KRAS<sup>G12C</sup> inhibitor suppresses mTOR and MAPK signaling and decreases proliferation



\*\*\* = p<0.001



- Combination of KO-2806 with adagrasib enhances the depth and duration of response compared with single-agent KRAS<sup>G12C</sup> inhibitor treatment

# FIT-001 Phase 1 First-in-Human Clinical Trial of KO-2806 in Patients with Advanced Solid Tumors



PART 1A (MONOTHERAPY)  
DOSE ESCALATION

PART 1A (COMBINATIONS)  
DOSE ESCALATION

PART 1B (COMBINATIONS)  
DOSE EXPANSION

## OBJECTIVES

### Primary

- Evaluate the safety and tolerability of KO-2806 (dose escalation)
- Determine the MTD/HPDD and/or the OBAD of KO-2806 (dose escalation)
- Define the RP2D of KO-2806 (dose expansion)
- Evaluate the antitumor activity of KO-2806 in combination with cabozantinib in ccRCC and adagrasib in KRAS<sup>G12C</sup>-mutant NSCLC (dose expansion)

### Secondary

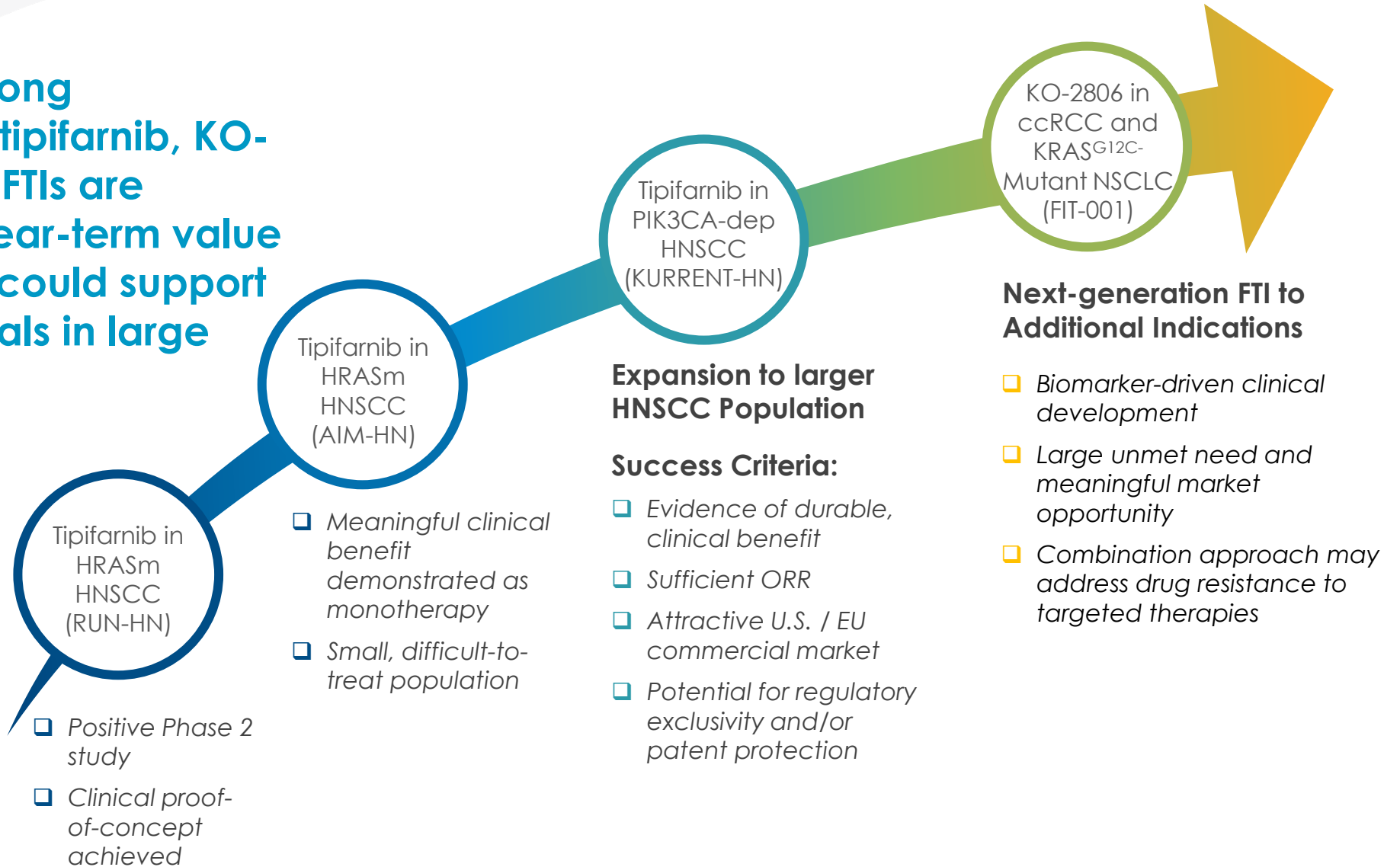
- Evaluate the safety and tolerability of KO-2806 (dose expansion)
- Evaluate the preliminary antitumor activity of KO-2806 (dose escalation and dose expansion)
- Characterize the PK of KO-2806 when administered as monotherapy, and the PK of KO-2806 and the combination agents when administered in combination therapy (dose escalation and expansion)

**Ongoing enrollment in FIT-001 Phase 1 dose-escalation trial of KO-2806 as a monotherapy and in combination**



# FTI Franchise Development Strategy

Building on a strong foundation with tipifarnib, KO-2806 and future FTIs are positioned for near-term value inflections, and could support multiple approvals in large indications





# Forecasted Milestones & Financial Highlights



ESTIMATED TIME OF  
ACHIEVEMENT

PROGRAM	MILESTONE	ESTIMATED TIME OF ACHIEVEMENT
<b>ZIFTOMENIB</b> Menin Inhibitor	Dose first patients in non-NPM1m/non-KMT2A-r AML expansion cohort	✓
	Dose first patients in KOMET-008 combination trial	✓
	Initiate post-transplant maintenance program	✓
	Receive Breakthrough Therapy Designation for R/R NPM1-mutant AML	✓
	Complete enrollment of 85 patients in KOMET-001 registration-directed trial	Mid-2024
	Identify recommended Phase 2 dose in combination with ven/aza	Mid-2024
	Identify recommended Phase 2 dose in combination with 7+3	Mid-2024
	Initiate dose validation/expansion with ven/aza in 1L AML	2H 2024
	Investigational new drug application for solid tumor indication	2H 2024
	Nominate next-generation menin inhibitor development candidate	End of 2024
<b>TIPIFARNIB</b> Farnesyl Transferase Inhibitor (FTI)	Identify OBAD in combination with alpelisib in PIK3CA-dependent HNSCC	End of 2024
	Present data from KURRENT-HN trial in combination with alpelisib in PIK3CA-dependent HNSCC	1H 2025
<b>KO-2806</b> Next-Generation FTI	Dose first patients in FIT-001 trial in combination with cabozantinib in ccRCC	✓
	Dose first patients in FIT-001 trial in combination with adagrasib in KRAS <sup>G12C</sup> -mutated NSCLC	Mid-2024
<b>Financial Highlights</b> Nasdaq: KURA	\$527M in pro forma cash as of March 31, 2024* provides runway into 2027	
	Shares outstanding as of March 31, 2024: 76.2M basic; 25.2M options, RSUs, PSUs, warrants & pre-funded warrants	

OBAD = optimal biologically active dose

\* Cash, cash equivalents and short-term investments



# DEVELOPING PRECISION MEDICINES FOR THE TREATMENT OF CANCER

Corporate Presentation – May 2024