



# Decoding Biology To Radically Improve Lives

NOVEMBER 2024



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# Post-Combination portfolio poised for value creation from a unified, AI-powered Operating System

~10

Clinical and pre-clinical programs<sup>1</sup>

Oncology, Rare diseases, and other High Unmet Need Diseases

~10

Additional advanced discovery programs

10+

Partnered programs

Oncology, Immunology, and other High Unmet Need Diseases

4

Large pharma collaborations

~10

Clinical program milestones over the next 18 months<sup>2</sup>

1

Unified Operating System (OS) with both First & Best-in-Class capabilities

~\$450M

Upfront and milestone payments earned to-date

~\$20B potential milestone payments



sanofi



Merck KGaA  
Darmstadt, Germany

<sup>3</sup> 1. Includes preclinical programs (programs expected to enter the clinic within the next 18 months).  
2. Program milestones includes data readouts, preliminary data updates, regulatory submissions, trial initiation, etc.




# Pipeline of ~10 clinical and preclinical technology-enabled programs

	Candidate	Target	Indication	Preclinical	IND-Enabling	Phase 1/2	Pivotal / Phase 3	Next Anticipated Milestone
ONCOLOGY	REC-617 <sup>1</sup>	<b>CDK7</b>	Advanced Solid Tumors <sup>2</sup>	ELUCIDATE				• Initial Ph1 monotherapy safety & PK / PD data expected on Dec 9th, 2024
	REC-1245	<b>RBM39</b>	Biomarker-Enriched Solid Tumors & Lymphoma	DAHLIA				• Ph 1 update on dose-escalation expected in H1, 2026
	REC-3565 <sup>3</sup>	<b>MALT1</b>	B-Cell Malignancies	EXCELERIZE				• Ph 1 FPD expected in Q1, 2025
	REC-4539 <sup>4</sup>	<b>LSD1</b>	Small-Cell Lung Cancer (SCLC)					• Ph 1 FPD expected in H1, 2025
RARE	REC-994	<b>Superoxide</b>	Cerebral Cavemous Malformations (CCM)	SYCAMORE				• Ph 2 data to be shared via a congress / publication / webinar in H1, 2025 • Regulatory update by H2, 2025
	REC-4881	<b>MEK1/2</b>	Familial Adenomatous Polyposis (FAP)	TUPELO				• Ph 2 safety / early efficacy data expected in H1, 2025
	REC-2282	<b>HDAC</b>	Neurofibromatosis Type 2 (NF2)	POPLAR				• PFS maturing – PFS6 futility analysis anticipated in H1, 2025
	REV102 <sup>5</sup>	<b>ENPP1</b>	Hypophosphatasia (HPP)					• Development candidate nomination expected in Q4, 2024
OTHER	REC-3964	<b>TcdB</b>	Prevention of Recurrent <i>C. difficile</i> (rCDI)	ALDER				• Ph 2 update expected in Q1, 2026
	REC-4209	<b>Undisclosed</b>	Idiopathic Pulmonary Fibrosis (IPF)					• IND-enabling studies ongoing
	~10 advanced discovery programs							

REC-4881 in APC/AXIN1 indications has been deprioritized as part of a disciplined, strategic portfolio prioritization as part of the integration

4 1. Formerly GTAEXS617 2. Includes non-small cell lung cancer (NSCLC), colorectal cancer, breast cancer, pancreatic cancer, ovarian cancer, head and neck cancer  
3. Formerly EXS73565 4. Formerly EXS74539 5. Joint venture with Rallybio

# Robust pipeline of partnered programs

Therapy Area (Partner)	Discovery	Hit-to-Lead	Lead Optimization	Preclinical	Clinical
Neuroscience  Sumitomo Pharma	5-HT2A / 5-HT7				
Neuroscience  Sumitomo Pharma	5-HT1A / 5-HT2A				
Immunology  Bristol Myers Squibb™	PKC-θ				
Immunology					
Other					
Oncology					
Oncology					
Other					
Oncology					
Immunology					
Immunology					
Immunology					
Immunology					
Oncology					
Oncology					
Other					
Immunology					
Oncology					
Oncology					
Oncology					

Partners



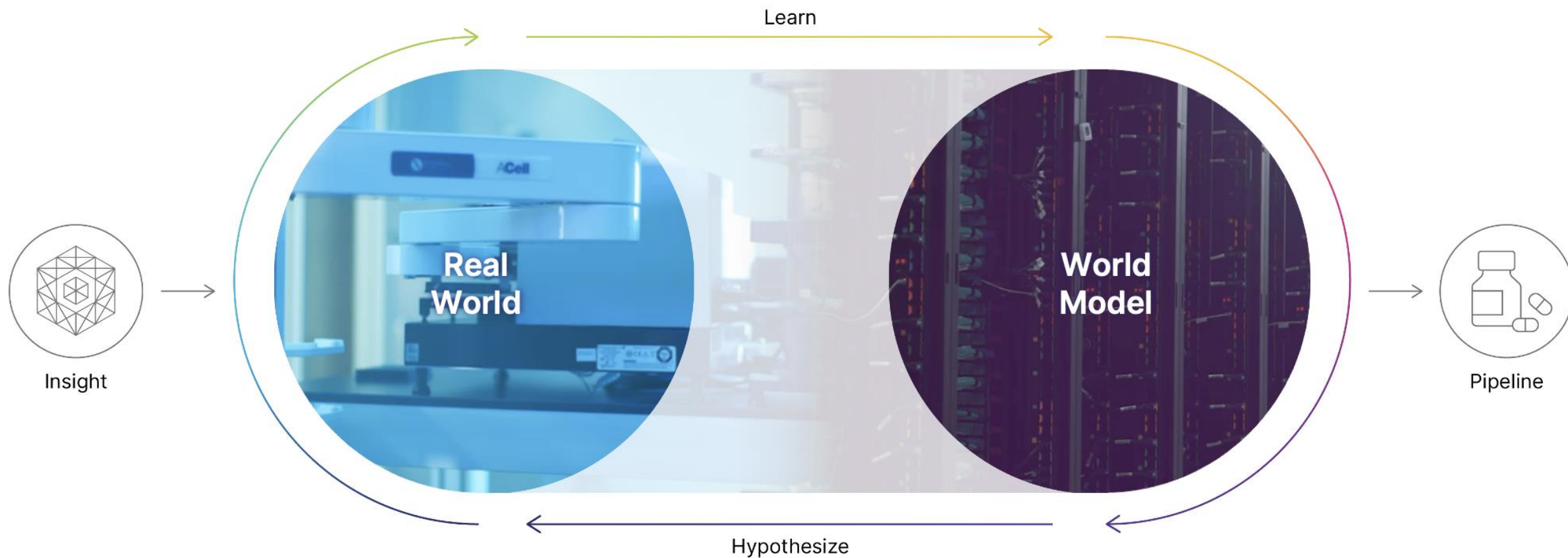


Merck KGaA  
Darmstadt, Germany

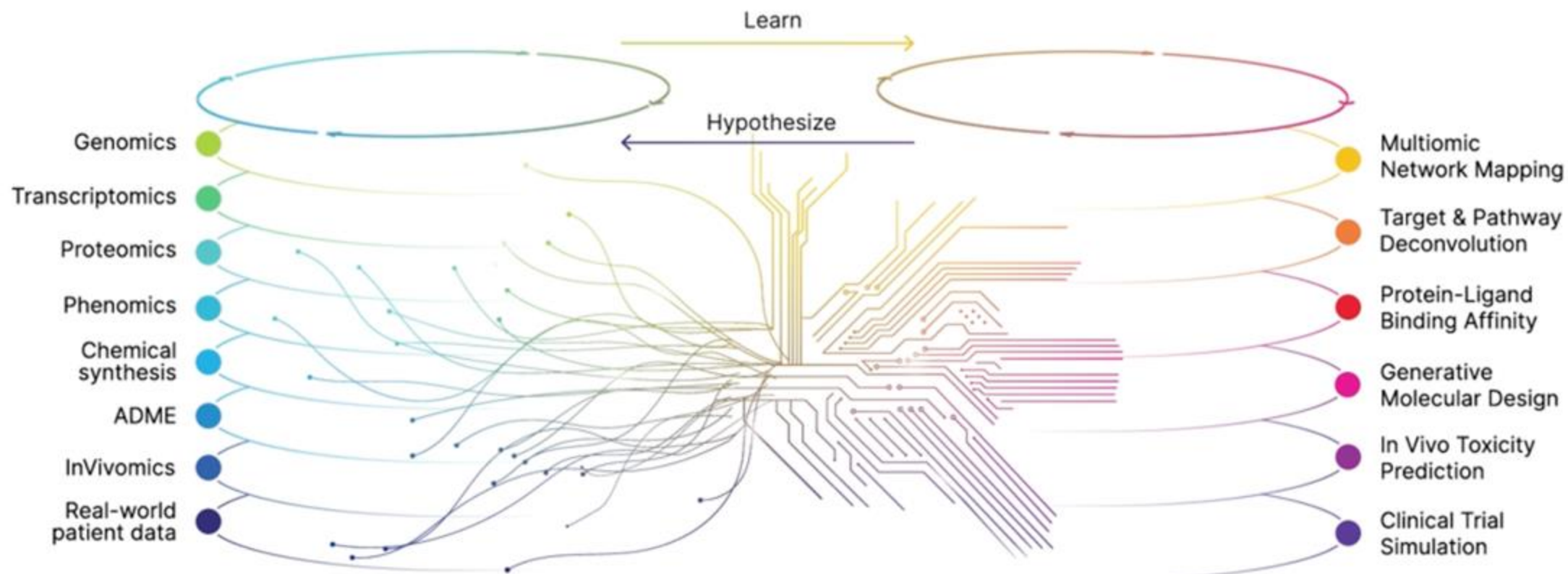
BILL & MELINDA\*  
GATES foundation

Note: Respective partner and disclosed target are noted for each clinical program  
\*Bill & Melinda Gates Foundation (BMGF) is a funder of anti-infective programs

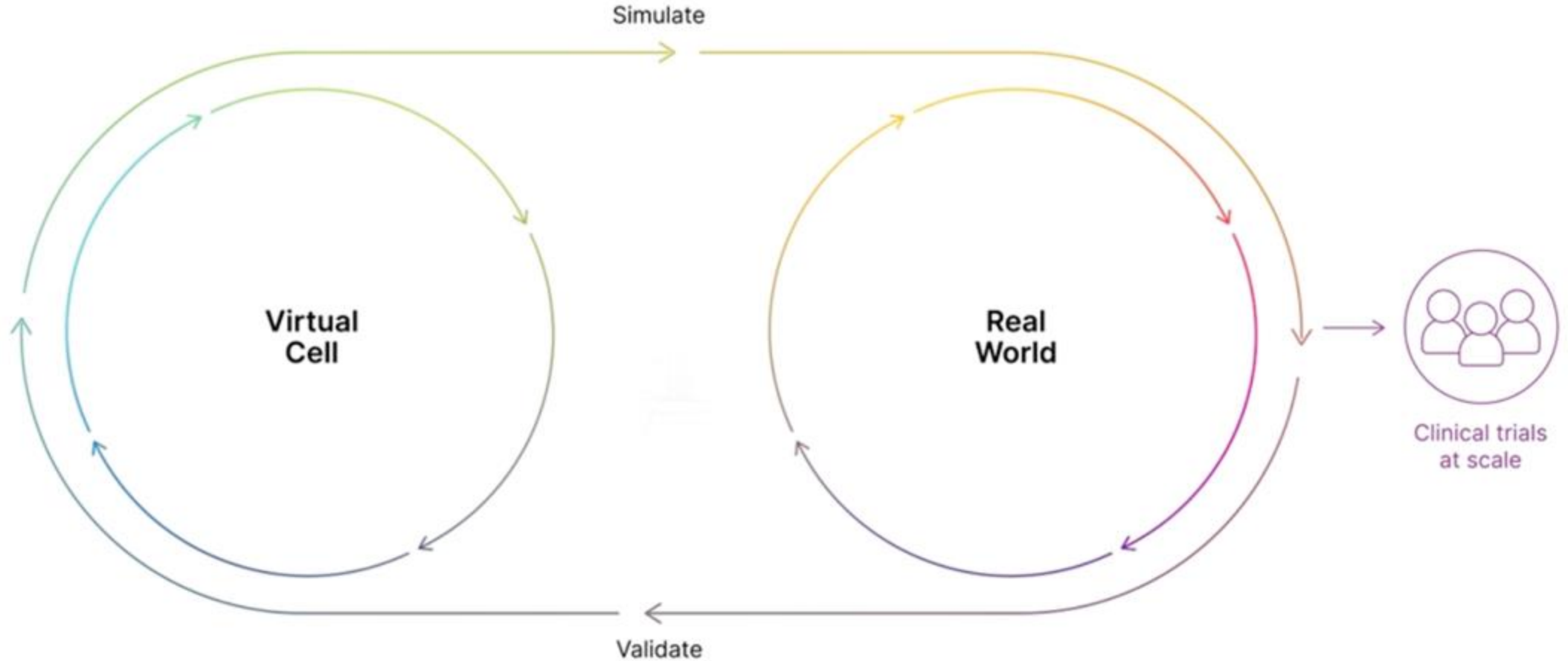
# Unified Recursion OS with First-in-Class & Best-in-Class capabilities



# Unified Recursion OS with First-in-Class & Best-in-Class capabilities



# Unified Recursion OS with First-in-Class & Best-in-Class capabilities





# We harness value from the Recursion OS with a multi-pronged capital efficient business strategy



## PIPELINE

### Pipeline strategy

Build internal pipeline in indications with potential for **advance transformational medicines for patients**

- Oncology
- Rare disease
- Other areas of high unmet need

## PARTNERSHIP

### Partnership strategy

Partner in **complex therapeutic areas** requiring large financial commitment or competitive arbitrage  
**Leverage partner knowledge** and clinical development capabilities

- Neuroscience
- Oncology
- Immunology
- Other large, intractable areas of biology

## DATA

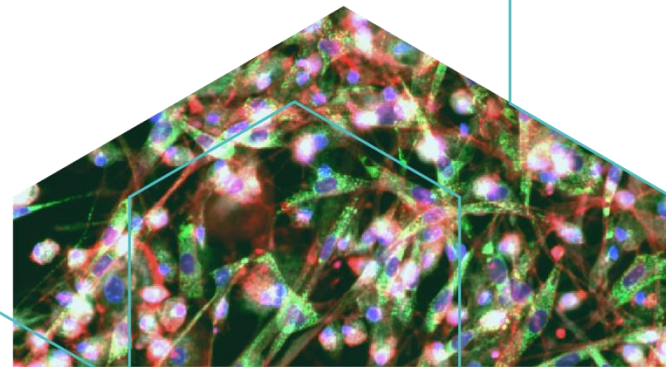
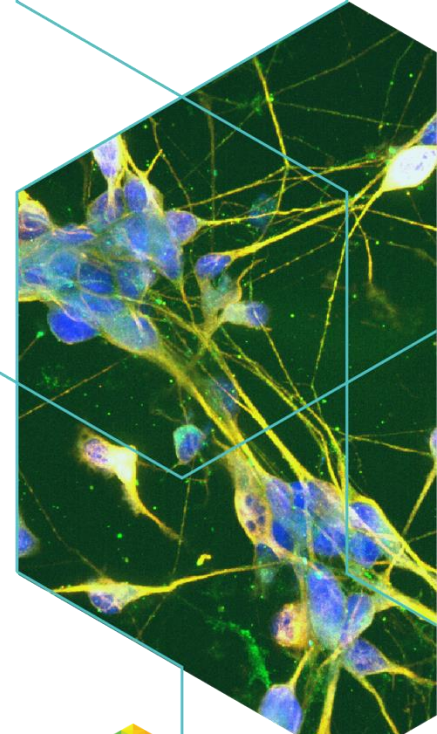
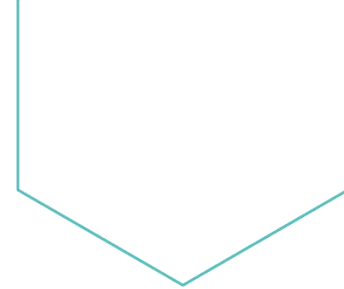
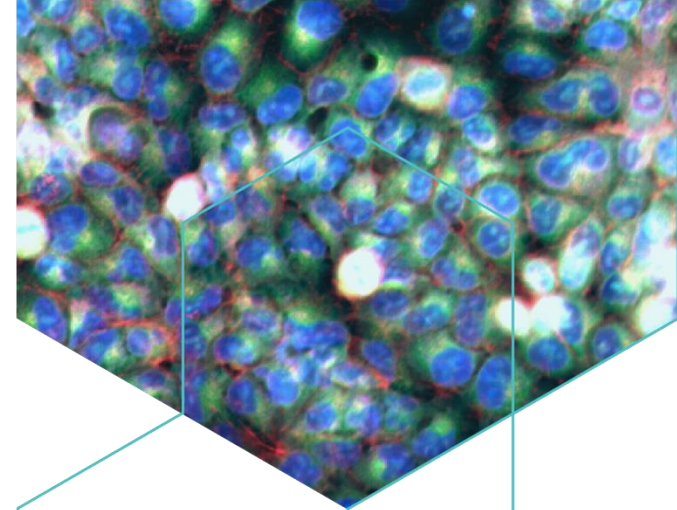
### Data strategy

**License** subsets of data and key tools  
Direct generation of new data internally **to maximize pipeline and partnership value-drivers**

- Licensing
- Augment Recursion OS
- LOWE

VALUE CREATION

# Pipeline



# We harness value from the Recursion OS with a multi-pronged capital efficient business strategy



## PIPELINE

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PIPELINE

# Oncology

# Advanced Solid Tumors (CDK7 Inhibitor): REC-617\*

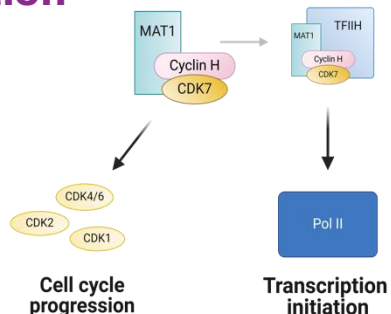
## Unmet Need

- **Aberrant CDK7** overexpression common in advanced **transcriptionally-addicted** solid tumors
- Potential to address **multiple indications**, including post CDK4/6 population patients

~185,000  
Treatable US + EU<sup>1</sup>

## Mechanism of Action

- **Reversible** CDK7 inhibitor
- **Dual function** that targets both cell cycle progression and transcriptional regulation



## Development Strategy

CURRENT STAGE

ELUCIDATE

**Phase 1**  
Monotherapy  
Dose Escalation

EXPECTED STAGES

Q1 2025

**Phase 1**  
Combo (SERD)  
Dose Escalation

H2 2025

**Phase 2**  
Combination  
Dose Expansion

## Differentiation

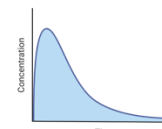
- Potential **Best-in-Class** and **First-in-Class** CDK7 Inhibitor
- Designed with **reduced transporter interactions** to **minimize GI adverse events** seen with competitor molecules



Noncovalent, Potent  
& Selective



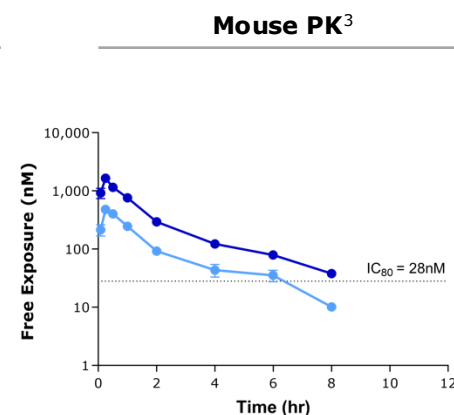
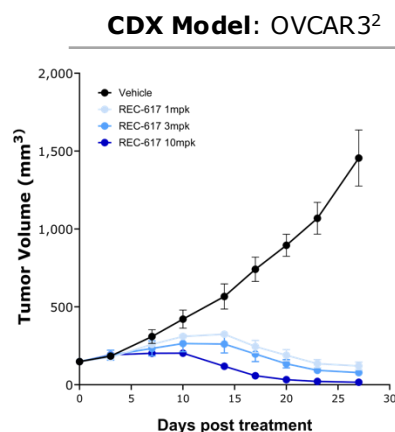
Shorter  
Half-Life



Rapid  
Absorption

## Key Preclinical Data

- REC-617 demonstrates **potent tumor regression** with <10 hours of exposure above IC<sub>80</sub> to **optimize benefit-risk**



## Recursion Approach

- **AI-powered precision design** to optimize PK/PD and **maximize potential therapeutic index**

136

Novel compounds  
synthesized to candidate ID

## What's Next

- Initial Phase 1 monotherapy safety, PK/PD update expected at **AACR Special Conference in Cancer Research on December 9th**

13 \* Formerly GTAEXS617

1. Advanced solid tumors including breast, NSCLC, ovarian, pancreatic, colorectal and head & neck. US and EU5 treatable incidence, 2022. 2. Besnard et al, AACR (2022).

3. PK studies conducted in CD1 mice, single-dose administration. >10 hr IC<sub>80</sub> results in significant body weight loss

# Solid Tumors & Lymphoma (RBM39 Degradar): REC-1245

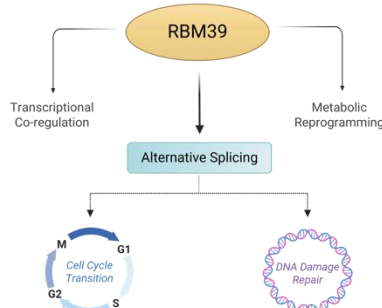
## Unmet Need

- Solid tumor and lymphoma patients experience disease progression while on frontline therapies
- Potential as a single agent or in combination with chemo/IO

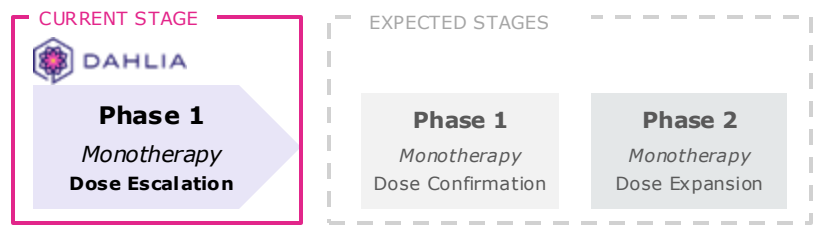
> 100,000  
Treatable US + EU<sup>1</sup>

## Mechanism of Action

- **Molecular glue** RBM39 degrader via E3 ligase adaptor DCAF15
- **Disrupts RNA splicing** to downregulate cell cycle checkpoints, DDR networks, triggering cell stress, apoptosis

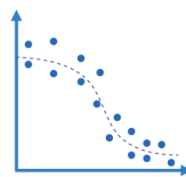


## Development Strategy



## Differentiation

- Potential **First-in-Class** RBM39 Degradar
- **No significant** in vitro safety concerns (hERG, CEREP)



Highly Potent & Selective



Minimal Off-Target Liabilities

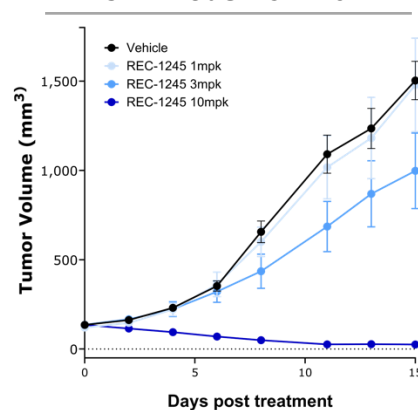


Biomarker Defined Population

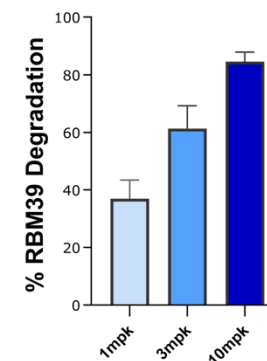
## Key Preclinical Data

- REC-1245 shows significant **monotherapy regressions**
- **Dose-dependent** anti-tumor activity correlates with PD

CDX Model: OVK18<sup>2</sup>



PD: Target Engagement<sup>3</sup>



## Recursion Approach

- Unbiased **ML-powered phenomap insight** to identify **novel DDR signature** and relate cellular phenotypes

204

Novel compounds synthesized to candidate ID

18 months

From Target ID to IND-Enabling studies

## What's Next

- Ph 1 initiation expected in **Q4 2024**
- Ph 1 update in dose-escalation expected in **H1 2026**

# B-Cell Malignancies (MALT1 Inhibitor): REC-3565\*

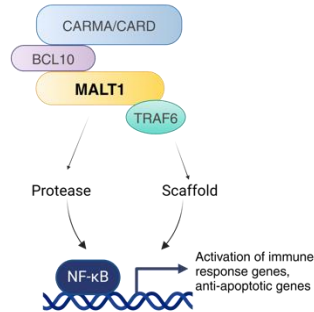
## Unmet Need

- Mutations causing **constitutive MALT1 protease activity** and MALT1-cIAP fusions are aggressive with **limited treatment options**
- Potential to enhance **NF-κB inhibition** with BTK inhibitors

~41,000  
Treatable US + EU<sup>1</sup>

## Mechanism of Action

- **Reversible** allosteric MALT1 inhibitor
- **Dampens NF-κB signaling** which drives survival and proliferation of B-cell tumors including ABC-DLBCL, MCL, FL, and CLL



## Development Strategy

Q1 2025

EXCELERIZE

**Phase 1**  
Monotherapy  
Dose Escalation

EXPECTED STAGES

**Phase 1**  
Combination  
Dose Escalation

**Phase 1**  
Combination  
Dose Expansion

## Differentiation

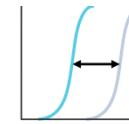
- Potential **Best-in-Class** MALT1 Inhibitor
- **Low UGT1A1** anticipated liability versus competitors
- **No significant off-target** safety concerns (CEREP, Kinome)



Lower Predicted  
Jaundice Risk



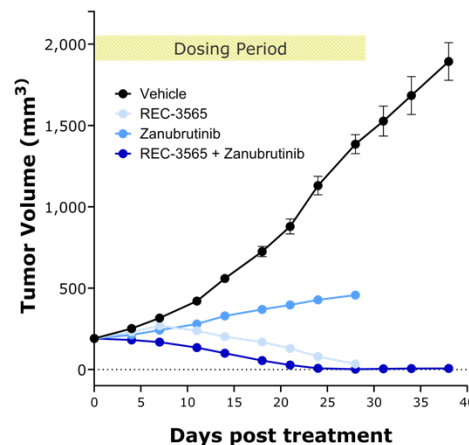
High Oral  
Bioavailability



Wider  
Therapeutic Index

## Key Preclinical Data

- REC-3565 monotherapy shows significant **tumor regression**
- **Sustained** anti-tumor activity **in combo** with zanubrutinib



CDX Model: OCI-Ly10<sup>2</sup>

**70%**

Of mice in combination arm (REC-3565 + zanu) had no palpable tumors 10-days post last dose

## Recursion Approach

- **AI-powered** precision designed **novel molecule** using molecular dynamics and hotspot analysis

**344**

Novel compounds synthesized to candidate ID

## What's Next

- **Phase 1 First Patient Dosed** in B-Cell Malignancies (e.g., chronic lymphocytic leukemia), expected **Q1 2025**

15 \* Formerly EXS73565.

1. Cerner Enviza Treatment Architecture Reports 2023, rounded to nearest 1,000 patients per year. 2. Payne et al. ENA, (2024)

# Small-Cell Lung Cancer (LSD1 Inhibitor): REC-4539\*

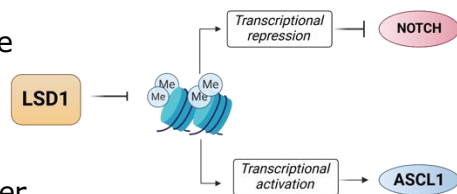
## Unmet Need

- **SCLC** is a highly progressive disease with **5-year OS ~3%** in the extensive stage
- Clinical trial enrollment **remains NCCN-recommended** after 1L chemo/IO, despite advancements with DLL3-targeting BiTEs<sup>2</sup>

**>45,000**  
Treatable US + EU<sup>1</sup>

## Mechanism of Action

- **Reversible** LSD1 inhibitor that can selectively upregulate NOTCH signaling
- **Promotes differentiation** of neuroendocrine cancer cells



## Development Strategy

H1 2025

**Phase 1**  
Monotherapy  
Dose Escalation

EXPECTED STAGES

**Phase 1**  
Combination  
Dose Escalation

**Phase 1**  
Combination  
Dose Expansion

## Differentiation

- Potential **Best-in-Class** LSD1 Inhibitor
- **Shorter-predicted half-life** plus **reversible MOA** to manage **on-target AEs**



Lower Predicted  
Thrombocytopenia



Shorter  
Half-Life

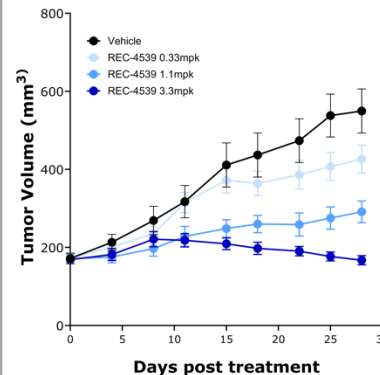


Optimal  
CNS Exposures

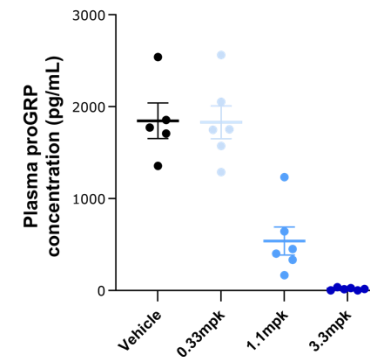
## Key Preclinical Data

- **Dose-dependent** efficacy in SCLC human xenograft model
- Well tolerated with limited impact on platelet levels

CDX Model: H1417<sup>3</sup>



Plasma ProGRP<sup>4</sup>



## Recursion Approach

- Precision design using **Active Learning**, combining reversibility with **CNS penetration**

**414**

Novel compounds  
synthesized to candidate ID

## What's Next

- Phase 1 **First Patient Dosed** in SCLC expected **H1 2025**



PIPELINE

# Rare disease

# Cerebral Cavernous Malformation (Superoxide Scavenger): REC-994

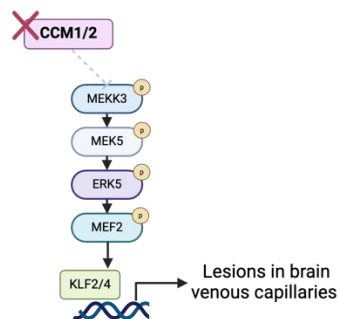
## Unmet Need

- **No approved therapy**
- **Surgical resection** or stereotactic radiosurgery is non curative and **not always feasible** because of location

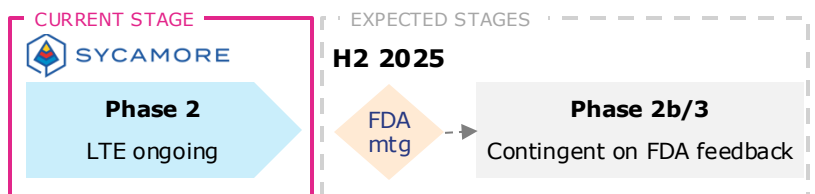
~360,000  
Symptomatic  
US + EU<sup>1</sup>

## Mechanism of Action

- **Selective**, orally bioavailable redox-cycling nitroxide
- Promotes the metabolism of ROS to **reduce oxidative stress** within cells
- **Stabilizes** endothelial barrier function



## Development Strategy



## Differentiation

- Potential **First-in-Disease** oral therapeutic for CCM
- **No TEAEs** leading to discontinuation up to **800 mg** in Ph 1<sup>3</sup>



Safe and well-tolerated MOA



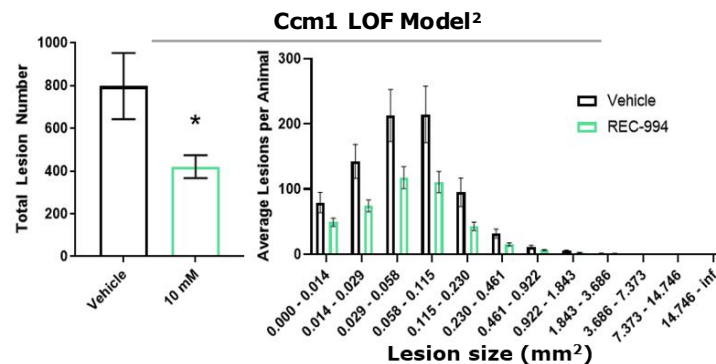
High oral bioavailability



Encouraging Ph 2 efficacy trends

## Key Preclinical Data

- **Reduces lesion number & size** in LOF mouse models
- Phase 2 **primary endpoint** of safety and tolerability met
- **Phase 2 encouraging trends in lesion volume reduction** consistent with *in vivo* POC



## Recursion Approach

- **Unbiased ML-aided** phenotypic drug screen to identify effective therapeutics driving CCM

**80%**  
Of Ph2 patients continued to LTE

**ODD**  
In US + EU

## What's Next

- **Phase 2** data expected to be shared at an upcoming medical congress / publication/webinar in **H1 2025**
- **FDA guidance** expected in **H2 2025**

# Familial Adenomatous Polyposis (MEK1/2 inhibitor): REC-4881

## Unmet Need

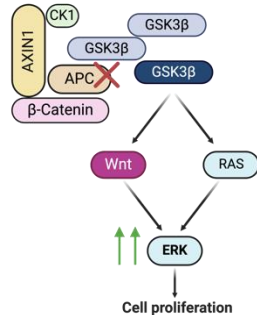
- **No approved therapy**
- **Colectomy** during adolescence is standard of care
- Patients at **significant risk of GI** cancer and suffer substantial decrease in **quality-of-life**

~50,000

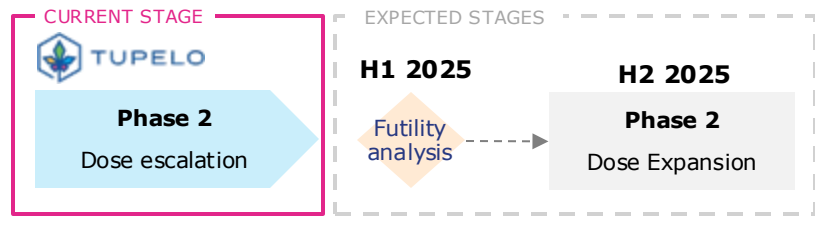
Diagnosed  
US + EU<sup>1</sup>

## Mechanism of Action

- **Loss of APC** drives FAP disease progression through aberrant pathway signaling (e.g., Wnt/B-catenin, MAPK signaling)
- REC-4881 **selectively blocks** the activation of ERK (MAPK pathway)

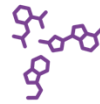


## Development Strategy



## Differentiation

- Potential **First-in-Disease** and **Best-in-Class** for FAP
- **Potent, non-competitive, allosteric** MEK1/2 inhibitor
- Oral 4 mg dose is **pharmacologically active**



Proof-of-mechanism  
in Phase 1b



Validated  
target

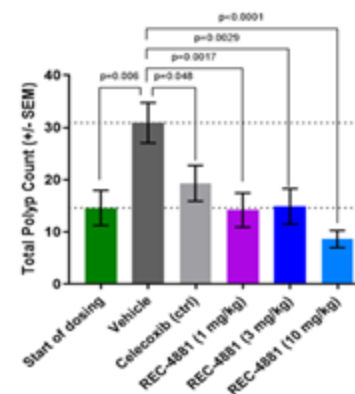


Preferential GI  
exposure

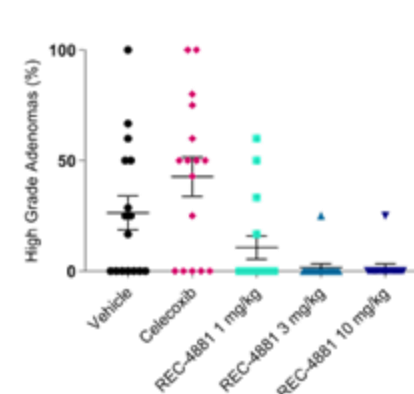
## Key Preclinical Data<sup>2</sup>

- **APC<sup>min/-</sup> mouse model:** Significantly reduces **polyp count** and **pre-cancerous adenoma**, outperforming celecoxib

Mean Polyps Per Group<sup>2</sup>



% Pre-Cancerous Polyps<sup>2</sup>



## Recursion Approach

- Unbiased **ML-aided phenomap insight** in human cancer cells

**FTD**  
In US

**ODD**  
In US + EU

## What's Next

- **Futility analysis** for reduction in polyp burden expected in **H1 2025**

# Hypophosphatasia (ENPP1 Inhibitor): REV102

## Unmet Need

- Opportunity to significantly **reduce costs & treatment burden**
- Many patients, particularly adults, may have difficulty accessing ERT
- Those who can access ERT face high treatment burden and tolerability hurdles

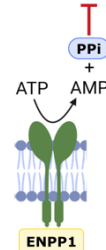
>7,800

Diagnosed prevalence  
US + EU<sup>1</sup>

## Mechanism of Action

- ENPP1 inhibition is a **genetically validated** target in HPP models
- Potent ENPP1 inhibitor that **restores Ppi balance** and enables bone mineralization

Pathologic soft tissue calcification



## Development Strategy

EXPECTED STAGES

2025

IND-Enabling Studies

2026

Phase 1

Healthy Volunteers

## Differentiation

- Potential **First-in-Class** and **Best-in-Class** ENPP1 Inhibitor
- **Non-immunogenic small molecule** offering potentially safer solution than ERT (3-6 injections per week)



No significant in vitro safety concerns



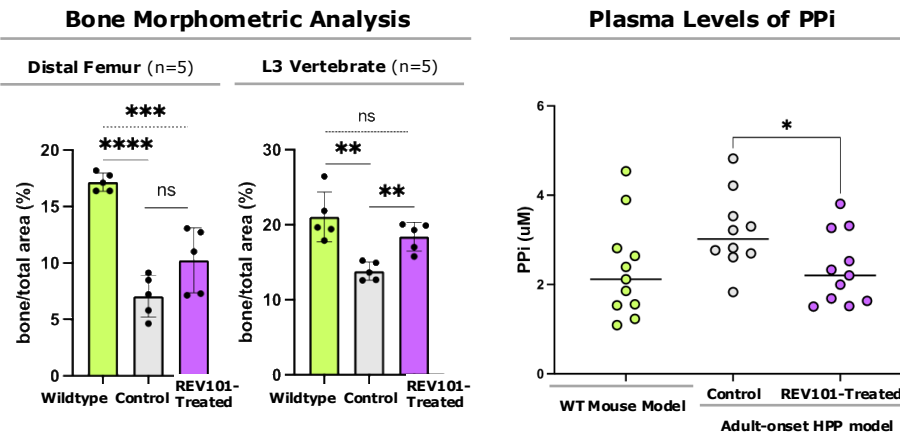
Affordable oral treatment option



Potential as mono or combo therapy

## Key Preclinical Data<sup>2</sup>

- Improvement in mineralization in mouse models of HPP
- Significantly reduced Ppi levels to that of wild-type mice



## Recursion Approach<sup>3</sup>

- **Precision designed for** both **high potency** and a lifetime of **chronic dosing**
- **Structurally distinct** differences vs competitor ENPP1 inhibitors
- **Maintain selectivity** and deliver a candidate with **high oral bioavailability** in the clinic

## What's Next

- **Development candidate nomination** expected in Q4 2024

# Neurofibromatosis Type 2 (HDAC Inhibitor): REC-2282

## Unmet Need

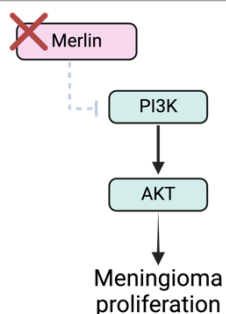
- **No approved therapy**
- Surgery/RT is standard of care (when feasible)<sup>2</sup>
- **Location** may make **complete resection untenable**, leading to hearing loss, facial paralysis, poor balance and visual difficulty

~33,000

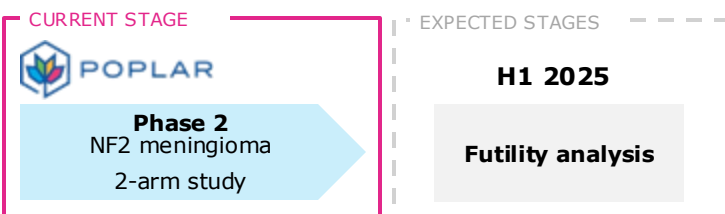
Treatable  
US + EU<sup>1</sup>

## Mechanism of Action

- **Loss of Merlin (NF2)** leads to PI3K signaling and meningioma proliferation
- **REC-2282** indirectly facilitates **AKT dephosphorylation** by disrupting the PP1-HDAC interaction



## Development Strategy



## Differentiation

- Potential **First-in-Disease** and **Best-in-Class** for NF2
- Potential to **rescue disease-inducing effects** of NF2 loss



High oral  
bioavailability



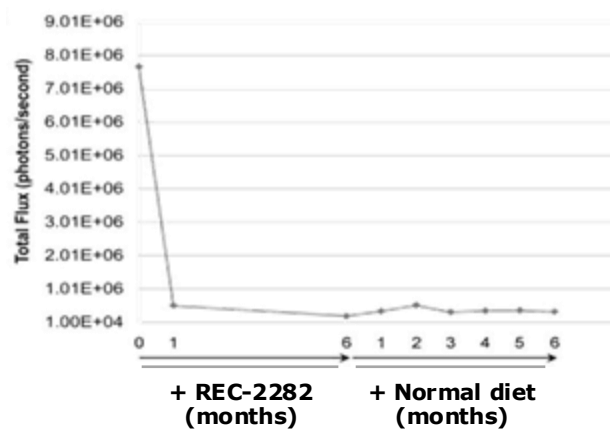
Improved CNS  
penetration



Reduced  
off-target effects

## Key Preclinical Data

- **Prevents growth & regrowth** of NF2-deficient meningioma model in mice<sup>3</sup>



## Recursion Approach

- Unbiased **ML-aided phenomap insight and drug screen** in human cells

**FTD**  
In US

**ODD**  
In US + EU

## What's Next

- **Phase 2 PFS data maturing**
- Futility analysis (PFS6) expected in **H1 2025**

PIPELINE

Other areas of high  
unmet need

# *C. difficile* (*C. diff* Toxin B Selective Inhibitor): REC-3964

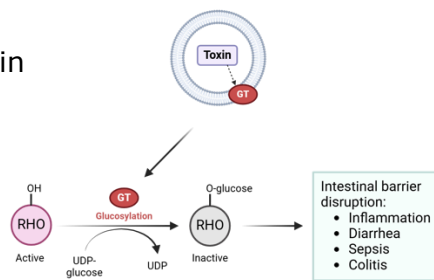
## Unmet Need

- **Limited treatment options** for high-risk population with recurrent CDI cases
- Ability to address populations not eligible for FMT or microbiome-based therapies

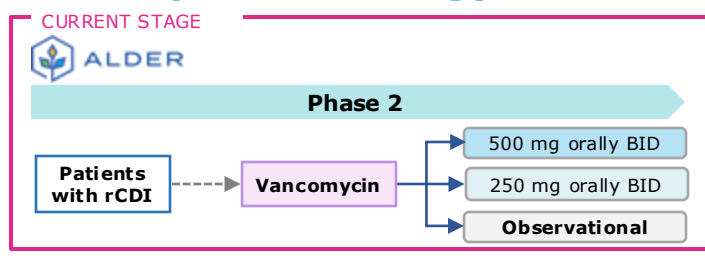
~175,000  
Recurrent *C. diff*  
cases US<sup>1</sup>

## Mechanism of Action

- **Highly potent**, orally bioavailable *C. diff* toxin B (TcdB) selective inhibitor
- **Selectively** inhibits catalytic activity of bacterial glucosyltransferase



## Development Strategy



## Differentiation

- Potential **First-in-Class** as non-antibiotic oral for rCDI
- **Highly potent** and **well-tolerated** with no reported DLTs, SAEs or treatment-related discontinuations in Phase 1



Safe and well-tolerated MOA



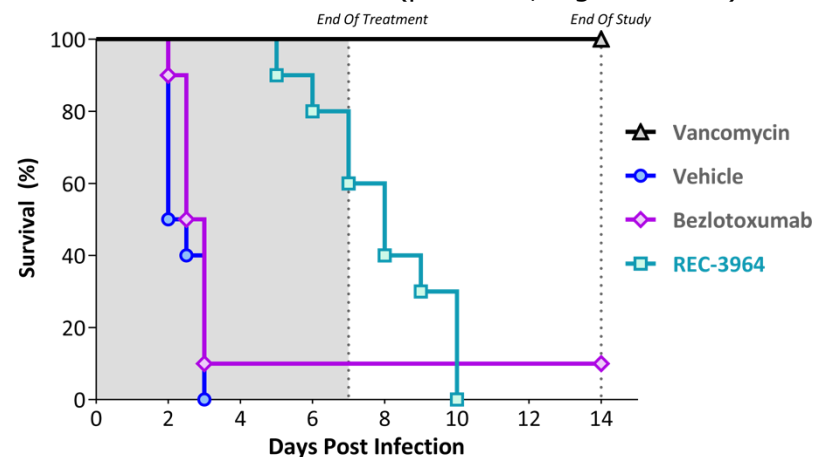
High oral bioavailability



Bacterial toxin selective

## Key Preclinical Data

- REC-3964 significantly extended survival vs bezlotoxumab alone at the end of treatment ( $p < 0.001$ , log rank test)<sup>2</sup>



## Recursion Approach

- Unbiased **ML-aided conditional phenotypic drug screen** in human cells

123

Novel compounds synthesized to candidate ID

## What's Next

- **First Patient Dosed in the Phase 2 ALDER trial** expected in Q4 2024
- Phase 2 update expected in **Q1 2026**

# Idiopathic Pulmonary Fibrosis (Target Epsilon - Undisclosed): REC-4209

## Unmet Need

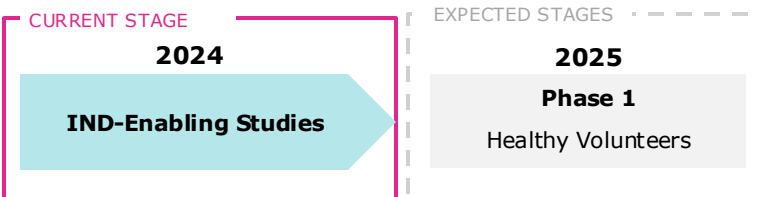
- **Approved therapies show modest slowing** of IPF progression
- **No improvement in survival (mOS 3-5 years) or quality of life** with current treatments

~130,000  
Diagnosed prevalence  
US<sup>1</sup>

## Mechanism of Action

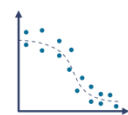
- **Reversible**, orally bioavailable, and potent Target Epsilon inhibitor
- Promotes **tissue repair and has potential to reverse fibrosis** likely by modulating TGF- $\beta$
- **Modulator of immuno-mesenchymal** populations in fibrosis, which **reduces fibrotic markers** in vivo and in vitro models of fibrotic disease

## Development Strategy



## Differentiation

- Potential **First-in-Class** treatment for IPF
- Potential for **safe** and **well-tolerated** novel treatment
- **In vitro models suggest** capability of reversing the fibrotic process driving IPF progression



Highly potent  
and selective



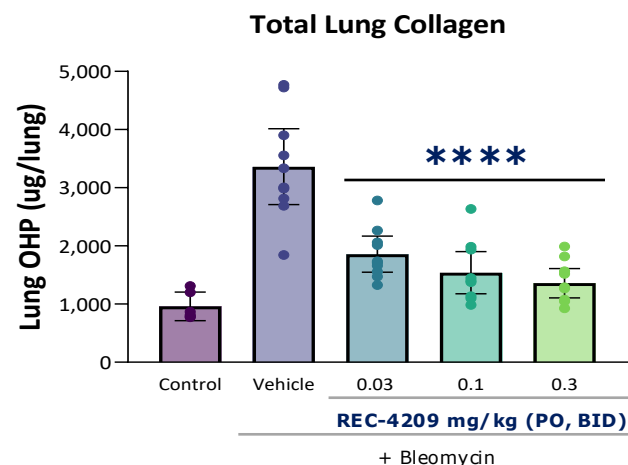
Novel mechanism  
of action



Potential fibrotic  
disease reversal

## Key Preclinical Data

- REC-4209 at low doses reduces total lung collagen by 45% to 60% versus vehicle mice<sup>2</sup>



## Recursion Approach

- Unbiased **ML-powered phenomap drug screen** in human cells

204

Novel compounds  
synthesized to candidate ID

## What's Next

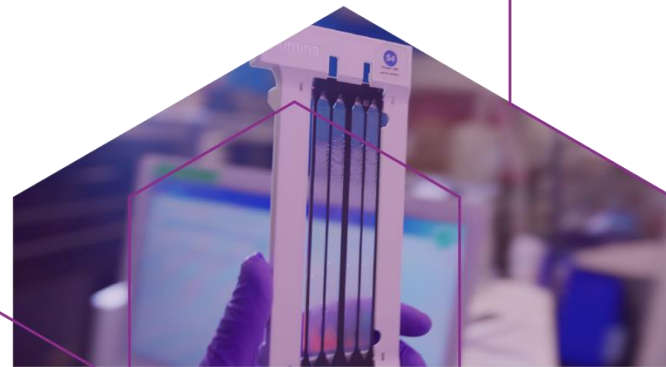
- **IND-enabling studies ongoing**

24 1. Global Data, Internal company estimates on IPF prevalence, Collard et al., Chest (2014).  
2. Groups compared against Vehicle. \*\*\*\*p<0.0001; one-way ANOVA with Tukey's multiple comparison test. Data reflects mean  $\pm$  95% CI

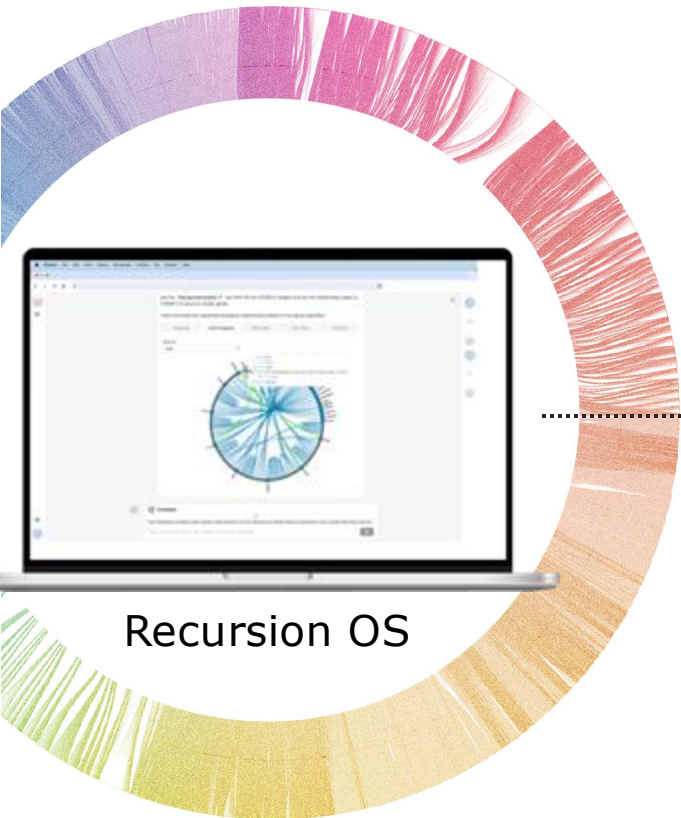


VALUE CREATION

# Partnerships & Data Strategy



# We harness value from the Recursion OS with a multi-pronged capital efficient business strategy



PIPELINE

## Pipeline strategy

Build internal pipeline in indications with potential for **advance transformational medicines for patients**

- Oncology
- Rare disease
- Other areas of high unmet need

PARTNERSHIP

## Partnership strategy

Partner in **complex therapeutic areas** requiring large financial commitment or competitive arbitrage  
**Leverage partner knowledge** and clinical development capabilities

- Neuroscience
- Oncology
- Immunology
- Other large, intractable areas of biology

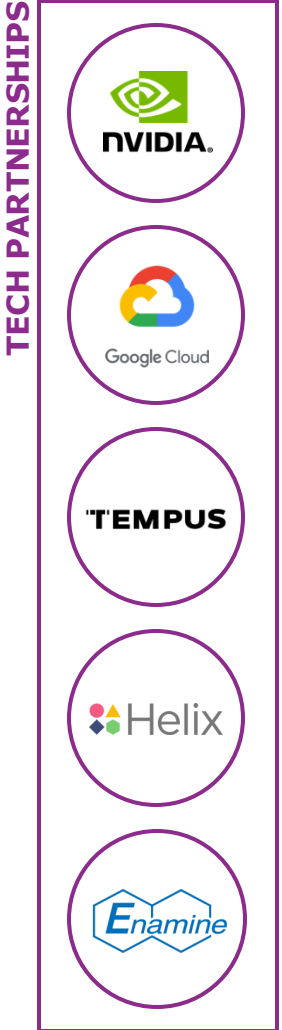
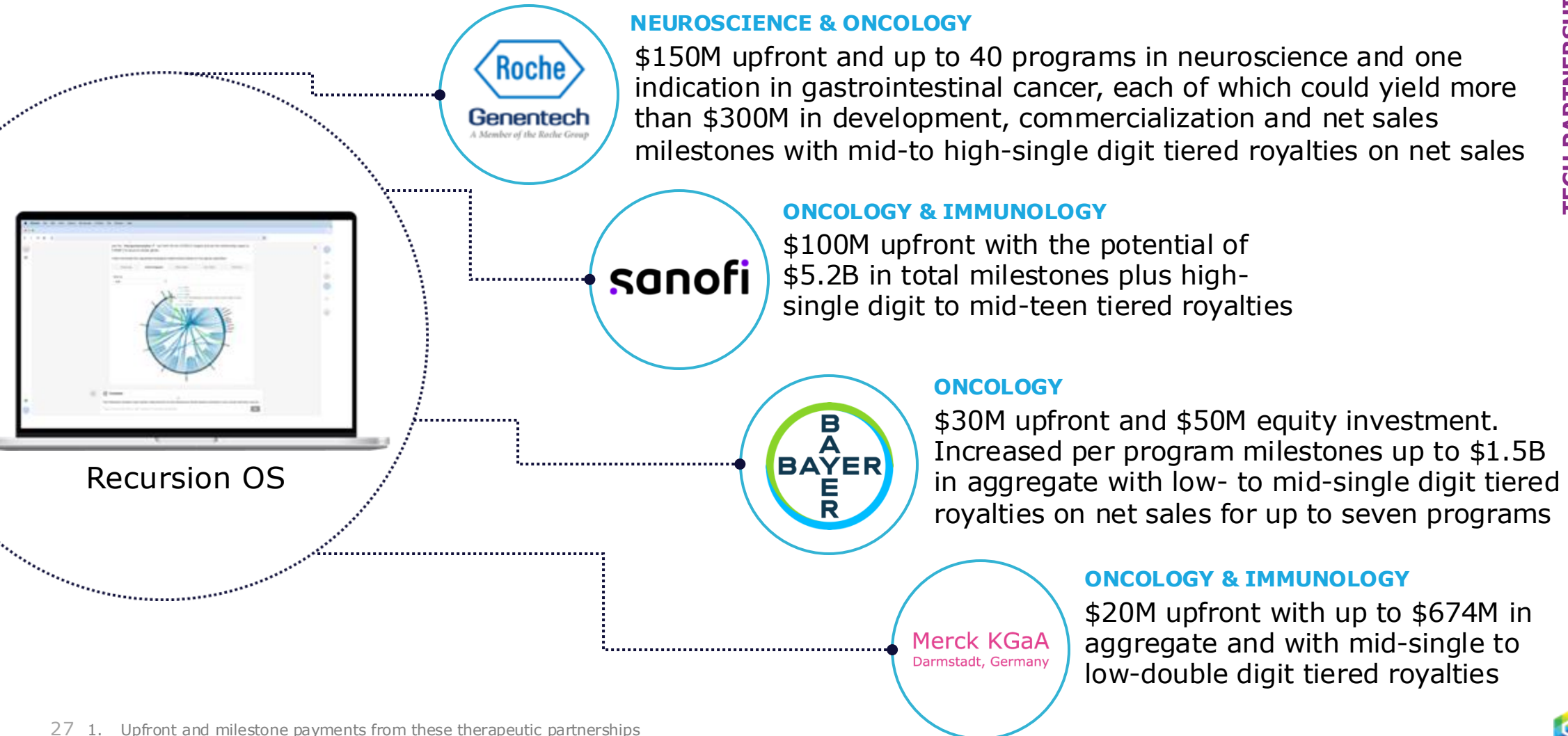
DATA

## Data strategy

**License** subsets of data and key tools  
Direct generation of new data internally **to maximize pipeline and partnership value-drivers**

- Licensing
- Augment Recursion OS
- LOWE

# Partnerships with approximately \$450M<sup>1</sup> earned to date and potential to receive more than \$20B<sup>2</sup> in additional milestones



27 1. Upfront and milestone payments from these therapeutic partnerships  
2. Additional milestone payments, excluding royalties

# We harness value from the Recursion OS with a multi-pronged capital efficient business strategy



PIPELINE

## Pipeline strategy

Build internal pipeline in indications with potential for **advance transformational medicines for patients**

- Oncology
- Rare disease
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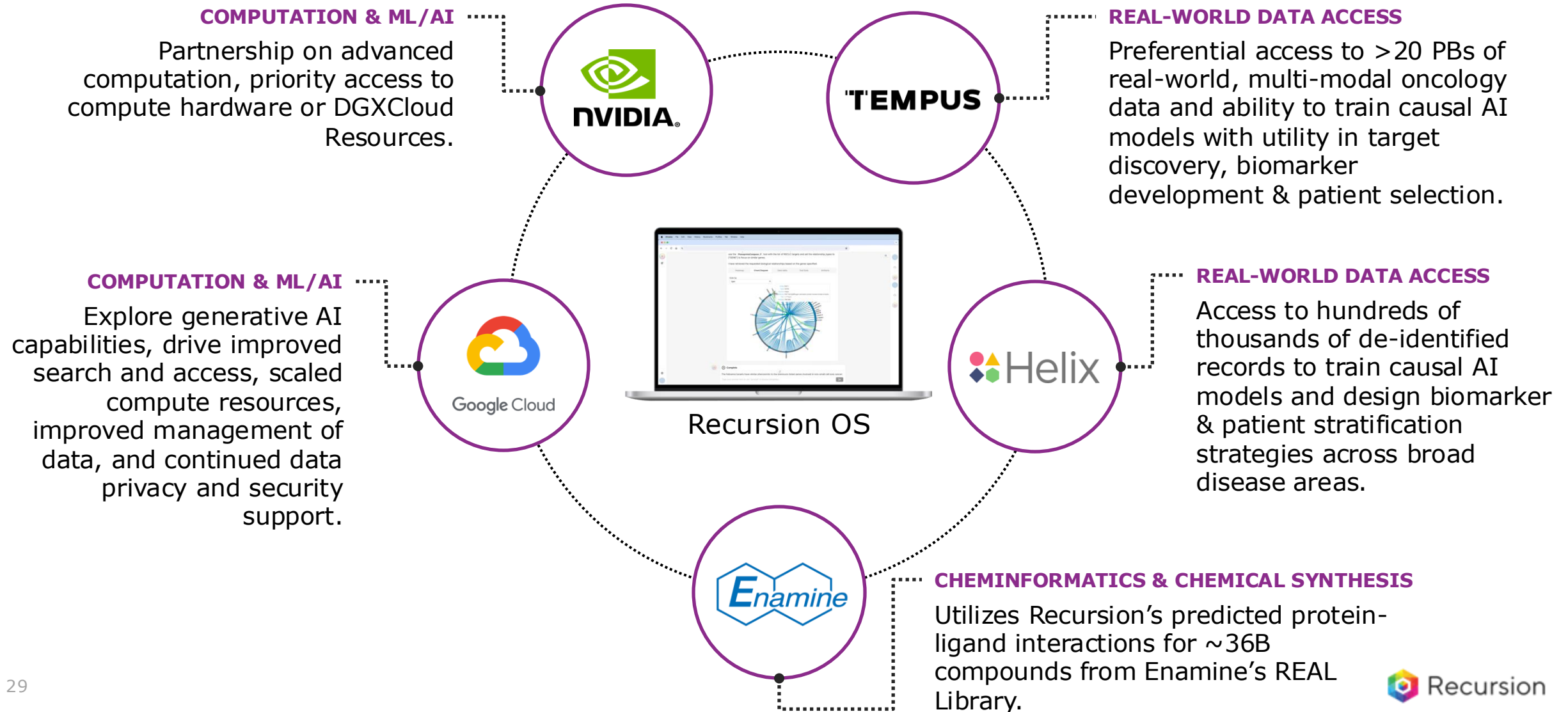
DATA

## Data strategy


**License** subsets of data and key tools  
Direct generation of new data internally **to maximize pipeline and partnership value-drivers**

- Licensing
- Augment Recursion OS
- LOWE

# We license subsets of data and key tools to generate new data to maximize pipeline and partnership value-drivers



# LOWE puts the Recursion OS at your fingertips via natural language without any coding expertise required



LOWE helps you orchestrate complicated workflows using natural language. Try one of the templates below or enter a prompt manually.

Listing Known Targets Associated with a Specific Phenotype [✎](#)

Identifying Top Compounds with Similar Phenoprints to a Given ... [✎](#)

Request for Novel Compounds with Same Scaffold as a Given M... [✎](#)

[Browse templates](#)

Type your prompt here or use "/prompt" to browse templates... [▶](#)

**D** Give me a list of targets involved in non small cell lung cancer


[Thought](#) [Complete](#)

The user has asked for a list of targets involved in non-small cell lung cancer (NSCLC). To provide this information, I will use the [PhenotypeToTarget](#) tool to retrieve a list of known biological entities (gene symbols) associated with NSCLC. The input will be the phenotype name "non small cell lung cancer."

I have retrieved the following targets for non small cell lung cancer.

Visualisation    Data table    Tool Data    Artifacts

Select Y Value:



Target	Overall Association Score
EGFR	0.90
ERBB2	0.83
ALK	0.82
KRAS	0.82
BRAF	0.80
MET	0.80
TP53	0.79
KDR	0.77
STK11	0.77
ERBB4	0.76

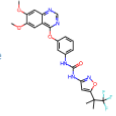
Type your prompt here or use "/prompt" to browse templates... [▶](#)

**D** Find the top 50 commercially available compounds that have a similar structure to COC1=CC2=NC=NC(OC3=CC(NC(=O)NC4=NO(C(C)(C)C(F)(F)F)=C4)=CC=C3)=C2C=C1OC and place an order for all 50

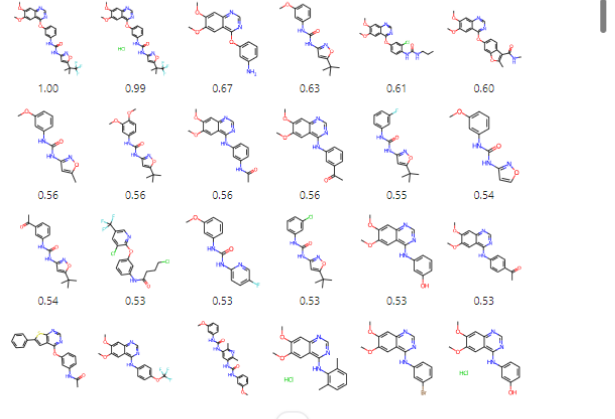
[Thought](#) [Complete](#)

The user has requested to find the top 50 commercially available compounds similar to a given SMILES string and then place an order for all 50. I will first use the [MolSimilaritySearch](#) tool to find the top 50 similar compounds and then use the [OrderCompounds](#) tool to place an order for them.

I have retrieved the most similar compounds starting from the following structure



Molecule grid    Data table    Tool Data    Artifacts



Similarity Score
1.00
0.99
0.67
0.63
0.61
0.60
0.56
0.56
0.56
0.56
0.55
0.54
0.54
0.53
0.53
0.53
0.53
0.53
0.53

Type your prompt here or use "/prompt" to browse templates... [▶](#)

# Culture and Team



# Our leadership brings together experience & innovation to advance TechBio

## Executive Team



**Chris Gibson, PHD**  
Co-Founder, &  
Chief Executive  
Officer



**Najat Khan, PHD**  
Chief R&D Officer &  
Chief Commercial  
Officer  
**Johnson&Johnson**



**Ben Taylor**  
Chief Financial  
Officer & President  
Recursion UK  
**Goldman Sachs** **AETION**



**David Mauro, MD PHD**  
Chief Medical Officer



**David Hallett, PHD**  
Chief Scientific Officer



**Ben Mabey**  
Chief Technology  
Officer



**Kristen Rushton**  
Chief Operations  
Officer  
**Vyriad genetics**



**Nathan Hatfield**  
Chief Legal  
Officer  
**WILSON  
SONSINI**



**Matt Kinn**  
Chief Business  
Officer



**Erica Fox**  
Chief People &  
Impact Officer



**Lina Nilsson, PHD**  
SVP, Head of  
Platform



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CSO, & CBO of Celgene



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Executive Officer



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Chief R&D Officer &  
Chief Commercial Officer



**Dean Li, MD PHD**  
Co-Founder of RXRX,  
President of Merck  
Research Labs

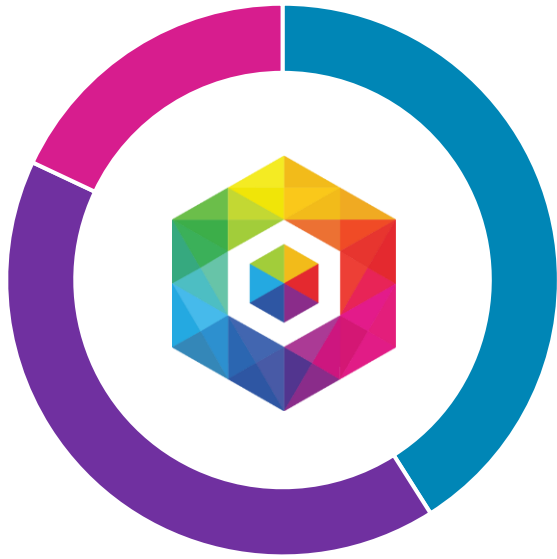


**Zavain Dar**  
Co-Founder & Partner  
of Dimension





# Our people are the most important ingredient for our mission



~800 employees

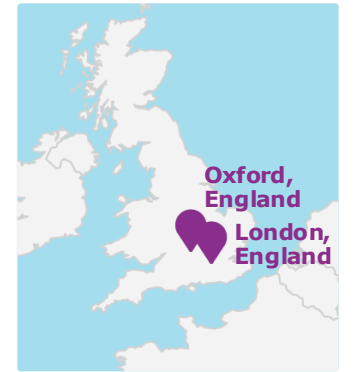
- Technology – data science, software engineering, automation, etc.
- Life Sciences – biology, chemistry, development, etc.
- Strategic Operations

*Parity Pledge Signer:*  
Gender parity and people of color parity



Headquartered in **Salt Lake City, Utah** with other primary locations in:

- Milpitas, California
- New York, New York
- Toronto, Ontario
- Montréal, Québec
- London, England
- Oxford, England



## ESG Highlights



Learn more about Recursion's ESG stewardship:  
[www.recursion.com/esg](http://www.recursion.com/esg)

## Community Impact

altitude ▲ lab

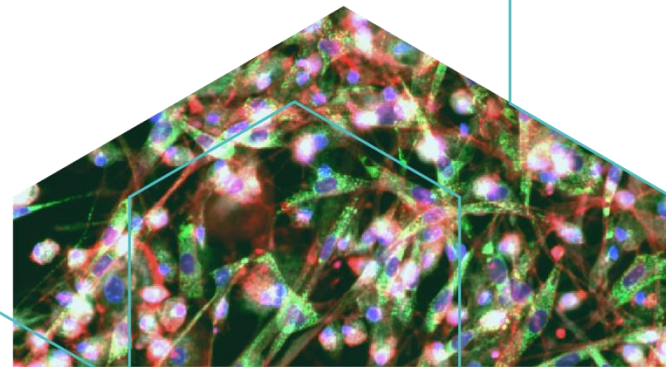
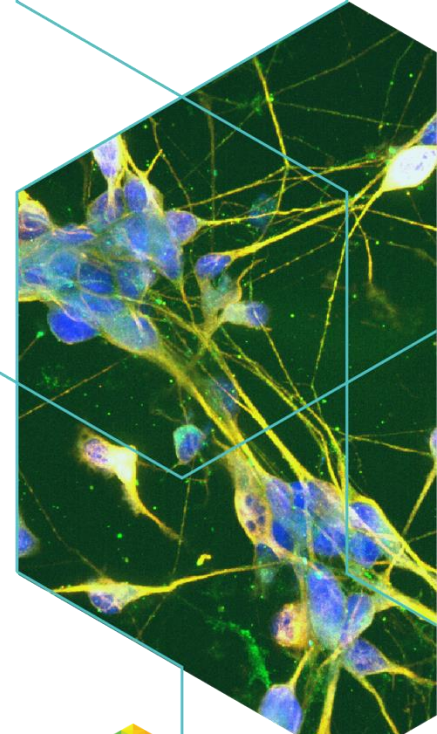
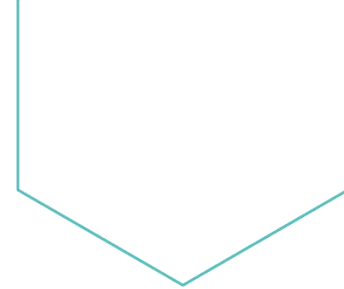
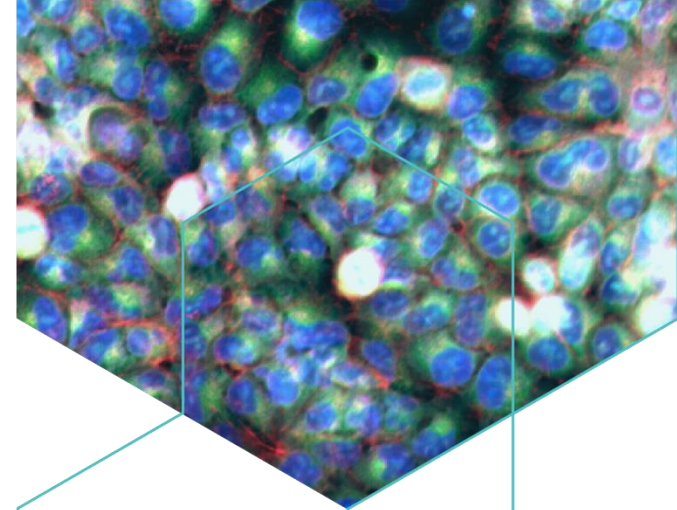
Founding Partner,  
Life Science Accelerator

biohive™

Founding Member,  
Life Science Collective

APPENDIX

# Pipeline Details



PIPELINE

# Oncology

# REC-617\*: CDK7 Inhibitor

A precision designed highly selective CDK7 inhibitor for Relapsed and/or Refractory (R/R) Solid Tumors

## Program Status

- Potential **Best-in-Class** and **First-in-Class** CDK7 inhibitor
- Phase 1/2 study in advanced solid tumors ongoing
- Initial Phase 1 monotherapy safety, PK/PD update expected at **AACR Special Conference in Cancer Research on December 9, 2024**

## Mechanism of Action

- **Reversible CDK7 inhibitor** that targets both cell cycle progression and transcriptional regulation

## Thesis & Differentiation

- **Non-covalent binding and improved selectivity** to decrease off-target toxicity
- 8-10 hours of therapeutic coverage at IC<sub>80</sub> with a **short half-life** to reduce on-target toxicity
- **Rapid absorption and permeability** at lowest possible dose

## Unmet Need<sup>1</sup>

- **Multiple cancer indications** that have the potential to address ~185,000 patients annually
- **R/R solid tumors** including breast, NSCLC, ovarian, pancreatic, colorectal, and head & neck

## Recursion Approach

- AI-powered precision design to optimize PK/PD to **maximize potential therapeutic index**
- 136 novel compounds synthesized to candidate ID

# REC-617: Robust anti-tumor activity demonstrated in disease relevant preclinical tumor models

Initial clinical safety and PK/PD update on track for Q4 2024

## Key Preclinical Data

REC-617 has Best-in-Class potential<sup>1</sup>

Designed to avoid efflux transporter substrate to minimize GI adverse events

Category	Assay	DC Criteria	Ph 1 Competitor	Ph 1/2 Competitor	REC-617
Potency & Selectivity	CDK7 IC50 (nM)	<10	Meets or exceeds criteria	Minor deviation	Meets or exceeds criteria
	CDK family selectivity	>100-fold	Meets or exceeds criteria	Major deviation	Meets or exceeds criteria
	HCC70 (breast cancer) IC50 (nM)	<100	Meets or exceeds criteria	Minor deviation	Meets or exceeds criteria
ADME	<b>Caco-2 A2B (efflux)</b> <b>10<sup>-6</sup> cm/s</b>	>5 (<3)	Major deviation	Major deviation	Meets or exceeds criteria
	Predicted human half-life (hr)	<15	Minor deviation	Major deviation	Meets or exceeds criteria

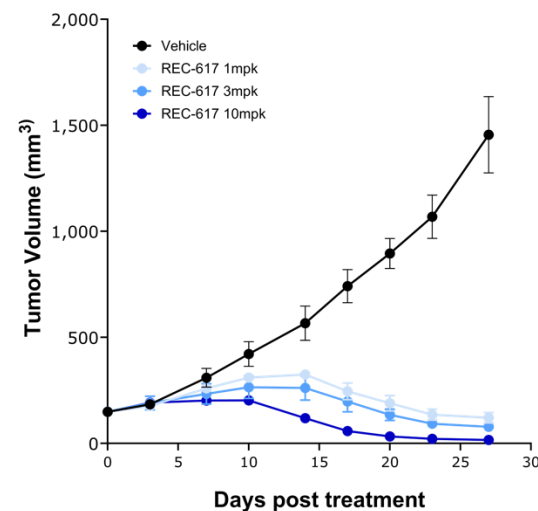
Meets or exceeds criteria    Minor deviation    Major deviation

Development Candidate (DC) Criteria:

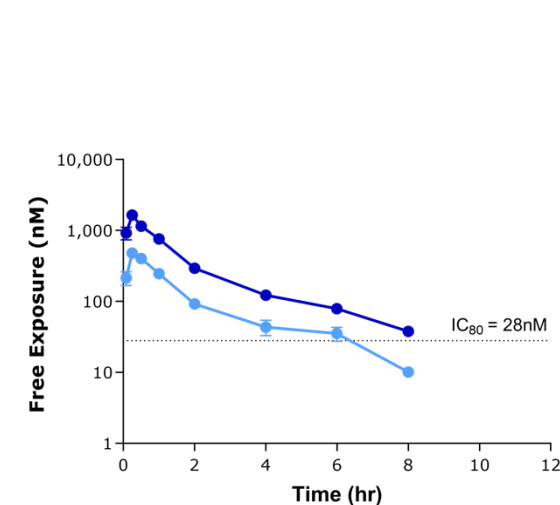
- **CDK7 IC50:** green <10nM; yellow 10-30nM; red >30nM
- **CDK7 selectivity:** green >100-fold; yellow 30-100-fold; red <30-fold
- **HCC70 IC50:** green <100nM; yellow 100-500nM; red >500 nM
- **Caco-2 A2B (efflux):** green >5(<3); yellow >1.5 (<10); red <1.5 (>30)
- **Half-life:** green <15, yellow <24, red >24

Potent tumor regression with minimal IC<sub>80</sub> exposure

CDX Model: OVCAR3<sup>2</sup>



Mouse PK<sup>3</sup>



- REC-617 demonstrates potent tumor regression with less than 10 hours of exposure above IC<sub>80</sub> to optimize benefit-risk

# REC-617 (CDK7 inhibitor): Study Design and Next Steps

## Development Strategy

### ELUCIDATE Phase 1/2 study design

#### CURRENT STAGE

**Phase 1**  
Dose Escalation (N≤60)

#### **Part A**

- Monotherapy

#### **Part B**

- Initial combination with SERD in HR+/HER2- post CDK4/6 inhibitor population

#### **Primary Endpoint:**

- Safety and Tolerability
- Recommended Dose

RP2D

**Phase 2**  
Dose Expansion

- N = 30-60 patients in combination
- N will depend on number of disease specific cohorts

#### **Primary Endpoint:**

- ORR

### REC-617 Competitive Profile

- Potential **Best-in-Class** CDK7 inhibitor
- **Reduced risk** of off-target toxicity
- **Highly selective & potent**

### Trial Update

- Phase 1 monotherapy preliminary safety and PK/PD data update expected **Dec 9, 2024 (AACR Special Conference in Cancer Research)**

# REC-1245: RBM39 Degradator

A highly selective RBM39 degrader for Biomarker-Enriched Solid Tumors and Lymphoma

## Program Status

- Potential **First-in-Class** RBM39 degrader in solid tumors
- Phase 1/2 study **initiation** expected in **Q4 2024**
- **Phase 1 monotherapy** update on dose-escalation expected in **H1 2026**

## Mechanism of Action

- **Molecular glue** that degrades RBM39 via E3 ligase adaptor DCAF15
- **Disrupts RNA splicing** to downregulate cell cycle checkpoints and DDR networks

## Thesis & Differentiation

- **RBM39 phenotypically mimics CDK12** and is **distinct** from **CDK13** in Recursion OS
- **Novel approach** to target DDR biology via RBM39 **avoids on-target toxicities** associated with cell cycle checkpoint inhibitors (e.g., CDK12, WEE1, ATR, ATM, PLK1)
- **Selective** RBM39 degrader with **minimal ITGA2 liability** to limit thrombocytopenia

## Unmet Need<sup>1</sup>

- **>100,000 patients** with solid tumor or lymphoma experience disease progression while on frontline therapies
- Potential to be used as a **single agent or in combination** with chemo/IO

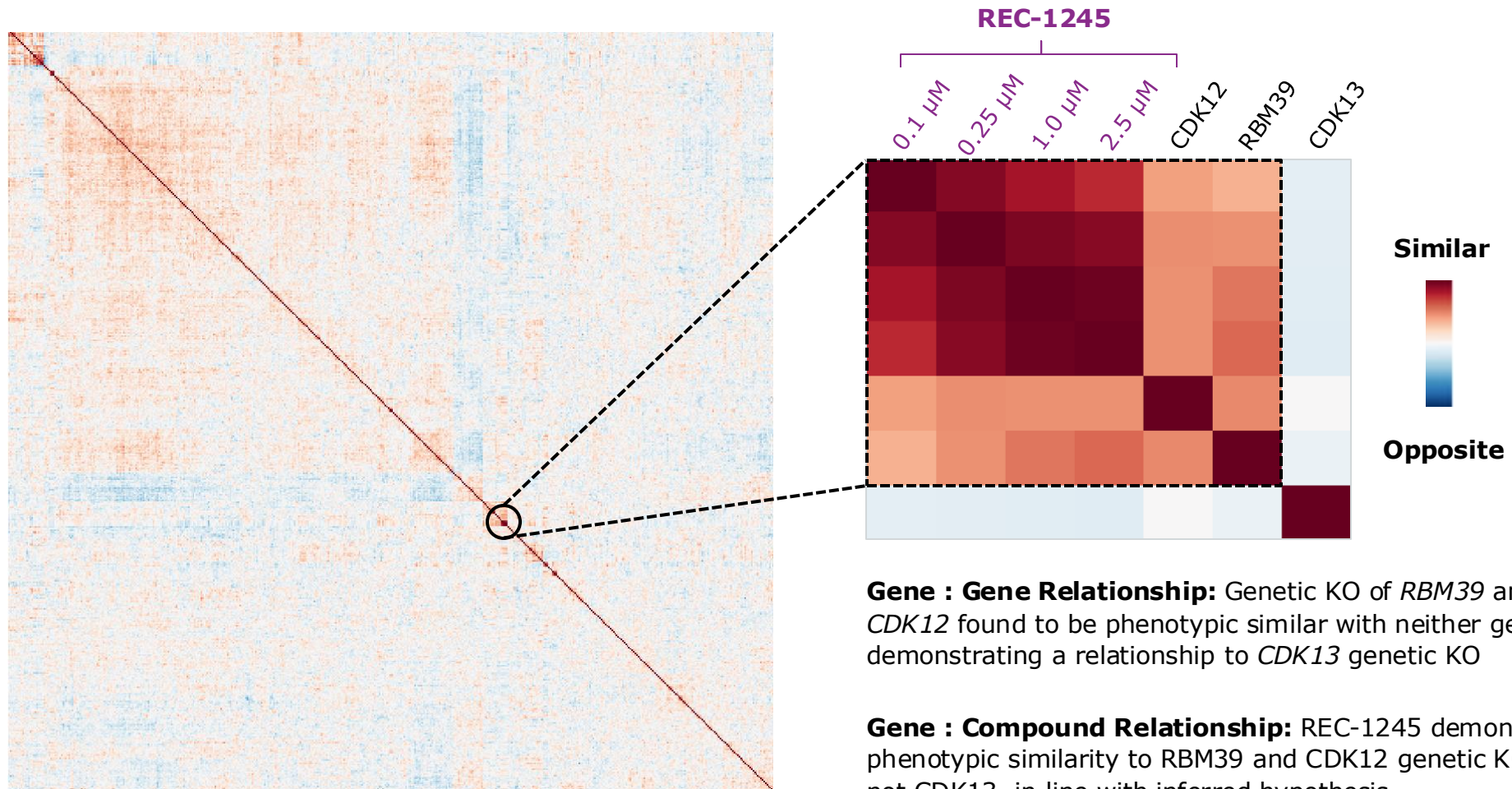
## Recursion Approach

- **Unbiased ML-aided genomics screen** to identify biological signature and relate cellular phenotypes
- Progressed REC-1245 from target biology to IND-Enabling studies in **under 18 months (vs. 42 months in industry<sup>2</sup>)**

39 1. Internal company estimates. Assumes US+EU5 addressable incidence with biomarker-enriched solid tumors and other select histologies.  
2. Paul et al, Nat Rev Drug Discov (2010)

# REC-1245 (RBM39 degrader): Platform inferred a functional similarity between RBM39 and CDK12 biology suggesting a novel approach to potential DDR modulation

## Recursion OS Novel Insight





# REC-1245 (RBM39 degrader): Robust efficacy/PK/PD in biomarker-positive disease relevant preclinical tumor models with Phase 1 initiation expected Q4 2024

## Key Preclinical Data<sup>1</sup>

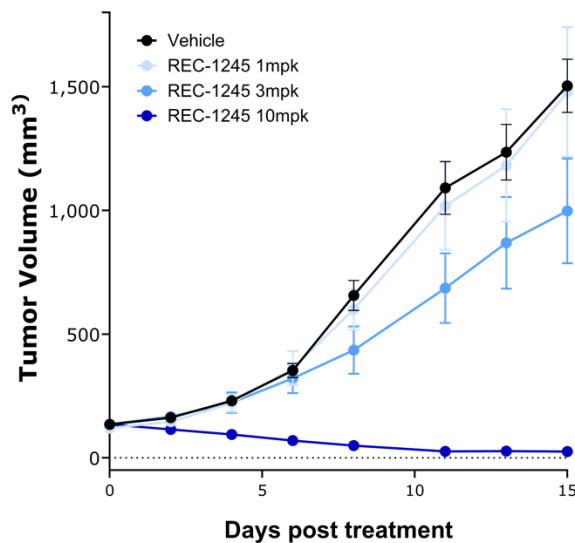
REC-1245 is highly selective and potent

Category	Assay	DC Criteria	REC-1245
Potency / Selectivity	RBM39 Degradation DC <sub>50</sub>	<100 nM	Meets or exceeds criteria
	CDK12 Kinase	No sig. activity	Meets or exceeds criteria
In Vitro Safety	CEREP Safety Panel	No sig. activity	Meets or exceeds criteria
	hERG IC <sub>50</sub> (μM)	>30	Meets or exceeds criteria
Pharmacokinetics	Oral Bioavailability (%F)	>30	Meets or exceeds criteria

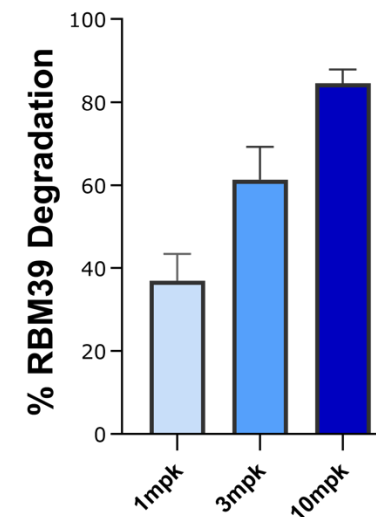
Meets or exceeds criteria    Minor deviation    Major deviation

REC-1245 has compelling efficacy and PK/PD in preclinical models

CDX Model: OVK18<sup>2</sup>



PD: Target Engagement<sup>3</sup>



- REC-1245 shows significant monotherapy regressions
- Dose-dependent antitumor activity correlates with PD

41 1. Data on File. 2. N=8 mice per group in TV portion. REC-1245 administered BID PO. 3. PD evaluated after 5 days BID oral of REC-1245 at doses noted ; N=3 mice per group in PD portion

# REC-1245 (RBM39 degrader): Study Design and Next Steps

## Development Strategy



DAHLIA Phase 1/2 study design

### CURRENT STUDY

**Phase 1**  
Dose Escalation (N~55)

#### **Part A**

- Monotherapy dose-finding

#### **Part B**

- Monotherapy dose-confirmation

RP2D

**Phase 2**  
Dose Expansion

### Key Study Characteristics

#### Study objectives:

- Safety/Tolerability
- RP2D
- ORR

#### Eligibility:

- Select histologies including a biomarker population and R/R lymphomas

### REC-1245 Competitive Profile

- **Highly potent**, potential **First-in-Class** RBM39 degrader (<100nM DC50)
- No significant in vitro safety concerns (CEREP, hERG)
- No significant activity in CDK12 kinase assay
- Minimal ITGA2 liability to **limit thrombocytopenia**
- High oral bioavailability

### Trial Update

- Monotherapy dose escalation trial initiation expected **Q4 2024**
- Trial **active and enrolling** at 5 US sites

# REC-3565\*: MALT1 Inhibitor

A precision designed selective MALT1 inhibitor for B-Cell Malignancies

## Program Status

- Potential **Best-in-Class** MALT1 inhibitor
- Phase 1 initiation in B-Cell Malignancies (e.g., chronic lymphocytic leukemia), expected **Q1 2025**

## Mechanism of Action

- **Reversible allosteric MALT1 inhibitor** that can dampen NF-κB signaling
- **Selectively** inhibits CLL proliferation with limited impact on T-Cell viability

## Thesis & Differentiation

- **Low UGT1A1 liability** with potential for reduced risk of hyperbilirubinemia
- **Potential for reduced liver toxicity and enhanced efficacy** in combination with BTK and BCL2 inhibitors
- Low predicted human clearance and **high oral bioavailability**

## Unmet Need<sup>1</sup>

- **Current monotherapy treatments** in B-cell malignancies not curative and prone to resistance
- ~41,000 patients with R/R B-cell malignancies (treatable in US and EU5) – targeting CLL combination therapy

## Recursion Approach

- **AI powered** precision-designed novel molecule using **molecular dynamics and hotspot analysis**
- 344 novel compounds synthesized to candidate ID
- Maintain selectivity and deliver a candidate with lower predicted safety risk in the clinic

43 \*Formerly EXS73565.

1. Cerner Enviza Treatment Architecture Reports 2023, rounded to nearest 1,000 patients per year.

# REC-3565 (MALT1 inhibitor): Minimal UGT1A1 liability vs competitors and significant tumor regression observed in vivo with Phase 1 initiation anticipated on Q1 2025

## Key Preclinical Data

REC-3565 has Best-in-Class potential<sup>1</sup>

Category	Assay	DC Criteria	Ph 1 large pharma	Ph1 biotech	REC-3565
Potency & Selectivity	MALT1 IC <sub>50</sub> (nM)	<100	Yellow	Green	Green
	OCI-Ly3 proliferation IC <sub>50</sub> (nM)	<400	Yellow	Green	Green
ADME	<b>UGT1A1 IC<sub>50</sub> (μM)</b>	>10	Red	Red	Green
	Caco-2 A2B (efflux) 10 <sup>-6</sup> cm/s	>5 (<3)	Green	Yellow	Green

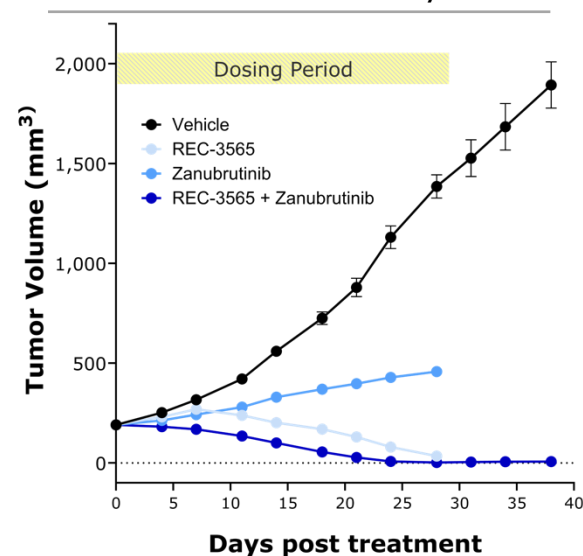
■ Meets or exceeds criteria   
 ■ Minor deviation   
 ■ Major deviation

### Development Candidate (DC) Criteria:

- **MALT1 IC<sub>50</sub> nM:** green <100 nM; yellow >100-<300 nM; red >300 nM
- **OCI-Ly3 IC<sub>50</sub> nM:** green <400 nM; yellow >400-<1000 nM; red >1000 nM
- **UGT1A1 IC<sub>50</sub> μM:** green >10 μM; yellow <10->1 μM; red <1 μM
- **Caco-2 A2B (efflux):** green >5(<3); yellow >1-<5(>3-<10); red <1(>10)

Single-agent and synergistic activity in vivo<sup>2</sup>

### CDX Model : OCI-Ly10<sup>2</sup>



**70%**

Of mice in combination arm (REC-3565 + zanu) had no palpable tumors 10-days post last dose

- OCI-Ly10 and Rec-1 cells are sensitive to both MALT1i and zanutrutinib *in vitro*
- Administration of REC-3565 as a single agent showed tumor growth regression
- Durable tumor growth regression observed when REC-3565 was combined with zanutrutinib

# REC-3565 (MALT1 inhibitor): Study Design and Next Steps

## Development Strategy

### EXCELERIZE Phase 1 study design

#### Phase 1 Dose Escalation

Q1 2025

#### Part A Monotherapy

- N ~30
- R/R B-Cell Malignancies
- REC-3565 PO QD or BID

RD

#### Part B Combination

#### Primary Endpoint:

- Safety / tolerability
- RD for combination

### REC-3565 Competitive Profile

- **Low** predicted human clearance and **high oral bioavailability**
- **No unexpected** in vitro or in vivo **safety concerns** identified
- **Well tolerated** in rat/dog dose range finding (DRF) studies
- GLP-tox studies completed with **suitable no-observed-adverse-effect level (NOAEL)** enabling clinical trials

### Trial Update

- Trial initiation expected **Q1 2025**

# REC-4539\*: LSD1 Inhibitor

A precision designed unique LSD1 inhibitor with CNS penetrance

## Program Status

- Potential **Best-in-Class** LSD1 inhibitor
- **Phase 1 initiation** in SCLC expected **1H 2025**

## Mechanism of Action

- **Reversible LSD1 inhibitor** that can selectively upregulate NOTCH signaling
- Promotes **differentiation of neuroendocrine cancer cells**
- **Impairs DNA repair pathways** sensitizing SCLC cells to immune checkpoint inhibitors

## Thesis & Differentiation

- LSD1 inhibitor designed to be **reversible** and **brain penetrant**
- **Shorter-predicted half life** versus competitors to manage **on-target toxicity**
- **Highly selective** to reduce **off-target toxicity**
- Preclinical data shows therapeutic exposures have minimal effects on platelets, suggesting potential **reduced risk of thrombocytopenia**

## Unmet Need<sup>1</sup>

- **>45,000 patients** with treatable Stage III/IV SCLC
- Limited treatment options post progression on frontline therapies

## Recursion Approach

- **Precision design** using **active learning to select most information rich compounds**
- 414 novel compounds synthesized to candidate ID
- Used multiparameter optimization to design a unique candidate combining reversibility with CNS penetration

# REC-4539 (LSD1 inhibitor): Sufficient CNS exposures vs competitors and compelling dose-response demonstrated in vivo with Phase 1 initiation anticipated in H1 2025

## Key Preclinical Data

REC-4539 has Best-in-Class potential<sup>1</sup>

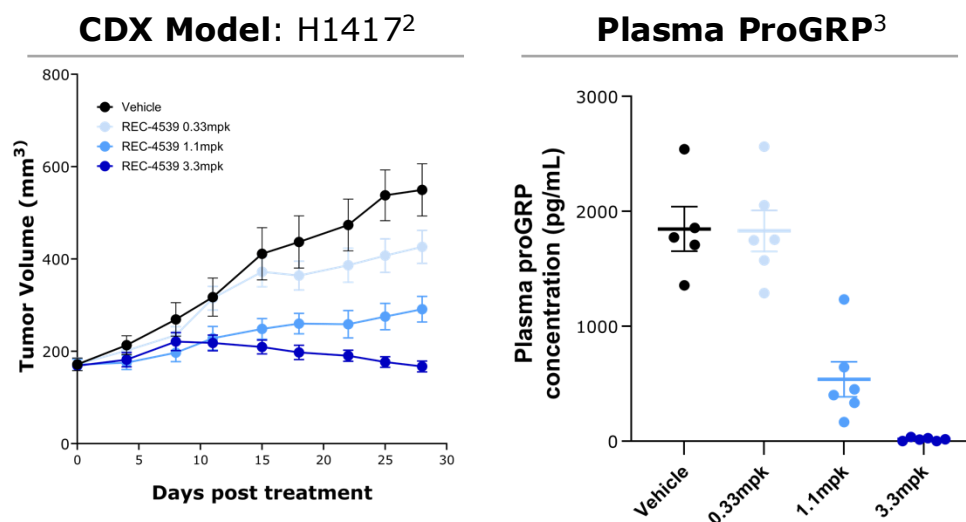
Assay	DC Criteria	Competitor 1	Competitor 2	REC-4539
<b>Brain : Plasma Ratio</b>	>0.5	Major deviation	Major deviation	Meets or exceeds criteria
MDCK-MDR1 Efflux Ratio (Pgp)	<2	Minor deviation	Minor deviation	Meets or exceeds criteria
Predicted Human Half-life	QD dosing	Major deviation	Major deviation	Meets or exceeds criteria

■ Meets or exceeds criteria   
 ■ Minor deviation   
 ■ Major deviation

Development Candidate (DC) Criteria:

- **Brain:plasma ratio:** green >0.5; red <0.5
- **MDCK-MDR1 efflux ratio (Pgp):** green <2; yellow >2-<10; red >10
- **Predicted half-life:** green <24 hours; yellow 24-48h hours; red >48 hours

REC-4539 highly efficacious in SCLC xenograft model<sup>2</sup>



- Dose-dependent regression
- Well-tolerated with limited impact on platelet levels

### Trial Update

- Phase 1 **First Patient Dosed** in SCLC expected **H1 2025**

PIPELINE

# Rare disease



# REC-994: Superoxide Scavenger

A safe and well tolerated superoxide scavenger for the treatment of Cerebral Cavernous Malformation (CCM)

## Program Status

- **First therapeutic candidate** advanced to an industry-sponsored Phase 2 trial
- **Phase 2 primary endpoint** of safety **met** with similar AE profile across arms
- Meeting with FDA anticipated in **H2 2025** to discuss plans for additional clinical study

## Mechanism of Action

- **Selective**, orally bioavailable, redox-cycling nitroxide
- Promotes the metabolism of ROS to **reduce oxidative stress** within cells
- **Stabilizes** endothelial barrier function

## Thesis & Differentiation

- Develop the **first oral therapy** for the treatment of symptomatic CCM
- Target the **underlying genetic mechanisms** that drive the disease pathophysiology of CCM

## Unmet Need<sup>1</sup>

- ~360,000 symptomatic CCM patients with **no approved therapies**
  - ~**63,000 patients** harboring **brainstem lesions** and elevated bleeding risk
  - ~**36,000 patients** with **cavernoma-related epilepsy**<sup>2,3</sup>

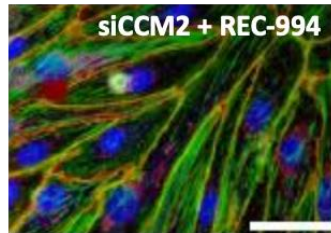
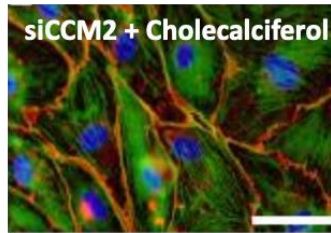
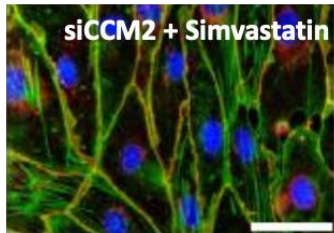
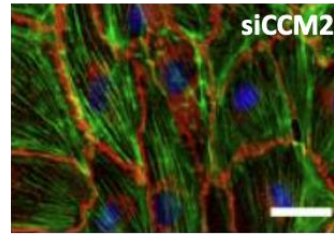
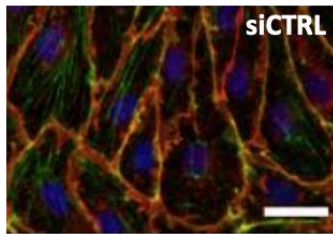
## Recursion Approach

- **Unbiased ML-aided** phenotypic drug screen to identify effective therapeutics driving CCM
- In vivo POC demonstrated lesion reductions that were also observed in the Ph2 trial

# REC-994 (Superoxide Scavenger): Preclinical studies showing reduction of lesion burden de-risked the first industry-sponsored Phase 2 study in CCM

## Recursion OS Insight

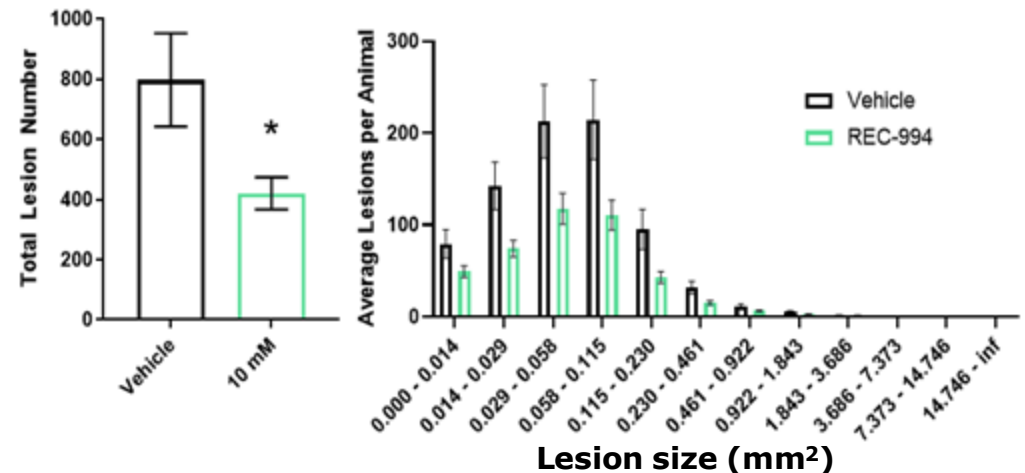
Identified REC-994 as potential rescue molecule in phenotype associated with CCM2 loss of function



## Key Preclinical Data<sup>1</sup>

Reduces lesion number & size in *Ccm1* and *Ccm2*<sup>2</sup> loss of function (LOF) mouse models

### Ccm1 LOF Model



# REC-994 (Superoxide Scavenger): Topline Phase 2 data in September demonstrated encouraging signals of efficacy

## Trial Update



- Randomized, double-blind, placebo-controlled Phase 2 study
- **Primary endpoint** of safety and tolerability **met** September 2024
- **Encouraging trends** observed in objective MRI-based exploratory efficacy measures observed
- **Time- and dose-dependent trends in reduced lesion volume** and **hemosiderin ring size** compared to placebo
- **80% of Phase 2 study participants** remain on the long-term extension phase of the study

## Next Steps

- **Meeting with FDA** to define regulatory path and Phase 2/3 study under development
- Data expected to be presented at **forthcoming meeting in 2025**

# REC-4881: MEK1/2 Inhibitor

A highly selective and potent MEK1/2 inhibitor for chemoprevention of Familial Adenomatous Polyposis (FAP)

## Program Status

- **First-in-Disease** and **Best-in-Class** potential for the treatment of FAP
- **Phase 1b** safety and futility analysis (polyp burden) anticipated in **H1 2025**

## Mechanism of Action

- **Loss of APC** drives FAP disease progression through **aberrant MAPK signaling**
- **REC-4881 is a highly potent, non-competitive, allosteric** MEK1 and MEK2 inhibitor
- Selectively blocks the activation of ERK (MAPK pathway)

## Thesis & Differentiation

- **Develop the first oral therapy** for the treatment of FAP
- Target **underlying genetic mechanisms** that drive the FAP disease progression
- Preferential distribution to GI tissues vs competitors which may enable greater activity at lower doses

## Unmet Need<sup>1</sup>

- **No approved systemic therapies and significant unmet need** for ~50,000 FAP patients beyond colectomy
  - Includes ~7,000<sup>2</sup> **advanced duodenal polyposis** patients in the US at high-risk of developing cancer

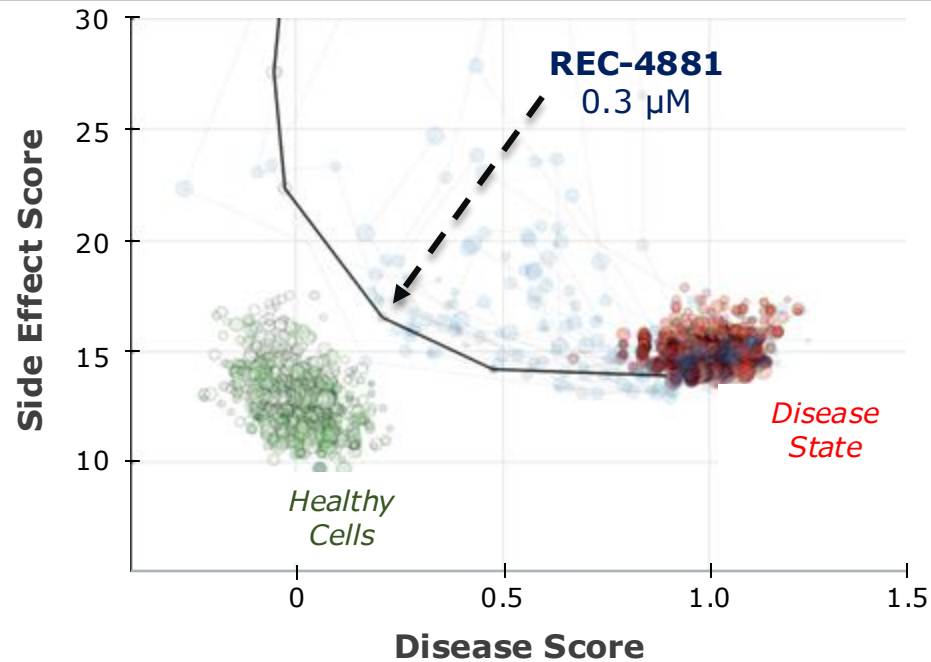
## Recursion Approach

- **Unbiased ML-aided phenotypic** drug screen in **human cancer cells**
- **Validated findings** in vivo demonstrating significant reductions in polyps and adenomas

# REC-4881 (MEK1/2 Inhibitor): Highly selective and potent molecule demonstrated superior in vivo efficacy versus celecoxib

## Recursion OS Insight

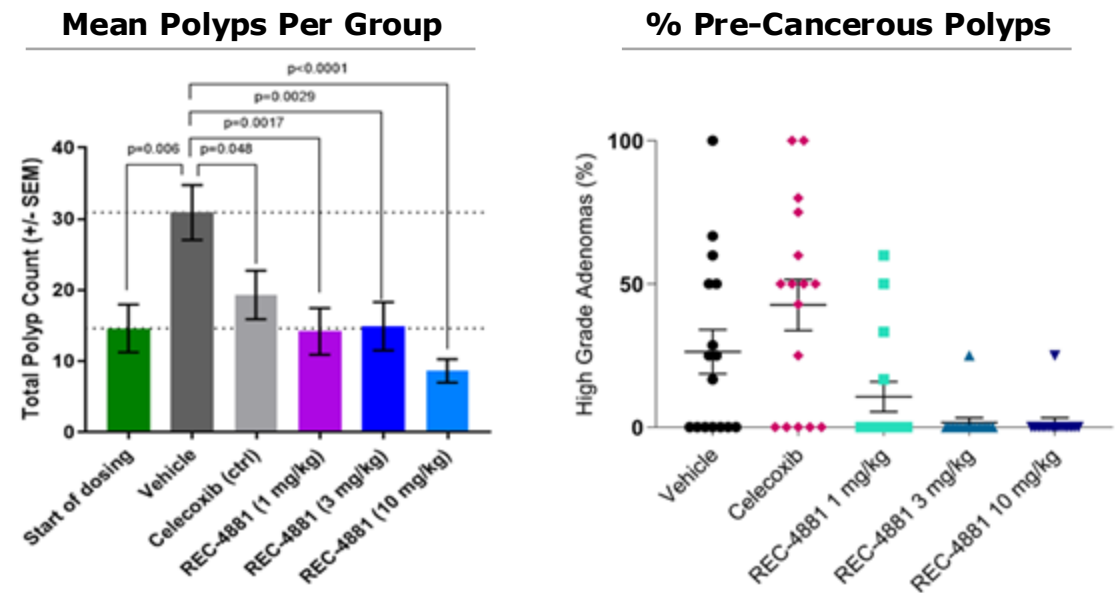
REC-4881 suppresses disease-inducing effects of APC mutations



- AI/ML extracts morphological features to distinguish “diseased” vs. “healthy” states
- Compounds co-treated with APC siRNA for 24 hours to find hits that reverse disease state back to healthy in a concentration-dependent manner

## Key Preclinical Data<sup>1</sup>

REC-4881 Decreases Polyp Count and Pre-Cancerous Adenomas



- Significantly reduces polyp count at all dose levels, outperforming celecoxib in *APC<sup>min/-</sup>* mouse
- Unlike celecoxib, REC-4881 reduces both polyp numbers and % of adenomas
- Meaningful efficacy seen at lowest dose tested (1mg/kg) – suggests potential for therapeutic activity at reduced systemic exposures

# REC-4881 (MEK1/2 Inhibitor): Study Design and Next Steps

## Development Strategy



TUPELO Phase 1b/2 study design

### REC-4881 Phase 1b Dose Escalation (N≤18)

4mg

8mg

12mg

RP2D

### Phase 2

Dose Expansion (N=30)

### Key Eligibility

- Confirmed *APC* mutation
- ≥ 55 years old
- Post-colectomy/proctocolectomy
- No cancer present
- Polyps in upper and lower GI

### Study Objectives:

- Identify RP2D
- Safety/tolerability
- Reduction in polyp burden at week 12

### REC-4881 Competitive Profile

- Early PD data indicates **4 mg dose** is pharmacologically active and well-tolerated
- **Fast Track Designation** in FAP granted by FDA in 2022
- **ODD** in US and EU

### Trial Update

- Futility – reduction in polyp burden; assessed after **10 evaluable** patients at the RP2D
- Futility analysis expected in **H1 2025**

# REV102: ENPP1 Inhibitor

A safe and highly selective ENPP1 inhibitor for Hypophosphatasia (HPP)

## Program Status

- Potential **First-in-Class** and **Best-in-Class** ENPP1 inhibitor for the treatment of patients with HPP
- Development candidate nomination expected in Q4 2024

## Mechanism of Action

- **Potent ENPP1** inhibitor is a **non-immunogenic** small molecule that restores PPI balance
- **Highly selective** ENPP1 inhibitor with low nM potency

## Thesis & Differentiation

- **ENPP1** inhibition is a **genetically validated** target in HPP models
- Potential for **first oral disease-modifying therapy (compared to multiple weekly injections)** without dose-limiting adverse events
- **Non-immunogenic** small molecule approach offering potentially safer solution than enzyme replacement therapy (ERT)
- REV102 offers a **more tolerable and affordable** option to ERTs

## Unmet Need<sup>1</sup>

- **~7,800 diagnosed prevalence** of HPP across US and EU5
- Many patients, particularly adults, may have difficulty accessing ERT
- Those who can access ERT face high treatment burden and tolerability hurdles
- Opportunity to **significantly reduce costs** and **treatment burden**

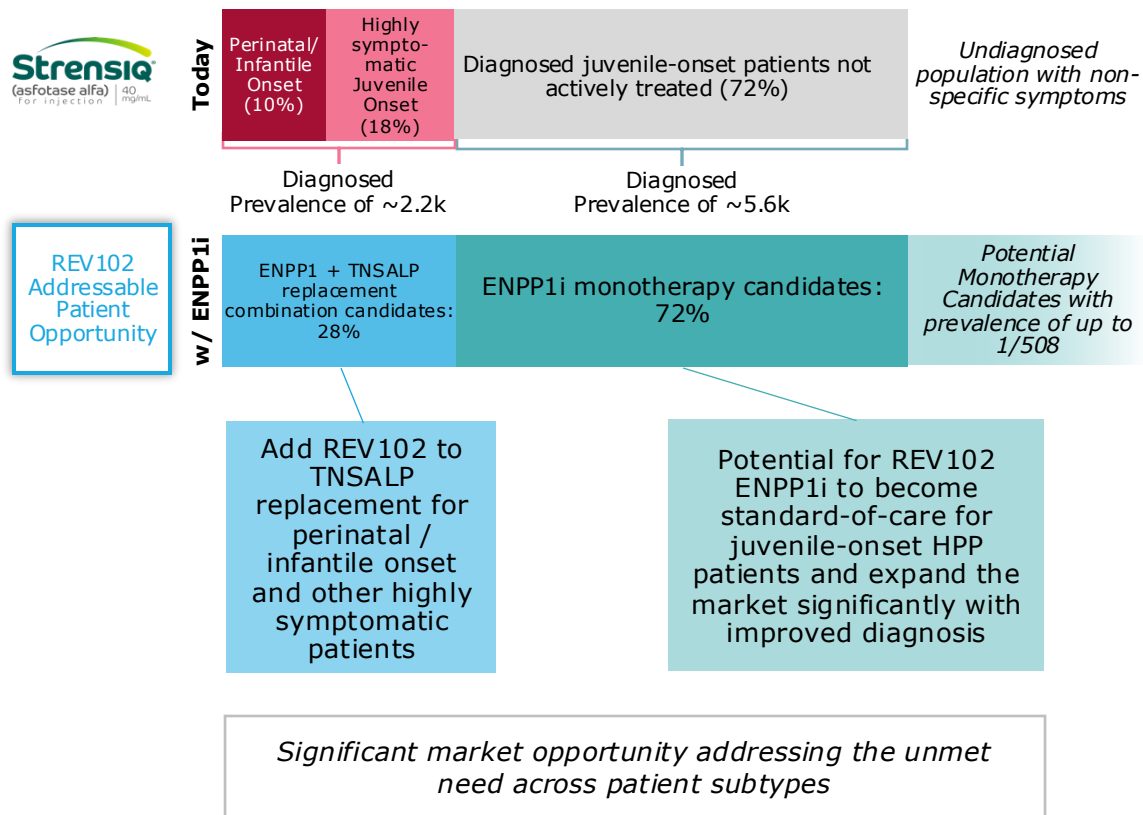
## Recursion Approach<sup>2</sup>

- **Precision designed for both high potency** and a lifetime of **chronic dosing**
- **Structurally distinct** differences vs competitor ENPP1 inhibitors
- **Maintain selectivity** and deliver a candidate with **high oral bioavailability** in the clinic

# REV102 (ENPP1 Inhibitor): OS insights validated using in vivo mouse model showing significant difference in restoring HPP biomarker that promotes bone mineralization

## Market Opportunity<sup>1</sup>

### Estimated Diagnosed Prevalence Hypophosphatasia Patients in US



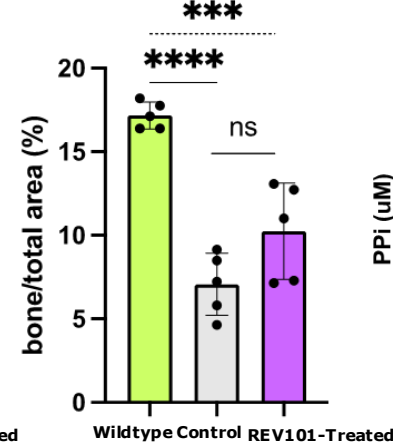
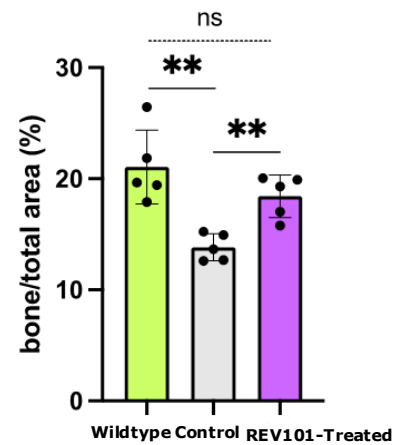
## Key Preclinical Data<sup>2</sup>

### Bone Morphometry

2D Analysis of Trabecular Bones

**L3 Vertebrae**  
(n=5, females)

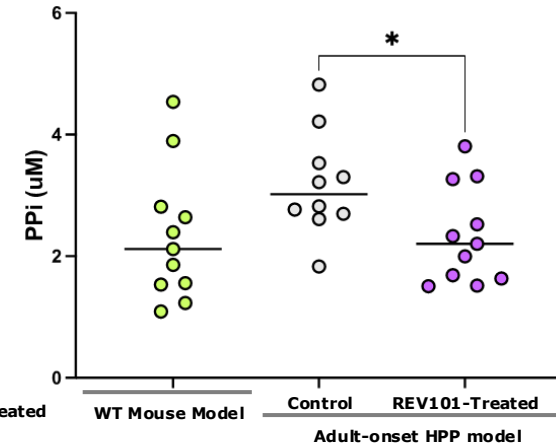
**Distal Femur**  
(n=5, males)



### Plasma Levels of PPI

After 100-Day Dosing

*PPI lowered by ~30% in adult HPP mice when treated with REV101*



Data is for REV101 (1st gen tool compound); compound being developed is REV102

### What's Next

- **Development candidate nomination** expected in Q4 2024



# REC-2282: Pan-HDAC Inhibitor

CNS-penetrating pan-HDAC inhibitor for the first oral therapeutic to treat Neurofibromatosis Type 2 (NF2)

## Program Status

- Potential **First-in-Disease and Best-in-Class** therapy for NF2 mutant meningioma
- **Data maturing** with PFS6 results expected H1 2025

## Mechanism of Action

- **Orally bioavailable, CNS penetrant, and potent** pan-HDAC inhibitor
- **Loss of Merlin** (*NF2*) leads to PI3K signaling and meningioma proliferation REC-2282 indirectly facilitates AKT dephosphorylation by disrupting the PP1-HDAC interaction

## Thesis & Differentiation

- **Develop the first therapeutic** for NF2 meningioma
- Highly selective molecule with favorable brain exposure and reduced risk of cardiac toxicity

## Unmet Need<sup>1</sup>

- **No approved therapy for ~33,000** NF2 meningioma patients beyond surgery
- Surgery only feasible in a limited number of patients and carries high rate of recurrence<sup>2</sup>

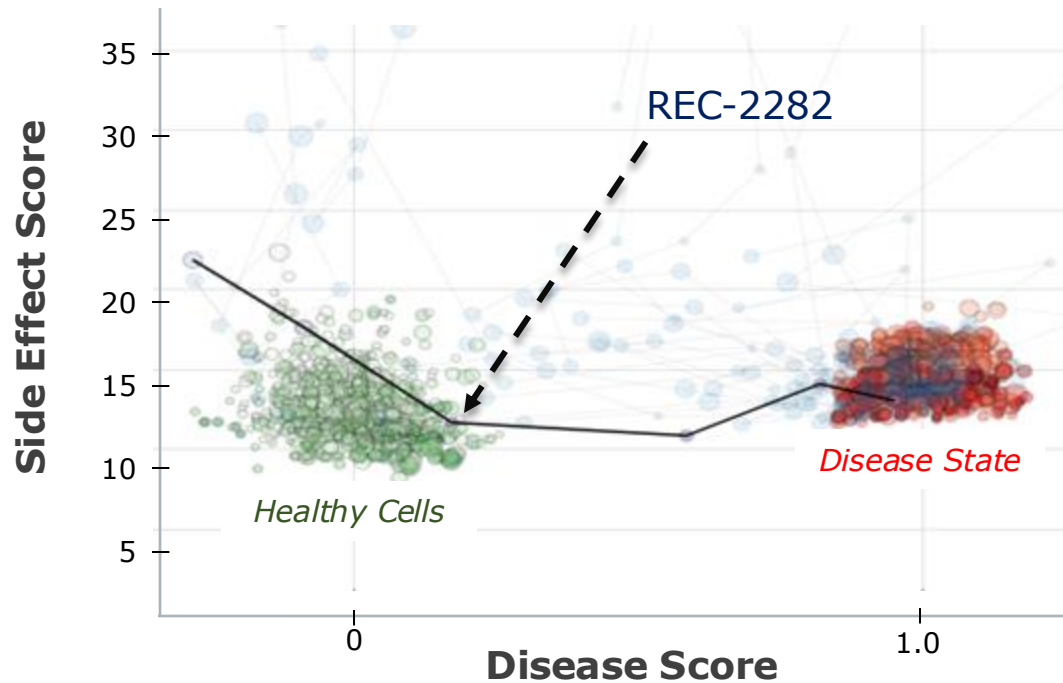
## Recursion Approach

- Unbiased **ML-aided phenomap insight and drug screen** in human cells
- Identify effective therapeutics that **rescue disease-inducing effects of *NF2*** loss

# REC-2282 (Pan-HDAC Inhibitor): Identified as a unique HDAC inhibitor in Recursion's unbiased screen modeling NF2 loss-of-function

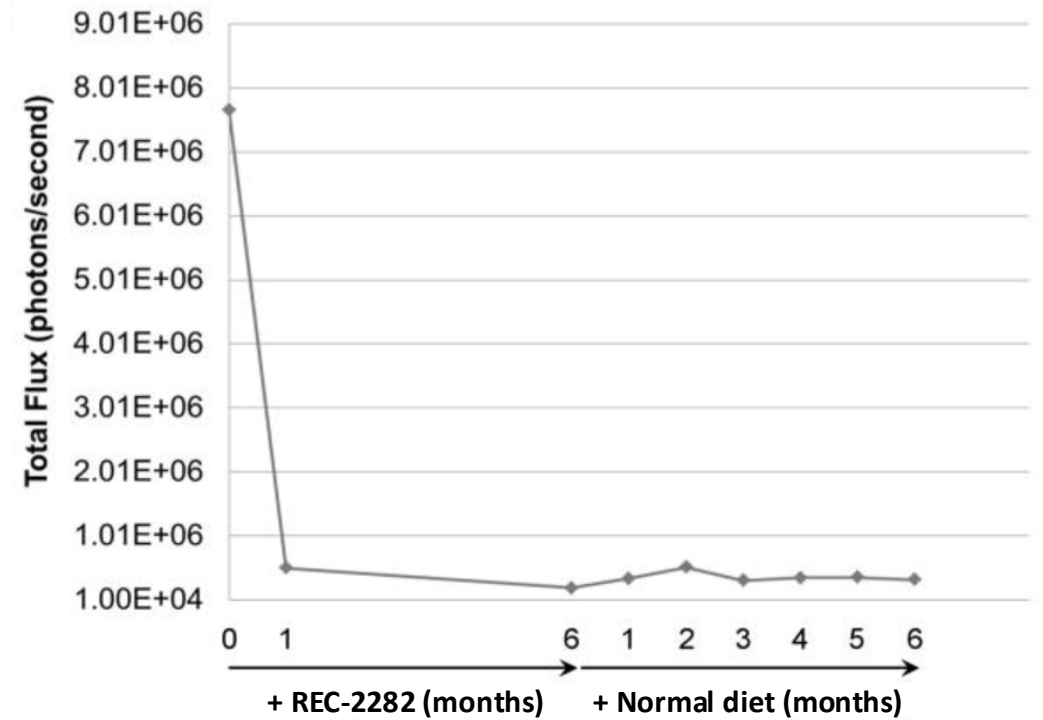
## Recursion OS Insight

REC-2282 demonstrates concentration-dependent reversal of NF2 loss



## Key Preclinical Data<sup>1</sup>

Prevents growth & regrowth of NF2-deficient meningioma model in mice



# REC-2282 (Pan-HDAC Inhibitor): Study Design and Next Steps

## Development Strategy



POPLAR Phase 2/3 study design

### REC-2282 Phase 2 Portion Open-label, 2-arm study

**24 Patients**  
Allocated 1:1

40 mg

60 mg

#### Primary Endpoint:

- PFS6

#### Secondary Endpoint:

- Safety, ORR

#### Key Eligibility

- MRI-confirmed progressive meningioma
- Sporadic meningioma with confirmed NF2 mutation
- Familial NF2 meningioma
- Have documented progression with past 24 months

### REC-2282 Competitive Profile

- Orally bioavailable and CNS penetrant
- **Fast Track Designation** in NF2 granted by FDA in 2021
- **ODD** in US and EU

### Trial Update

- Phase 2 **Data maturing**
- Futility analysis (PFS6) expected in **H1 2025**

PIPELINE

# Other areas of high unmet need

# REC-3964: *C. difficile* Toxin B Selective Inhibitor

Non-antibiotic selective toxin-inhibitor for the prevention of recurrent *C. difficile* infection (rCDI)

## Program Status

- **First-in-Class** therapy for prevention rCDI
- **First Patient Dosed** in the Phase 2 ALDER trial expected **in Q4 2024**
- Phase 2 update expected **in Q1 2026**

## Mechanism of Action

- **Highly potent, orally bioavailable** *C. diff* toxin B (TcdB) selective inhibitor
- **Selectively inhibits** catalytic activity of **bacterial** glucosyltransferase

## Thesis & Differentiation

- Develop the **first non-antibiotic oral therapy** that is safe and convenient
- **Selectively targets bacterial toxin** while sparing the host to minimize adverse events
- Preclinical efficacy demonstrates **superiority** in survival **versus bezlotoxumab**

## Unmet Need<sup>1</sup>

- **~175,000 cases of rCDI** with limited treatment options for high-risk population
- Ability to address populations **not eligible** for **FMT** or **microbiome-based therapies**

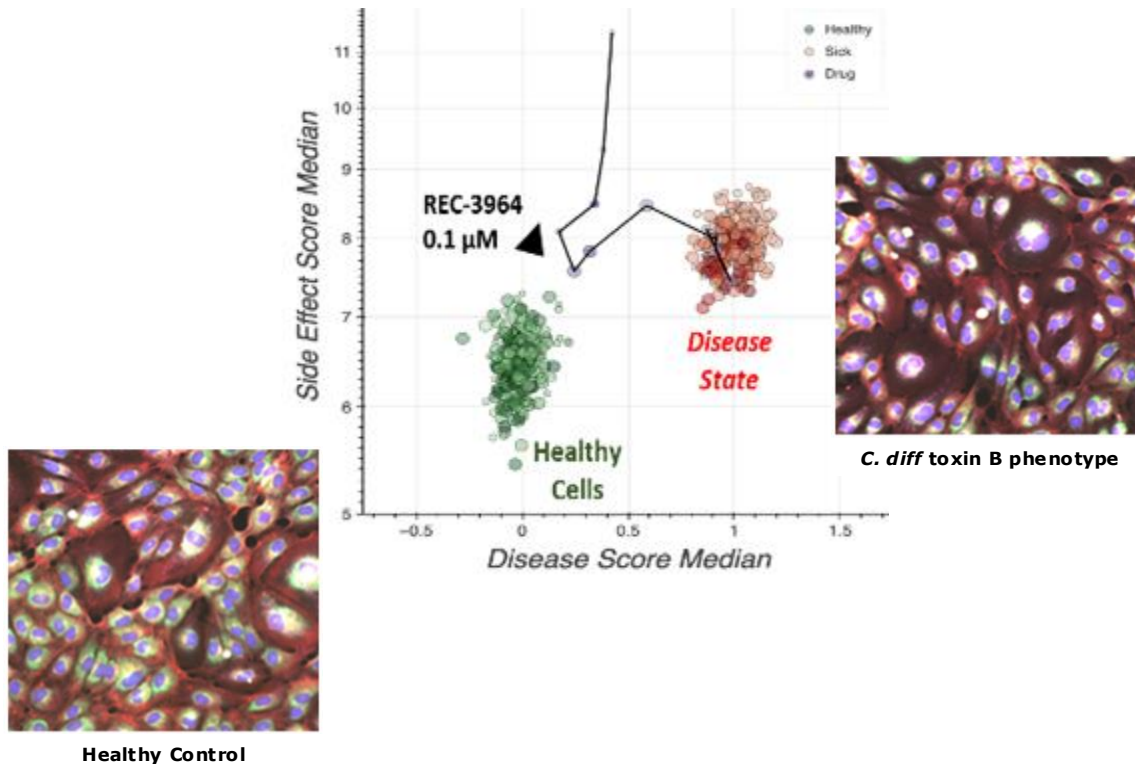
## Recursion Approach

- Unbiased **ML-aided conditional phenotypic** drug screen in **human cells**
- Identified **novel mechanisms** that mitigated the effect of *C. diff.* toxin B treatment

# REC-3964 (CDI TcdB Inhibitor): Identified as potential superior inhibitor compared to SOC in in vitro and in vivo preclinical studies

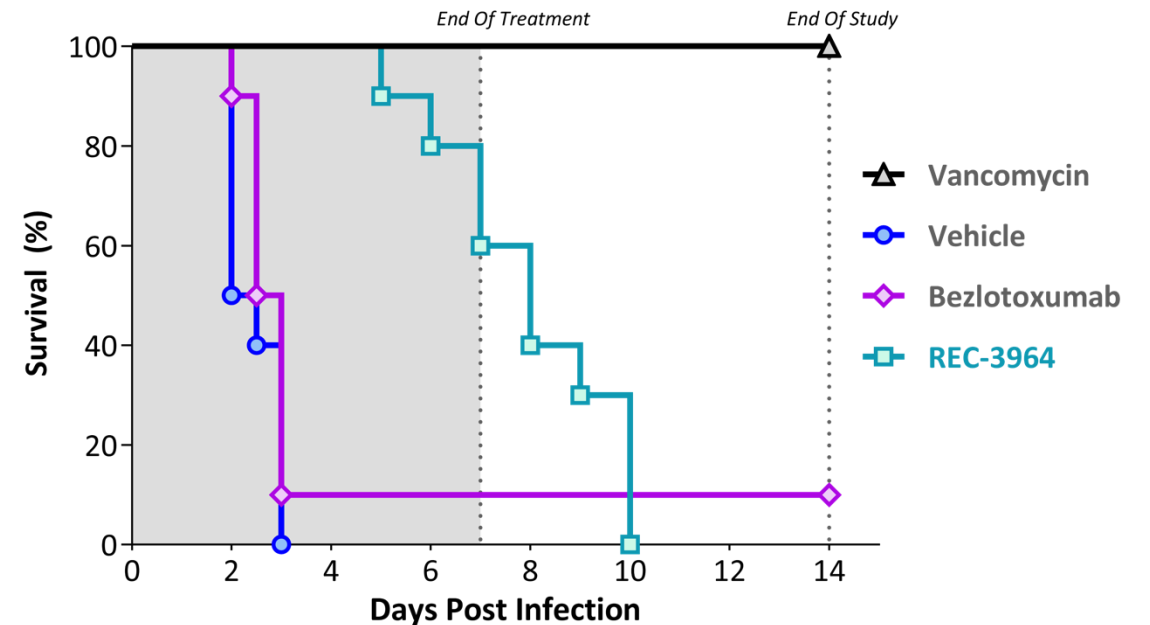
## Recursion OS Insight

REC-3964 potently inhibits Toxin B with some activity against Toxin A, while bezlotoxumab is specific to Toxin B



## Key Preclinical Data<sup>1</sup>

REC-3964 significantly extended survival vs. bezlotoxumab alone at the end of treatment (p<0.001, log rank test)



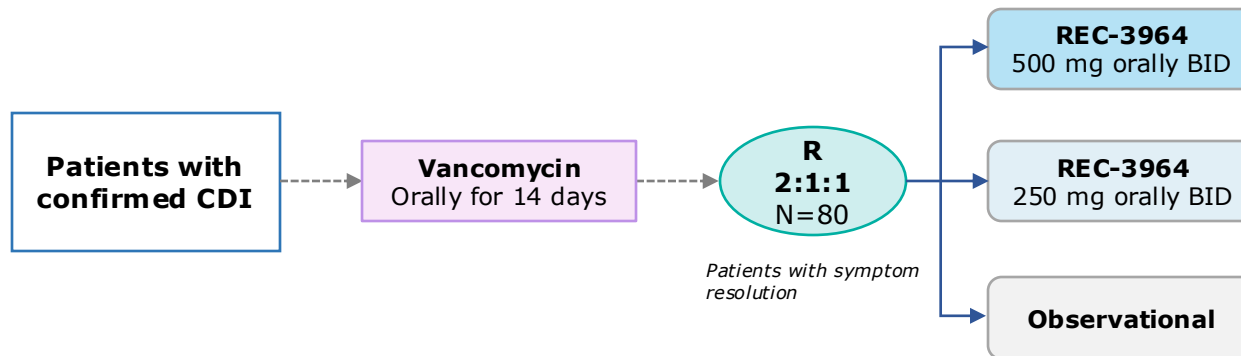
# REC-3964 (CDI TcdB Inhibitor): Study Design and Next Steps

## Development Strategy



ALDER Phase 2 study design

### REC-3964 Phase 2 Open-label, 2-arm study



#### Key Eligibility

- High-risk of CDI
- $\geq 3$  bowel movements in 24 hours
- Confirm CDI using EIA (toxin)
- No fulminant CDI
- No history of chronic diarrheal illness due to other causes

#### Primary Endpoint:

- Rate of recurrence

#### Secondary Endpoint:

- Additional efficacy measures
- Safety / tolerability
- PK

### REC-3964 Competitive Profile

- **Highly potent**, orally bioavailable
- Potential **First-in-Class** therapy for prevention of rCDI
- First non-antibiotic oral therapy

### Trial Update

- First Patient Dosed expected in **Q4 2024**
- Program update expected **Q1 2026**

# REC-4209: Target Epsilon

Highly potent and potential First-in-Class medicine for the treatment of Idiopathic Pulmonary Fibrosis (IPF)

## Program Status

- **First-in-Class** therapeutic for treatment of IPF
- **IND submission** expected in **2025**
- **Phase 1** study in healthy volunteers expected to initiate in **2025**

## Mechanism of Action

- Reversible, orally bioavailable, and potent Target Epsilon inhibitor
- Promotes tissue repair and reverses fibrosis by potentially modulating TGF- $\beta$

## Thesis & Differentiation

- Develop a novel preferred treatment option that is **safe** and **well-tolerated**
- **In vitro models suggest** capability of reversing the fibrotic process driving IPF progression

## Unmet Need<sup>1</sup>

- **~130,000** patients with IPF in the US
- Approved therapies show **modest slowing of IPF progression**
- **No improvement** in **survival** (mOS 3-5 years) or **quality of life** with current treatments

## Recursion Approach

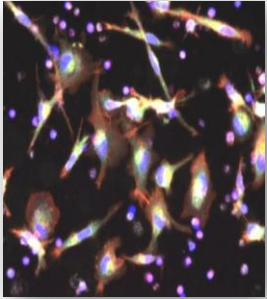
- Unbiased **ML-powered phenomap drug screen** in human cells
- Identify **novel mechanisms** that reversed the differentiation of fibrocytes



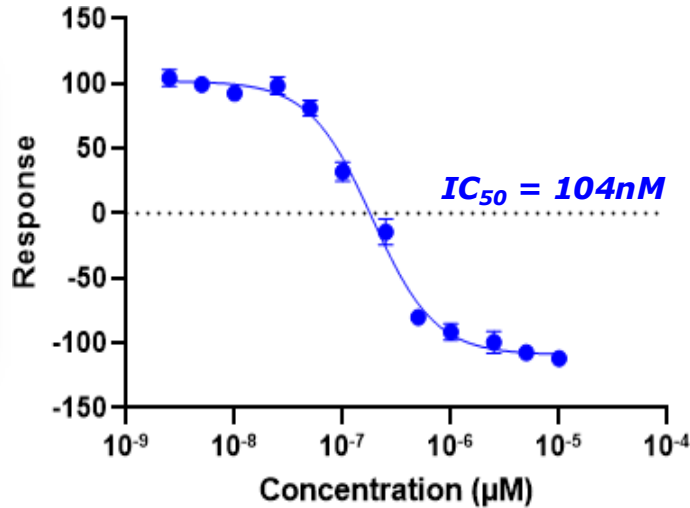
# REC-4209 (Target Epsilon): Identified as a novel mechanism in Recursion's screen with compelling preclinical efficacy demonstrated in bleomycin lung fibrosis mouse model

## Recursion OS Insights<sup>1</sup>

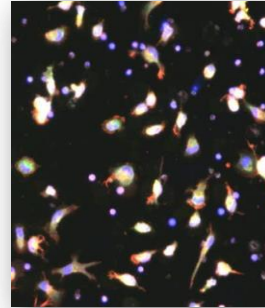
**Disease State**  
(Fibrocytes)



**In Vitro Fibrocyte Rescue Assay**

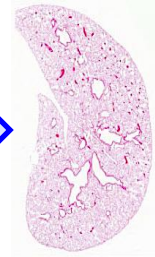


**Control State**  
(Pentraxin-2)



**Fibrotic Lung**

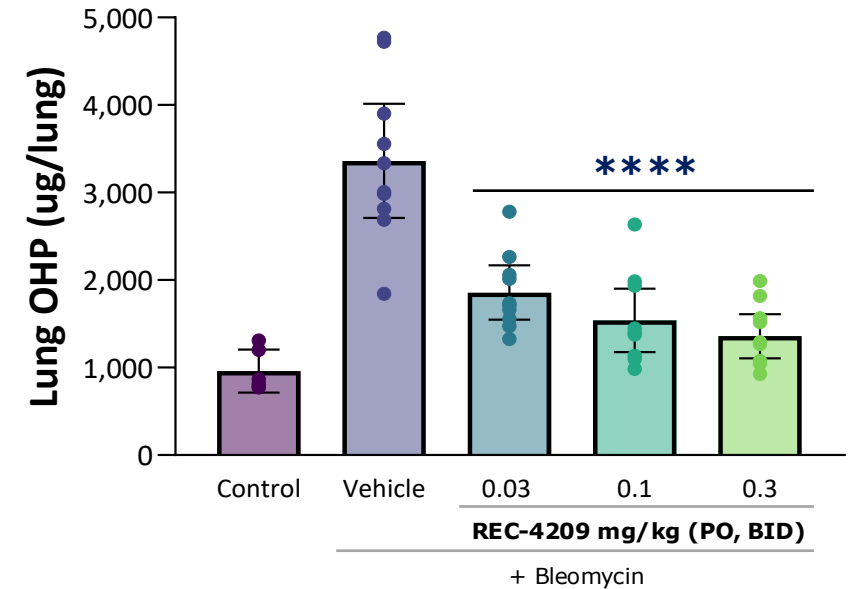
+ REC-4209



**Normal Lung**

## Key Preclinical Data<sup>2</sup>

**Total Lung Collagen**



- REC-4209 at low doses reduces total lung collagen by 45% to 60% versus vehicle mice

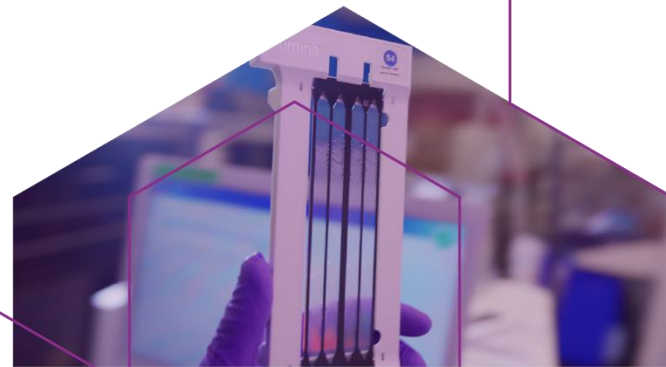
## What's Next

- IND-enabling studies ongoing**

65 1. Data on File  
2. Groups (n=10 per group; n=6 in control) compared against Vehicle. \*\*\*\*p<0.0001; one-way ANOVA with Tukey's multiple comparison test. Data reflects mean ± 95% CI


APPENDIX


# Partnerships & Data Strategy Details





# Exciting scientific collaborations span biopharma, tech & data

## Therapeutic discovery partnerships

 <p>Announced Dec. 2021</p>	<ul style="list-style-type: none"><li>• Up to or exceeding <b>\$300M in possible program milestones</b> for up to <b>40 programs</b></li><li>• <b>One program</b> and <b>one map</b> already <b>optioned</b></li><li>• <b>Mid- to high-single digit tiered royalties</b> on net sales</li></ul>
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 <p>Announced Jan. 2022</p>	<ul style="list-style-type: none"><li>• <b>\$100M upfront</b> with the potential of <b>\$5.2B in total milestones plus high-single digit to mid-teen tiered royalties</b></li><li>• Up to <b>15 novel small molecule candidates</b> across <b>oncology and immunology</b></li><li>• <b>New discovery stage program added</b> identified and initially advanced by Exscientia in Dec. 2023</li><li>• <b>3 programs advanced</b> through initial milestones</li></ul>
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 <p>Announced Sept. 2020 Updated Nov. 2023</p>	<ul style="list-style-type: none"><li>• <b>\$30M upfront</b> and <b>\$50M equity investment</b></li><li>• Increased per program milestones which may be <b>up to \$1.5B</b> in aggregate for up to 7 oncology programs</li><li>• <b>Low- to mid-single digit royalties</b> on net sales</li><li>• <b>Recursion owns all algorithmic improvements</b></li><li>• <b>First beta-user of LOWE</b></li></ul>
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 <p>Announced Sept. 2023</p>	<ul style="list-style-type: none"><li>• <b>\$20M upfront</b> at initiation for three projects with up to <b>\$674M in discovery, development, regulatory and sales-based milestones</b></li><li>• <b>Mid-single to low-double digit tiered royalties</b></li></ul>
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Platform, technology, and data partnerships


## Computation and ML/AI

 <p><b>NVIDIA</b></p> <p>Announced July 2023</p>	<ul style="list-style-type: none"><li>• <b>\$50M equity investment</b></li><li>• Partnership on <b>advanced computation</b> (e.g., foundation model development)</li><li>• <b>Priority access</b> to compute hardware or <b>DGXCloud Resources</b></li><li>• <b>BioHive-2</b>: helped design and build <b>next generation supercomputer</b></li></ul>
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 <p>Google Cloud</p> <p>Announced Oct. 2024</p>	<ul style="list-style-type: none"><li>• Includes exploring generative AI capabilities (including <b>Gemini models</b>) and driving improved search and access with <b>BigQuery</b></li><li>• Scaled compute resources, improved management of petabytes of RX data, and continued data privacy and security support</li><li>• Recursion will also explore <b>making some of its AI models available on Google Cloud</b></li></ul>
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## Real-world data access

<p><b>"TEMPUS</b></p> <p>Announced Nov. 2023</p>	<ul style="list-style-type: none"><li>• <b>Preferential access to &gt;20 PBs of real-world, multi-modal oncology data</b>, including DNA &amp; RNA sequencing and clinical outcome data for &gt;100,000 patients</li><li>• Ability to train <b>causal AI models</b> with utility in <b>target discovery, biomarker development &amp; patient selection</b></li><li>• <b>Opportunity to accelerate clinical trial enrollment</b> through broad clinical network</li></ul>
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 <p>Announced May 2024</p>	<ul style="list-style-type: none"><li>• Access to hundreds of thousands of de-identified records, including Helix's Exome+(R) <b>genomics &amp; longitudinal health data</b>, to train <b>causal AI models</b> and design <b>biomarker &amp; patient stratification strategies</b> across broad disease areas</li></ul>
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## Cheminformatics and chemical synthesis

 <p>Announced Dec. 2023</p>	<ul style="list-style-type: none"><li>• Utilizes Recursion's <b>predicted protein-ligand interactions for ~36B compounds</b> from Enamine's REAL Library</li><li>• Aim to generate <b>enriched screening libraries</b> &amp; co-brand customer offerings</li></ul>
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# Announcing OpenPhenom for non-commercial use

**AVAILABLE ON:**



**Hugging Face**



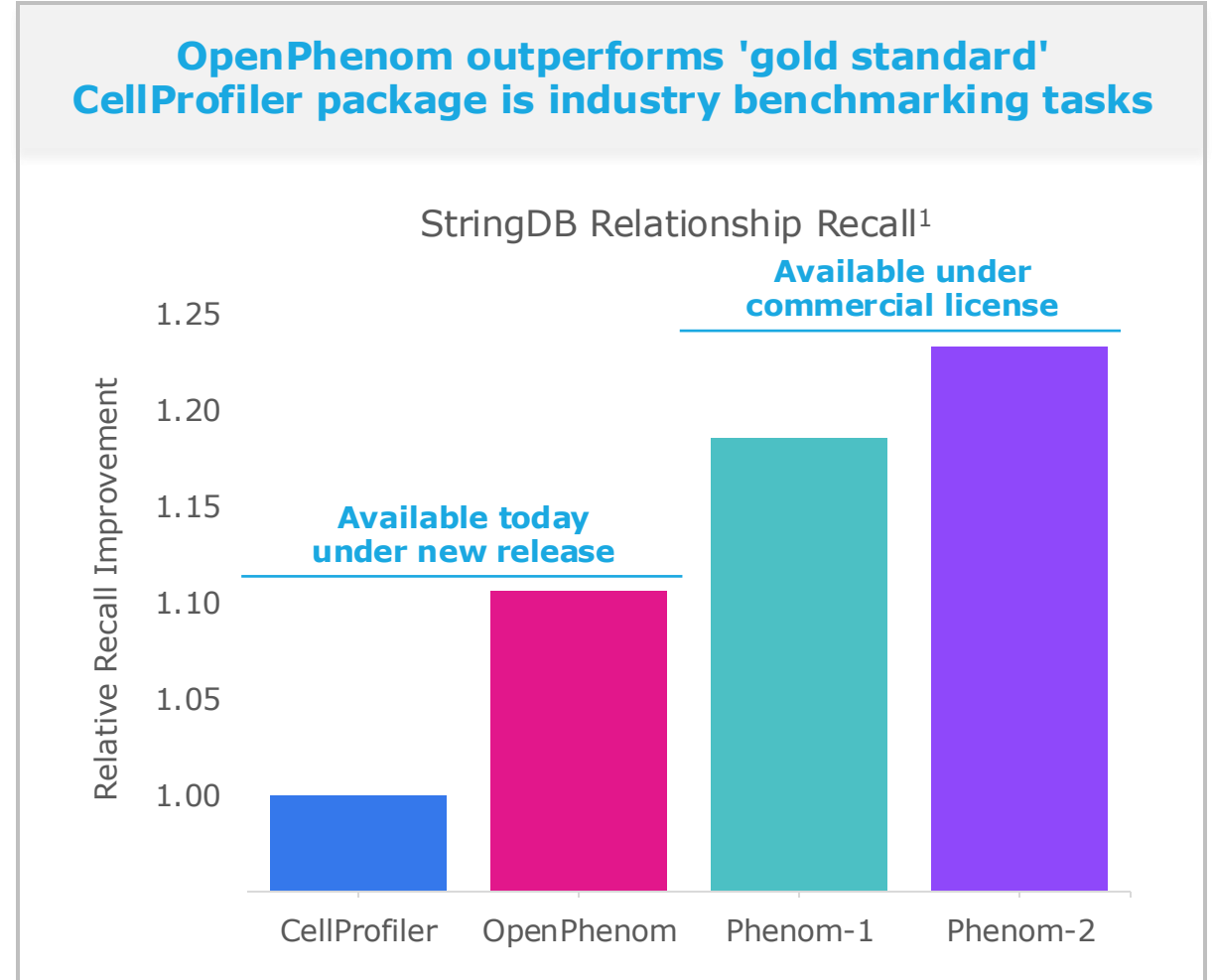
**Google Cloud**

**COMING SOON TO:**



**NVIDIA**

- Publicly accessible Foundation Model for microscopy data workflows
- Replaces legacy image segmentation and feature extraction software packages for non-commercial applications





Recursion<sup>®</sup>