

UNITED STATES  
SECURITIES AND EXCHANGE COMMISSION  
Washington, D.C. 20549

FORM 8-K

CURRENT REPORT  
Pursuant to Section 13 or 15(d)  
of the Securities Exchange Act of 1934

Date of Report (Date of earliest event reported): May 8, 2023

**Foghorn Therapeutics Inc.**  
(Exact name of registrant as specified in its charter)

Delaware  
(State or other jurisdiction of incorporation)

001-39634  
(Commission  
File Number)

47-5271393  
(IRS Employer Identification No.)

500 Technology Square, Ste 700  
Cambridge, MA  
(Address of principal executive offices)

02139  
(Zip Code)

(Registrant's telephone number, including area code): (617) 586-3100

Not Applicable  
(Former name or former address, if changed since last report)

Check the appropriate box below if the Form 8-K filing is intended to simultaneously satisfy the filing obligation of the registrant under any of the following provisions:

- Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425)
- Soliciting material pursuant to Rule 14a-12 under the Exchange Act (17 CFR 240.14a-12)
- Pre-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b))
- Pre-commencement communications pursuant to Rule 13e-4(c) under the Exchange Act (17 CFR 240.13e-4(c))

Securities registered pursuant to Section 12(b) of the Act:

Title of each class	Trading Symbol(s)	Name of each exchange on which registered
Common Stock, \$0.0001 par value per share	FHTX	The Nasdaq Global Market

Indicate by check mark whether the registrant is an emerging growth company as defined in Rule 405 of the Securities Act of 1933 (§230.405 of this chapter) or Rule 12b-2 of the Securities Exchange Act of 1934 (§240.12b-2 of this chapter).

Emerging growth company

If an emerging growth company, indicate by check mark if the registrant has elected not to use the extended transition period for complying with any new or revised financial accounting standards provided pursuant to Section 13(a) of the Exchange Act.

**Item 2.02 Results of Operations and Financial Condition.**

On May 8, 2023, Foghorn Therapeutics Inc. (the “Company”) issued a press release announcing certain of the Company’s financial results for the quarter ended March 31, 2023. A copy of the press release is furnished as Exhibit 99.1 to this Current Report on Form 8-K.

The information in this Item 2.02 (including Exhibit 99.1 attached hereto) is being furnished and shall not be deemed “filed” for purposes of Section 18 of the Securities Exchange Act of 1934, as amended (the “Exchange Act”), or otherwise subject to the liabilities of that section, nor shall it be deemed incorporated by reference into any filing by the Company under the Securities Act of 1933, as amended (the “Securities Act”), or the Exchange Act, except as expressly set forth by specific reference in such filing.

**Item 7.01 Regulation FD Disclosure.**

The Company is furnishing as Exhibit 99.2 to this Current Report on Form 8-K a presentation, dated May 2023, which the Company intends to use in meetings with or presentations to investors.

The information in this Item 7.01 (including Exhibit 99.2 attached hereto) is being furnished and shall not be deemed “filed” for purposes of Section 18 of the Exchange Act, or otherwise subject to the liabilities of that section, nor shall it be deemed incorporated by reference into any filing by the Company under the Securities Act or the Exchange Act, except as expressly set forth by specific reference in such filing.

**Item 9.01 Financial Statements and Exhibits.**

(d) Exhibits

Exhibit No.	Description
<a href="#">99.1</a>	<a href="#">Press Release issued on May 8, 2023</a>
<a href="#">99.2</a>	<a href="#">Investor Presentation dated May 2023</a>

**SIGNATURES**

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned hereunto duly authorized.

**FOGHORN THERAPEUTICS INC.**

By: /s/ Allan Reine  
Allan Reine, M.D.  
Chief Financial Officer

Date: May 8, 2023

### Foghorn Therapeutics Provides First Quarter 2023 Financial and Corporate Update

- Data from Phase 1 dose escalation study of FHD-286, a BRG1/BRM inhibitor, in metastatic uveal melanoma expected in the second quarter of 2023
  - Selective BRM, ARID1B, EP300 and CBP, targeting key regulators of gene expression, continue to advance towards IND
- Cash, cash equivalents and marketable securities of \$316.0 million, as of March 31, 2023, provides cash runway into the second half of 2025

**CAMBRIDGE, Mass. -- (GLOBE NEWSWIRE) -- May 8, 2023** -- Foghorn® Therapeutics Inc. (Nasdaq: FHTX), a clinical-stage biotechnology company pioneering a new class of medicines that treat serious diseases by correcting abnormal gene expression, today provided a financial and corporate update in conjunction with the Company's 10-Q filing for the quarter ended March 31, 2023. With an initial focus in oncology, Foghorn's Gene Traffic Control® Platform and resulting broad pipeline have the potential to transform the lives of people suffering from a wide spectrum of diseases.

"In the coming months, we anticipate the initial Phase 1 results for FHD-286 in metastatic uveal melanoma and we continue to advance our exciting early-stage oncology programs—including our BRM selective inhibitor, CBP, EP300 and ARID1B—toward the clinic while showcasing our ability to repeatedly generate selective chemical matter against important targets in oncology," said Adrian Gottschalk, President and Chief Executive Officer of Foghorn. "These programs have the potential to deliver novel therapies that hold tremendous value for large patient populations in a broad range of different cancers."

#### Key Recent Updates and Upcoming Milestones

- **FHD-286.** FHD-286 is a potent, selective inhibitor of the BRG1 and BRM subunits of the BAF chromatin remodeling complex where dependency on BRG1/BRM is well-established pre-clinically with multiple tumor types, including uveal melanoma, acute myelogenous leukemia (AML)/myelodysplastic syndrome (MDS), non-small cell lung cancer (NSCLC) and prostate cancer.
  - **mUM Update.** Phase 1 dose escalation of FHD-286 in metastatic uveal melanoma (mUM) continues to enroll patients per protocol. Top-line Phase 1 safety and efficacy data is expected in the second quarter of 2023.
  - **AML/MDS Update.** In August 2022, the U.S. Food and Drug Administration (FDA) placed a full clinical hold on the Phase 1 dose escalation study of FHD-286 in relapsed and/or refractory AML and MDS. The Company anticipates providing a regulatory update for FHD-286 in AML/MDS in the second quarter of 2023.
- **FHD-609 Update.** On April 24, 2023, Foghorn provided an update on the FHD-609 Phase 1 program in synovial sarcoma and SMARCB1-deleted tumors. (Link to press release [here](#)).

- **Differentiated Pipeline Advancement.** Foghorn continues to expand its platform and pipeline. The Company anticipates the potential for six new molecular investigational new drug (IND) applications in the next four years. The Company continues to progress programs for multiple targets which include chromatin remodeling complexes, transcription factors, helicases and other chromatin related factors. These targets include Selective BRM\* and wholly owned programs including CBP, EP300 and ARID1B, as well as other undisclosed targets, which combined could address more than 20 tumor types impacting more than 500,000 new patients annually.
- **Medical Conference Participation.** In April 2023, Foghorn participated at the 2023 American Association for Cancer Research Annual Meeting and the 18<sup>th</sup> Annual Drug Discovery Chemistry Meeting, highlighting preclinical data from its selective CBP and EP300 protein degrader programs, preclinical data for FHD-286 and its transcription factor and protein degradation capabilities. To access the presentations, please visit the “[Our Data](#)” section of the Foghorn website.
- **Strategic Collaborations.** During the first quarter of 2023, Foghorn continued to progress the Company’s strategic collaborations with two world-leading pharmaceutical companies, which validate the rigor of our science, highlight the importance of the targets we are tackling and confirm the relevance of the biology on which we are focused.
  - In December 2021, Foghorn entered into a strategic collaboration with Loxo@Lilly. In 2023, Foghorn anticipates continued progress across the collaboration including a co-development and co-commercialization agreement on the Selective BRM program\*, an additional undisclosed oncology target and three additional discovery programs. The Selective BRM program is on track to transition to Loxo@Lilly in the second half of 2023.
  - In July 2020, Foghorn entered into a strategic collaboration with Merck Sharp & Dohme. In 2023, Foghorn will continue to utilize its Gene Traffic Control platform to discover and develop novel therapeutics under the collaboration based on disruptors of a specified transcription factor target.

\*In December 2021, Foghorn announced a strategic collaboration with Loxo@Lilly to create novel oncology medicines. The collaboration includes a co-development and co-commercialization agreement for Foghorn’s Selective BRM oncology program and an additional undisclosed oncology target. In addition, the collaboration includes three discovery programs using Foghorn’s proprietary Gene Traffic Control platform.

#### **First Quarter 2023 Financial Highlights**

- **Strong Balance Sheet and Cash Runway.** As of March 31, 2023, the Company had \$316.0 million in cash, cash equivalents and marketable securities, which provides a cash runway into the second half of 2025.
- **Collaboration Revenues.** Collaboration revenue was \$5.3 million for the three months ended March 31, 2023, compared to \$3.9 million for the three months ended March 31, 2022. The increase year-over-year was primarily driven by revenue recognized under the Lilly collaboration agreement.

- **Research and Development Expenses.** Research and development expenses were \$30.0 million for the three months ended March 31, 2023, compared to \$24.5 million for the three months ended March 31, 2022. This increase was primarily due to costs associated with continued investment in R&D personnel and platform and early-stage research investments.
- **General and Administrative Expenses.** General and administrative expenses were \$8.6 million for the three months ended March 31, 2023, compared to \$7.2 million for the three months ended March 31, 2022. This increase was primarily due to an increase in investments to support the growing business which included increases in personnel-related costs and stock-based compensation expense.
- **Net Loss.** Net loss was \$30.5 million for the three months ended March 31, 2023, compared to a net loss of \$26.9 million for the three months ended March 31, 2022.

#### **About FHD-286**

FHD-286 is a highly potent, selective, allosteric and orally available, small-molecule, enzymatic inhibitor of BRG1 and BRM, two highly similar proteins that are the ATPases, or the catalytic engines across all forms of the BAF complex, one of the key regulators of the chromatin regulatory system. In preclinical studies, FHD-286 has shown anti-tumor activity across a broad range of malignancies including both hematologic and solid tumors. To learn more about these studies, please visit [ClinicalTrials.gov](https://ClinicalTrials.gov). (Link [here](#) for metastatic uveal melanoma and [here](#) for AML and MDS).

#### **About Uveal Melanoma**

Uveal (intraocular) melanoma (UM) is a rare eye cancer that forms from cells that make melanin in the iris, ciliary body and choroid. It is the most common eye cancer in adults. It is diagnosed in about 2,000 adults every year in the United States and occurs most often in lightly pigmented individuals with a median age of 55 years. However, it can occur in all races and at any age. UM metastasizes in approximately 50% of cases, leading to very poor prognosis.

#### **About AML**

Adult acute myeloid leukemia (AML) is a cancer of the blood and bone marrow and the most common type of acute leukemia in adults. AML is a diverse disease associated with multiple genetic mutations. It is diagnosed in about 20,000 people every year in the United States.

#### **About Foghorn Therapeutics**

Foghorn\* Therapeutics Inc. is discovering and developing a novel class of medicines targeting genetically determined dependencies within the chromatin regulatory system. Through its proprietary scalable Gene Traffic Control® platform, Foghorn is systematically studying, identifying and validating potential drug targets within the chromatin regulatory system. The Company is developing multiple product candidates in oncology. Visit our website at [www.foghornrx.com](http://www.foghornrx.com) for more information on the company, and follow us on [Twitter](#) and [LinkedIn](#).

#### **Forward-Looking Statements**

This press release contains “forward-looking statements” regarding the Company’s clinical programs for FHD-286 and FHD-609, including its efforts to resolve the full clinical hold relating to FHD-286 in AML and MDS, the anticipated timing of release of clinical data, its

collaborations with Lilly and Merck and its research pipeline, including the status of its Selective BRM program, the filing of INDs and its protein degrader efforts. Forward-looking statements include statements regarding the Company's clinical trials, product candidates and research efforts and other statements identified by words such as "could," "may," "might," "will," "likely," "anticipates," "intends," "plans," "seeks," "believes," "estimates," "expects," "continues," "projects" and similar references to future periods. Forward-looking statements are based on our current expectations and assumptions regarding capital market conditions, our business, the economy and other future conditions. Because forward-looking statements relate to the future, by their nature, they are subject to inherent uncertainties, risks and changes in circumstances that are difficult to predict. As a result, actual results may differ materially from those contemplated by the forward-looking statements. Important factors that could cause actual results to differ materially from those in the forward-looking statements include regional, national or global political, economic, business, competitive, market and regulatory conditions, including risks relating to our clinical trials and other factors set forth under the heading "Risk Factors" in the Company's Annual Report on Form 10-K for the year ended December 31, 2022, as filed with the Securities and Exchange Commission. Any forward-looking statement made in this press release speaks only as of the date on which it is made.

**Condensed Consolidated Balance Sheets**  
(In thousands)

	March 31, 2023	Dec. 31, 2022
Cash, cash equivalents and marketable securities	\$ 315,970	\$ 345,798
All other assets	56,913	59,085
<b>Total assets</b>	<b>\$ 372,883</b>	<b>\$ 404,883</b>
Deferred revenue, total	\$ 331,511	\$ 336,820
All other liabilities	65,958	67,951
<b>Total liabilities</b>	<b>397,469</b>	<b>404,771</b>
<b>Total stockholders' equity (deficit)</b>	<b>(24,586)</b>	<b>112</b>
<b>Total liabilities and stockholders' equity</b>	<b>\$ 372,883</b>	<b>\$ 404,883</b>

**Condensed Consolidated Statements of Operations**  
(In thousands, except share and per share amounts)

	Three Months Ended March 31,	
	2023	2022
Collaboration revenue	\$ 5,309	\$ 3,920
Operating expenses:		
Research and development	29,985	24,508
General and administrative	8,641	7,216
<b>Total operating expenses</b>	<b>38,626</b>	<b>31,724</b>
<b>Loss from operations</b>	<b>(33,317)</b>	<b>(27,804)</b>
<b>Total other income, net</b>	<b>3,389</b>	<b>890</b>
<b>Provision for income taxes</b>	<b>(560)</b>	<b>—</b>
<b>Net loss</b>	<b>\$ (30,488)</b>	<b>\$ (26,914)</b>
Net loss per share attributable to common stockholders—basic and diluted	\$ (0.73)	\$ (0.65)
<b>Weighted average common shares outstanding—basic and diluted</b>	<b>41,811,087</b>	<b>41,370,186</b>

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# FCGHORN<sup>®</sup>

## THERAPEUTICS

### CORPORATE OVERVIEW

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Leveraging unique insights into the chromatin regulatory system to pioneer a new class of precision therapies in oncology and beyond

May 2023

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## FORWARD LOOKING STATEMENTS

This presentation contains forward-looking statements. All statements other than statements of historical facts contained in this presentation are forward-looking statements. In some cases, you can identify forward-looking statements by terms such as "could," "may," "might," "will," "likely," "anticipates," "intends," "plans," "seeks," "believes," "estimates," "expects," "continues," "projects" or the negative of these terms or other similar expressions, although not all forward-looking statements contain these words. Forward-looking statements include, but are not limited to, statements concerning: the potential outcomes from our collaboration agreements with Lilly and Merck; the initiation, timing, progress and results of our research and development programs and preclinical and clinical trials, including the potential resolution of the full clinical hold and anticipated timing of release of clinical data; our ability to advance product candidates that we may develop and to successfully complete preclinical and clinical studies; our ability to leverage our initial programs to develop additional product candidates using our Gene Traffic Control Platform; the impact of the COVID-19 pandemic and other exogenous factors on our and our collaborators' business operations, including our research and development programs and preclinical studies; developments related to our competitors and our industry; our ability to expand the target populations of our programs and the availability of patients for clinical testing; our ability to obtain regulatory approval for FHD-286 and any future product candidates from the FDA and other regulatory authorities; our ability to identify and enter into future license agreements and collaborations; our ability to continue to rely on our CDMOs and CROs for our manufacturing and research needs; regulatory developments in the United States and foreign countries; our ability to attract and retain key scientific and management personnel; the scope of protection we are able to establish, maintain and enforce for intellectual property rights covering FHD-286, our future products and our Gene Traffic Control Platform; and our use of proceeds from capital-raising transactions, estimates of our expenses, capital requirements and needs for additional financing. Any forward-looking statements represent the Company's views only as of today and should not be relied upon as representing its views as of any subsequent date. The Company explicitly disclaims any obligation to update any forward-looking statements. The Company's business is subject to substantial risks and uncertainties.

# FIRST-IN-CLASS PRECISION MEDICINES TARGETING MAJOR UNMET NEEDS IN CANCER



## LEADER IN NEW AREA OF CANCER BIOLOGY

Foghorn is a **leader in targeting chromatin biology**, which has unique potential to address underlying dependencies of many genetically defined cancers

Broad pipeline of **over 15 programs** across a range of targets and modalities



## LARGE MARKET POTENTIAL

Chromatin biology is implicated in up to **50% of tumors**, potentially impacting **~2.5 million patients**

Foghorn's current pipeline potentially addresses **more than 500,000** of these patients



## WELL-FUNDED

**\$316.0 million** in cash and equivalents

(as of 03/31/2023)

Provides **runway into H2'25**



## SIGNIFICANT VALUE DRIVERS IN 2023

Initial clinical data in uveal melanoma with **FHD-286** expected **Q2'23**

AML/MDS study with FHD-286 on full clinical hold, development **clarity anticipated in Q2'23**



## COLLABORATIONS WITH MAJOR ONCOLOGY PLAYERS

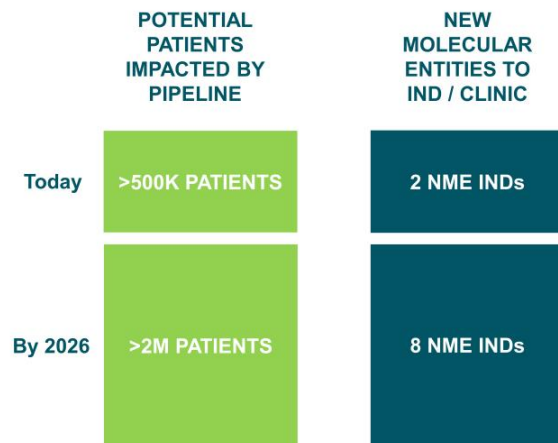
Strategic collaboration with Loxo Oncology at Lilly; **\$380 million upfront**; 50/50 U.S. economic split on two lead programs

Merck collaboration to drug single specified transcription factor target; **\$15 million upfront** and up to **\$410 million** in milestones

## FOGHORN: SIGNIFICANT VALUE CREATION OPPORTUNITIES

Potential Impact in >500K Patients Across More Than 20 Tumor Types with 6 Potential New INDs by 2026

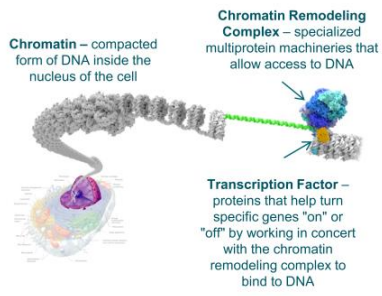
- Validated platform with first-in-class target in the clinic (FHD-286), with Phase 1 dose escalation data expected in Q2 2023
- At least **6** additional potential NME **INDs** by 2026
- **>20** genetically defined tumor types in **over 500K** patients – includes lung, prostate, bladder, ovarian, colorectal, breast
- Opportunity for additional partnerships



# UNIQUE INSIGHTS INTO CHROMATIN BIOLOGY

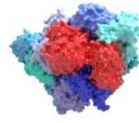
Untapped Area for Novel Targets and Therapeutics

## CHROMATIN REGULATORY SYSTEM CRITICAL FOR GENE EXPRESSION



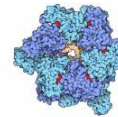
## NOVEL TARGETS GUIDED BY GENETIC DEPENDENCIES

**Chromatin Remodeling Complex** Mutations / Overexpression

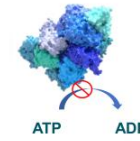


**Transcription Factor** Mutations / Overexpression

**Helicases & Other Chromatin Binding Proteins** involved in gene expression / function



## TAILORED DRUGGING APPROACHES



**Enzymatic Inhibitors**  
Highly selective and allosteric small molecule inhibitors

**Targeted Protein Degradation**  
Molecular glue and bi-functional protein degraders

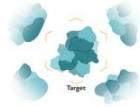


**Transcription Factor Disruptors**  
Disrupt interactions between chromatin remodeling complexes and transcription factors



# FOGHORN'S VALIDATED GENE TRAFFIC CONTROL® PLATFORM

Integrated, Scalable, Efficient – Repeatable Paradigm



## UNIQUE TARGETS

Deep Mechanistic Understanding of the Chromatin Regulatory System

### What to Drug:

Identify disease dependencies



## SPECIALIZED APPROACH

Biochemistry, Biophysics and Assays of Large Complexes and Proteins

### Where to Drug:

Engineer selectivity via unique assays and protein capabilities



## SELECTIVE THERAPEUTICS

Small Molecule and Degradation Platform

### How to Drug:

Biology first - small molecule modality agnostic

Enzymatic Inhibitors

Targeted Protein Degraders

Transcription Factor Disruptors

# BROAD PIPELINE ACROSS A RANGE OF TARGETS AND MODALITIES

Precision Oncology / Breadth and Depth / Over 15 Programs

Modality	Program	Discovery	Phase 1	Phase 2	Phase 3	Commercial Rights	Patient Population*
Enzyme Inhibitors	FHD-286 (BRG1/BRM)*	AML & MDS	[Progress bar]			FCGHORN THERAPEUTICS	Over 27,000
	FHD-286 (BRG1/BRM)	Uveal Melanoma	[Progress bar]			FCGHORN THERAPEUTICS	Over 5,000
	Selective BRM	BRG1 Mutated Cancers, e.g., NSCLC & Bladder	[Progress bar]			LOXO THERAPEUTICS, FCGHORN THERAPEUTICS	Over 100,000
Protein Degraders	Selective BRM	BRG1 Mutated Cancers, e.g., NSCLC & Bladder	[Progress bar]			LOXO THERAPEUTICS, FCGHORN THERAPEUTICS	Over 100,000
	Selective ARID1B	ARID1A Mutated Cancers, e.g., Ovarian, Endometrial & Colorectal	[Progress bar]			FCGHORN THERAPEUTICS	Over 175,000
	Selective CBP	EP300 Mutated Cancers, e.g., Prostate, Bladder, Colorectal, Breast	[Progress bar]			FCGHORN THERAPEUTICS	Over 100,000
	Selective EP300	CBP Mutated & Subsets of EP300 Dependent Cancers	[Progress bar]			FCGHORN THERAPEUTICS	Over 100,000
	FHD-609 (BRD9)^	Synovial Sarcoma & SMARCB1-Loss Tumors	[Progress bar]			FCGHORN THERAPEUTICS	Over 2,800
Transcription Factor Disruptors	Undisclosed	Undisclosed	[Progress bar]			FCGHORN THERAPEUTICS	
	Undisclosed	Undisclosed	[Progress bar]			MERCK	
Partnered Program	Undisclosed	Undisclosed	[Progress bar]			LOXO THERAPEUTICS, FCGHORN THERAPEUTICS	
	3 Discovery Programs	3 Undisclosed Programs	[Progress bar]			LOXO THERAPEUTICS, FCGHORN THERAPEUTICS	

\* Per year incidence in the U.S., EU5, Japan | \* On full clinical hold | ^ On partial clinical hold



# Inhibition of the BRG1 and BRM Subunits of the BAF Complex

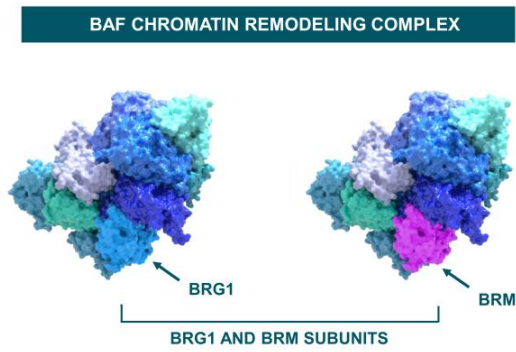
## IN PHASE 1 DOSE ESCALATION FOR METASTATIC UVEAL MELANOMA & AML/MDS

FHD-286 is a Potent, Selective, Allosteric, Small Molecule Inhibitor of the BRG1 and BRM Subunits of the BAF Complex

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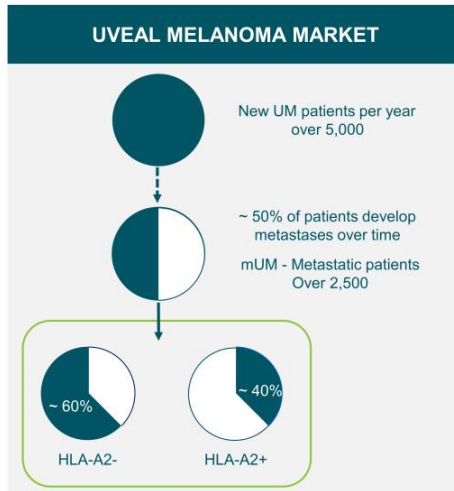
## TARGETING BAF DEPENDENCY IN CANCER



- BRM / BRG1 is the engine (ATPase) of the BAF chromatin remodeling complex
- Dependency on BRM / BRG1 is **well-established with multiple tumor types**, including uveal melanoma, AML / MDS, NSCLC and prostate
- Foghorn's lead asset targeting BRM / BRG1, **FHD-286, is a potent, selective, allosteric, small molecule inhibitor of the BRG1 and BRM subunits** of the BAF complex
- In Phase 1 dose escalation for uveal melanoma & AML / MDS

# SIGNIFICANT UNMET NEED IN UVEAL MELANOMA

Most Common Form of Eye Cancer



## UVEAL MELANOMA OVERVIEW

### Market Opportunity:

- Over 2,500 new metastatic UM patients impacted per year in the U.S. / over 5,000 U.S. and E.U.
- Potential additional opportunity in the adjuvant and neo-adjuvant settings

### Limited Treatment Options:

- Treatment options include enucleation, checkpoint inhibitors, KIMMTRAK and chemotherapy/radiation
- KIMMTRAK is indicated for HLA-A2+ haplotype (~40% of the metastatic patient population)

# FHD-286 FOR METASTATIC UVEAL MELANOMA

## Clinical Development Plan

### PHASE 1 DOSE ESCALATION STUDY

- 3+3 cohort design
- Retrospective biomarker analysis to further evaluate safety and efficacy
- Assess safety, PK, biomarkers and therapeutic activity
- Identify dose(s) for expansion

### PHASE 1 EXPANSION STUDIES

- Evaluate identified dose(s)
- Consider refined patient population, if necessary
- Consider exploration of combination partners
- Assess safety, PK, biomarkers and therapeutic activity

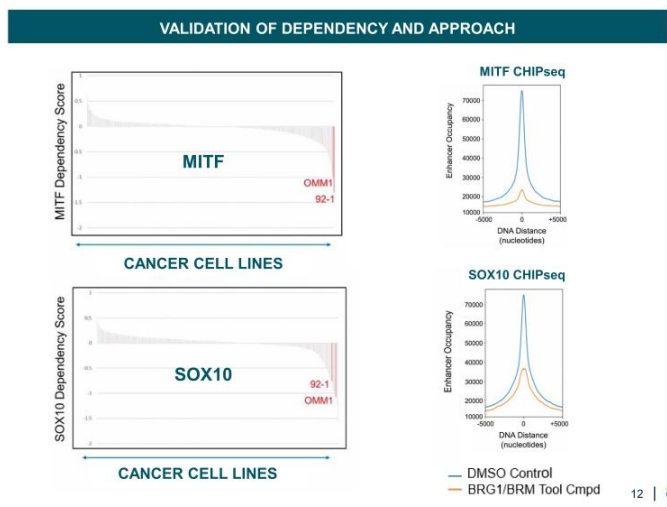
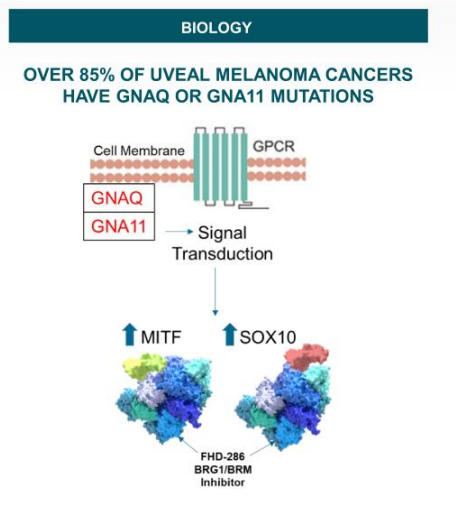
### POTENTIAL DEFINITIVE EFFICACY TRIALS & INDICATION EXPANSION

- Potential for entry into definitive efficacy trials in metastatic UM
- Potential for indication expansion

Initial clinical data in uveal melanoma with FHD-286 expected Q2'23

# THERAPEUTIC RATIONALE FOR UVEAL MELANOMA

Dependency on Two Lineage Transcription Factors: MITF / SOX10

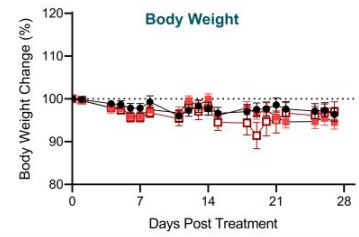
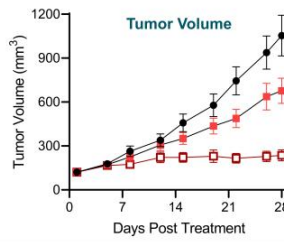


# DOSE-DEPENDENT TUMOR REGRESSION IN UVEAL MELANOMA CDX MODELS AT TOLERATED DOSES WITH FHD-286

## MP-46 UVEAL MELANOMA CDX MODEL

Dose-dependent tumor growth inhibition

Well-tolerated

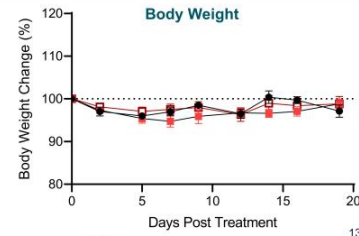
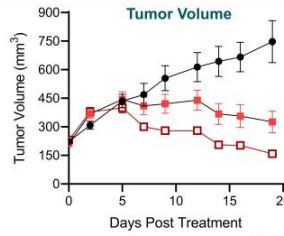


## 92-1 UVEAL MELANOMA CDX MODEL

Dose-dependent tumor growth inhibition

Tumor regression at 1.5 mg / kg, PO, QD

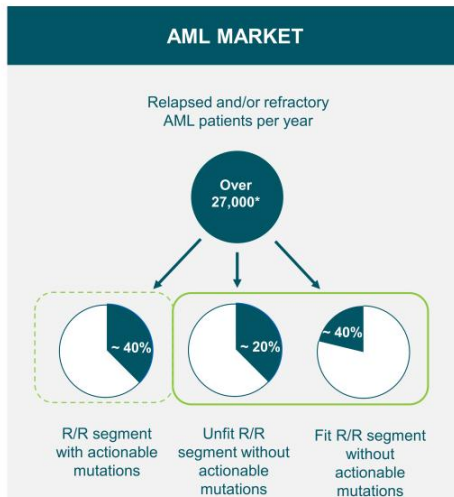
Well-tolerated



● Vehicle ■ FHD-286, 0.5 mg/kg, PO, QD □ FHD-286, 1.5 mg/kg, PO, QD

# SIGNIFICANT UNMET NEED REMAINS IN R/R AML & MDS

Most Common Type of Acute Leukemia in Adults



## AML OVERVIEW

### Mutation:

- Elevated BRG1-BAF / TF activity in AML blast cells

### Market Opportunity:

- Over 27,000 relapsed and/or refractory patients impacted per year\*

### Treatment Options:

- Limited options for relapsed and/or refractory patients without actionable mutations

\* Per year incidence in the U.S., EU5, Japan

# FHD-286 FOR RELAPSED/REFRACTORY AML & MDS

## Clinical Development Plan

### PHASE 1 DOSE ESCALATION STUDY

- 3+3 cohort design
- Retrospective biomarker analysis to further evaluate safety and efficacy
- Assess safety, PK, biomarkers and therapeutic activity
- Identify dose(s) for expansion

### PHASE 1 EXPANSION STUDIES

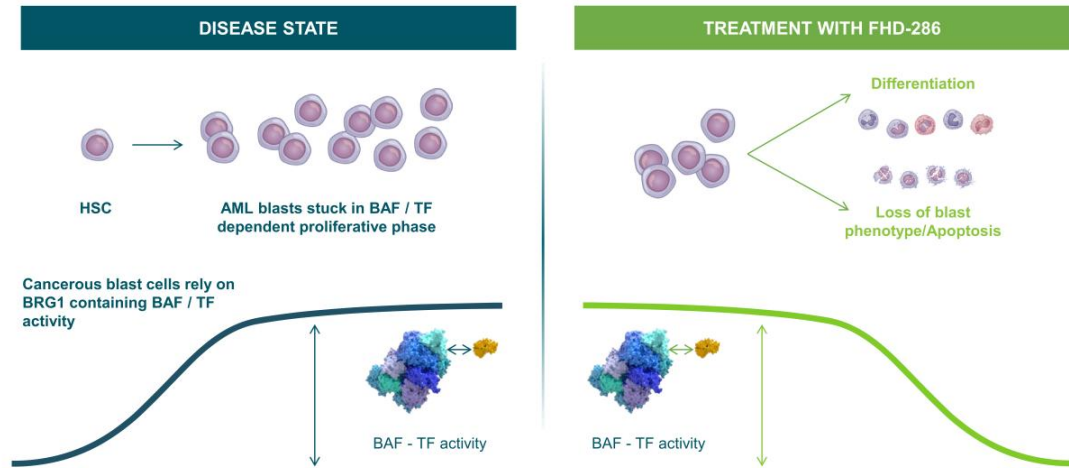
- Evaluate identified dose(s)
- Consider refined patient population if necessary
- Consider exploration of combination partners
- Assess safety, PK, biomarkers and therapeutic activity

### POTENTIAL DEFINITIVE EFFICACY TRIALS & INDICATION EXPANSION

- Potential for entry into definitive efficacy trials in AML / MDS
- Potential for indication expansion

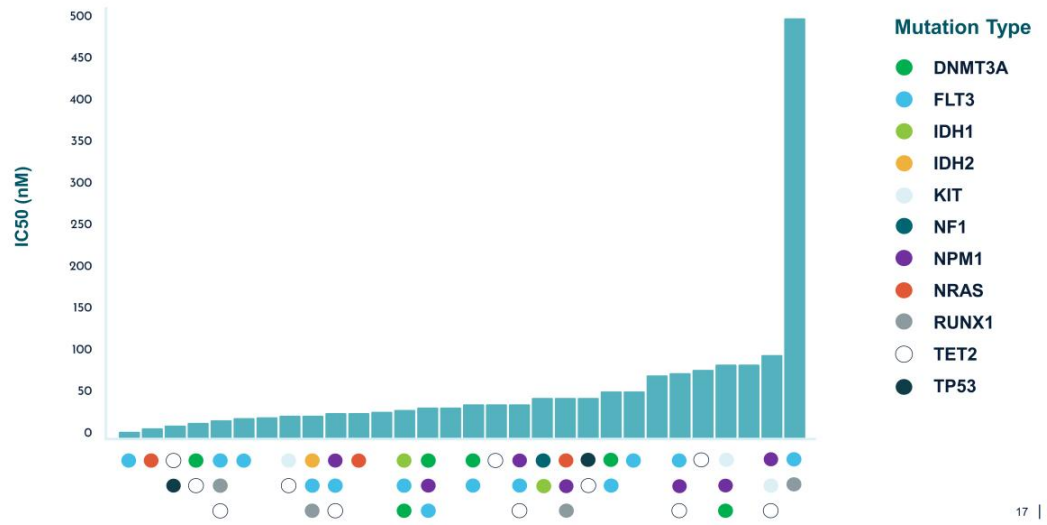
AML / MDS study with FHD-286 on full clinical hold, development clarity anticipated in Q2'23

# AML: DEPENDENCY ON BRG1 / LINEAGE TF INTERACTIONS





## FHD-286 SHOWS EFFECT ACROSS A BROAD RANGE OF MUTATIONS IN AML PATIENT-DERIVED SAMPLES



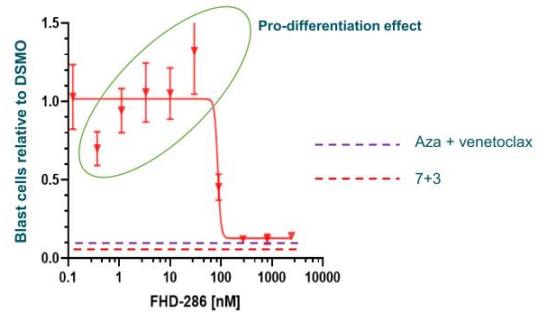
## PRECLINICAL FHD-286 DATA SHOWS EFFICACY ACROSS AML PATIENT-DERIVED SAMPLES

Notable Patient ID	Deep Response	Pathology Review	Disease Status
1690AML1	Y	AML	Secondary
1695AML1	Y	AML/MDS	Secondary
1696AML1	Y	AML	Secondary
1701AML1	Y	AML	Secondary
1893AML1	Y	AML	R/R
1899AML1	Y	AML	R/R
1990pAML1	Y	AML	R/R
1991pAML1	Y	AML	de novo
2041AML1	Y	N/A	de novo
2043pAML1	Y	AML	R/R
2059AML1	Y	AML	R/R
1682AML1	~	N/A	N/A
1689AML1	~	AML/MDS	de novo
1684AML1	N	CML	R/R
1924AML1	N	AML/MDS	R/R

Y = Deep reduction in blast cells    ~ = Partial reduction    N = No response



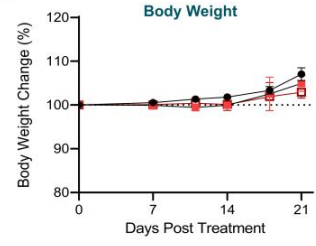
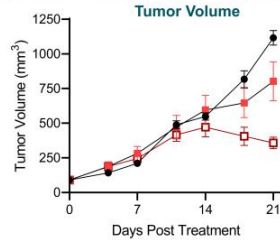
1695AML1 – BM-secondary AML



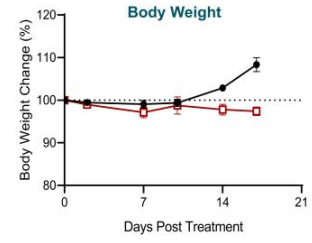
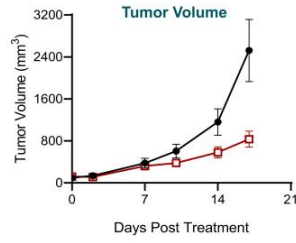
- Response observed in a majority of primary AML samples, irrespective of prior treatment or disease stage
- Additional data set from patient-derived samples demonstrates mutation-agnostic responses

## DOSE-DEPENDENT TUMOR GROWTH INHIBITION OBSERVED WITH FHD-286 TREATMENT IN AML CDX MODELS

**MV4-11 CDX Model**  
(FLT3 ITD, MLL-AF4)



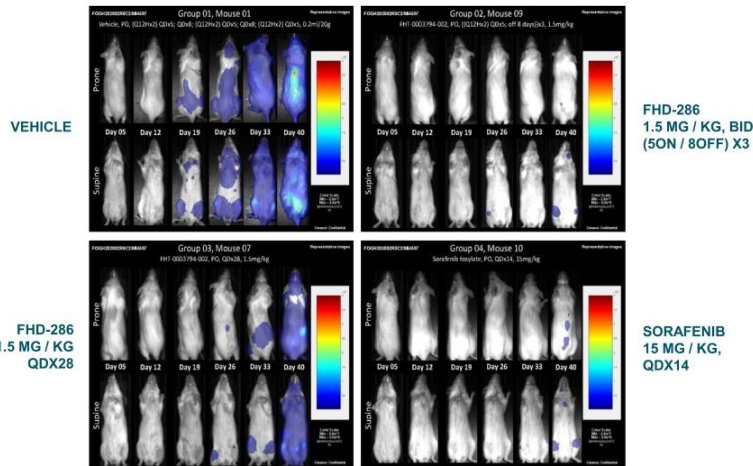
**OCI-AML2 CDX MODEL**  
(MII-AF6, DNMT3A MUT.)



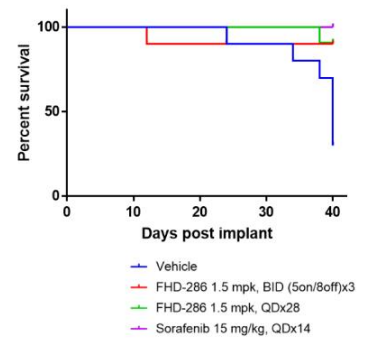
● Vehicle    ■ FHD-286, 0.5 mg/kg, PO, QD    ■ FHD-286, 1.5 mg/kg, PO, QD

# TUMOR GROWTH INHIBITION WITH FHD-286 TREATMENT OBSERVED BY BIOLUMINESCENCE

Imaging in a Disseminated AML Model



## FHD-286 SURVIVAL ADVANTAGE IN DISSEMINATED AML MODEL



# SELECTIVE BRM MODULATORS FOR BRG1 MUTATED CANCERS

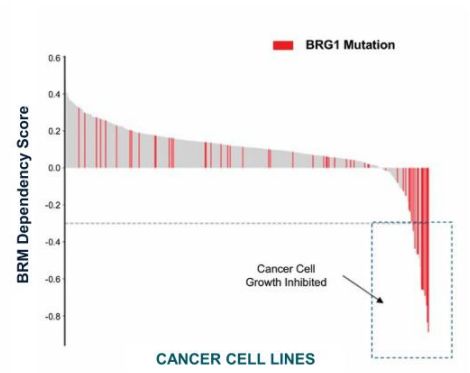
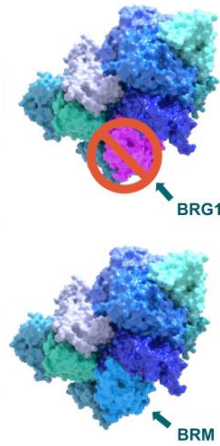
Enzymatic Inhibitor and Protein Degradation Programs Targeting BRG1 Mutated Cancers (e.g., NSCLC), 30+ Cancers with BRG1 Mutations

---

# BRG1 MUTATIONS CREATE A GENETIC DEPENDENCY ON BRM

## Selective BRM Modulators Overview

<b>Target / Approach</b>	<ul style="list-style-type: none"> <li>BRM</li> <li>Enzymatic inhibitor</li> <li>Targeted protein degrader</li> </ul>
<b>Indications</b>	<ul style="list-style-type: none"> <li>BRG1 mutated cancers (e.g., NSCLC), 30+ cancers with BRG1 mutations</li> </ul>
<b>Mutation / Aberration</b>	<ul style="list-style-type: none"> <li>BRG1</li> </ul>
<b>Stage</b>	<ul style="list-style-type: none"> <li>Pre-clinical</li> </ul>
<b>New Patients Impacted / Year*</b>	<ul style="list-style-type: none"> <li>&gt; 100,000</li> </ul>
<b>Economics of Lilly Collaboration</b>	<ul style="list-style-type: none"> <li>50/50 U.S. economics</li> <li>Tiered ex-U.S. royalties starting in the low double-digit range and escalating into the twenties</li> </ul>

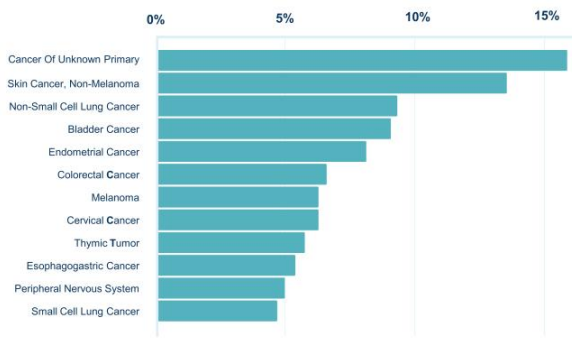


\* Per year incidence in the U.S., EU5, Japan

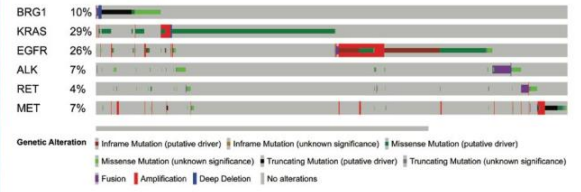
# BRG1 MUTATED IN ~5% OF ALL TUMORS

Broad Addressable Patient Population

## BRG1 MUTATED ACROSS RANGE OF TUMORS

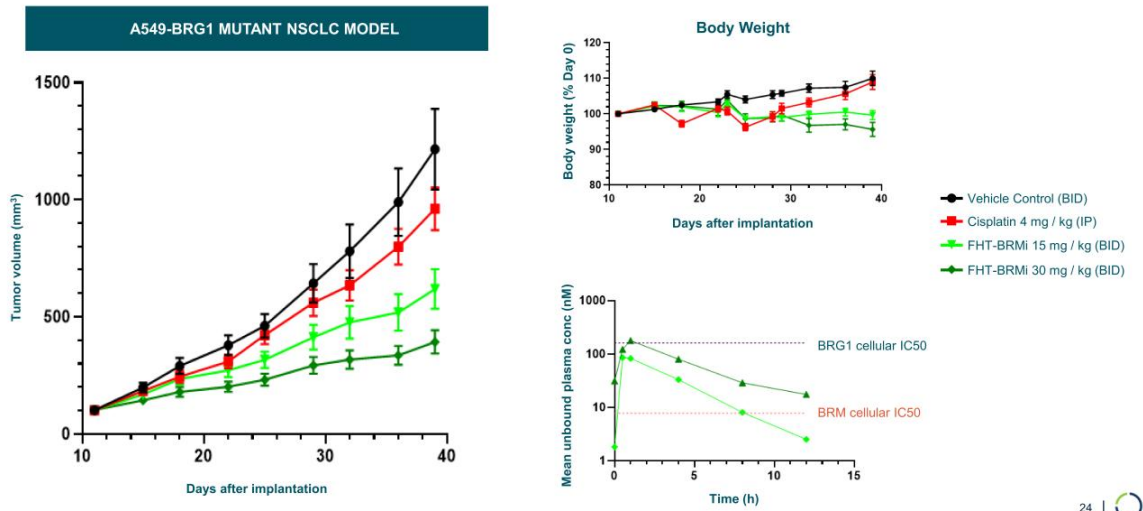


## BRG1 MUTATED IN UP TO 10% OF NSCLC TUMORS, MINIMAL OVERLAP WITH OTHER MUTATIONS



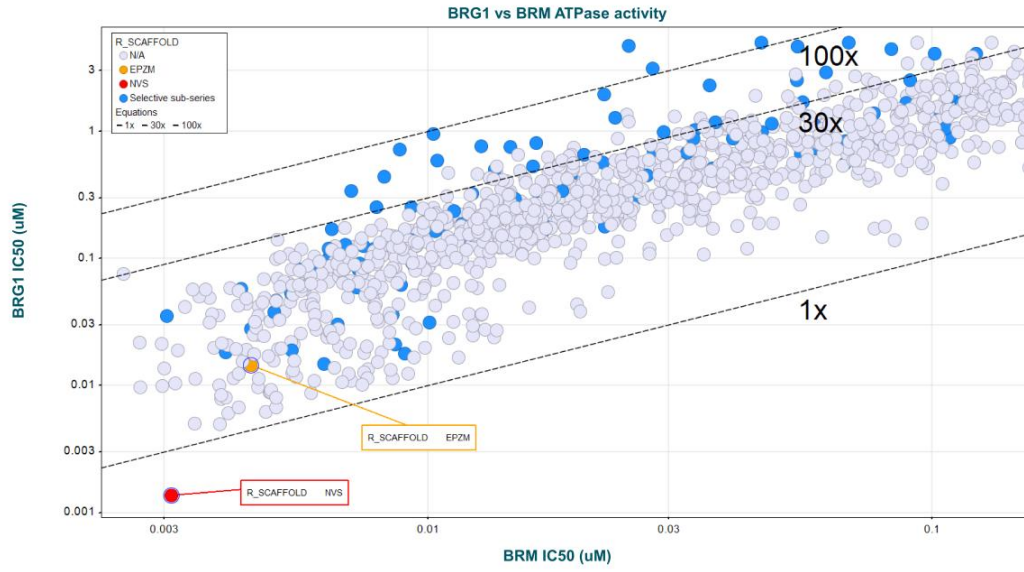
# BRM SELECTIVE INHIBITOR *IN VIVO* EFFICACY

Demonstrates PK / PD and *In Vivo* Efficacy in a BRG1 Mutant Lung CDX Model





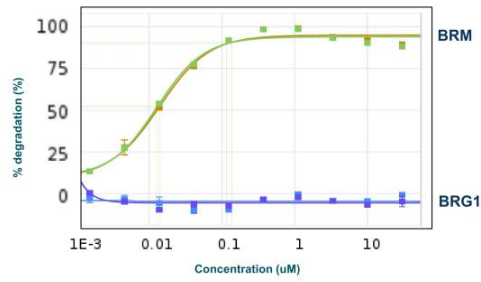
## ENZYMATIC SELECTIVITY APPROACHING 200X ACHIEVED



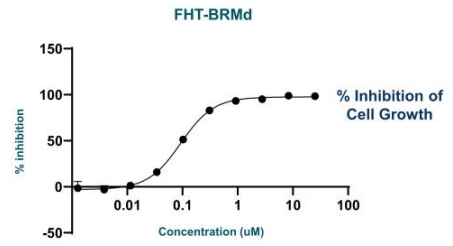
# ADVANCING BRM SELECTIVE DEGRADERS

Achieving Complete BRM Degradation

BRM / BRG1 HIBIT DATA



A549 TEN-DAY PROLIFERATION ASSAY



DEGRADERS CAUSE TIME- AND DOSE-DEPENDENT BRM DEGRADATION, ANTIPROLIFERATIVE EFFECTS IN A549 BRG1 MUTANT NSCLC LUNG MODEL

# SELECTIVE CBP PROTEIN DEGRADER FOR EP300 MUTATED CANCERS

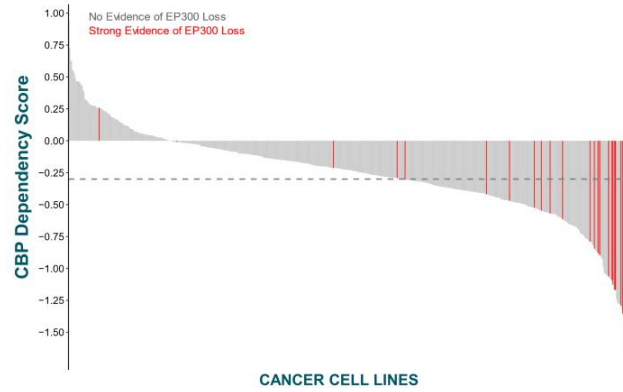
Implicated in Subsets of Cancers Including Bladder, Colorectal, Breast, Gastric and Lung

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# ADVANCING HIGHLY SELECTIVE CBP PROTEIN DEGRADER FOR EP300 MUTATED CANCERS

## Selective CBP Protein Degradation Overview

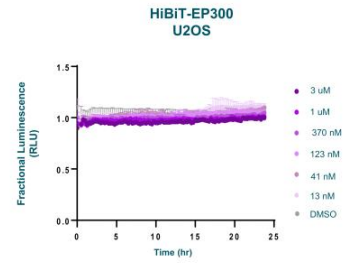
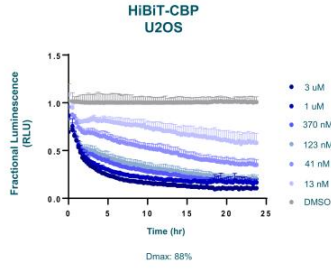
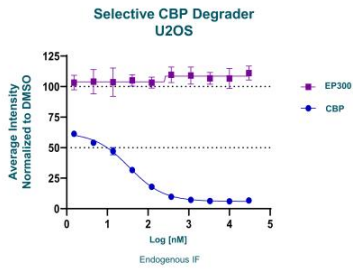
<b>Target / Approach</b>	<ul style="list-style-type: none"><li>CREB binding protein (CBP)</li><li>Targeted protein degrader</li></ul>
<b>Initial Indication</b>	<ul style="list-style-type: none"><li>EP300 mutated cancers (e.g., subsets of prostate, bladder, colorectal, breast, gastric and lung cancers)</li></ul>
<b>Mutation / Aberration</b>	<ul style="list-style-type: none"><li>EP300 mutated cancers</li></ul>
<b>Stage</b>	<ul style="list-style-type: none"><li>Pre-clinical</li></ul>
<b>New Patients Impacted / Year*</b>	<ul style="list-style-type: none"><li>Over 100,000</li></ul>



\* Per year incidence in the U.S., EU5, Japan

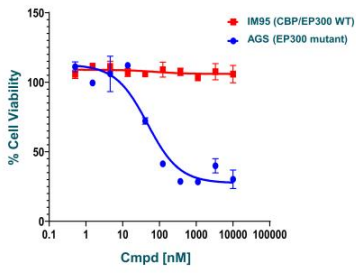
# ADVANCEMENT OF HIGHLY SELECTIVE CBP DEGRADERS

## SELECTIVE CBP DEGRADATION Osteosarcoma Cell Line

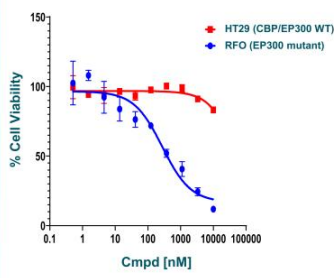


# HIGHLY SELECTIVE DEGRADER OF CBP DEMONSTRATES CBP-DEPENDENT CELL KILLING ACROSS MULTIPLE CANCERS

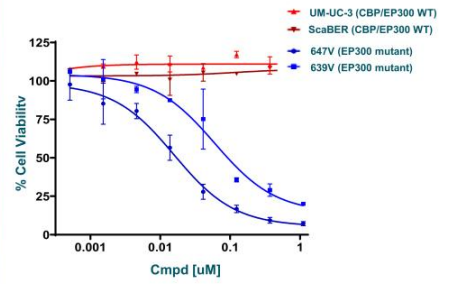
### Gastric Cancer



### Colorectal Cancer



### Bladder Cancer



CELL PROLIFERATION ASSAYS



# SELECTIVE EP300 PROTEIN DEGRADER FOR CBP MUTANT CANCERS & EP300 DEPENDENT MALIGNANCIES

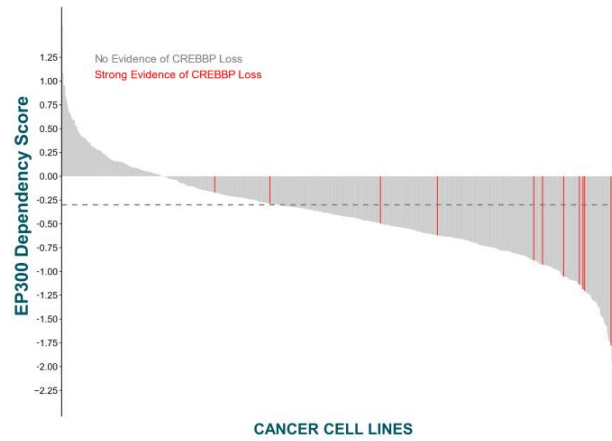
Implicated in CBP Mutated Cancers and Subsets of EP300 Dependent Malignancies (e.g., Bladder, NSCLC, Various Lymphomas and Leukemias)

---

# ADVANCING HIGHLY SELECTIVE EP300 PROTEIN DEGRADER FOR CBP MUTANT CANCERS & EP300 DEPENDENT MALIGNANCIES

## Selective EP300 Protein Degradation Overview

<b>Target / Approach</b>	<ul style="list-style-type: none"> <li>E1A binding protein p300 (EP300)</li> <li>Targeted protein degrader</li> </ul>
<b>Initial Indication</b>	<ul style="list-style-type: none"> <li>CBP mutant cancers and subsets of EP300 dependent malignancies (e.g., bladder, NSCLC, various lymphomas and leukemias)</li> </ul>
<b>Mutation / Aberration</b>	<ul style="list-style-type: none"> <li>CBP mutant cancers</li> <li>EP300 dependent malignancies</li> </ul>
<b>Stage</b>	<ul style="list-style-type: none"> <li>Pre-clinical</li> </ul>
<b>New Patients Impacted / Year*</b>	<ul style="list-style-type: none"> <li>Over 100,000</li> </ul>

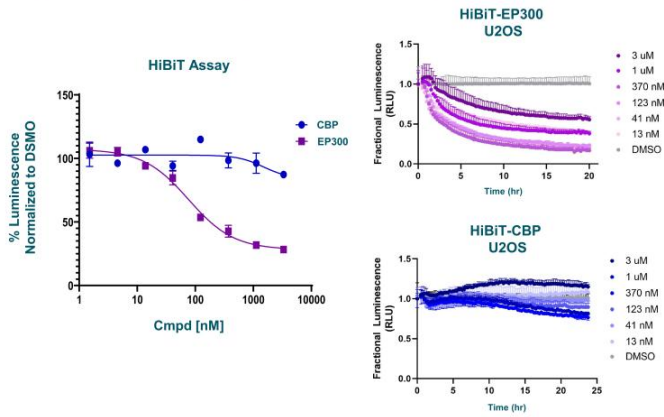


\* Per year incidence in the U.S., EU5, Japan

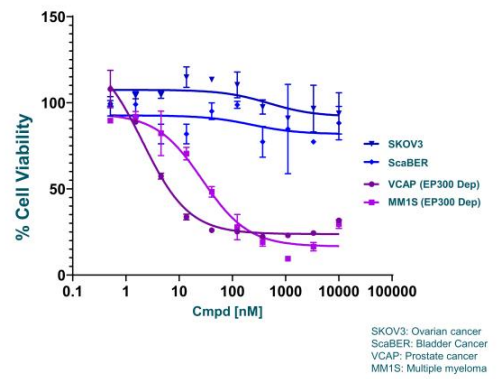


# ADVANCEMENT OF HIGHLY SELECTIVE EP300 DEGRADERS

## SELECTIVE EP300 DEGRADATION (Osteosarcoma Cell Line)



## CELL PROLIFERATION ASSAYS (EP300 Dependent vs. Non-Dependent Cell Lines)





# SELECTIVE ARID1B PROTEIN DEGRADER FOR ARID1A MUTATED CANCERS

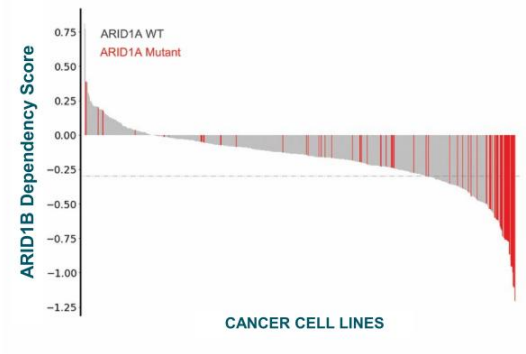
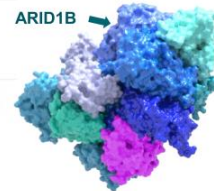
Protein Degradator Targeting ARID1A Mutated Cancers, the Most Mutated Subunit in the BAF Complex  
(e.g., Ovarian, Endometrial, Colorectal, Bladder and Other Cancers)

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# ARID1A: MOST MUTATED SUBUNIT IN BAF COMPLEX – CREATES DEPENDENCY ON ARID1B

## Selective ARID1B Protein Degradation Overview

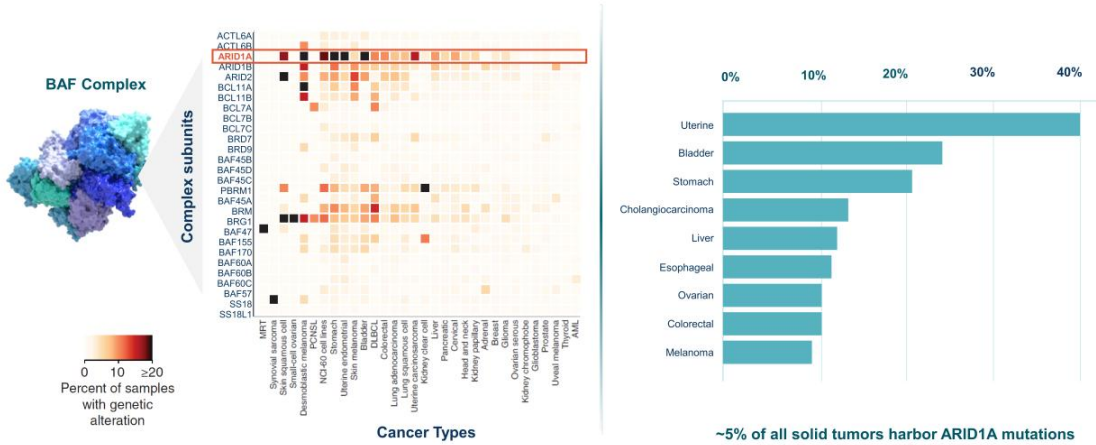
<b>Target / Approach</b>	<ul style="list-style-type: none"> <li>ARID1B</li> <li>Targeted protein degrader</li> </ul>
<b>Initial Indication</b>	<ul style="list-style-type: none"> <li>ARID1A mutated cancers</li> </ul>
<b>Mutation / Aberration</b>	<ul style="list-style-type: none"> <li>ARID1A mutations (e.g., ovarian, endometrial, colorectal, bladder and other cancers)</li> </ul>
<b>Stage</b>	<ul style="list-style-type: none"> <li>Pre-clinical</li> </ul>
<b>New Patients Impacted / Year*</b>	<ul style="list-style-type: none"> <li>&gt; 175,000</li> </ul>



\* Per year incidence in the U.S., EU5, Japan

# ARID1A MUTATED CANCERS: SIGNIFICANT OPPORTUNITY

ARID1A Mutated Across Range of Tumors



Hodges et al. 2017

# TARGETING ARID1A MUTATED CANCERS: ARID1B PROTEIN DEGRADER

Advantaged by Gene Traffic Control Platform and Protein Degradation Capabilities

## GENE TRAFFIC CONTROL PLATFORM

- Platform produces BAF complexes and subcomplexes containing either ARID1A or ARID1B at scale
- Enables proprietary screens against ARID1B

## PROTEIN DEGRADER CAPABILITIES

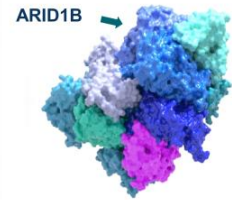
- Utilize protein degrader toolbox to create ARID1B hetero-bifunctional degraders

## PROGRAM STATUS

- Validated selective chemical binders of ARID1B
- In process of expanding binders into novel selective protein degraders
- Assessing outcomes of ARID1B degradation and impact on BAF complex formation



Highly purified ARID1B /  
BAF complex





**TRANSCRIPTION FACTORS**  
A NOVEL APPROACH

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# A NEW APPROACH TO DRUGGING TRANSCRIPTION FACTORS

Enabled by Proprietary Ability to Purify and Synthesize Chromatin Regulatory System Components

## TFS ARE COMPELLING DRUG TARGETS...

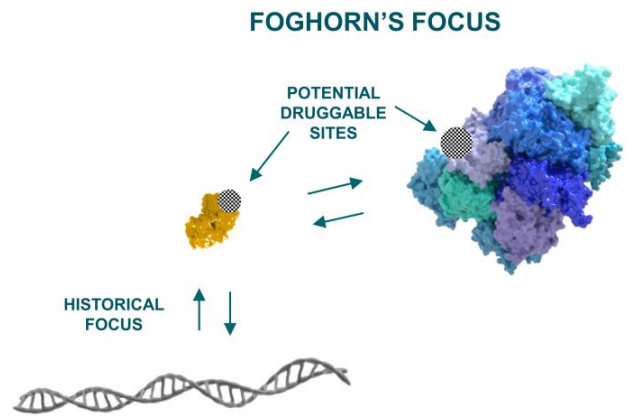
- Highly involved in gene expression
- Implicated in range of cancers and other diseases

## ...BUT HISTORICALLY DIFFICULT TO TARGET

- Featureless surface: no druggable binding pocket
- Tight interactions with DNA: undruggable affinities

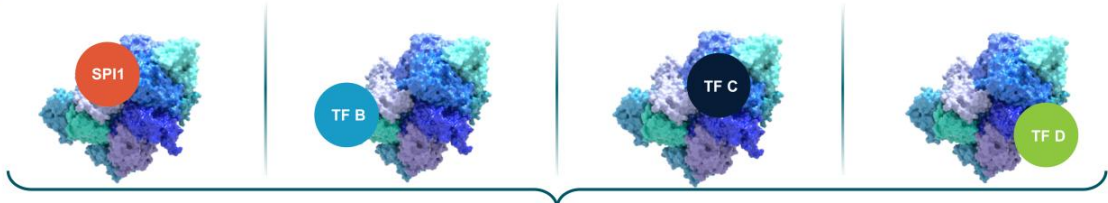
## FOGHORN HAS A NEW APPROACH FOCUSING ON INTERACTION WITH BAF

- Druggable binding pockets
- Druggable affinities

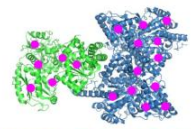


# TRANSCRIPTION FACTORS BIND TO BAF DIRECTLY WITH HIGH DEGREE OF SPECIFICITY

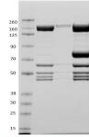
Unique Insights into Where and How Transcription Factors Bind



## MAPPING THE TF-BAF INTERACTION



MASS SPEC. FOOT-PRINTING

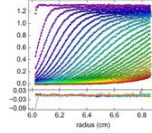


PULL-DOWN ASSAYS

Foghorn's collection of BAF sub-complexes and domains

## VALIDATING THE TF-BAF INTERACTION

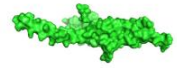
**BIOPHYSICAL**  
AUC / SPR / ITC



**BIOCHEMICAL**  
TR-FRET / FP



**STRUCTURAL**  
Crystal / NMR

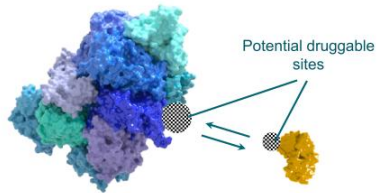




## HIGHLY SCALABLE APPROACH TO ADDRESS SIGNIFICANT UNMET MEDICAL NEED DRIVES MERCK COLLABORATION

Potential to Drug > 100 TFs Associated with BAF

### TRANSCRIPTION FACTOR DISRUPTORS



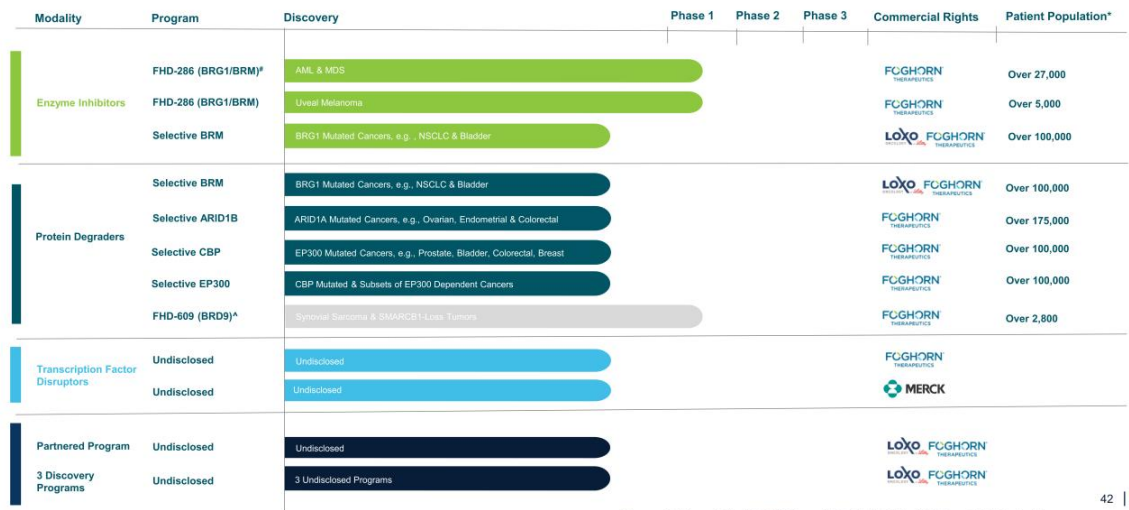
- >100 TFs estimated associated with BAF
- Foghorn pursuing multiple TFs in parallel
- Approach highly scalable and potential broad application – other chromatin remodeling complexes and other diseases



- Merck collaboration to drug single specified transcription factor target
- \$15 million upfront; up to \$410 million in research, development, regulatory and sales-based milestones
- Up to low double-digit royalties on product sales

# BROAD PIPELINE ACROSS A RANGE OF TARGETS AND MODALITIES

Precision Oncology / Breadth and Depth / Over 15 Programs



\* Per year incidence in the U.S., EU, Japan | \* On full clinical hold | ^ On partial clinical hold

# FIRST-IN-CLASS PRECISION MEDICINES TARGETING MAJOR UNMET NEEDS IN CANCER



## LEADER IN NEW AREA OF CANCER BIOLOGY

Foghorn is a **leader in targeting chromatin biology**, which has unique potential to address underlying dependencies of many genetically defined cancers

Broad pipeline of **over 15 programs** across a range of targets and modalities



## LARGE MARKET POTENTIAL

Chromatin biology is implicated in up to **50% of tumors**, potentially impacting **~2.5 million patients**

Foghorn's current pipeline potentially addresses **more than 500,000** of these patients



## WELL-FUNDED

**\$316.0 million** in cash and equivalents

(as of 03/31/2023)

Provides **runway into H2'25**



## SIGNIFICANT VALUE DRIVERS IN 2023

Initial clinical data in uveal melanoma with **FHD-286** expected **Q2'23**

AML/MDS study with FHD-286 on full clinical hold, development **clarity anticipated in Q2'23**



## COLLABORATIONS WITH MAJOR ONCOLOGY PLAYERS

Strategic collaboration with Loxo Oncology at Lilly; **\$380 million upfront**; 50/50 U.S. economic split on two lead programs

Merck collaboration to drug single specified transcription factor target; **\$15 million upfront** and up to **\$410 million** in milestones

# APPENDIX



**STRATEGIC PARTNERSHIP**  
LOXO ONCOLOGY AT LILLY

---

## STRATEGIC COLLABORATION WITH LOXO ONCOLOGY AT LILLY

Foghorn to Lead Discovery and Research Activities



### \$380 MILLION UPFRONT

\$300 million cash payment

\$80 million investment in Foghorn common stock at a price of \$20 per share



### 50/50 U.S. ECONOMICS ON TWO PROGRAMS

50/50 U.S. economic split on BRM-Selective and another undisclosed program

Tiered ex-U.S. royalties starting in the low double-digit range and escalating into the twenties based on revenue levels



### THREE UNDISCLOSED DISCOVERY PROGRAMS

Option to participate in a percentage of the U.S. economics

Tiered ex-U.S. royalties from the mid-single digit to low-double digit range

\$1.3 billion in potential milestones

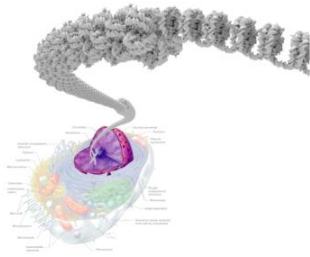


**THE CHROMATIN REGULATORY SYSTEM**  
ORCHESTRATES GENE EXPRESSION

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# THE CHROMATIN REGULATORY SYSTEM ORCHESTRATES GENE EXPRESSION

Two Major Components Work in Concert: Chromatin Remodeling Complexes and Transcription Factors

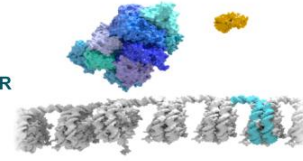


## CHROMATIN

Chromatin – compacted form of DNA inside the nucleus of the cell

## 1 | CHROMATIN REMODELING COMPLEX AND TRANSCRIPTION FACTOR

Work together to orchestrate gene expression



## 2 | RIGHT GENES

TFs guide chromatin remodeling complexes to the right locations

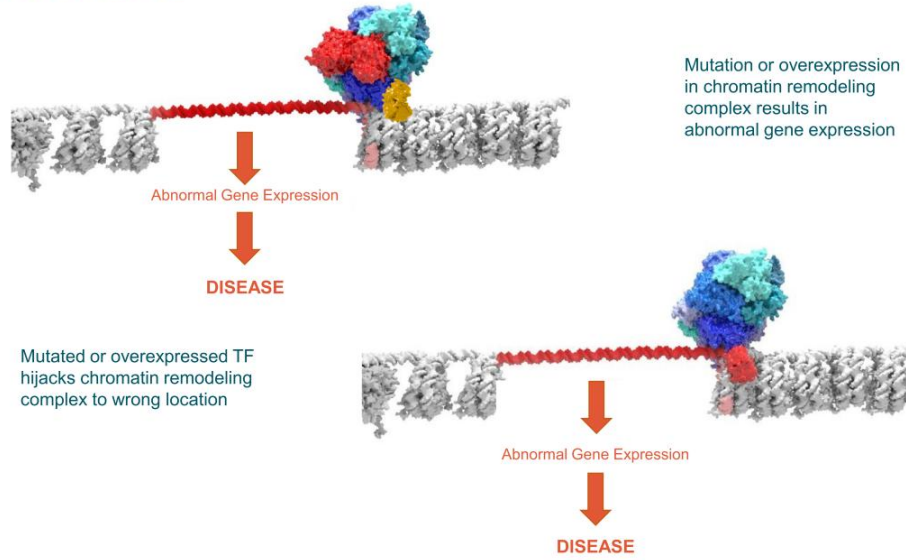


## 3 | NORMAL GENE EXPRESSION

Once chromatin is unpacked, gene expression can occur



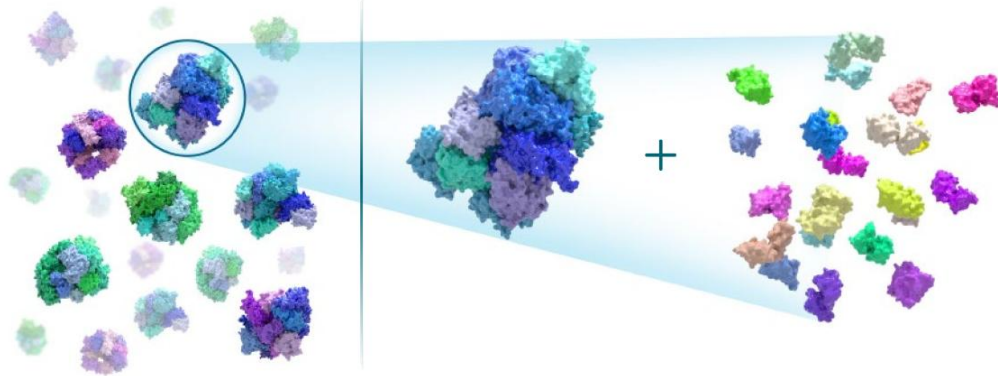
## BREAKDOWNS IN THE CHROMATIN REGULATORY SYSTEM CAN LEAD TO DISEASE



# CHROMATIN REGULATORY SYSTEM

Abundance of Targets within the BAF Complex

## BAF COMPLEX AND ASSOCIATED TRANSCRIPTION FACTORS



28 Chromatin Remodeling  
Complexes and >1,000 TFs

BAF Complex Subunits Mutated  
and Dysregulated in Cancer

Estimate >100  
Transcription Factors  
Associated with Just  
the BAF Complex



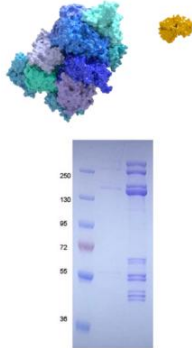
**PLATFORM &  
DRUGGING CAPABILITIES**


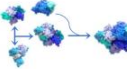
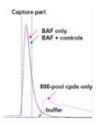


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# PLATFORM IS POWERED BY ABILITY TO PRODUCE COMPONENTS AT SCALE

Drives Drug Discovery Pipeline with Cutting Edge Technology

## PRODUCTION OF CHROMATIN REGULATORY SYSTEM COMPONENTS



	FEATURES	BENEFITS
	Surface Mapping	Characterize TF / BAF Binding Sites
	Assembly	Synthesize subcomplexes to enable drug discovery
	Affinity Screening & Validation	ASMS on full complex to yield novel degraders
	HTS	Multiple screening options with full complex
	Biophysics/SPR	Validation of novel small molecule binders

# PROTEIN DEGRADER PLATFORM

## CURRENT APPROACH

- A leader in developing heterobifunctional degraders for clinical evaluation in oncology
- Employing PROTAC and non-CRBN based molecular glue degradation approaches

## DEGRADER CHEMICAL TOOLBOX

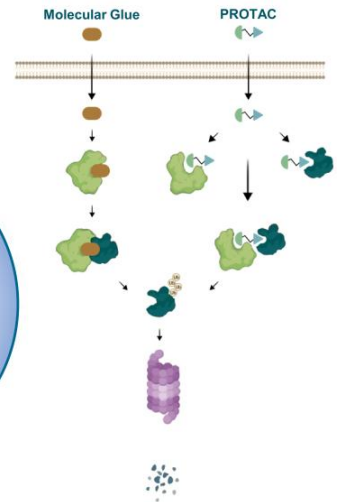
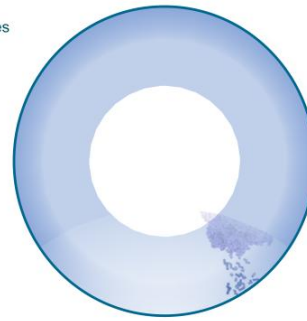
- Proprietary library of drug-like linkers, E3 ligase binders and potential glues
- Chemistry to rapidly identify and optimize degraders

## ADVANCED MECHANISTIC CHARACTERIZATION

- Native target turnover understanding
- Cellular degradation kinetics and rates
- Structural, biochemical and cellular ternary complex characterization
- Global proteomics and ubiquitination studies
- Computational modeling of degraders
- Degradation efficacy across multiple cell types

## OPTIMIZATION OF DEGRADER DRUG PROPERTIES

- Guidelines for both of oral and IV-administered degraders
- PK / PD, efficacy and safety modeling to optimize dosing and scheduling



## **Leadership Team, Board & Advisors**

**EXPERTISE ACROSS DRUG DISCOVERY, CLINICAL  
DEVELOPMENT AND COMMERCIALIZATION**

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