Corporate Presentation

October 2024



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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. 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

- Data in ~20 pts in R/R FLT3m AML (ASH 2024)
- Data in ~15-20 pts with at least 1 post treatment scan in R/R PCNSL (Q1 2025)
- o Initial safety data for triplet combination in frontline AML (Q1 2025)

Demonstrated safety and single-agent activity

Demonstrated synergy with BTKi, HMA, BCL2i

Broad opportunity in NHL, AML, and Solid Tumors

Abbreviations: Interleukin 1 Receptor Associated Kinase 4 (IRAK4), Acute Myeloid Leukemia (AML), Non-Hodgkin Lymphoma (NHL), Primary Central Nervous System Lymphoma (PCNSL), Bruton's Tyrosine Kinase inhibitors (BTKi), Hypomethylating agents (HMA) and B-Cell Lymphoma 2 inhibitor (BCL2i)

Broad Market Opportunity in NHL, AML, and Solid Tumors

Significant market opportunities in current development programs

	PCNSL	FLT3m	AML
US Incidence per 100K	0.5 ¹	1.3 ²	4.2 ³
	Newly Diagnosed Per Year		
US	1,700 ¹	6,000 ²	20,000 ³
Big 5 Europe/Canada	1,800 ¹	5,200 ⁴	17,000 ⁴
Japan/China	<u>7,700</u> 1	<u>12,700⁴</u>	<u>41,200⁴</u>
Total	11,200	23,900	78,200

Additional opportunities with NHL expansion

WM	MCL	MZL	ABC-DLBCL	
0.5 ⁵	0.5 ⁶	1.5 ⁷	2.0 ⁸	
	Newly Diagnosed Per Year			
1,700 ⁵	1,700 ⁶	5,000 ⁷	6,800 ⁸	
1,800 ⁵	1,800 ⁶	5,500 ⁷	7,500 ⁸	
<u>7,700⁵</u>	<u>7,700⁶</u>	<u>23,000⁷</u>	<u>31,400⁸</u>	
11,200	11,200	33,500	45,700	

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

5 – Derived from incident rate in <u>https://rarediseases.org/rare-diseases/waldenstroms-macroglobulinemia/#affected</u> and 2022 country population [data.worldbank.org].

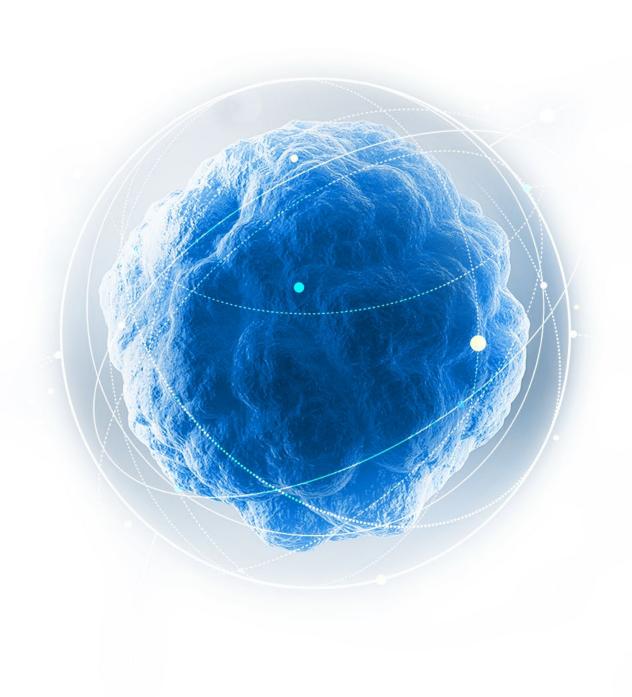
6 – Derived from incident rate in <u>https://www.ncbi.nlm.nih.gov/books/NBK536985/</u> and 2022 country population [data.worldbank.org].

7 – Derived from incident rate in Kalashnikov, Blood Cancer Journal, April 2023 and 2022 country population [data.worldbank.org].

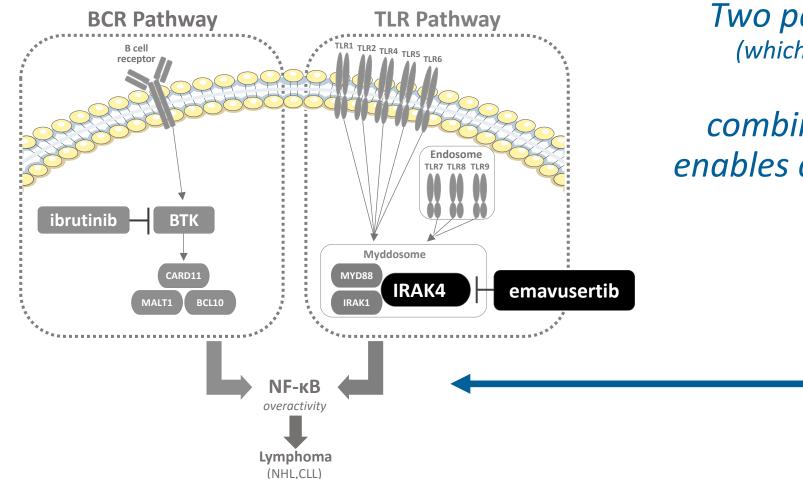
8 – Derived from incident rates in NHL incident rate of 18.6 per 100,000 (seer.cancer.gov) with DLBCL representing 25% of NHL per https://www.ncbi.nlm.nih.gov/books/NBK557796/. ABC represents 44% per letters to the editor, haematologica, 2011, 96 and 2022 country population [data.worldbank.org].

Emavusertib in NHL





Emavusertib's Mechanism in NHL



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

combining IRAK4i and BTKi enables dual blockade of NF-кВ

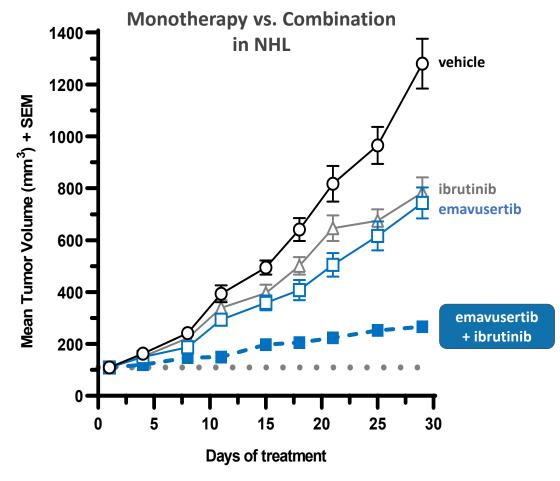


Emavusertib is synergistic with BTKi in NHL

emavusertib + BTKi

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

- IRAK4i synergizes with BTKi to promote killing of ABC-DLBCL¹
- Concurrent treatment with IRAKi and BTKi was significantly more potent in patient CLL cells than either drug alone²
- Data suggest IRAK4 as a novel treatment target for CLL; inhibition of IRAK4 blocks survival and proliferation of CLL cells³



Preclinical data for emavusertib and ibrutinib in OCI-Ly10 model (Booher et al., IWWM 2018)

Strategy in NHL



Demonstrate safety

31 patients¹ treated in TakeAim Lymphoma Ph 1b study, acceptable safety profile established, no overlapping dose-limiting toxicity with ibrutinib



Demonstrate single-agent activity

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



Pursue fastest path to 1st label in R/R patients

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



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 NHL¹

- 31 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 doselimiting CNS toxicities have been observed
- No dose-limiting myelosuppression has been observed

Grade 3+ TRAE in >1 Patient	100 mg BID Ema + Ibr (N=6)	200 mg BID Ema + Ibr (N=18)	300 mg BID Ema + Ibr (N=7)	Total (N=31)
	n (%)	n (%)	n (%)	n (%)
# patients having grade 3+ TRAEs	4 (67)	8 (44)	6 (86)	18 (58)
Lipase increased	2 (33)	1 (6)		3 (10)
Neutropenia	2 (33)	1 (6)		3 (10)
Platelet count decreased		2 (11)	1 (14)	3 (10)
Alanine aminotransferase increased		1 (6)	1 (14)	2 (6.5)
Amylase increased	2 (33)			2 (6.5)
Aspartate aminotransferase increased		1 (6)	1 (14)	2 (6.5)
Fatigue		1 (6)	1 (14)	2 (6.5)
Hyponatraemia		2 (11)		2 (6.5)

1 – As of July 10, 2024

Strategy in NHL



Demonstrate safety

31 patients¹ treated in TakeAim Lymphoma Ph 1b study, acceptable safety profile established, no overlapping dose-limiting toxicity with ibrutinib



Demonstrate single-agent activity

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



Pursue fastest path to 1st label in R/R patients

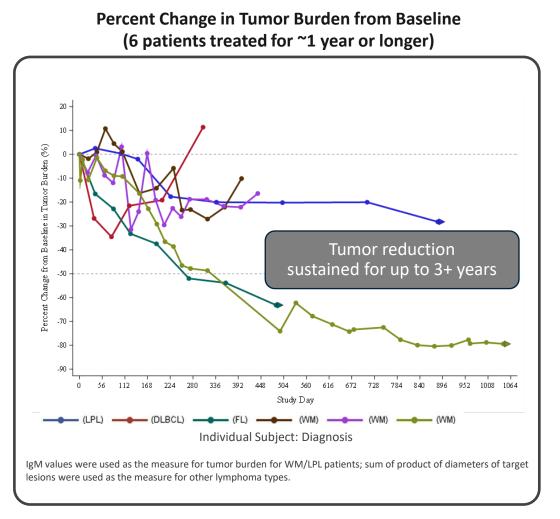
Identify orphan indication with clear unmet need that is addressable with emavusertib's novel mechanism of action

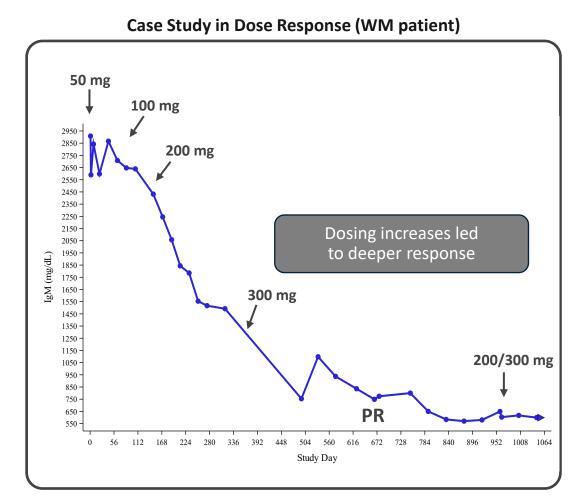


Pursue partnership to expand across NHL

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





2022 IWWM Conference Presentation

Strategy in NHL



Demonstrate safety

31 patients¹ treated in TakeAim Lymphoma Ph 1b study, acceptable safety profile established, no overlapping dose-limiting toxicity with ibrutinib



Demonstrate single-agent activity

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



Pursue fastest path to 1st label in R/R line patients

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



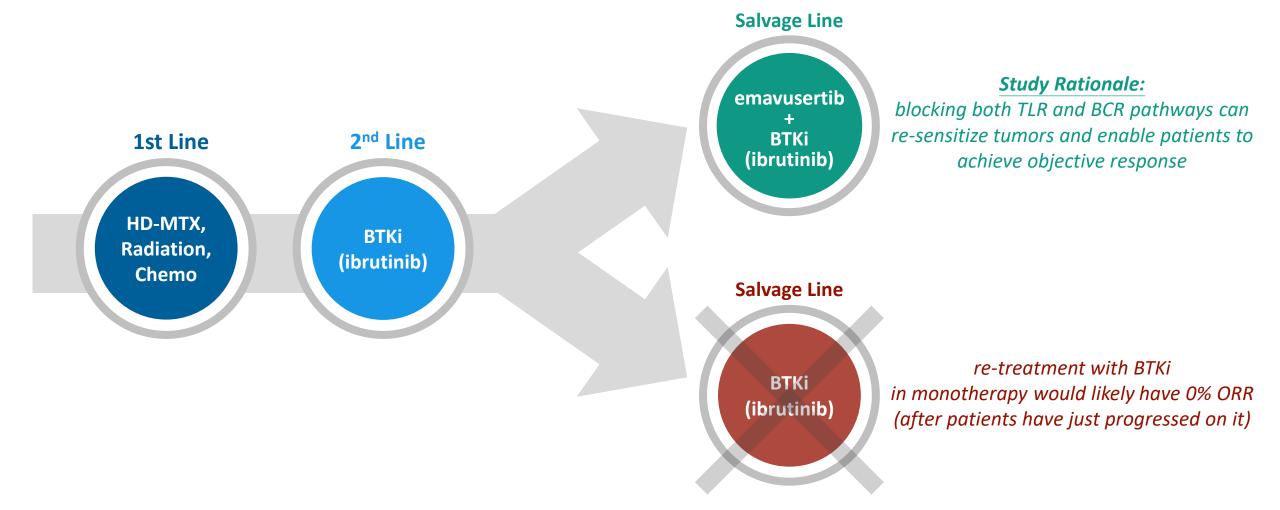
Pursue partnership to expand across NHL

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



R/R PCNSL selected for 1st NHL indication

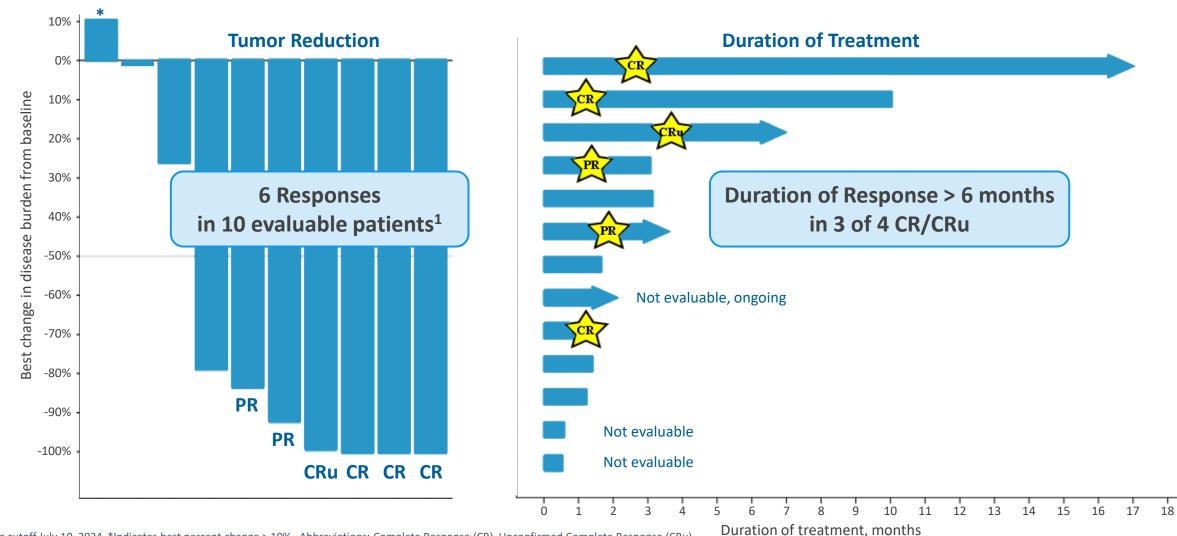
PCNSL is an orphan NHL indication, where no drugs are approved in salvage-line therapy





Encouraging clinical data in R/R PCNSL

Results for patients treated with emavusertib + ibrutinib, after they have progressed on prior BTKi



Data cutoff July 10, 2024. *Indicates best percent change > 10%. Abbreviations: Complete Response (CR), Unconfirmed Complete Response (CRu) ¹Evaluable patients are those who have completed at least one cycle of treatment and received at least one post-treatment assessment.

Strategy in NHL



Demonstrate safety

31 patients¹ treated in TakeAim Lymphoma Ph 1b study, acceptable safety profile established, no overlapping dose-limiting toxicity with ibrutinib



Demonstrate single-agent activity

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



Pursue fastest path to 1st label in R/R line patients

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



Pursue partnership to expand across NHL

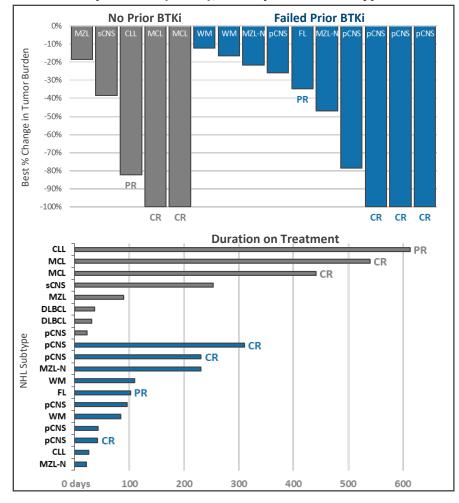
Significant resources will be required to execute clinical studies across multiple NHL subtypes and prepare for potential commercial launch

Anti-cancer activity shown across multiple NHL subtypes



Supports the mechanism of emavusertib + BTKi combination in additional NHL subtypes

- Heavily pre-treated patients (1-10 prior lines)
- Ongoing study with median treatment of 96 days (range 21-613 days)
- 7 of 19 patients achieved objective responses, including patients who failed prior BTKi
- 15 of 19 patients saw a reduction in tumor burden



All patients (n=19), multiple NHL subtypes



Opportunities in additional NHL subtypes

Emavusertib's inhibition of NF-kB via IRAK4, enables broad therapeutic potential in NHL

Tumor	(US only) Incidence	Key Targets of Interest	SOC
ABC-DLBCL	2 per 100,000 ~ 6,800 patients	IRAK4, MYD88, CD79, NF-kB	R-CHOP
PCNSL	0.5 per 100,000 ~ 1,700 patients	IRAK4, MYD88, CD79, NF-kB	Chemotherapy, HDMTX
WM	0.5 per 100,000 ~ 1,700 patients	IRAK4, MYD88, CD79, NF-kB	Chemotherapy, αCD20
MCL	0.5 per 100,000 ~ 1,700 patients	BCR and TLR pathway activation	BTKi, αCD20
MZL	1.5 per 100,000 ~ 5,000 patients	IRAK4, MYD88, CARD11, NF-kB	Chemotherapy, αCD20, RT
CLL	5 per 100,000 ~ 18,700 patients	NF-kB	BTKi, αCD20

Emavusertib's inhibition of IRAK4 suppresses NF-kB pathway activity, offering a therapeutic strategy for other NHL indications driven by NF-kB overactivity

Abbreviations: NF-kB, Nuclear factor-kB; IR; CLL, Chronic lymphocytic leukemia; NHL

Sources: 1. Vermaat, J. S., et al. (2019). MYD88 mutations identify a molecular subgroup of diffuse large B-cell lymphoma with an unfavorable prognosis. Haematologica, 105(2), 424–434 (<u>Link</u>); 2. Zhou, Y., et al (2019). Analysis of genomic alteration in primary central nervous system lymphoma and the expression of some related genes. Neoplasia, 20(10), 1059–1069. (<u>Link</u>); 3. Alcoceba, M., et al (2022). MYD88 mutations: Transforming the landscape of IGM monoclonal gammopathies. International Journal of Molecular Sciences, 23(10), 5570. (<u>Link</u>); 4. Shekhar, R., et al. (2021). Frequency of MYD88 L256P mutation and its correlation with clinico-hematological profile in mature B-cell neoplasm. Hematology/Oncology and Stem Cell Therapy, 14(3), 231–239 (<u>Link</u>); 5. Insusati-Beltran, G., et al. (2015). Significance of MYD88 L256P mutations attus in the subclassification of Low-Grade B-Cell Lymphoma/Leukemia. Archives of Pathology & Laboratory Medicine, 139(8), 1035–1041 (<u>Link</u>); 6. Shuai, W., et al. (2020). Clinicopathological characterization of chronic lymphocytic leukemia with MYD88 mutations: L265P and and non-L265P mutations are associated with different features. Blood Cancer Journal, 10(8) (<u>Link</u>); HILES://www.ncbi.nlm.nih.gov/books/NBK557796/. ABC represents 24% per letters to the editor, haematologica, 2011; Lv Ther Adv Hematol 2022; <u>https://arediseases.org/rarediseases/org/rarediseases/cal0-chronic-lymphocytic-leukemia</u> April 2023; <u>https://mww.ncbi.nlm.nih.gov/books/NBK557796</u>/. ABC represents 44% per letters to the editor, haematologica, 2011; Lv Ther Adv Hematol 2022; <u>https://arediseases.org/rarediseases/cal0-chronic-lymphocytic-leukemia</u>



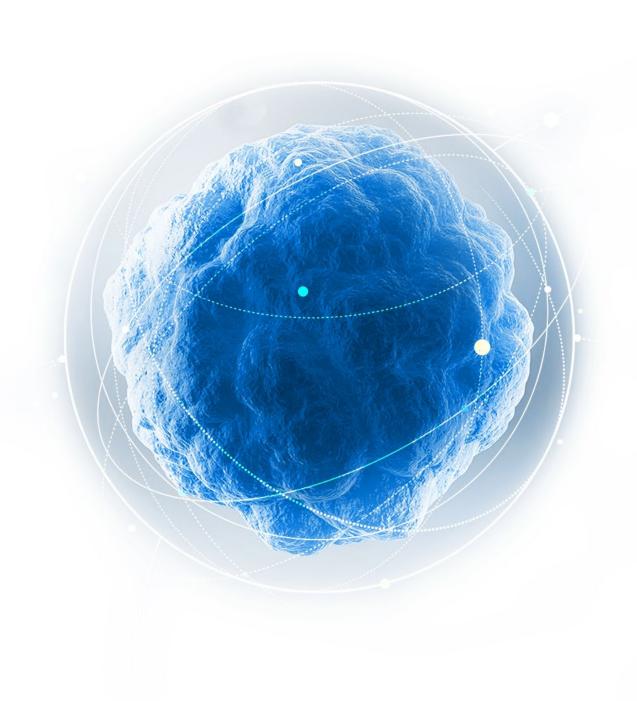
Summary in NHL



- Emavusertib continues to demonstrate clear anti-cancer activity in R/R PCNSL
- Next steps:
 - Work with FDA to explore the potential for an accelerated approval path
 - Prioritize next NHL indications (after PCNSL) that could benefit from the dual-blockade of NF-κB (blocking the TLR pathway with emavusertib and blocking the BCR pathway with a BTKi)

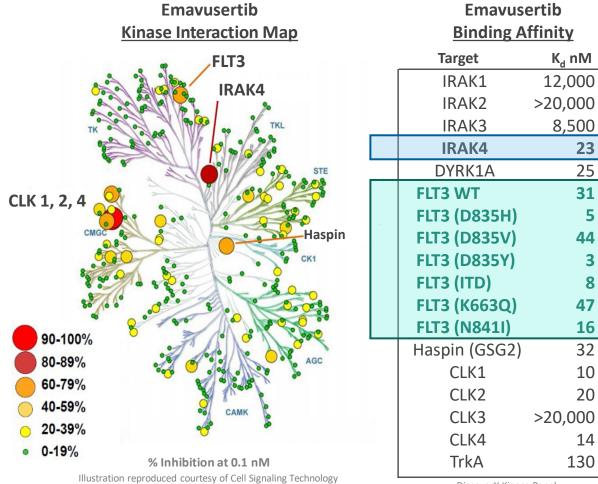
Emavusertib in AML







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



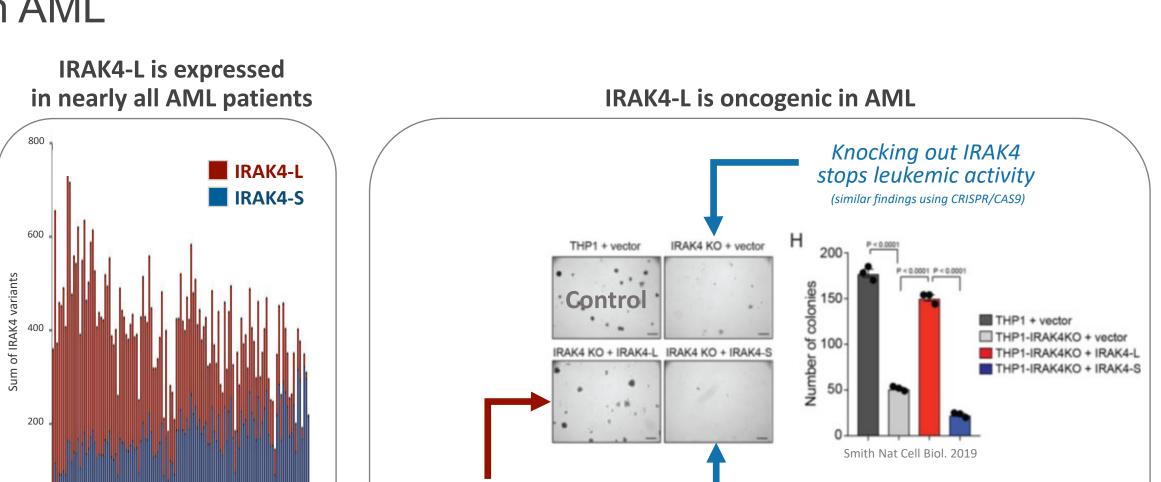
Binds tightly to IRAK4

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

DiscoverX Kinase Panel (378 kinases screened)

3

8



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

Smith Nat Cell Biol 2019

AML Patient Population

(ordered by ratio of IRAK4-L to IRAK-S)

0

adding back IRAK4-L

restarts activity

adding back IRAK4-S

has no effect

CURIS

Strategy in AML



Demonstrate safety

123 patients¹ treated in TakeAim AML Ph 1/2 study, acceptable safety profile established



Demonstrate single-agent activity

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



Pursue fastest path to 1st label in R/R patients

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



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



Pursue partnership to maximize potential commercial opportunity

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



Safety Profile of Emavusertib as Monotherapy in AML¹

- 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 ²	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



Demonstrate safety

123 patients¹ treated in TakeAim AML Ph 1/2 study, acceptable safety profile established



Demonstrate single-agent activity

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



Pursue fastest path to 1st label in R/R patients

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Explore frontline opportunity with combination

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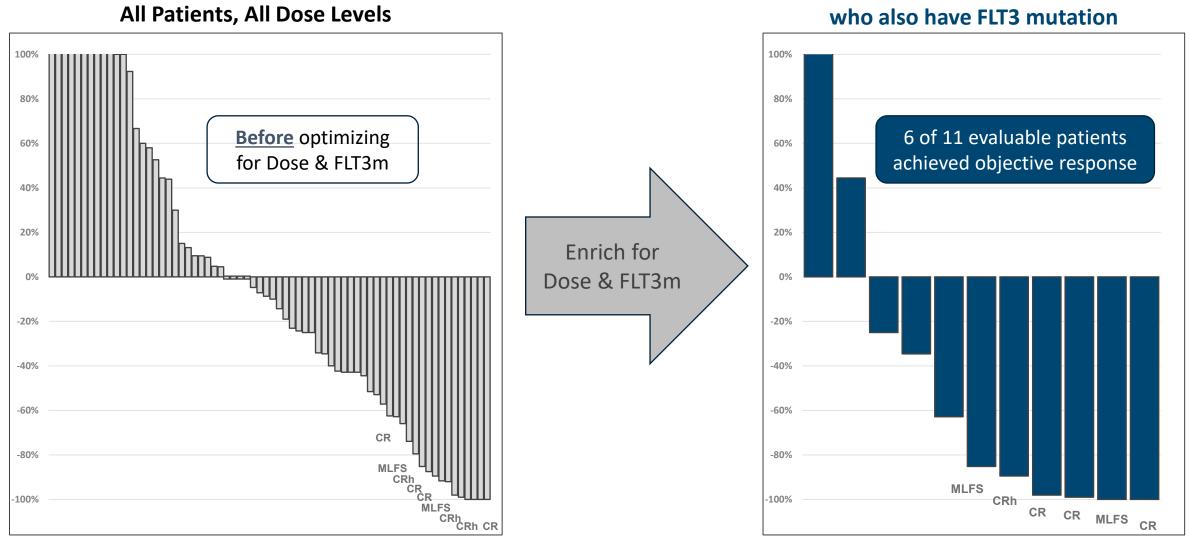


Pursue partnership to maximize potential commercial opportunity

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

Patients treated at 300mg BID

Single-agent activity demonstrated in AML



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



Demonstrate safety

123 patients¹ treated in TakeAim AML Ph 1/2 study, acceptable safety profile established



Demonstrate single-agent activity

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



Pursue fastest path to 1st label in R/R patients

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



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

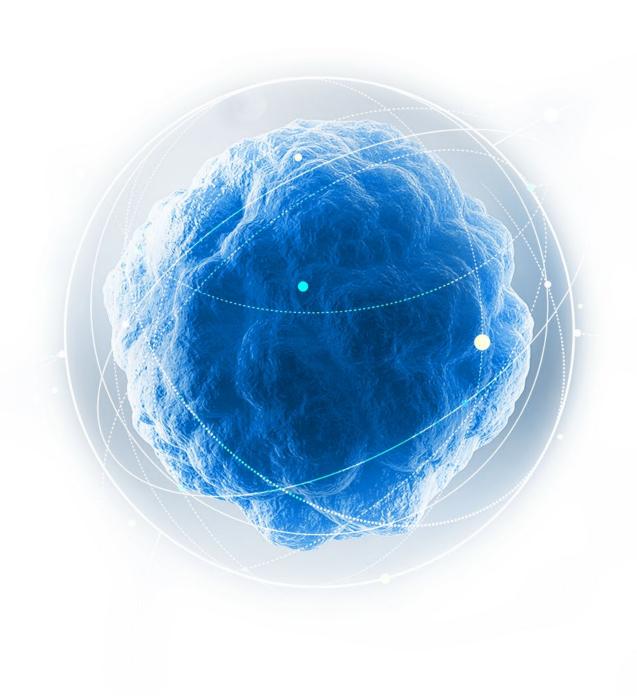


Pursue partnership to maximize potential commercial opportunity

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

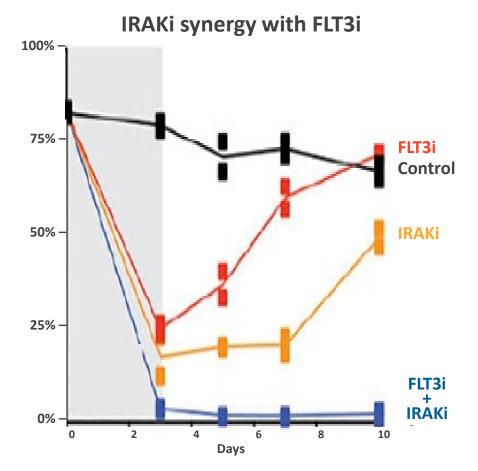
Emavusertib in FLT3m AML







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



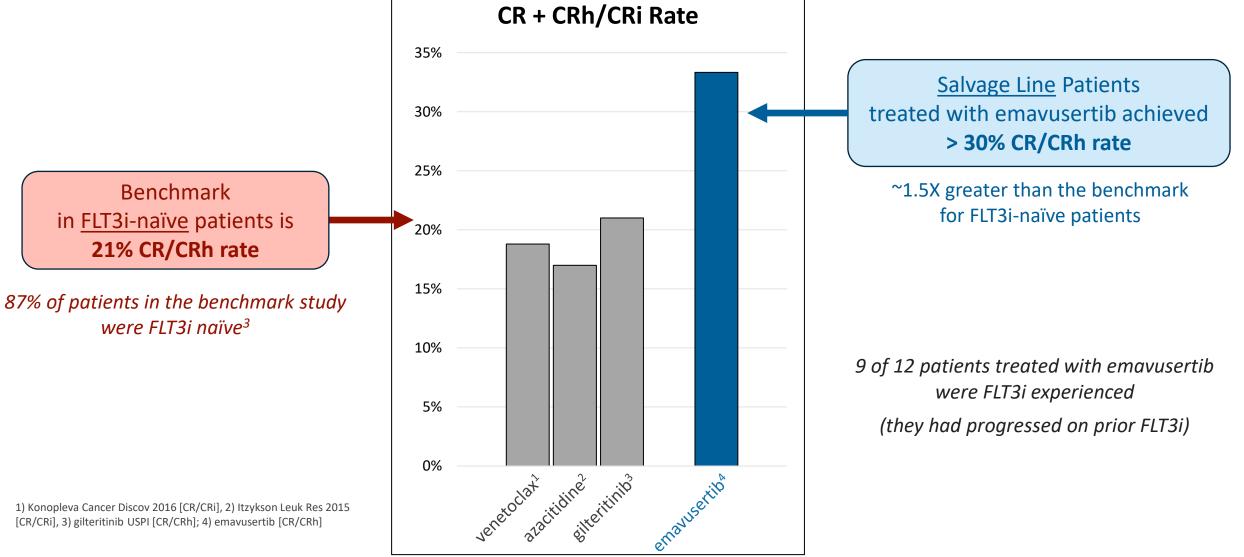
IRAK4 inhibition overcomes adaptive resistance to FLT3i

Concomitant targeting of IRAK1 or IRAK4, alongside FLT3, is the most effective means to overcome the adaptive resistance incurred when targeting FLT3¹

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

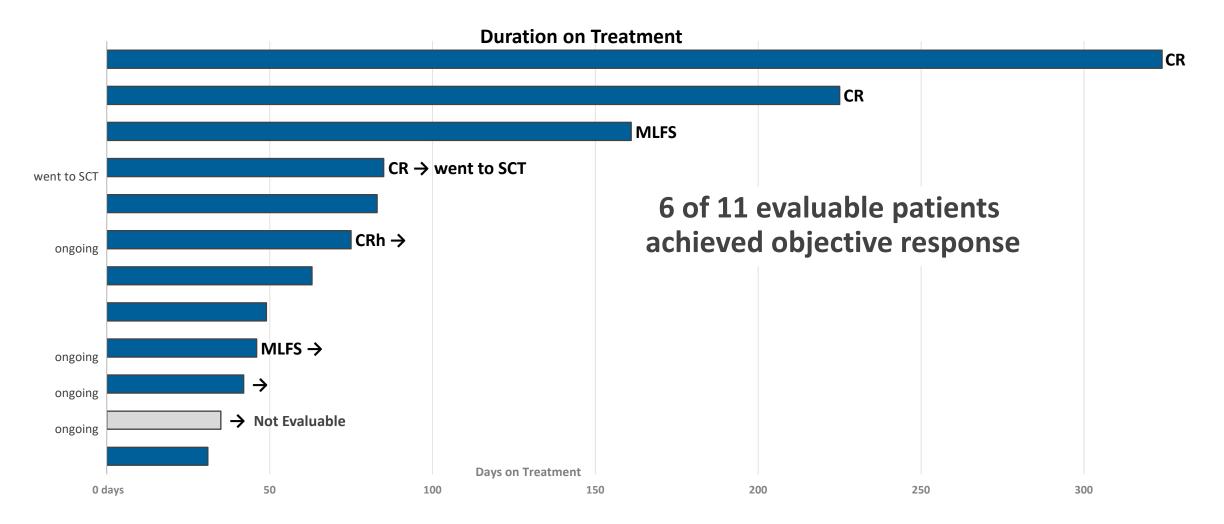


Emavusertib is a potential best-in-class therapy in FLT3m AML





Encouraging updated data in FLT3m AML 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

ightarrow Denotes ongoing with treatment

Strategy in AML



Demonstrate safety

123 patients¹ treated in TakeAim AML Ph 1/2 study, acceptable safety profile established



Demonstrate single-agent activity

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



Pursue fastest path to 1st label in R/R patients

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



Explore frontline opportunity with combination

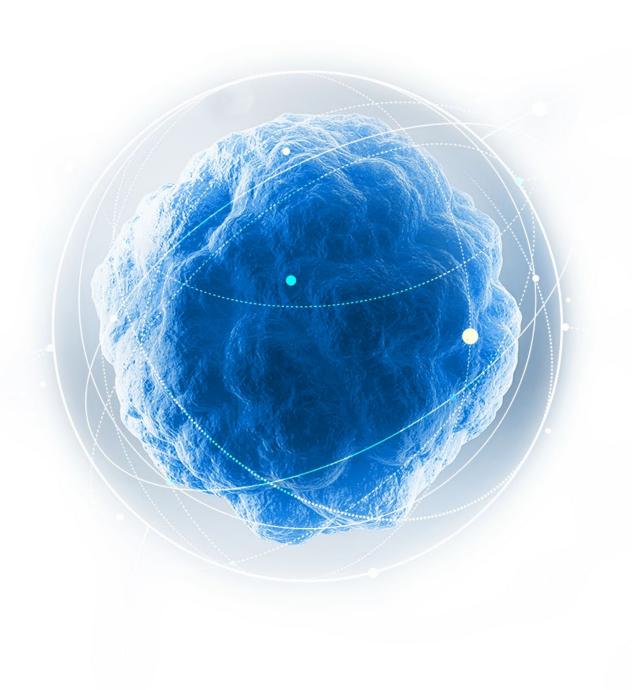
IRAK4-L is expressed in nearly all AML patients; preclinical "all comer" models suggest emavusertib is synergistic with azacitidine and venetoclax



Pursue partnership to maximize potential commercial opportunity

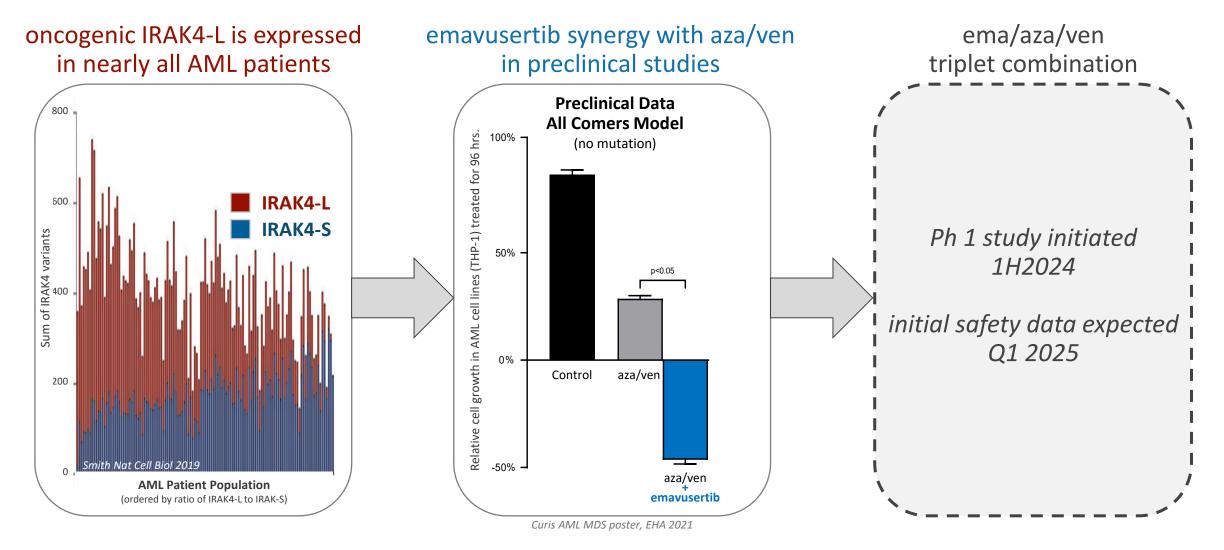
Significant resources will be required to execute a large clinical study and prepare for potential commercial launch

Emavusertib in All Comers





Emavusertib combination with aza/ven targets all comers in frontline AML



TLRIS



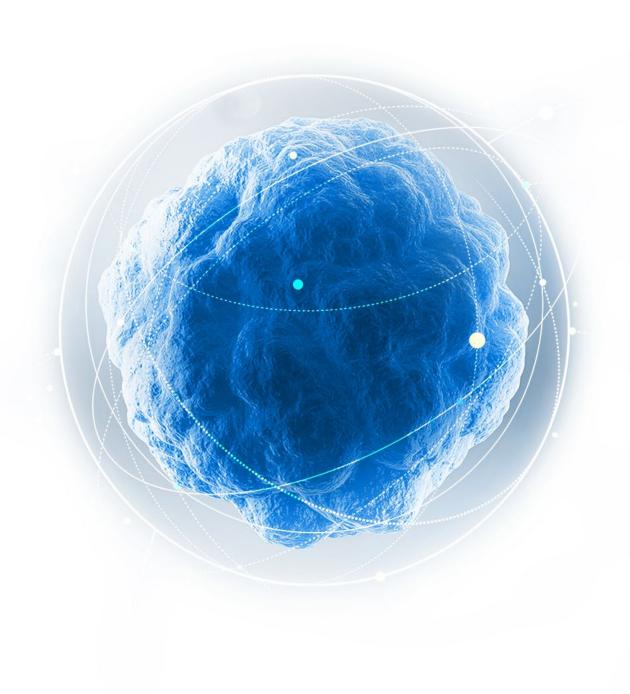
Summary in AML



- 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





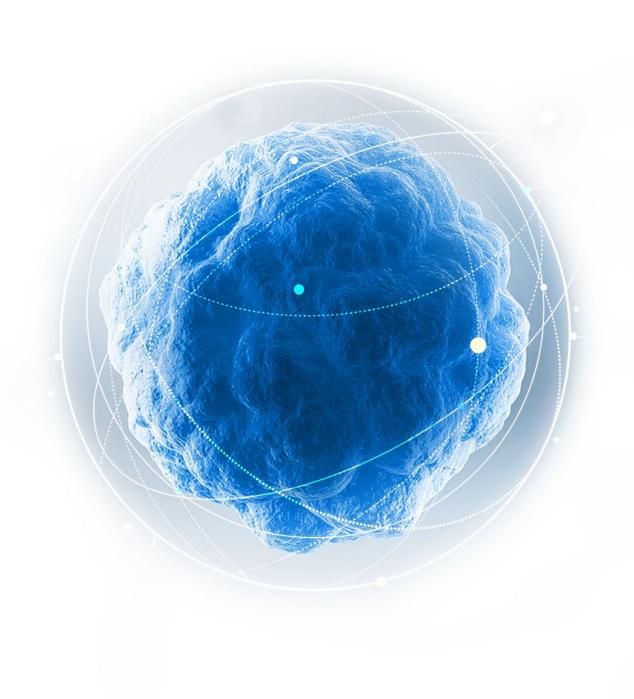


Ongoing studies (ISTs) of emavusertib in Solid Tumors

Tumor Type	Institution (Investigator)	Emavusertib Combination Partner
Pancreatic	CRADA Washington University (Grierson) Washington University (Lim)	gemcitabine, (nab)-paclitaxel
Colorectal	CRADA Oklahoma University (Ulahannan) Washington University (Lim)	FOLFOX + bevacizumab
Gastro/Esophageal	Washington University (Grierson)	FOLFOX/PD1 +/- trastuzumab
Melanoma	University of Florida (Doonan)	pembrolizumab
Urothelial	CRADA Mount Sinai (Galsky)	pembrolizumab

Other Information





Financials and IP



<u>As of June 30, 2024</u>

- \$28.4M Cash and investments¹
- ~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 near-term milestones

- Updated PCNSL data 15-20 patients (4Q24/1Q25)
- AML triplet safety data (4Q24/1Q25)