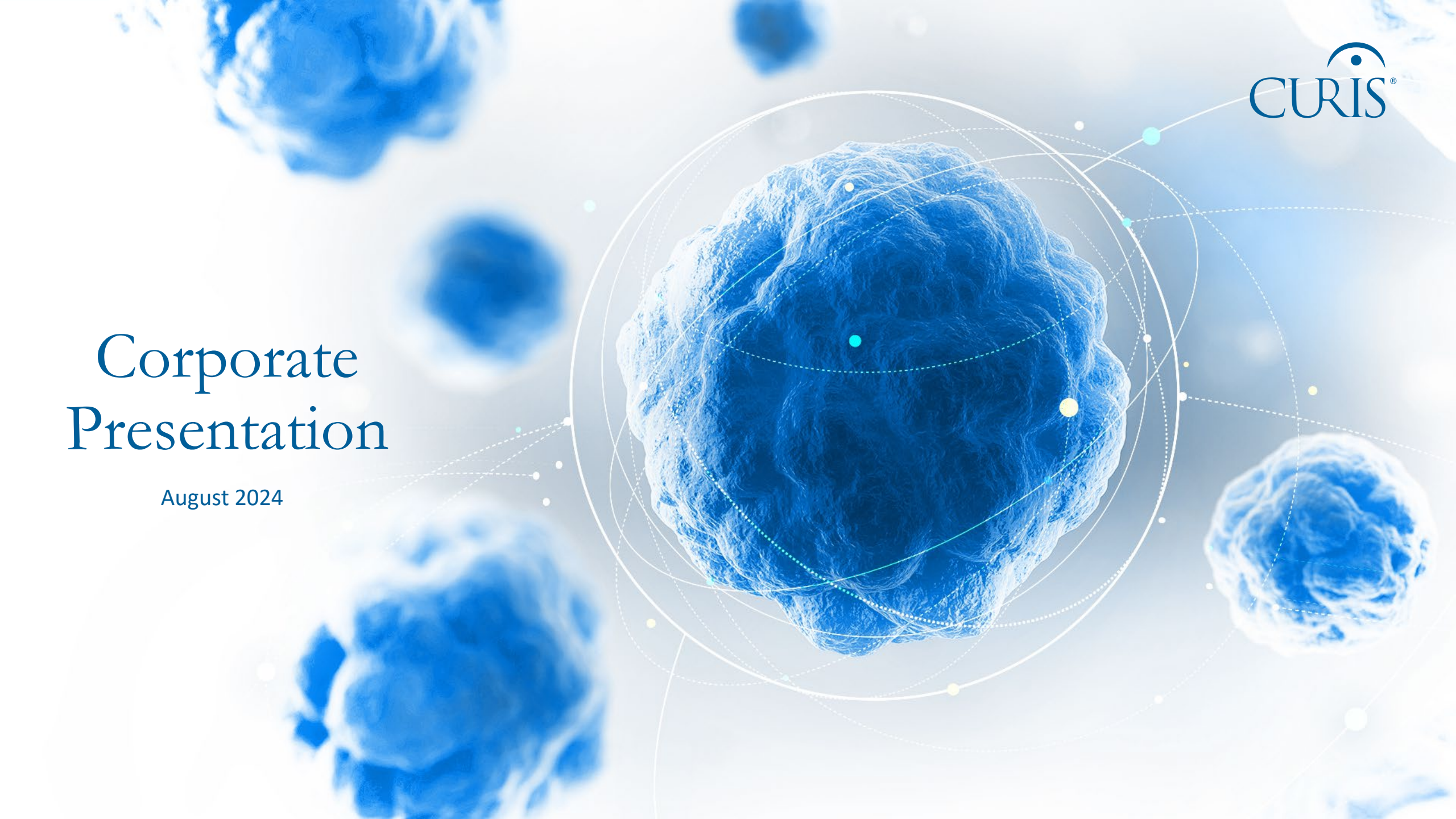


# Corporate Presentation

August 2024



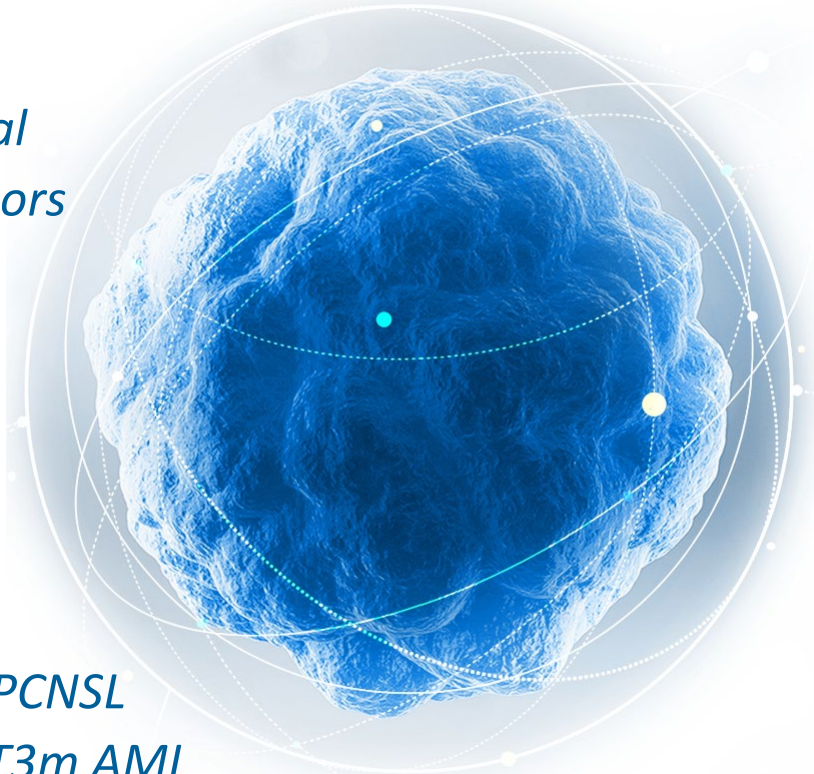
# Cautionary note regarding forward looking statements and disclaimers

This presentation contains certain forward-looking statements about Curis, Inc. (“we,” “us,” or the “Company”) within the meaning of the Private Securities Litigation Reform Act of 1995, as amended. Words such as “expect(s),” “believe(s),” “will,” “may,” “anticipate(s),” “focus(es),” “plans,” “mission,” “strategy,” “potential,” “estimate(s),” “opportunity,” “intend,” “project,” “seek,” “should,” “would” and similar expressions are intended to identify forward-looking statements. Forward-looking statements are statements that are not historical facts, reflect management’s expectations as of the date of this presentation, and involve important risks and uncertainties. Forward-looking statements herein include, but are not limited to, statements with respect to the timing and results of future clinical and pre-clinical milestones; the timing of future preclinical studies and clinical trials and results of these studies and trials; the clinical and therapeutic potential of our drug candidates; our cash runway; the proposed focus on emavusertib and management’s ability to successfully achieve its goals. These forward-looking statements are based on our current expectations and may differ materially from actual results due to a variety of important factors including, without limitation, risks relating to: whether and when the U.S. Food and Drug Administration (the “FDA”) may take further regulatory action with regard to our trials, whether any of our drug candidates will advance further in the clinical development process and whether and when, if at all, they will receive approval from the FDA or equivalent foreign regulatory agencies; whether historical preclinical results will be predictive of future clinical trial results; whether historical clinical trial results will be predictive of future trial results; whether any of our drug candidate development efforts will be successful; whether any of our drug candidates will be successfully marketed if approved; our ability to achieve the benefits contemplated by our collaboration agreements; management’s ability to successfully achieve its goals; the sufficiency of our cash resources; our ability to raise additional capital to fund our operations on terms acceptable to us and the use of proceeds of any offering of securities or other financing; general economic conditions; competition; and the other risk factors contained in our periodic reports filed with the Securities and Exchange Commission, including the Company's Annual Report on Form 10-K for the fiscal year ended December 31, 2023 and the Company's Quarterly Reports on Form 10-Q for the quarters ended March 31, 2024 and June 30, 2024, which are available on the SEC website at [www.sec.gov](http://www.sec.gov). You are cautioned not to place undue reliance on these forward-looking statements that speak only as of the date hereof, and we do not undertake any obligation to revise and disseminate forward-looking statements to reflect events or circumstances after the date hereof, or to reflect the occurrence of or non-occurrence of any events, except as required by law.

This presentation includes statistical and other industry and market data that we obtained from industry publications and research, surveys, and studies conducted by third parties as well as our own estimates. All of the market data used in this presentation involves a number of assumptions and limitations, and you are cautioned not to give undue weight to such data. Industry publications and third-party research, surveys, and studies generally indicate that their information has been obtained from sources believed to be reliable, although they do not guarantee the accuracy or completeness of such information. Our estimates of the potential market opportunities for our product candidates include several key assumptions based on our industry knowledge, industry publications, third-party research, and other surveys, which may be based on a small sample size and may fail to accurately reflect market opportunities. While we believe that our internal assumptions are reasonable, no independent source has verified such assumptions.

# Emavusertib is a potential first-to-market IRAK 4 inhibitor

- *Being evaluated in Phase 1/2 clinical studies in NHL, AML, and Solid Tumors*
- *Potential for monotherapy and combination use in NHL, AML and Solid Tumors*
- *Near-Term Milestones (late 2024)*
  - *POC combination data in R/R PCNSL*
  - *Data in ~20 patients in R/R FLT3m AML*
  - *Initial safety data for triplet combination in frontline AML*



*Demonstrated safety and single-agent activity*

*Demonstrated synergy with BTKi, HMA, BCL2i*

*Broad opportunity in NHL, AML, and Solid Tumors*



# Broad Market Opportunity in NHL, AML, and Solid Tumors

## Significant market opportunities in current development programs

	PCNSL	FLT3m	AML
<b>US Incidence per 100K</b>	<b>0.5<sup>1</sup></b>	<b>1.3<sup>2</sup></b>	<b>4.2<sup>3</sup></b>
	<u>Newly Diagnosed Per Year</u>		
US	1,700 <sup>1</sup>	6,000 <sup>2</sup>	20,000 <sup>3</sup>
Big 5 Europe/Canada	1,800 <sup>1</sup>	5,200 <sup>4</sup>	17,000 <sup>4</sup>
Japan/China	<u>7,700<sup>1</sup></u>	<u>12,700<sup>4</sup></u>	<u>41,200<sup>4</sup></u>
<b>Total</b>	<b>11,200</b>	<b>23,900</b>	<b>78,200</b>

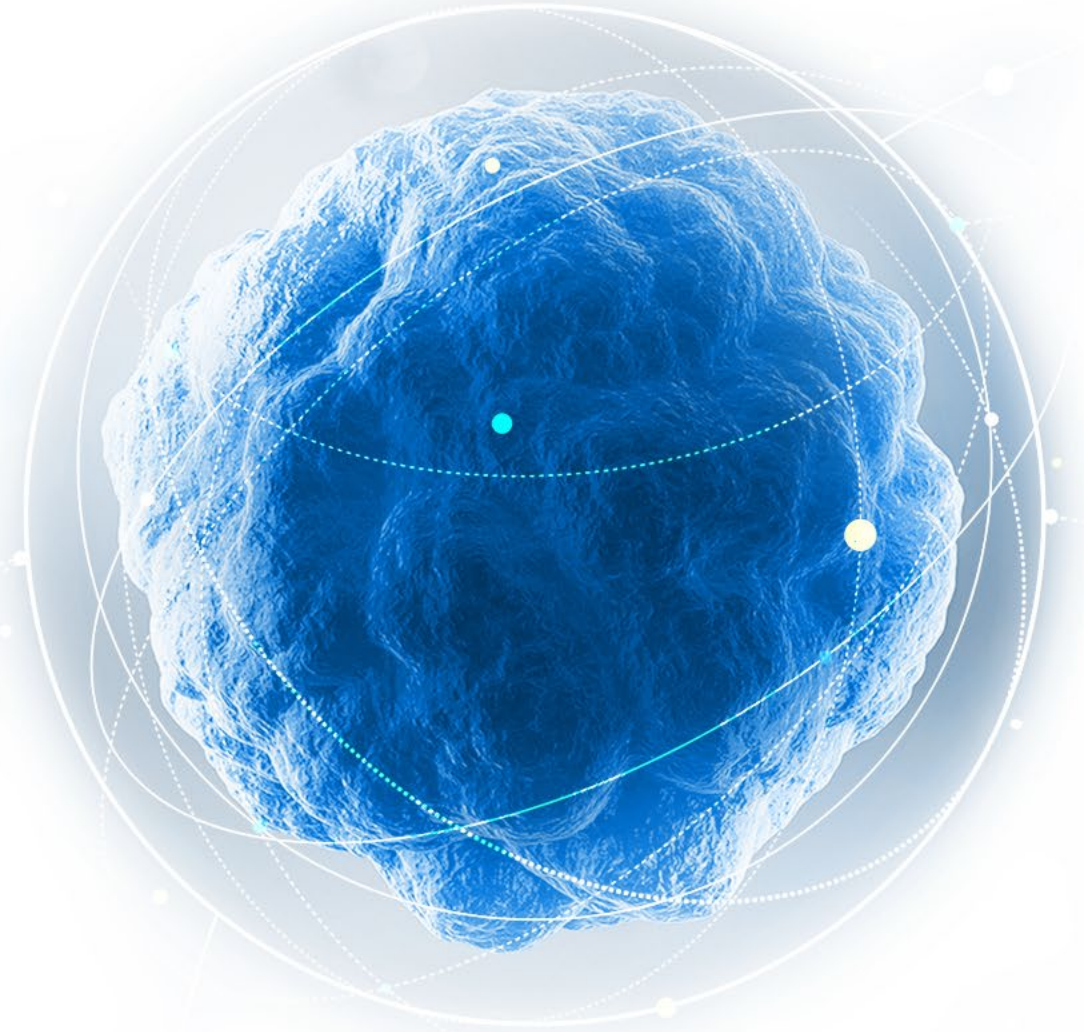
1 – Derived from incident rate in Lv Ther Adv Hematol 2022 and 2022 country population [data.worldbank.org]  
 2 – Derived from total AML cases (see footnote 4); FLT3m represents 30% of newly diagnosed AML cases [Daver Leukemia 2019]  
 3 – Vakiti Acute Myeloid Leukemia 2023 [www.ncbi.nlm.nih.gov]  
 4 – Clarivate DRG, March 2024

## Additional opportunities with NHL expansion

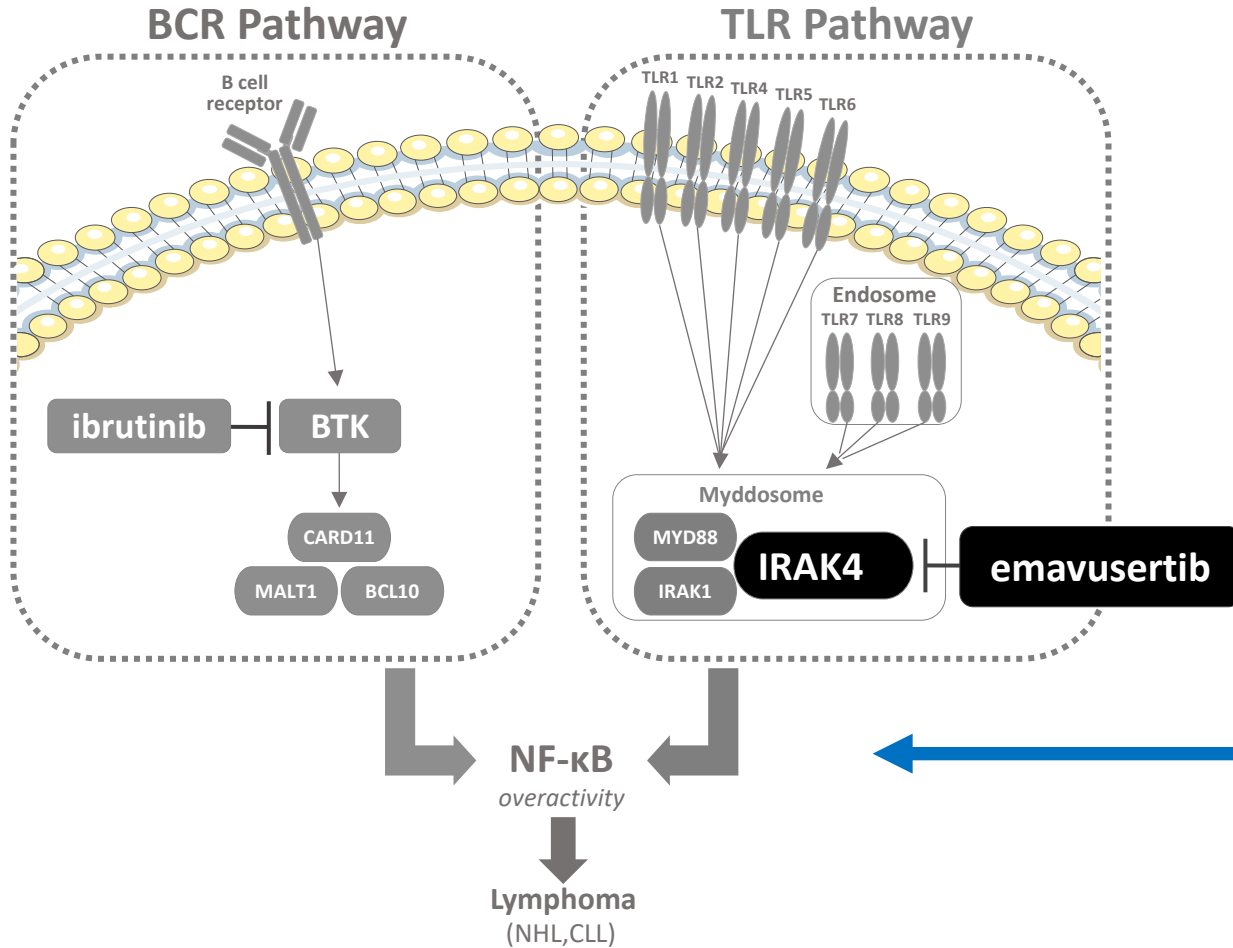
WM	MCL	MZL	ABC-DLBCL
<b>0.5<sup>5</sup></b>	<b>0.5<sup>6</sup></b>	<b>1.5<sup>7</sup></b>	<b>2.0<sup>8</sup></b>
	<u>Newly Diagnosed Per Year</u>		
1,700 <sup>5</sup>	1,700 <sup>6</sup>	5,000 <sup>7</sup>	6,800 <sup>8</sup>
1,800 <sup>5</sup>	1,800 <sup>6</sup>	5,500 <sup>7</sup>	7,500 <sup>8</sup>
<u>7,700<sup>5</sup></u>	<u>7,700<sup>6</sup></u>	<u>23,000<sup>7</sup></u>	<u>31,400<sup>8</sup></u>
<b>11,200</b>	<b>11,200</b>	<b>33,500</b>	<b>45,700</b>

5 – Derived from incident rate in <https://rarediseases.org/rare-diseases/waldenstroms-macroglobulinemia/#affected> and 2022 country population [data.worldbank.org].  
 6 – Derived from incident rate in <https://www.ncbi.nlm.nih.gov/books/NBK536985/> and 2022 country population [data.worldbank.org].  
 7 – Derived from incident rate in Kalashnikov, Blood Cancer Journal, April 2023 and 2022 country population [data.worldbank.org].  
 8 – Derived from incident rates in NHL incident rate of 18.6 per 100,000 (seer.cancer.gov) with DLBCL representing 25% of NHL per <https://www.ncbi.nlm.nih.gov/books/NBK557796/>. ABC represents 44% per letters to the editor, haematologica, 2011, 96 and 2022 country population [data.worldbank.org].

# Emavusertib in NHL



# Mechanism in NHL



*Two pathways drive NF-κB  
(which drives B-cell lymphomas)*

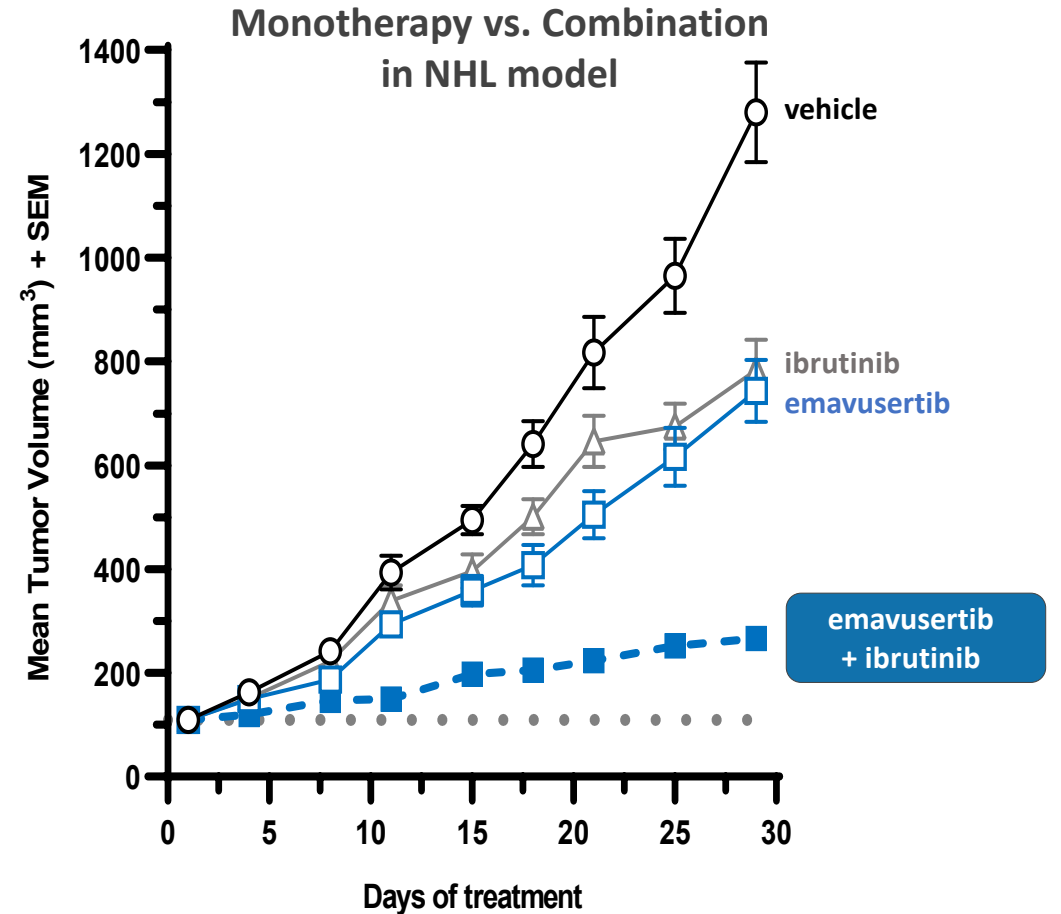
*combining IRAK4i and BTKi  
enables dual blockade of NF-κB*

# Emavusertib is synergistic with BTKi in NHL

## emavusertib + ibrutinib

blocking both BCR and TLR pathways has been demonstrated to be more effective than blocking either one alone

- IRAK4 inhibition synergizes with BTK inhibition to promote killing of **ABC-DLBCL**<sup>1</sup>
- Concurrent treatment with IRAKi and ibrutinib was significantly more potent in patient **CLL** cells than either drug alone<sup>2</sup>
- Data suggest IRAK4 as a novel treatment target for **CLL**; inhibition of IRAK4 blocks survival and proliferation of CLL cells<sup>3</sup>



Preclinical data in OCI-Ly10 model from Boohar et al. Waldenström Roadmap Symposium 2019

# Strategy in NHL

**1****Demonstrate safety**

19 patients<sup>1</sup> treated in TakeAim Lymphoma Ph 1b study, acceptable safety profile established, no overlapping dose-limiting toxicity with ibrutinib

**2****Demonstrate single-agent activity**

Single-agent activity demonstrated, with patients remaining on study up to 4 years

**3****Pursue fastest path to 1<sup>st</sup> label in R/R patients**

Identify orphan indication with clear unmet need that is addressable with emavusertib's novel mechanism of action

**4****Pursue partnership to expand across NHL**

Significant resources will be required to execute large clinical studies across multiple NHL subtypes and prepare for potential commercial launch



# Emavusertib safety profile in NHL<sup>1</sup>

- 19 patients treated with emavusertib in combination with ibrutinib in multiple NHL subtypes
- Shown to be well tolerated with an acceptable safety profile
  - No DLTs observed at 100mg or 200mg
  - 2 reversible DLTs observed at 300mg (stomatitis and syncope)
- Emavusertib crosses the BBB and no dose-limiting CNS toxicities have been observed
- No dose-limiting myelosuppression has been observed

Grade 3+ TRAE in >1 Patient	100 mg BID+IBR (N=2)	200 mg BID+IBR (N=10)	300 mg BID+IBR (N=7)	Total (N=19)
	n (%)	n (%)	n (%)	n (%)
# patients having grade 3+ TRAEs	1 (50)	7 (70)	6 (86)	14 (74)
Platelet count decreased		2 (20)	1 (14)	3 (16)
Alanine aminotransferase increased		1 (10)	1 (14)	2 (11)
Aspartate aminotransferase increased		1 (10)	1 (14)	2 (11)
Fatigue		1 (10)	1 (14)	2 (11)
Hyponatraemia		2 (20)		2 (11)
Lipase increased	1 (50)	1 (10)		2 (11)

1 – Curis Emavusertib TakeAim Lymphoma ASH 2023 poster

Abbreviation: Treatment Related Adverse Event (TRAE), ibrutinib (IBR), Dose Limiting Toxicity (DLT), Blood Brain Barrier (BBB), Central Nervous System (CNS), twice daily (BID)

# Strategy in NHL

**1****Demonstrate safety**

19 patients<sup>1</sup> treated in TakeAim Lymphoma Ph 1b study, acceptable safety profile established, no overlapping dose-limiting toxicity with ibrutinib

**2****Demonstrate single-agent activity**

Single-agent activity demonstrated, with patients remaining on study up to 4 years

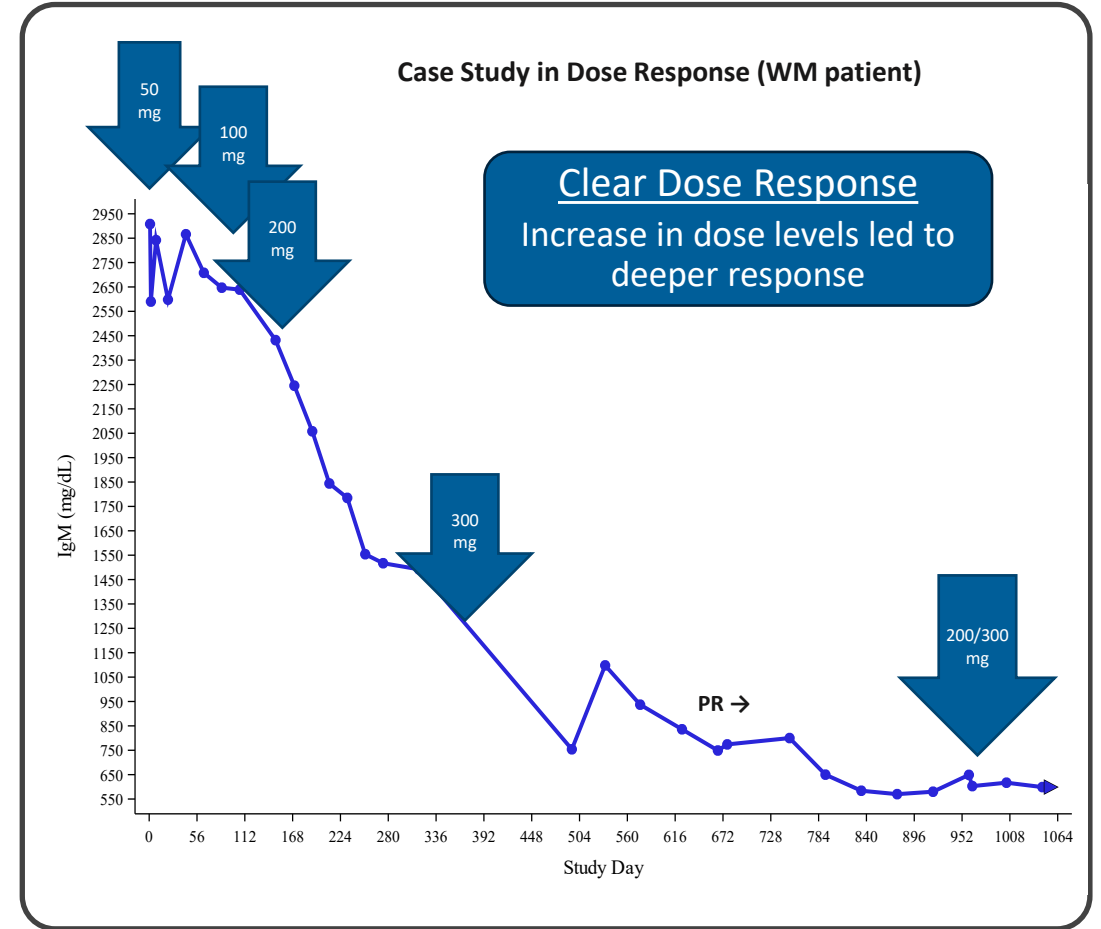
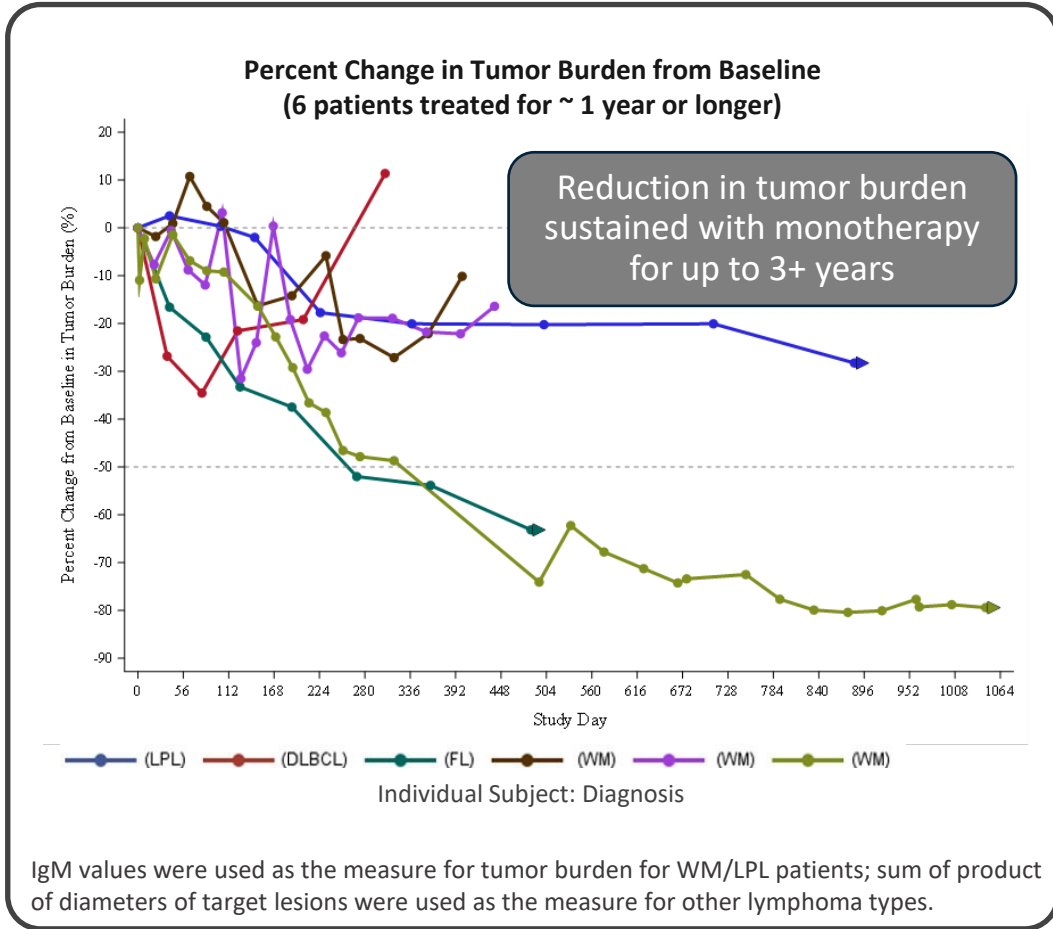
**3****Pursue fastest path to 1<sup>st</sup> label in R/R patients**

Identify orphan indication with clear unmet need that is addressable with emavusertib's novel mechanism of action

**4****Pursue partnership to expand across NHL**

Significant resources will be required to execute large clinical studies across multiple NHL subtypes and prepare for potential commercial launch

# Single-agent activity demonstrated in NHL



# Strategy in NHL

1

## Demonstrate safety

19 patients<sup>1</sup> treated in TakeAim Lymphoma Ph 1b study, acceptable safety profile established, no overlapping dose-limiting toxicity with ibrutinib

2

## Demonstrate single-agent activity

Single-agent activity demonstrated, with patients remaining on study up to 4 years

3

## Pursue fastest path to 1<sup>st</sup> label in R/R line patients

Identify orphan indication with clear unmet need that is addressable with emavusertib's novel mechanism of action

4

## Pursue partnership to expand across NHL

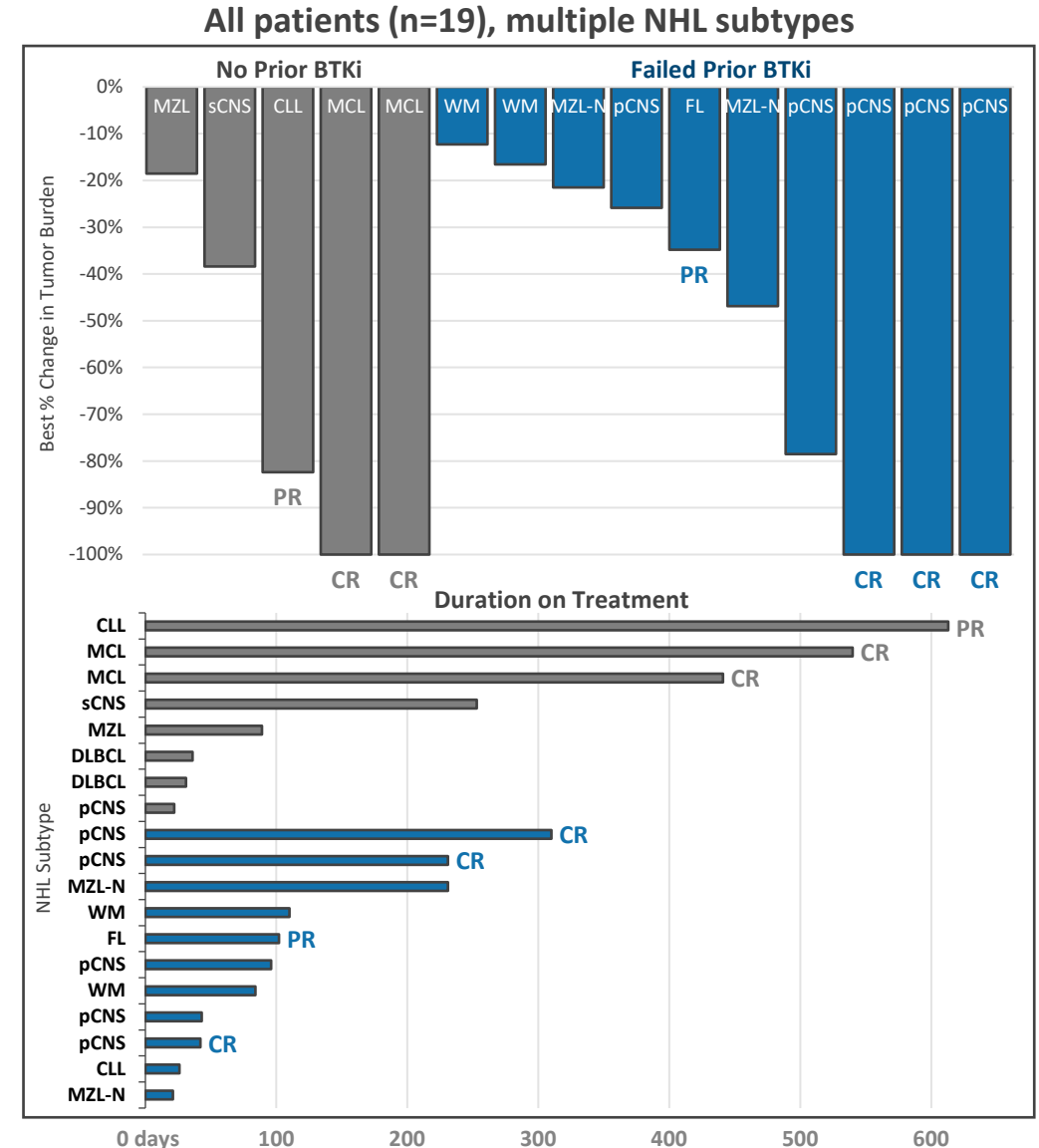
Significant resources will be required to execute clinical studies across multiple NHL subtypes and prepare for potential commercial launch



# Combination data from the Ph 1 basket study in NHL presented at ASH 2023

- Heavily pre-treated patients (1-10 prior lines)
- 7 of 19 patients achieved objective responses, **including in patients who failed prior BTKi**
- 15 of 19 patients saw a reduction in tumor burden
- Ongoing study with median treatment duration of 96 days (range 21-613 days)
- **PCNSL emerged as the lead indication in NHL with 3 of 5 objective responses**

Abbreviations: Complete Response (CR), Partial Response (PR)



# Strategy in NHL

**1****Demonstrate safety**

19 patients<sup>1</sup> treated in TakeAim Lymphoma Ph 1b study, acceptable safety profile established, no overlapping dose-limiting toxicity with ibrutinib

**2****Demonstrate single-agent activity**

Single-agent activity demonstrated, with patients remaining on study up to 4 years

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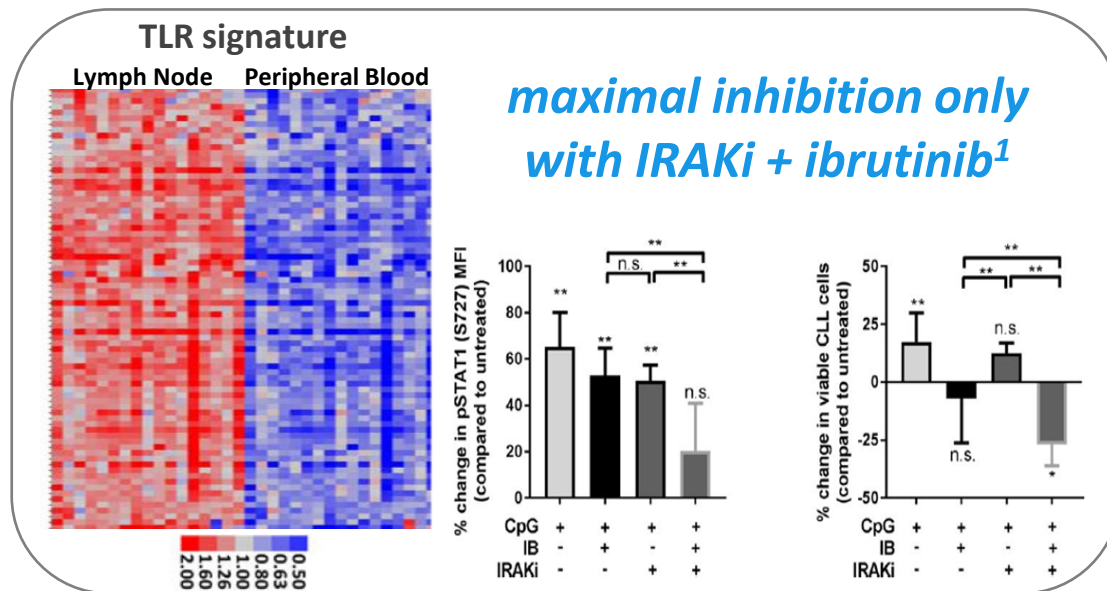
**4****Pursue partnership to expand across NHL**

Significant resources will be required to execute clinical studies across multiple NHL subtypes and prepare for potential commercial launch

# Additional NHL indications of interest

## CLL

- TLR signaling is highly activated in lymph node–resident CLL cells<sup>1</sup>



## Waldenström's Macroglobulinemia

- MYD88 and CXCR4 mutations activate NF- $\kappa$ B through the TLR pathway
- Recurring mutations in innate immune signaling and TLR/MYD88 pathway regulators are characteristic of ibrutinib-resistant WM patients<sup>2</sup>

## ABC-DLBCL

- Mutations in MYD88 activate NF- $\kappa$ B through the TLR pathway

## MCL

- TLR signaling is highly active in MCL, inducing proliferation and immune evasion in a MYD88-dependent fashion<sup>3</sup>

1) Dadashian Tumor Biol and Immunol 2019, 2) Jiménez Brit Jour Haem 2020, 3) Wang Cancer 2013

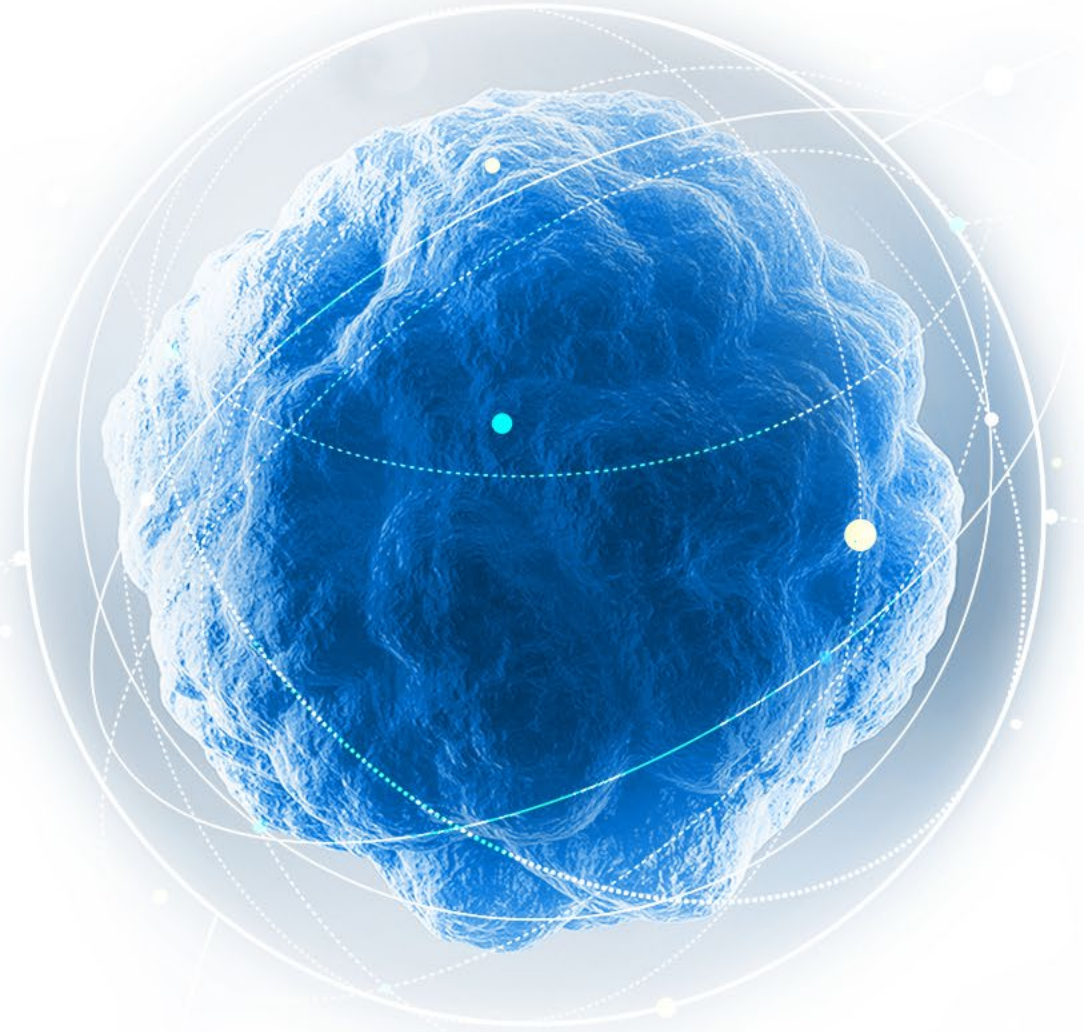
# Summary in NHL



- Emavusertib continues to demonstrate clear anti-cancer activity in R/R PCNSL
- Work with FDA to clarify registrational path and prepare for potential amendment of the current Ph 1/2 study into a study that would support an NDA filing for accelerated approval
- Prioritize additional NHL indications beyond PCNSL that could benefit from the dual-blockade of NF- $\kappa$ B (blocking the TLR pathway with emavusertib and blocking the BCR pathway with a BTKi)



# Emavusertib in AML



# Emavusertib binds to IRAK4 and FLT3, blocking both the TLR and FLT3 pathways

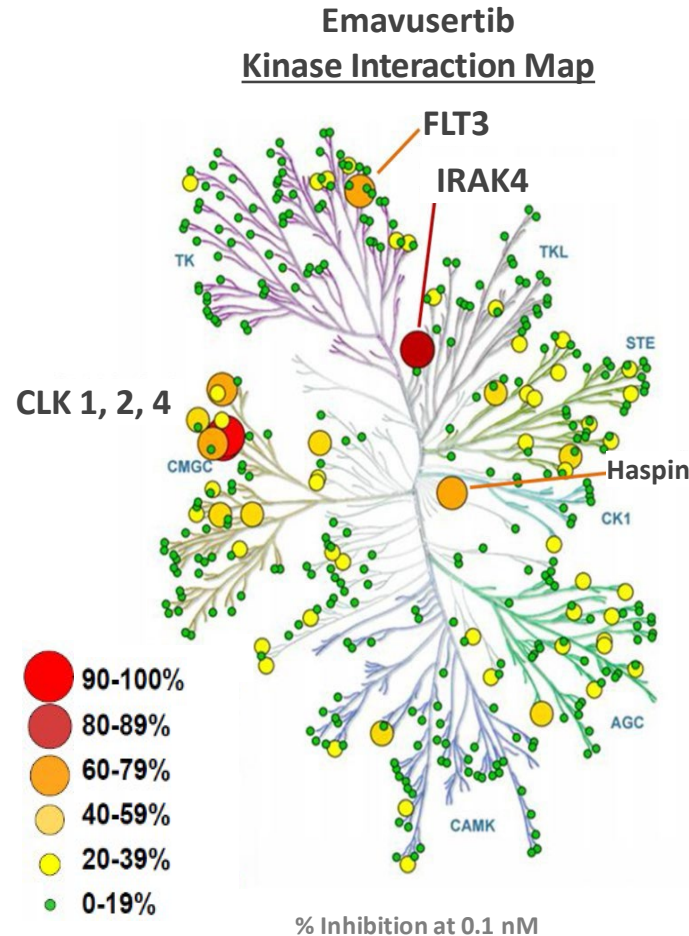


Illustration reproduced courtesy of Cell Signaling Technology

**Emavusertib Binding Affinity**

Target	K <sub>d</sub> nM
IRAK1	12,000
IRAK2	>20,000
IRAK3	8,500
<b>IRAK4</b>	<b>23</b>
DYRK1A	25
<b>FLT3 WT</b>	<b>31</b>
<b>FLT3 (D835H)</b>	<b>5</b>
<b>FLT3 (D835V)</b>	<b>44</b>
<b>FLT3 (D835Y)</b>	<b>3</b>
<b>FLT3 (ITD)</b>	<b>8</b>
<b>FLT3 (K663Q)</b>	<b>47</b>
<b>FLT3 (N841I)</b>	<b>16</b>
Haspin (GSG2)	32
CLK1	10
CLK2	20
CLK3	>20,000
CLK4	14
TrkA	130

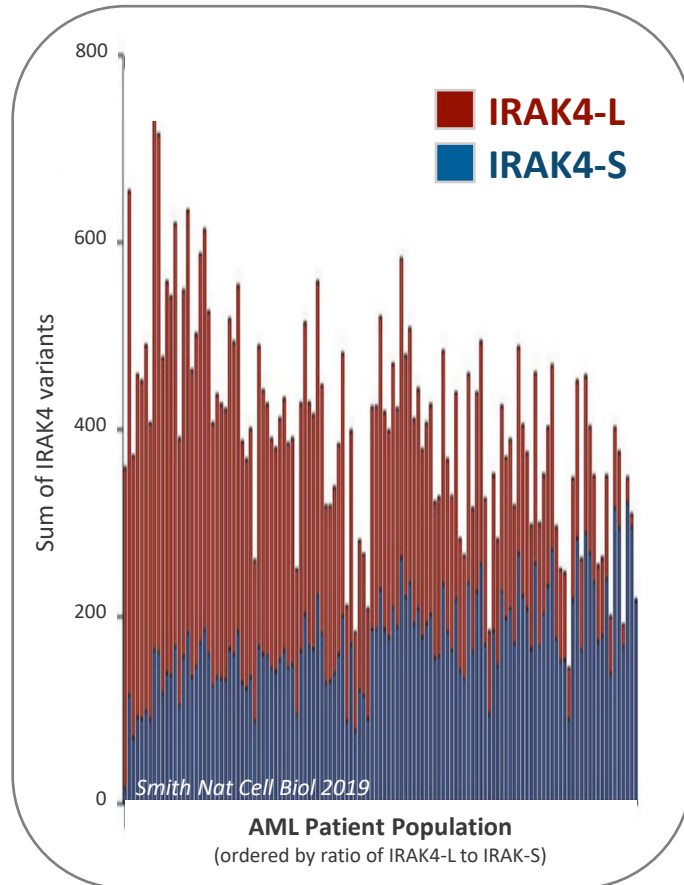
DiscoverX Kinase Panel (378 kinases screened)

**Binds tightly to IRAK4**

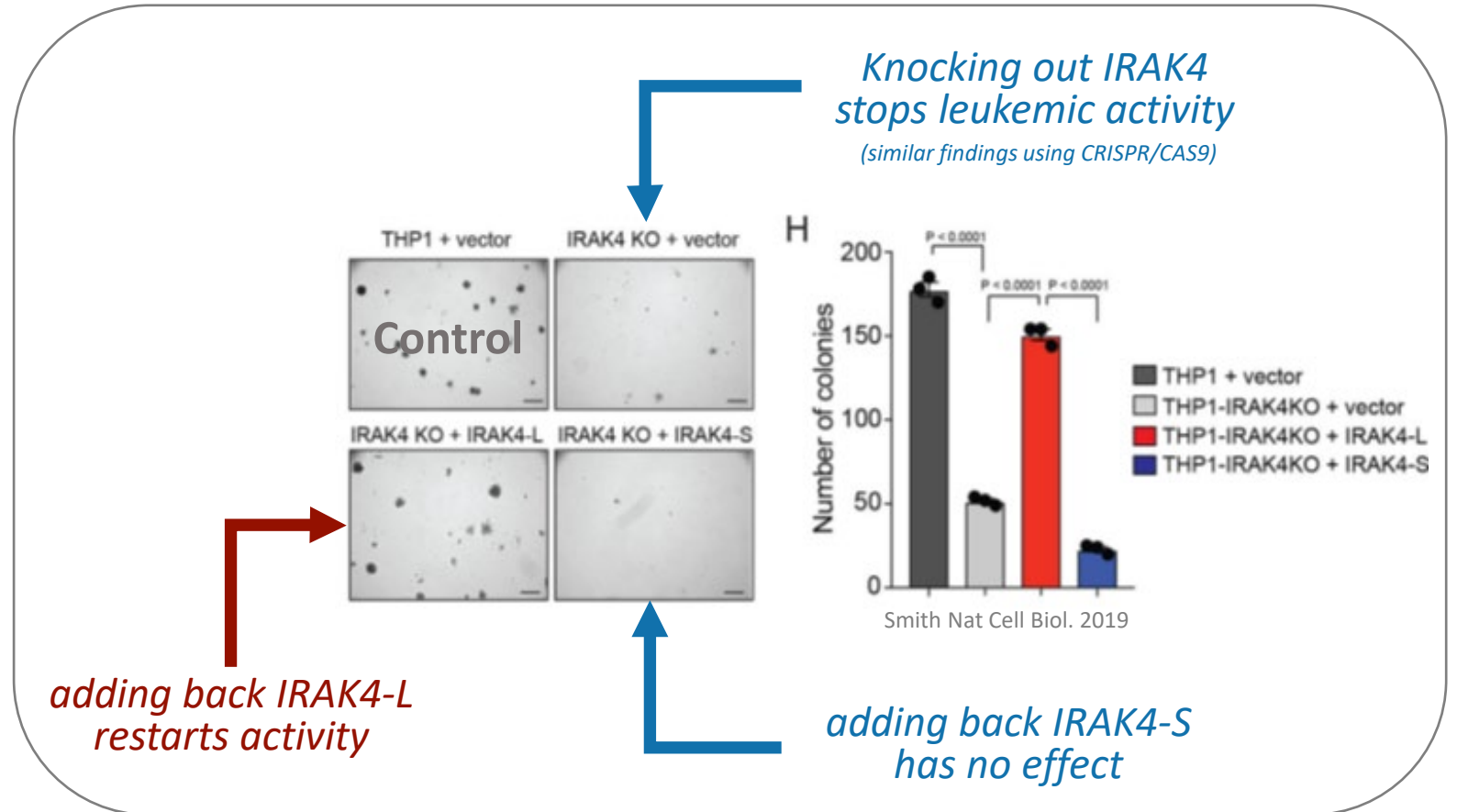
**Engineered to hit multiple targets of interest in oncology, including FLT3**

# IRAK4-L is an independent and powerful driver of disease in AML

**IRAK4-L is expressed in nearly all AML patients**



**IRAK4-L is oncogenic in AML**



# Strategy in AML

**1****Demonstrate safety**

123 patients<sup>1</sup> treated in TakeAim AML Ph 1/2 study, acceptable safety profile established

**2****Demonstrate single-agent activity**

Single-agent activity observed; next step is to confirm these initial findings in a larger number of patients

**3****Pursue fastest path to 1<sup>st</sup> label in R/R patients**

Address genetically-defined AML population with emavusertib's novel mechanism of action

**4****Explore frontline opportunity with combination**

IRAK4-L is expressed in nearly all AML patients; preclinical "all comer" models suggest emavusertib is synergistic with azacitidine and venetoclax

**5****Pursue partnership to maximize potential commercial opportunity**

Significant resources will be required to execute a large clinical study and prepare for potential commercial launch



# Emavusertib safety profile in AML<sup>1</sup>

- 123 patients treated with emavusertib in TakeAim Leukemia Study
- Shown to be well tolerated with an acceptable safety profile
- No dose-limiting myelosuppression has been observed

Grade 3+ TRAE > 1 patients	200 mg BID (n = 27)	300 mg BID (n = 78)	400 mg BID (n = 15)	500 mg BID (n = 3)	Total (n=123)
	n (%)	n (%)	n (%)	n (%)	n (%)
Number of patients having grade 3+ TRAEs	4 (14.8)	21 (26.9)	7 (46.7)	2 (66.7)	27 (27.6)
Blood creatine phosphokinase increased	0	6 (7.7)	0	0	6 (4.9)
Platelet count decreased	1 (3.7)	3 (3.8)	2 (13.3)	0	6 (4.9)
Rhabdomyolysis <sup>2</sup>	0	2 (2.6)	1 (6.7)	1 (33.3)	4 (3.3)
Anaemia	0	3 (3.8)	0	0	3 (2.4)
Aspartate aminotransferase increased	1 (3.7)	2 (2.6)	0	0	3 (2.4)

1 – Data as of February 26, 2024

2 – One patient with an event of Rhabdomyolysis met laboratory-defined criteria, defined as creatine phosphokinase > 10 × ULN with concurrent serum creatinine ≥ 1.5 × ULN. The remaining 3 patients experienced investigator-reported events of Rhabdomyolysis that did not meet laboratory-defined criteria.

# Strategy in AML

1

## Demonstrate safety

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## Demonstrate single-agent activity

Single-agent activity observed; next step is to confirm these initial findings in a larger number of patients

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## Explore frontline opportunity with combination

IRAK4-L is expressed in nearly all AML patients; preclinical "all comer" models suggest emavusertib is synergistic with azacitidine and venetoclax

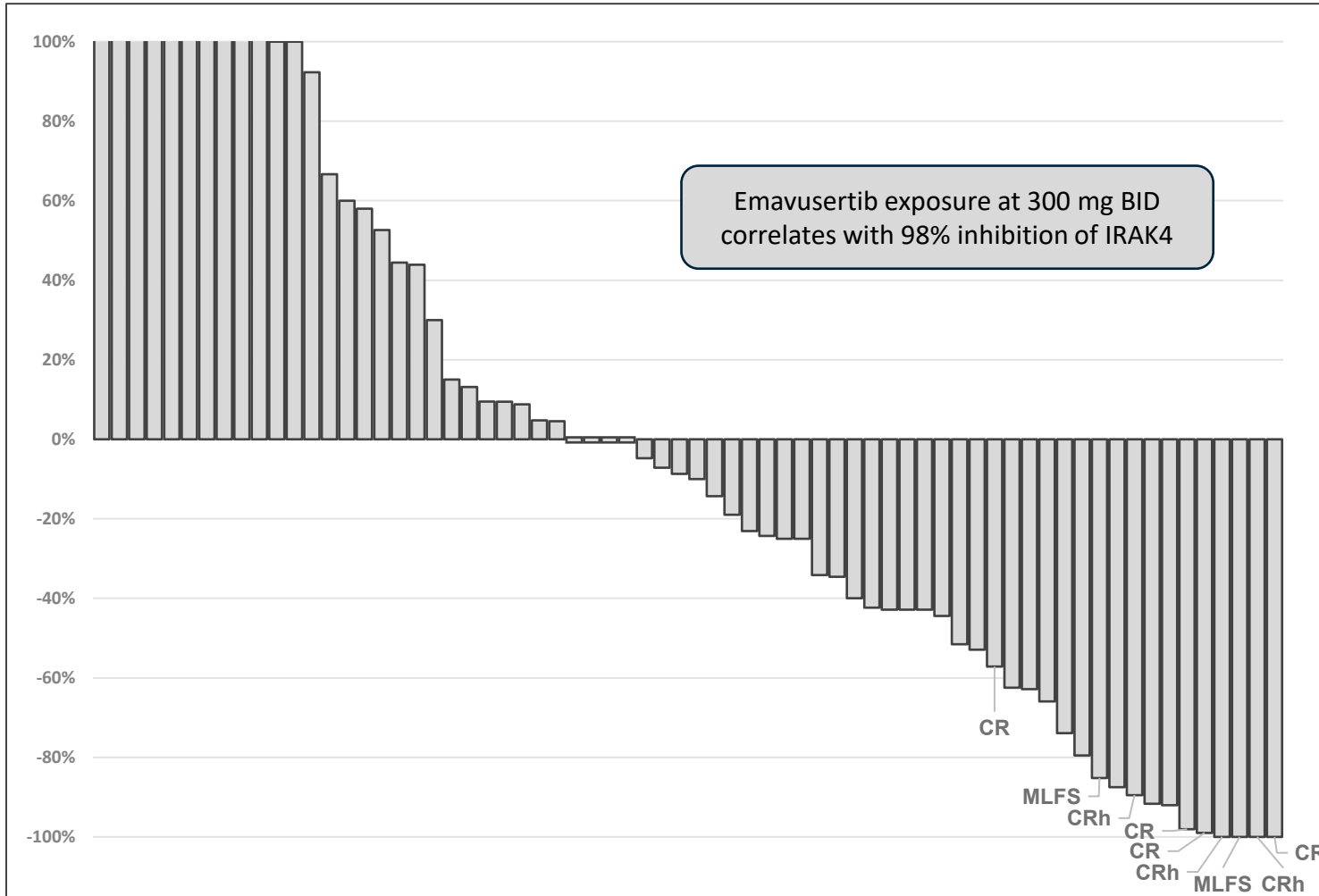
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## Pursue partnership to maximize potential commercial opportunity

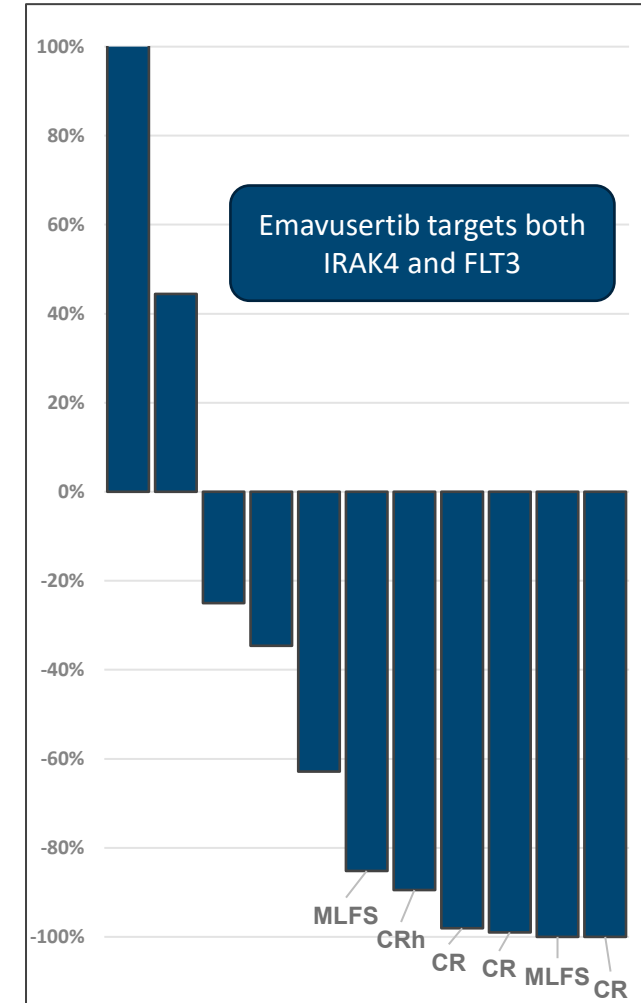
Significant resources will be required to execute a large clinical study and prepare for potential commercial launch

# Single-agent activity demonstrated in AML

**Total AML Patients, all dose levels**  
 nearly all AML patients express the oncogenic isoform of IRAK4



**Subset of AML Patients treated at 300mg BID who also have FLT3 mutation**



Data include all R/R AML patients determined to be evaluable for objective response using baseline and post-treatment marrow assessments as of Feb 26, 2024

# Strategy in AML

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## Demonstrate safety

123 patients<sup>1</sup> treated in TakeAim AML Ph 1/2 study, acceptable safety profile established

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## Demonstrate single-agent activity

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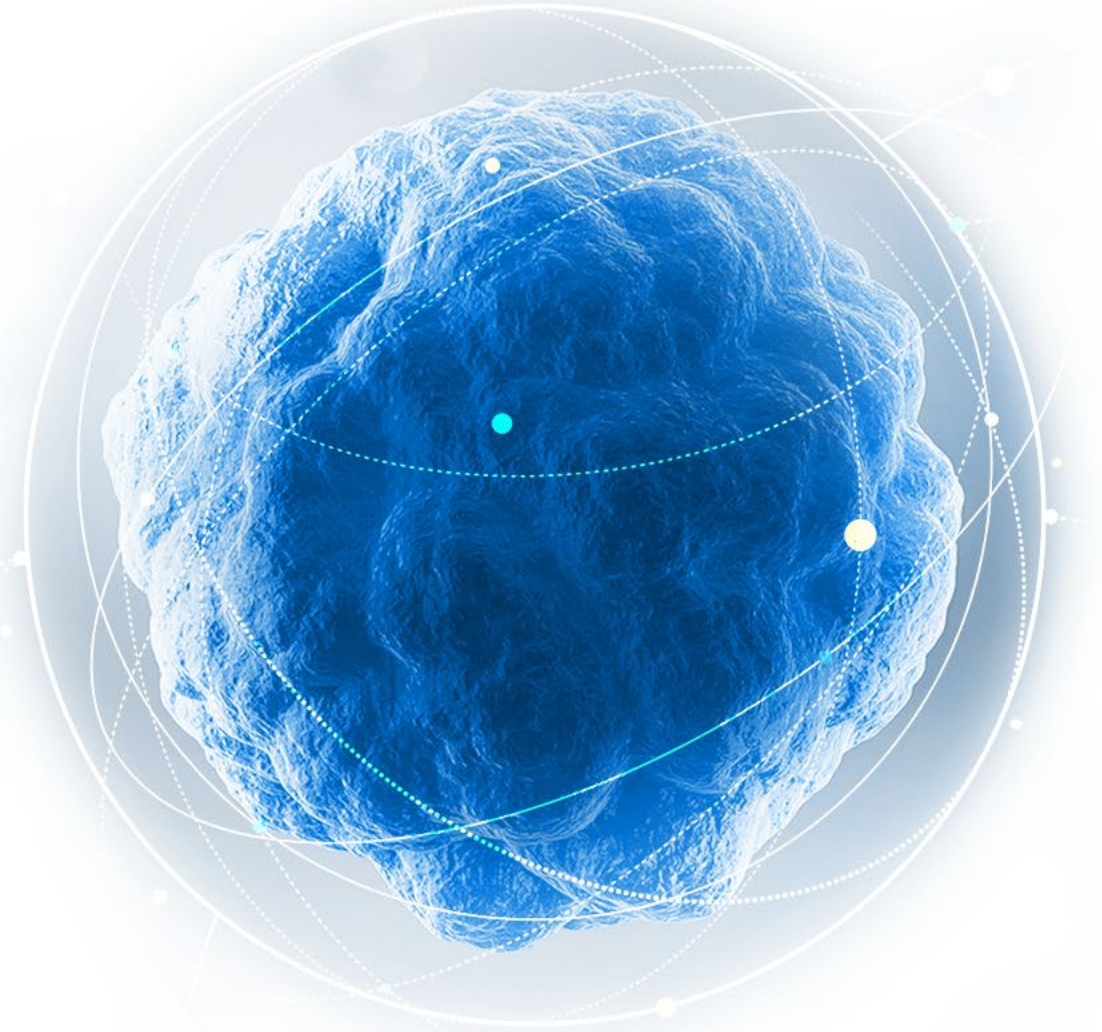
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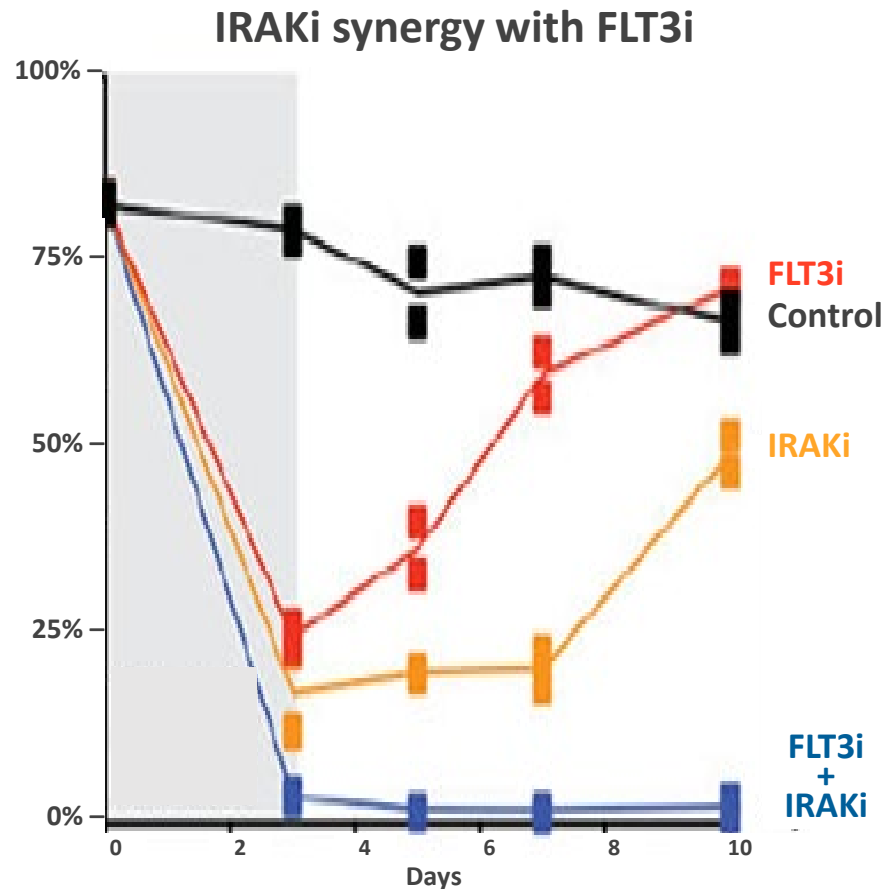
Significant resources will be required to execute a large clinical study and prepare for potential commercial launch



# Emavusertib in FLT3m AML



# Emavusertib's dual-targeting of IRAK4 and FLT3 enables monotherapy opportunity in FLT3m AML



**IRAK4 inhibition  
overcomes adaptive resistance  
to FLT3i**

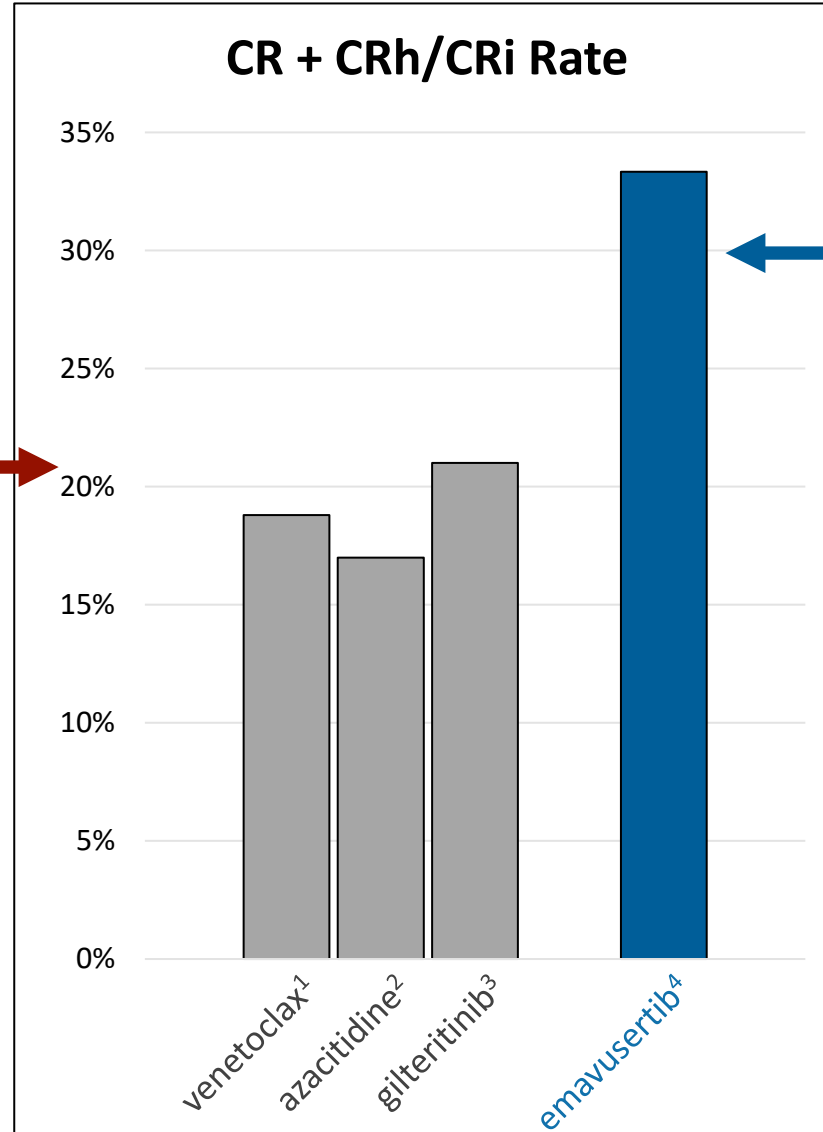
*Concomitant targeting of IRAK1 or IRAK4, alongside FLT3, is the most effective means to overcome the adaptive resistance incurred when targeting FLT3<sup>1</sup>*

Percent viable cells in preclinical AML cell lines (FLT3-ITD) treated for 72 hrs  
<sup>1</sup> Melgar Sci Transl Med 2019

# Emavusertib is a potential best-in-class therapy in FLT3m AML

Benchmark in FLT3i-naïve patients is **21% CR/CRh rate**

*87% of patients in the benchmark study were FLT3i naïve<sup>3</sup>*



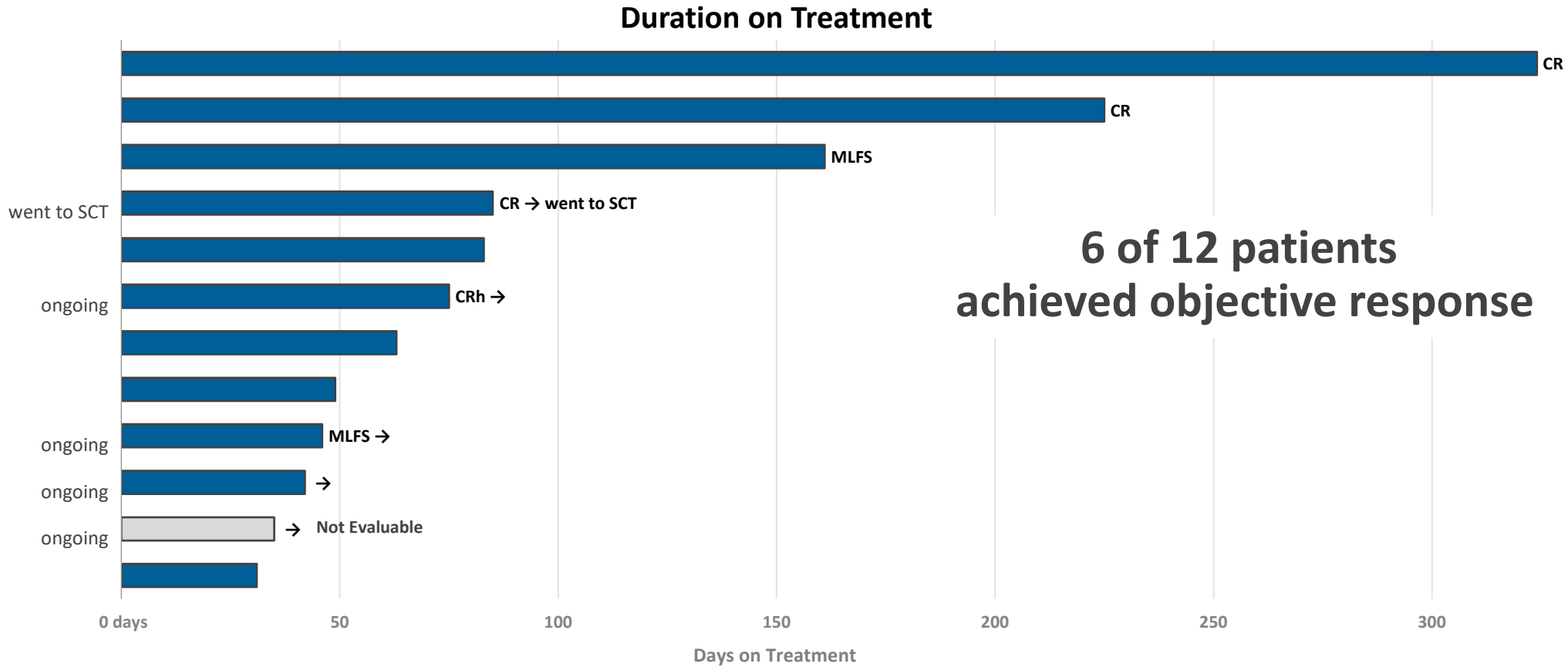
Salvage Line Patients treated with emavusertib achieved **> 30% CR/CRh rate**

~1.5X greater than the benchmark for FLT3i-naïve patients

*9 of 12 patients treated with emavusertib were FLT3i experienced (and had progressed on that prior FLT3i)*

1) Konopleva Cancer Discov 2016 [CR/CRi], 2) Itzykson Leuk Res 2015 [CR/CRi], 3) gilteritinib USPI [CR/CRh]; 4) emavusertib [CR/CRh]

# FLT3m AML data presented at ASCO/EHA 2024



*Data include all patients in target population (R/R AML patients with FLT3 mutation and < 3 prior lines of therapy) treated with 300 mg BID as of Feb 26, 2024; 1 patient w/CR and 1 patient w/MLFS had dual FLT3 and SF mutation  
 → Denotes ongoing with treatment*

# Strategy in AML

1

## Demonstrate safety

123 patients<sup>1</sup> treated in TakeAim AML Ph 1/2 study, acceptable safety profile established

2

## Demonstrate single-agent activity

Single-agent activity observed; next step is to confirm these initial findings in a larger number of patients

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## Pursue fastest path to 1<sup>st</sup> label in R/R patients

Address genetically-defined AML population with emavusertib's novel mechanism of action

4

## Explore frontline opportunity with combination

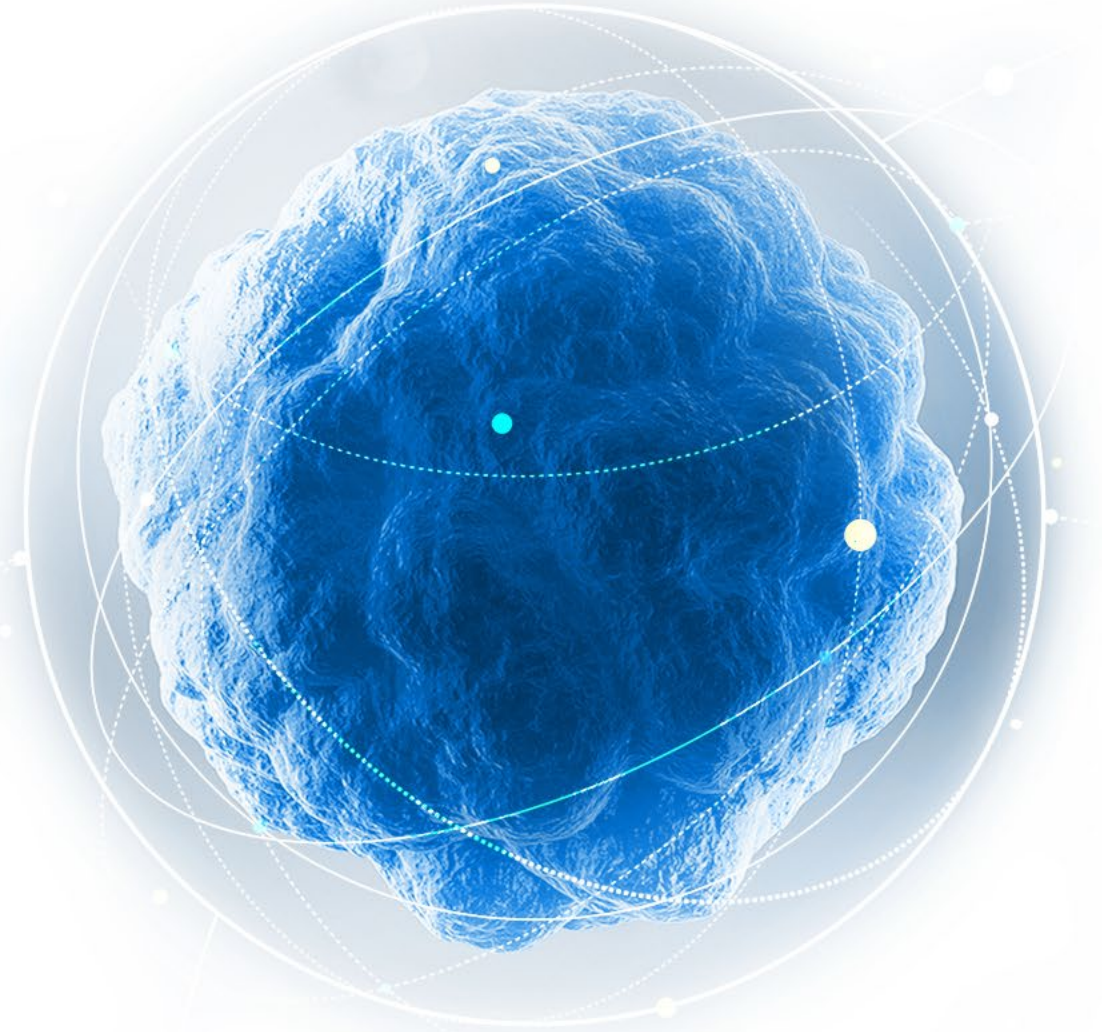
IRAK4-L is expressed in nearly all AML patients; preclinical "all comer" models suggest emavusertib is synergistic with azacitidine and venetoclax

5

## Pursue partnership to maximize potential commercial opportunity

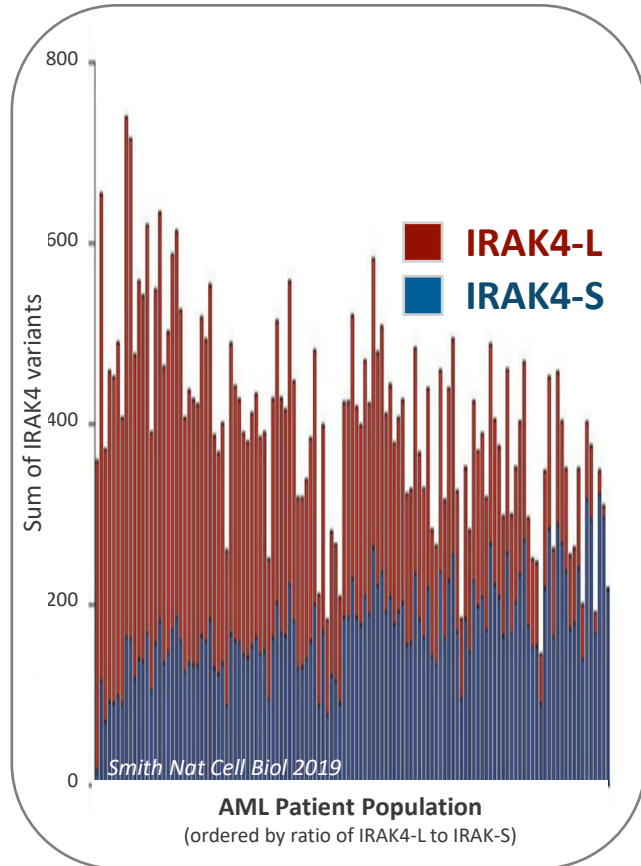
Significant resources will be required to execute a large clinical study and prepare for potential commercial launch

# Emavusertib in All Comers

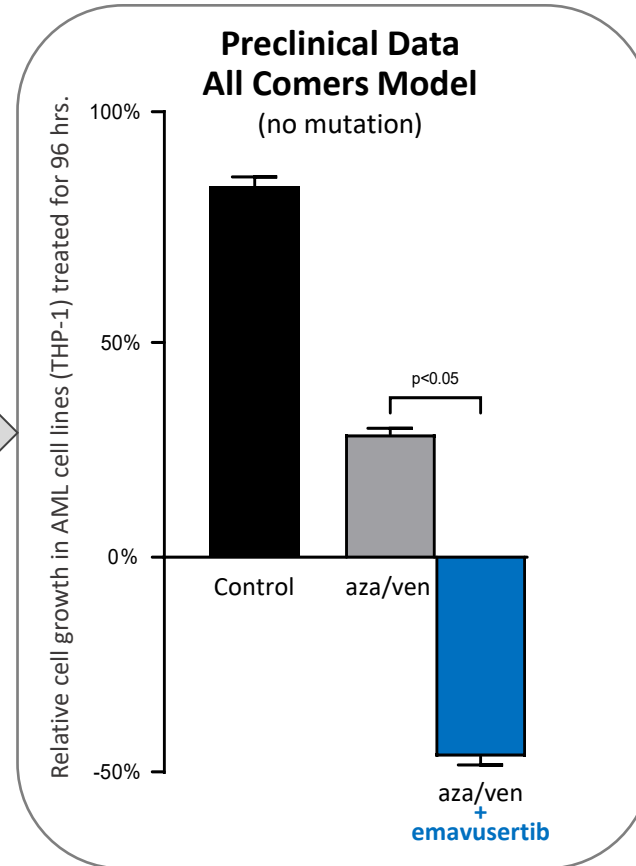


# Triplet Combination targets all comers in frontline AML

oncogenic IRAK4-L is expressed in nearly all AML patients



emavusertib synergy with aza/ven in preclinical studies



ema/aza/ven triplet combination

*Ph 1 study initiated 1H2024*  
*initial safety data expected late 2024*

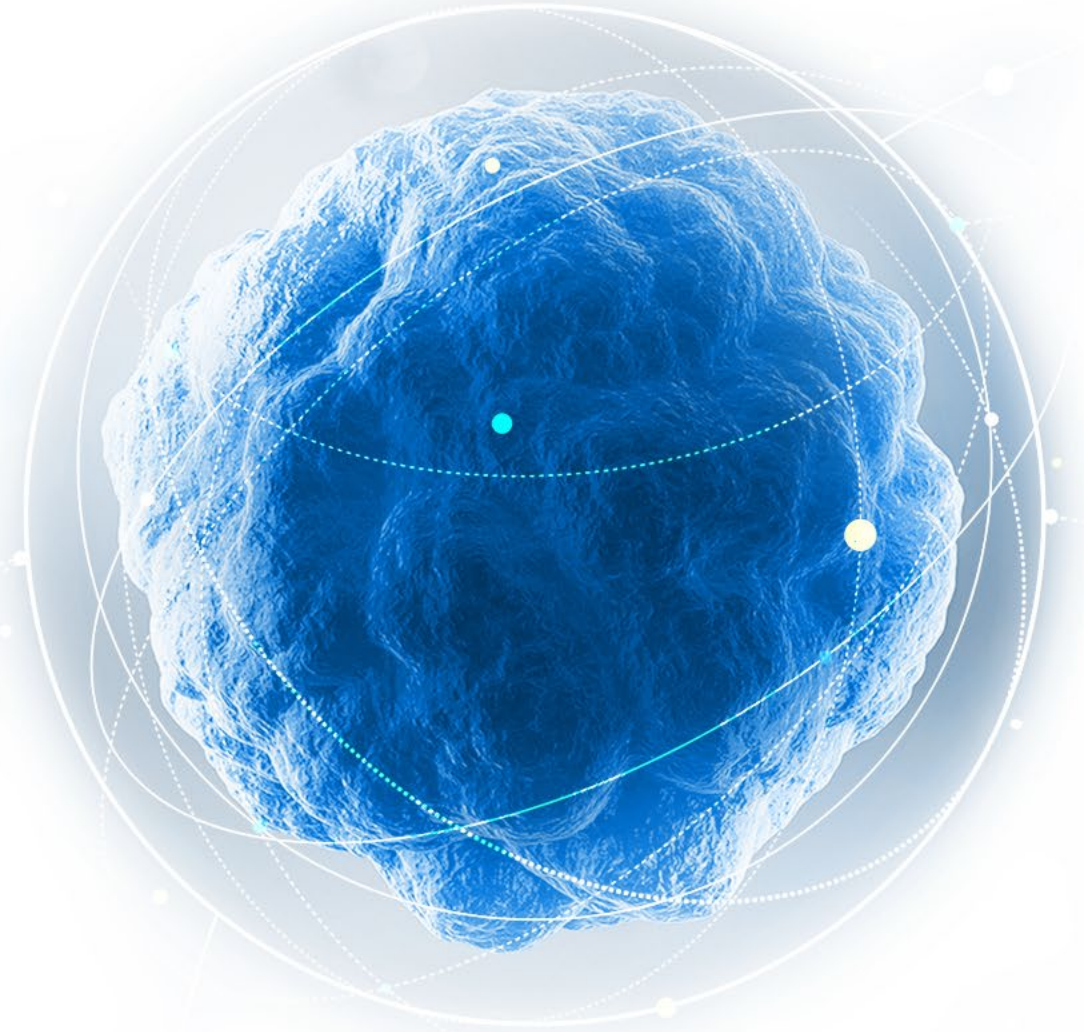


# Summary in AML



- Emavusertib targets both FLT3 and IRAK4
- Emavusertib offers potential for best-in-class therapeutic in FLT3m AML (a genetically-defined population)
- Oncogenic IRAK4 is expressed in nearly all AML patients and is not addressed by current standard-of-care (azacitidine and venetoclax)
- Emavusertib, in combination with azacitidine and venetoclax, offers potential for broad commercial opportunity in frontline AML

# Solid Tumors



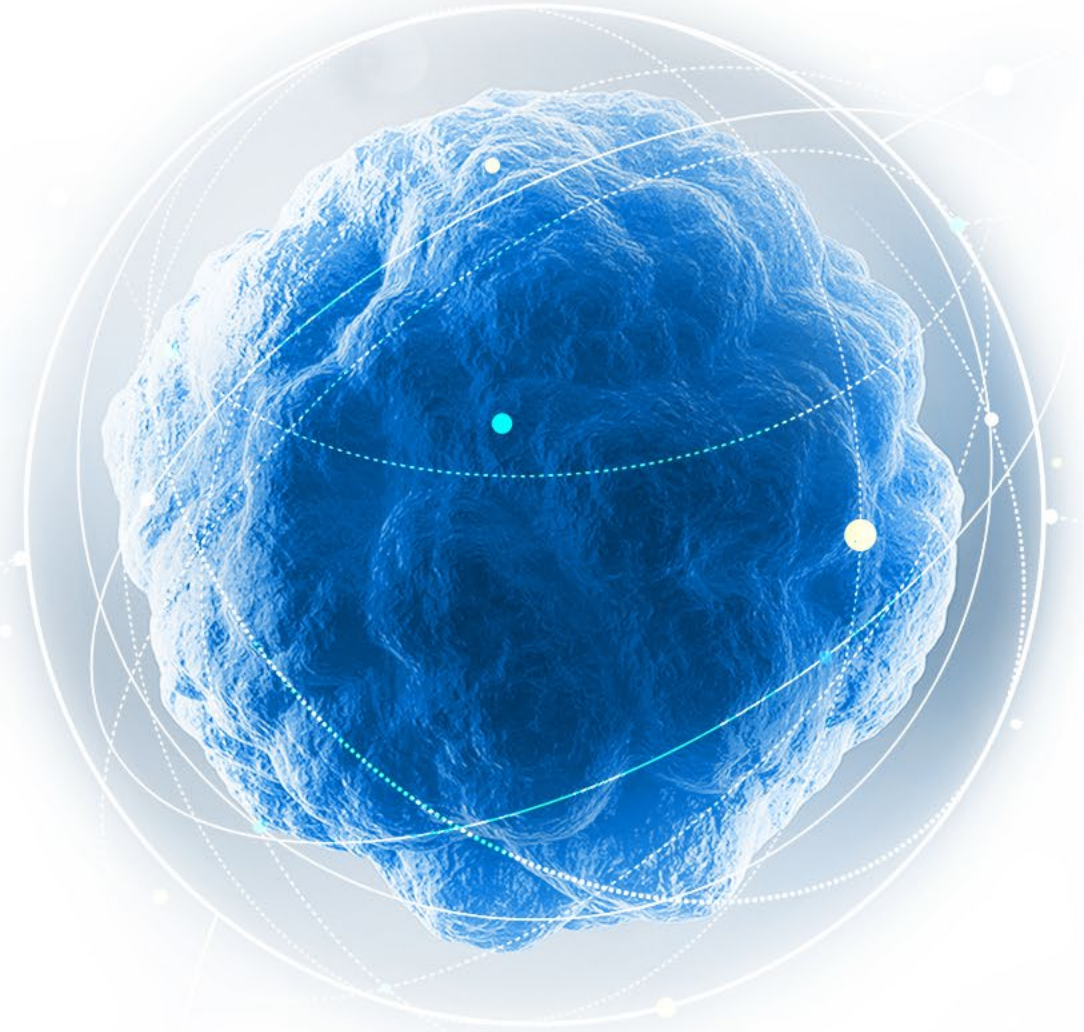
# Emavusertib is being evaluated in several solid tumor Investigator Sponsored Trials (ISTs)

Lead Investigator	Indication	Combination	Sponsor
Washington Univ St Louis – Grierson	Pancreatic	gemcitabine/nab-paclitaxel	NCI
Washington Univ St Louis – Grierson	Gastro-Esophageal	FOLFOX/PD1 +/- trastuzumab	Wash Univ St L
Univ of Florida – Doonan	Melanoma Brain Metastases	pembrolizumab	Univ of Florida*
Icahn School of Medicine at Mount Sinai – Galasky	Urothelial	pembrolizumab	NCI
Stephenson Cancer Center at OU Health – Ulahannan	Colorectal	FOLFOX/bevacizumab	NCI

\* Financial support is provided by Merck

Preliminary data are expected beginning in 2025; however, as these are ISTs, Curis does not control the timing of patient enrollment or data publication

# Other Information



# Financials and IP

## As of June 30, 2024

\$28.4M	Cash and investments <sup>1</sup>
~5.9M	Shares Outstanding
~7.0M	Shares Fully Diluted

2035 Composition of Matter IP on emavusertib  
(before extension)

***We believe cash is sufficient to achieve anticipated milestones***

- *Updated PCNSL data 15-20 patients (late 2024)*
- *AML triplet safety data (late 2024)*

<sup>1</sup> estimated cash runway into 2025