

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): August 13, 2024

Acumen Pharmaceuticals, Inc.

(Exact name of Registrant as Specified in Its Charter)

Delaware  
(State or Other Jurisdiction  
of Incorporation)

001-40551  
(Commission  
File Number)

36-4108129  
(IRS Employer  
Identification No.)

1210-1220 Washington Street, Suite 210  
Newton, Massachusetts  
(Address of Principal Executive Offices)

22902  
(Zip Code)

(434) 297-1000  
(Registrant's Telephone Number, Including Area Code)

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 (see General Instructions A.2. below):

- 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	ABOS	The Nasdaq Global Select 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 August 13, 2024, Acumen Pharmaceuticals, Inc. (the “Company”) reported financial results and business highlights for the quarter ended June 30, 2024. A copy of this press release (the “Earnings Press Release”) is furnished as Exhibit 99.1 to this Current Report on Form 8-K (this “Report”) and is incorporated by reference.

The information in this Item 2.02 of this Report (including Exhibit 99.1) 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 in any filing 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 a filing.

**Item 7.01 Regulation FD Disclosure.**

On August 13, 2024, the Company posted an updated corporate presentation to its website at <https://investors.acumenpharm.com/news-events/presentations>, which the Company may use from time to time in communications or conferences. The corporate presentation was updated to reflect the projected timing of the Company’s topline data in its Phase 1 subcutaneous trial of sabimetus and to update the Company’s cash position as of June 30, 2024. A copy of the corporate presentation is attached as Exhibit 99.2 to this Report.

The information in this Item 7.01 of this Report (including Exhibit 99.2), 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 in any filing under the Securities Act or the Exchange Act, except as expressly set forth by specific reference in such filing. The Company’s submission of this Report shall not be deemed an admission as to the materiality of any information required to be disclosed solely to satisfy the requirements of Regulation FD.

**Item 9.01 Financial Statements and Exhibits.****(d). Exhibits**

Exhibit No.	Description
99.1	<a href="#">Earnings Press Release, dated August 13, 2024</a>
99.2	<a href="#">Corporate Presentation, dated August 13, 2024</a>
104	Cover Page Interactive Data File (embedded within the Inline XBRL document)

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**SIGNATURES**

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

**Acumen Pharmaceuticals, Inc.**

Dated: August 13, 2024

By: /s/ Matthew Zuga  
Matthew Zuga  
*Chief Financial Officer and Chief Business Officer*



### Acumen Pharmaceuticals Reports Second Quarter 2024 Financial Results and Business Highlights

- Actively enrolling subjects in ALTITUDE-AD, a Phase 2 study to investigate sabirnetug (ACU193) for the treatment of early Alzheimer's disease
- Dosed the first subject in a Phase 1 study to support subcutaneous administration of sabirnetug in July 2024 with topline results anticipated in the first quarter of 2025
- Cash, cash equivalents and marketable securities of \$281.4 million as of Jun. 30, 2024, expected to support current clinical and operational activities into the first half of 2027
- Company to host conference call and webcast today at 8:00 a.m. ET

**NEWTON, Mass., Aug. 13, 2024** – Acumen Pharmaceuticals, Inc. (NASDAQ: ABOS) ("Acumen" or the "Company"), a clinical-stage biopharmaceutical company developing a novel therapeutic that targets toxic soluble amyloid beta oligomers (A $\beta$ Os) for the treatment of Alzheimer's disease (AD), today reported financial results for the second quarter of 2024 and provided a business update.

"Our team is highly focused on execution in 2024, and I'm very pleased with our progress in the first half of the year. We are actively enrolling subjects in our global Phase 2 ALTITUDE-AD study that we initiated this spring. We are highly encouraged by the level of interest from investigators and patients in sabirnetug's mechanism of action which has led to enrollment progressing faster than our expectations," said Daniel O'Connell, Chief Executive Officer of Acumen. "In addition to the progress with ALTITUDE-AD, we announced in July the initiation of a Phase 1 pharmacokinetic comparison study supporting subcutaneous administration of sabirnetug. Topline results from this healthy volunteer study are expected in the first quarter of 2025. With the momentum in our clinical program and sabirnetug's distinct selectivity for toxic amyloid beta oligomers, we believe that we are positioned to deliver a potential next-generation treatment for early Alzheimer's disease."

#### Recent Highlights and Anticipated Milestones

- **In May 2024, the Company announced the first patient dosed in ALTITUDE-AD, a Phase 2 study to investigate the clinical efficacy and safety of sabirnetug for the treatment of early AD.**
  - Currently, more than 50 sites are activated in the U.S., Canada, U.K. and EU.
- **In July 2024, the Company announced the first subject had been dosed with a subcutaneous formulation of sabirnetug in a Phase 1 pharmacokinetic (PK) comparison study.** The study will compare the PK profile between subcutaneous and intravenous administrations of sabirnetug in healthy volunteers.
  - Topline results are anticipated in the first quarter of 2025.

- **In July 2024, the Company presented additional biomarker and target engagement analyses, as well as insight into the patient experience from the Phase 1 INTERCEPT-AD study in early AD at the Alzheimer's Association International Conference (AAIC®) annual meeting.**
  - The research highlights the experiences of patients in the clinical trial to inform development of future trials, biomarker data to support sabirnetug's mechanism of action, and an ultra-sensitive method of detecting levels of sabirnetug in cerebrospinal fluid (CSF), given the small amounts of monoclonal antibodies that typically enter the brain from the blood. More details about the research are available [here](#).
- **The Company plans to host a virtual R&D Day on Oct. 2, 2024, providing a deep dive into the scientific rationale, Phase 1 clinical results and Phase 2 clinical plans for sabirnetug.** Registration details will be communicated prior to the event.

#### Second Quarter 2024 Financial Results

- **Cash Balance.** As of June 30, 2024, cash, cash equivalents and marketable securities totaled \$281.4 million, compared to cash, cash equivalents and marketable securities of \$306.1 million as of December 31, 2023. The decrease in cash is related to funding ongoing operations. Cash is expected to support current clinical and operational activities into the first half of 2027.
- **Research and Development (R&D) Expenses.** R&D expenses were \$19.5 million for the three-month period ended June 30, 2024, compared to \$9.1 million for the three-month period ended June 30, 2023. The increase in R&D expenses was primarily due to increased contract research organization and other clinical trial costs related to ALTITUDE-AD, as well as higher costs for personnel, license agreements, and shipping and packaging.
- **General and Administrative (G&A) Expenses.** G&A expenses were \$4.8 million for the three-month period ended June 30, 2024, compared to \$4.3 million for the three-month period ended June 30, 2023. The increase in G&A expenses was primarily due to increased costs related to personnel.
- **Loss from Operations.** Loss from operations was \$24.4 million for the three-month period ended June 30, 2024, compared to \$13.5 million for the three-month period ended June 30, 2023. This increase was due to the increased R&D and G&A expenses over the prior year period.
- **Net Loss.** Net loss was \$20.5 million for the three-month period ended June 30, 2024, compared to \$11.6 million for the three-month period ended June 30, 2023.

#### Conference Call Details

Acumen will host a conference call and live audio webcast today, August 13, 2024, at 8:00 a.m. ET.

To participate in the live conference call, please register using [this link](#). After registration, you will be informed of the dial-in numbers including PIN. Please register at least one day in advance.

The webcast audio will be available via [this link](#).

An archived version of the webcast will be available for at least 30 days in the Investors section of the Company's website at [www.acumenpharm.com](http://www.acumenpharm.com).

**About Sabirnetug (ACU193)**

Sabirnetug (ACU193) is a humanized monoclonal antibody (mAb) discovered and developed based on its selectivity for soluble amyloid beta oligomers (A $\beta$ O $_s$ ), which are a highly toxic and pathogenic form of A $\beta$ , relative to A $\beta$  monomers and amyloid plaques. Soluble A $\beta$ O $_s$  have been observed to be potent neurotoxins that bind to neurons, inhibit synaptic function and induce neurodegeneration. By selectively targeting toxic soluble A $\beta$ O $_s$ , sabirnetug aims to address the hypothesis that soluble A $\beta$ O $_s$  are an early and persistent underlying cause of the neurodegenerative process in Alzheimer's disease (AD). Sabirnetug has been granted Fast Track designation for the treatment of early AD by the U.S. Food and Drug Administration and is currently being evaluated in a Phase 2 study in patients with early AD.

**About ALTITUDE-AD (Phase 2)**

Initiated in 2024, ALTITUDE-AD is a Phase 2, multi-center, randomized, double-blind, placebo-controlled clinical trial designed to evaluate the efficacy and safety of sabirnetug (ACU193) infusions administered once every four weeks in slowing cognitive and functional decline as compared to placebo in participants with early Alzheimer's disease. The study will enroll approximately 540 individuals with early Alzheimer's disease (mild cognitive impairment or mild dementia due to AD). The global study is currently enrolling at multiple investigative sites located in the United States and Canada with plans for additional sites in Europe and the UK. More information can be found on [www.clinicaltrials.gov](http://www.clinicaltrials.gov), NCT identifier NCT06335173.

**About INTERCEPT-AD (Phase 1)**

Completed in 2023, INTERCEPT-AD was a Phase 1, U.S.-based, multi-center, randomized, double-blind, placebo-controlled clinical trial evaluating the safety and tolerability, and establishing clinical proof of mechanism, of sabirnetug in patients with early Alzheimer's disease (AD). Sixty-five individuals with early AD (mild cognitive impairment or mild dementia due to AD) enrolled in this first-in-human study of sabirnetug. The INTERCEPT-AD study consisted of single-ascending-dose (SAD) and multiple-ascending-dose (MAD) cohorts. Results showed sabirnetug to be well-tolerated with a favorable overall safety profile. The trial showed amyloid plaque reduction, effects on synaptic biomarkers, low overall rates of ARIA-E, and evidence of target engagement that validated proof of mechanism. More information can be found on [www.clinicaltrials.gov](http://www.clinicaltrials.gov), NCT identifier NCT04931459.

**About Acumen Pharmaceuticals, Inc.**

Acumen Pharmaceuticals is a clinical-stage biopharmaceutical company developing a novel therapeutic that targets toxic soluble amyloid beta oligomers (A $\beta$ O $_s$ ) for the treatment of Alzheimer's disease (AD). Acumen's scientific founders pioneered research on A $\beta$ O $_s$ , which a growing body of evidence indicates are early and persistent triggers of Alzheimer's disease pathology. Acumen is currently focused on advancing its investigational product candidate, sabirnetug (ACU193), a humanized monoclonal antibody that selectively targets toxic soluble A $\beta$ O $_s$ , in its ongoing Phase 2 clinical trial ALTITUDE-AD (NCT06335173) in early symptomatic Alzheimer's disease patients, following positive results in its Phase 1 trial INTERCEPT-AD. The company is headquartered in Newton, Mass. For more information, visit [www.acumenpharm.com](http://www.acumenpharm.com).

**Forward-Looking Statements**

This press release contains forward-looking statements within the meaning of The Private Securities Litigation Reform Act of 1995. Any statement describing Acumen's goals, expectations, financial or other projections, intentions or beliefs is a forward-looking statement and should be considered an at-risk statement. Words such as "believes," "expects," "anticipates," "could," "should," "would," "seeks," "aims," "plans," "potential," "will," "milestone" and similar expressions are intended to identify forward-looking statements, although not all forward-looking statements contain these identifying words. Forward-looking statements include statements concerning Acumen's business, and Acumen's ability to achieve its strategic and financial goals, including its projected use of cash, cash equivalents and marketable securities and the expected sufficiency of its cash resources into the first half of 2027, the therapeutic potential of Acumen's product candidate,



sabirnetug (ACU193), including against other antibodies, the anticipated enrollment progression of ALTITUDE-AD, and the anticipated timeline for results from the Phase 1 trial to support a subcutaneous dosing option of sabirnetug. These statements are based upon the current beliefs and expectations of Acumen management, and are subject to certain factors, risks and uncertainties, particularly those inherent in the process of discovering, developing and commercializing safe and effective human therapeutics. Such risks may be amplified by the impacts of geopolitical events and macroeconomic conditions, such as rising inflation and interest rates, supply disruptions and uncertainty of credit and financial markets. These and other risks concerning Acumen's programs are described in additional detail in Acumen's filings with the Securities and Exchange Commission ("SEC"), including in Acumen's most recent Annual Report on Form 10-K, and in subsequent filings with the SEC. Copies of these and other documents are available from Acumen. Additional information will be made available in other filings that Acumen makes from time to time with the SEC. These forward-looking statements speak only as of the date hereof, and Acumen expressly disclaims any obligation to update or revise any forward-looking statement, except as otherwise required by law, whether, as a result of new information, future events or otherwise.

**CONTACTS:**

**Investors:**

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abraun@acumenpharm.com

**Media:** AcumenPR@westwicke.com



**Acumen Pharmaceuticals, Inc.**  
**Condensed Balance Sheets**  
(in thousands, except share and per share data)

	June 30, 2024 (unaudited)	December 31, 2023
<b>ASSETS</b>		
Current assets		
Cash and cash equivalents	\$ 67,955	\$ 66,886
Marketable securities, short-term	192,517	176,636
Prepaid expenses and other current assets	6,443	3,093
Total current assets	266,915	246,615
Marketable securities, long-term	20,908	62,553
Right-of-use asset	325	381
Restricted cash	235	233
Property and equipment, net	105	122
Other assets	425	221
Total assets	\$ 288,913	\$ 310,125
<b>LIABILITIES AND STOCKHOLDERS' EQUITY</b>		
Current liabilities		
Accounts payable	\$ 4,211	\$ 1,379
Accrued clinical trial expenses	7,027	4,387
Accrued expenses and other current liabilities	4,004	6,339
Finance lease liability, short-term	—	756
Operating lease liability, short-term	125	110
Total current liabilities	15,367	12,971
Operating lease liability, long-term	219	284
Debt, long-term	29,380	29,897
Total liabilities	44,966	43,152
Commitments and contingencies		
Stockholders' equity		
Preferred stock, \$0.0001 par value; 10,000,000 shares authorized and no shares issued and outstanding as of June 30, 2024 and December 31, 2023	—	—
Common stock, \$0.0001 par value; 300,000,000 shares authorized as of June 30, 2024 and December 31, 2023; 60,079,778 and 57,910,461 shares issued and outstanding as of June 30, 2024 and December 31, 2023	6	6
Additional paid-in capital	502,313	489,453
Accumulated deficit	(258,208)	(222,798)
Accumulated other comprehensive income (loss)	(164)	312
Total stockholders' equity	243,947	266,973
Total liabilities and stockholders' equity	\$ 288,913	\$ 310,125





**Acumen Pharmaceuticals, Inc.**  
**Condensed Statements of Operations and Comprehensive Loss**  
(in thousands, except share and per share data)  
(unaudited)

	Three Months Ended June 30,		Six Months Ended June 30,	
	2024	2023	2024	2023
Operating expenses				
Research and development	\$ 19,533	\$ 9,133	\$ 31,982	\$ 17,846
General and administrative	4,848	4,345	10,173	8,767
Total operating expenses	24,381	13,478	42,155	26,613
Loss from operations	(24,381)	(13,478)	(42,155)	(26,613)
Other income (expense)				
Interest income	3,816	1,884	7,821	3,716
Interest expense	(1,004)	—	(2,004)	—
Change in fair value of embedded derivatives	1,100	—	1,050	—
Other expense, net	(68)	(16)	(122)	(20)
Total other income	3,844	1,868	6,745	3,696
Net loss	(20,537)	(11,610)	(35,410)	(22,917)
Other comprehensive gain (loss)				
Unrealized gain (loss) on marketable securities	(20)	(122)	(476)	105
Comprehensive loss	\$ (20,557)	\$ (11,732)	\$ (35,886)	\$ (22,812)
Net loss per common share, basic and diluted	\$ (0.34)	\$ (0.28)	\$ (0.59)	\$ (0.56)
Weighted-average shares outstanding, basic and diluted	60,079,778	41,025,062	59,945,889	41,025,062



**Acumen Pharmaceuticals, Inc.**  
**Condensed Statements of Cash Flows**  
**(in thousands)**  
**(unaudited)**

	<b>Six Months Ended June 30,</b>	
	<b>2024</b>	<b>2023</b>
<b>Cash flows from operating activities</b>		
Net loss	\$ (35,410)	\$ (22,917)
Adjustments to reconcile net loss to net cash used in operating activities:		
Depreciation	33	29
Stock-based compensation expense	4,954	2,911
Amortization of premiums and accretion of discounts on marketable securities, net	(3,222)	(634)
Change in fair value of embedded derivatives	(1,050)	—
Amortization of right-of-use asset	56	76
Realized gain on marketable securities	(2)	—
Non-cash interest expense	539	—
Changes in operating assets and liabilities:		
Prepaid expenses and other current assets	(3,350)	(1,933)
Other assets	(7)	(57)
Accounts payable	2,823	384
Accrued clinical trial expenses	2,640	1,385
Accrued expenses and other current liabilities	(2,335)	(1,013)
Finance lease liability	(23)	—
Operating lease liability	(50)	(76)
Net cash used in operating activities	<u>(34,404)</u>	<u>(21,845)</u>
<b>Cash flows from investing activities</b>		
Purchases of marketable securities	(57,093)	(52,131)
Proceeds from maturities and sales of marketable securities	85,605	21,268
Proceeds from sale of property and equipment	—	—
Purchases of property and equipment	(16)	—
Net cash provided by (used in) investing activities	<u>28,496</u>	<u>(30,863)</u>
<b>Cash flows from financing activities</b>		
Proceeds from issuance of common stock, net of issuance costs	7,938	—
Payment for financing lease	(739)	—
Proceeds from the exercise of stock options	—	—
Payments for deferred offering costs	(188)	(145)
Repurchase of common shares to pay employee withholding taxes	(32)	—
Net cash provided by (used in) financing activities	<u>6,979</u>	<u>(145)</u>
Net change in cash and cash equivalents and restricted cash	<u>1,071</u>	<u>(52,853)</u>
Cash and cash equivalents and restricted cash at the beginning of the period	<u>67,119</u>	<u>130,101</u>
Cash and cash equivalents and restricted cash at the end of the period	<u>\$ 68,190</u>	<u>\$ 77,248</u>



## Corporate Presentation

August 2024



## Forward-Looking Statements

This presentation contains forward-looking statements within the meaning of The Private Securities Litigation Reform Act of 1995. Any statement describing Acumen's goals, expectations, financial or other projections, intentions or beliefs is a forward-looking statement and should be considered an at-risk statement. Words such as "believes," "expects," "anticipates," "could," "would," "seeks," "aims," "plans," "potential," "will" and similar expressions are intended to identify forward-looking statements, although not all forward-looking statements contain these identifying words. Forward-looking statements include statements concerning Acumen's business, and Acumen's ability to achieve its strategic and financial goals, including its projected use of cash, cash equivalents and marketable securities and the expected sufficiency of its cash resources into the first half of 2027, the therapeutic potential of Acumen's product candidate, sabirnetug (ACU193), including against other antibodies, and the anticipated timeline for announcing the top-line results from our Phase 1 trial of a subcutaneous dosing option of ACU193. These statements are based upon the current beliefs and expectations of Acumen management, and are subject to certain factors, risks and uncertainties, particularly those inherent in the process of discovering, developing and commercializing safe and effective human therapeutics. Such risks may be amplified by the impacts of the COVID-19 pandemic. These and other risks concerning Acumen's programs are described in additional detail in Acumen's filings with the Securities and Exchange Commission ("SEC"), including in Acumen's most recent Annual Report Form 10-K and future filings and reports by Acumen. Copies of these and other documents are available from Acumen. Additional information will be made available in other filings that Acumen makes from time to time with the SEC. These forward-looking statements speak only as of the date hereof, and Acumen expressly disclaims any obligation to update or revise any forward-looking statement, except as otherwise required by law, whether, as a result of new information, future events or otherwise. In this presentation, references to cash also include cash equivalents.

## Advancing a Next Generation Antibody Targeting Toxic Amyloid Beta Oligomers (A $\beta$ O) for Early Alzheimer's Disease (AD)



Large market in need of additional treatment options



Sabirnetug (ACU193): monoclonal antibody (mAb) highly selective for toxic A $\beta$ O



Positive Phase 1 clinical trial results presented in 2H 2023



Experienced leadership team with extensive AD drug development experience

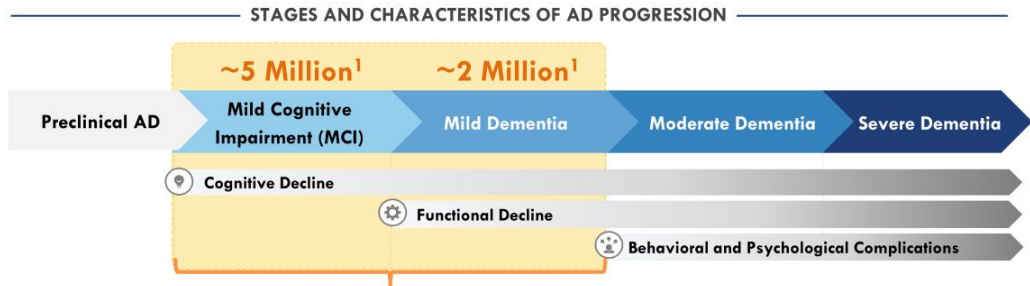


Strong balance sheet supporting clinical development plans for sabirnetug



Phase 2 (IV) initiated in May 2024; Phase 1 (subcutaneous) TLR expected in 1Q25

# Early AD Patient Population Represents Significant Market Opportunity



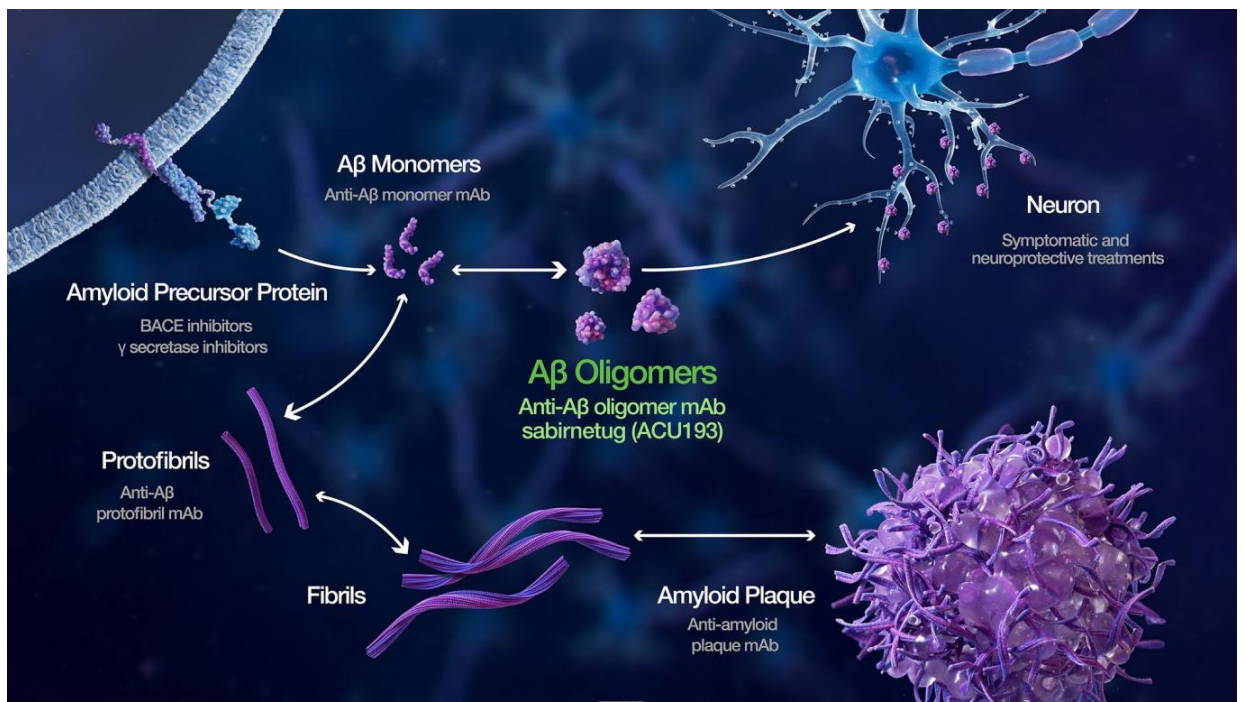
## Early Alzheimer's Disease in the U.S. *Acumen's commercial priority*

Uptake of first-generation, disease modifying, anti-amyloid beta treatment options is expected to increase, while significant unmet need and room for improvement will persist

1. 2021 Alzheimer's Association

# AD, Amyloid & Abeta Oligomers

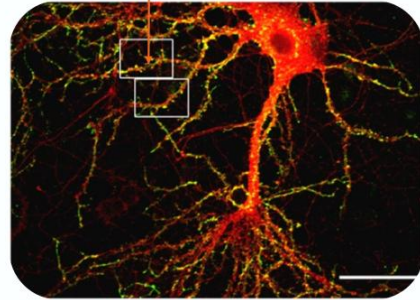
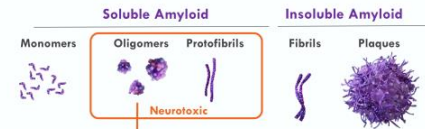






## Amyloid Beta Oligomers (A $\beta$ O<sub>s</sub>) are Widely Recognized as Highly Toxic Agents in AD Pathophysiology

- ➔ Impair synaptic function<sup>1</sup>
- ➔ Contribute to impairment of memory and cognition<sup>2</sup>
- ➔ Induce tau hyperphosphorylation<sup>3</sup>



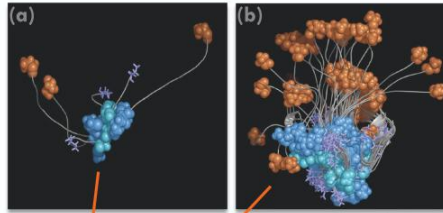
Mature hippocampal neuron and toxic A $\beta$ O<sub>s</sub> bound to dendritic spines

Image Lacor et al., 2004.

1. Lacor et al., 2004 & 2007; Townsend et al., 2006; Batista et al., 2018
2. Cleary et al., 2005; Foling et al., 2008; Cline et al., 2019
3. De Felice et al., 2008; Zempel et al., 2010

## What is an A $\beta$ Oligomer? A $\beta$ O<sub>s</sub> May Consist of 2 to >200 A $\beta$ Peptides

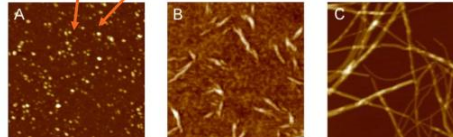
Figure 1. A $\beta$ O<sub>s</sub> composed of 3 (a) and 18 (b) A $\beta$  peptides are depicted below.



Source: Kelley et al. *J Chem Physics* 2008.

### Quaternary structures of A $\beta$ oligomers, protofibrils, and fibrils

Figure 2. Atomic force microscopy images of representative steps of amyloid aggregation: (A) oligomers; (B) protofibrils; (C) mature fibrils. Scan size 1.0  $\mu$ m. Z range (A) 8.0 nm; (B) 15 nm; (C) 20 nm.

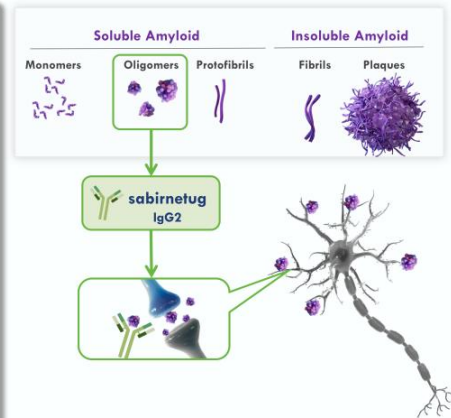


Source: Relini et al. *Biomolecules* 2014.

## Sabirnetug: Potential Best-in-Class Immunotherapy for Early AD

*Sabirnetug's High Selectivity for Toxic A $\beta$ O<sub>s</sub> May Provide Meaningful Cognitive Efficacy and Improved Safety*

<p><b>Rationally Designed for Improved Efficacy &amp; Safety</b></p>	<p><b>Humanized, affinity matured mAb developed to target toxic A<math>\beta</math> oligomers</b></p> <ul style="list-style-type: none"> <li>&gt; 500-fold greater selectivity for A<math>\beta</math>O<sub>s</sub> over A<math>\beta</math> monomers</li> <li>&gt; 85-fold greater selectivity for A<math>\beta</math>O<sub>s</sub> over A<math>\beta</math> fibrils</li> </ul> <p><b>IgG2 subclass mAb with reduced effector function</b></p>
<p><b>Large Pharma Discovery</b></p>	<p><b>Sabirnetug discovered in collaboration with Merck &amp; Co.</b></p> <p>Acumen holds exclusive program rights with no future financial or other obligations due to Merck</p>
<p><b>Encouraging FDA Interactions</b></p>	<p><b>FDA Fast Track designation for the treatment of early Alzheimer's disease</b></p> <p><b>FDA End of Phase 2 meeting in 4Q 2023</b></p>



## Sabirnetug: Value Proposition



# Positive INTERCEPT-AD Phase 1 Results for Sabirnetug



# INTERCEPT-AD: A Randomized Placebo Controlled Phase 1 in Early AD Patients

**PART A:**  
**SINGLE-ASCENDING DOSE**  
 n = 8 per cohort (32 total)  
 6:2 per cohort

**COHORT 1:**  
 2 mg/kg Sabirnetug  
 or Placebo

**COHORT 2:**  
 10 mg/kg Sabirnetug  
 or Placebo

**COHORT 3:**  
 25 mg/kg Sabirnetug  
 or Placebo

**COHORT 4:**  
 60 mg/kg Sabirnetug  
 or Placebo

**PART B:**  
**MULTIPLE-ASCENDING DOSE**  
 n = 10 per cohort (30 total)  
 3 administrations of drug or PBO  
 8:2 per cohort

**COHORT 5:**  
 10 mg/kg Sabirnetug  
 or Placebo (Q4W)

**COHORT 6:**  
 60 mg/kg Sabirnetug  
 or Placebo (Q4W)

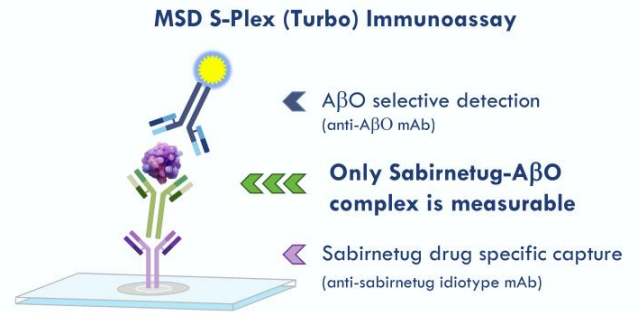
**COHORT 7:**  
 25 mg/kg Sabirnetug  
 or Placebo (Q2W)\*

Q2W: Dosing every two weeks; Q4W: Dosing every four weeks.

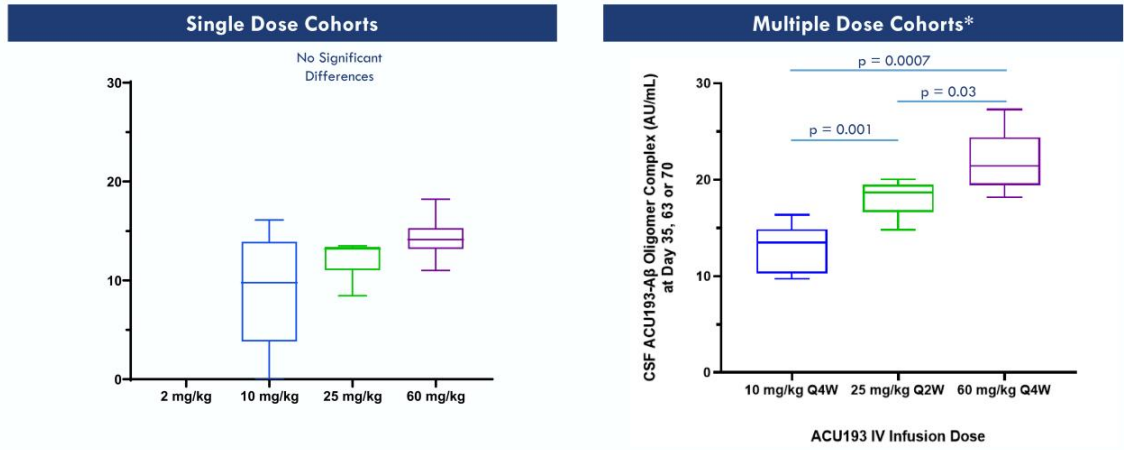
## Target Engagement Assessed by Measuring Sabirnetug-A $\beta$ O Complex in CSF

- Novel assay configuration tailored to selectively detect sabirnetug-A $\beta$ O complex in CSF as direct measure of target engagement
- Translated for clinical use from a preclinical assay developed by Merck that showed sabirnetug engages target A $\beta$ O in transgenic mouse brain (tg2576) in dose dependent manner

*(please see slide 41 for more information)*



## Target Engagement of Sabirnetug with A $\beta$ O $_s$ is Dose Proportional

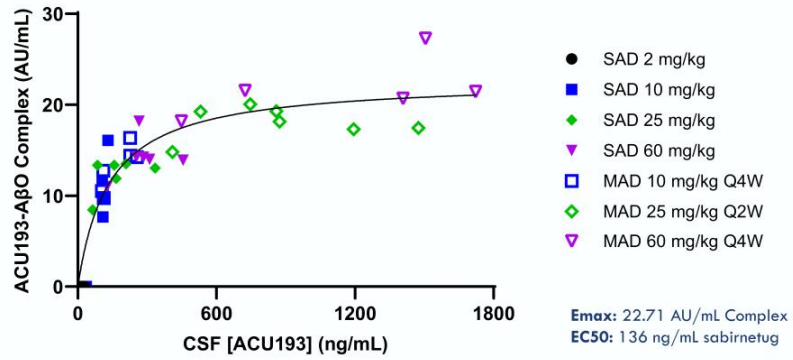


\*One patient from Cohort 5 (10 mg/kg Q4W) excluded because only received one administration of drug (study drug discontinued after lacunar infarct).



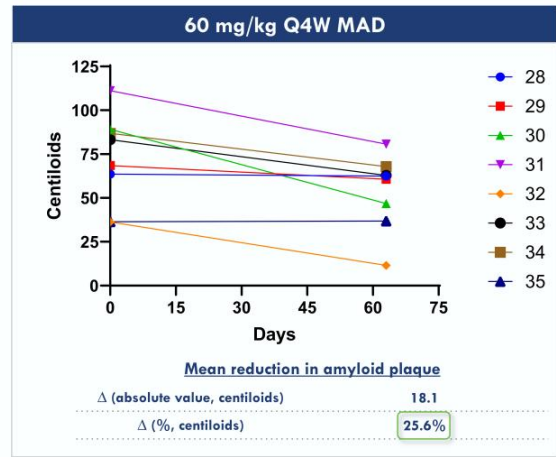
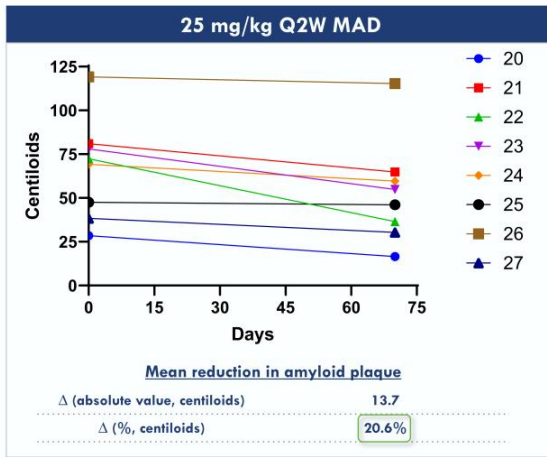
## Doses Approaching Maximal Target Engagement Support Sabirnetug A $\beta$ O Mechanism and Helped Guide Dose Selection for Next Study Phase

Single & Multiple Dose Cohorts - Exposure Response Relationship (Emax Model)



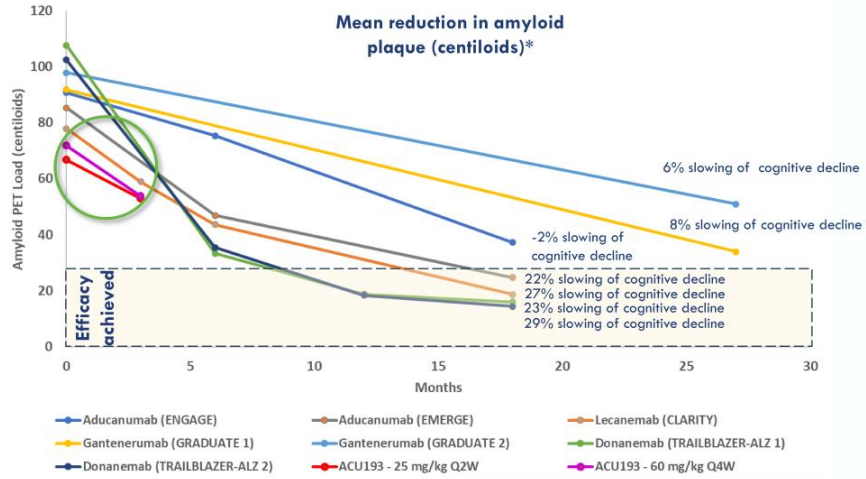
\*One patient from Cohort 5 (10 mg/kg Q4W) excluded because only received one administration of drug (study drug discontinued after lacunar infarct).

## Nearly All Sabirnetug-Treated Patients in High Dose MAD Cohorts Showed Reductions in Plaque Load After Three Doses at 63 or 70 days



Plaque load based on florbetapir PET

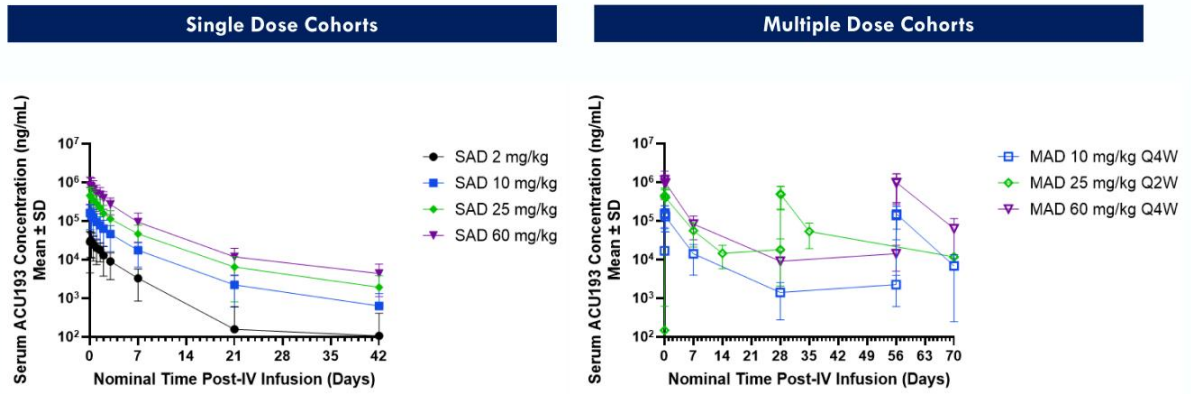
## Highest Doses of INTERCEPT-AD Reduced Amyloid Plaque at Similar Rate and Magnitude to Lecanemab at Comparable Timepoints



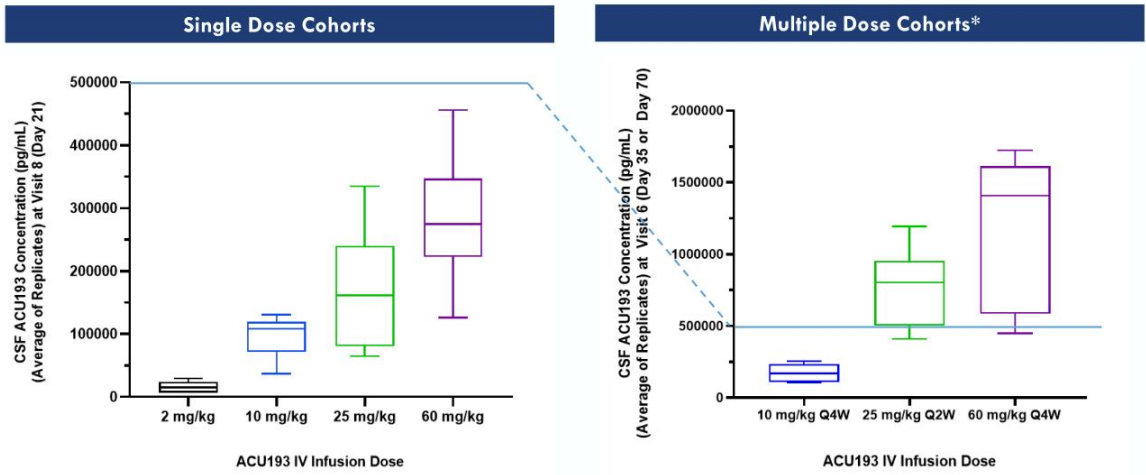
Acumen Pharmaceuticals, data on file; van Dyck (2023), NEJM (amyloid PET reduction estimated from graphs).

\*There have been no head-to-head clinical trials between the product candidates listed above. Study designs and protocols for each product candidate were different, and as a result, results may not be comparable between product candidates.

## Sabirnetug Serum Exposure is Dose Proportional Without Accumulation



## Sabirnetug CSF Exposure is Dose and Dose-Regimen Proportional

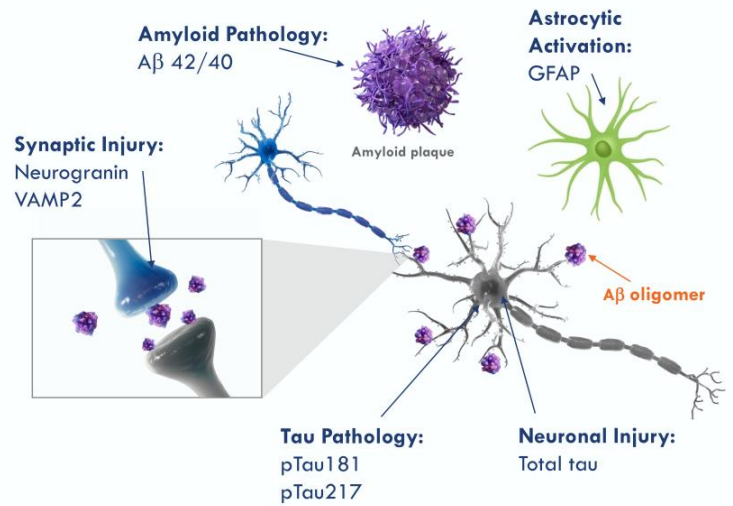


\*One patient from Cohort 5 (10 mg/kg Q4W) excluded because only received one administration of drug (study drug discontinued after lacunar infarct).

## Importance of Key Fluid Biomarkers Associated with AD Pathology

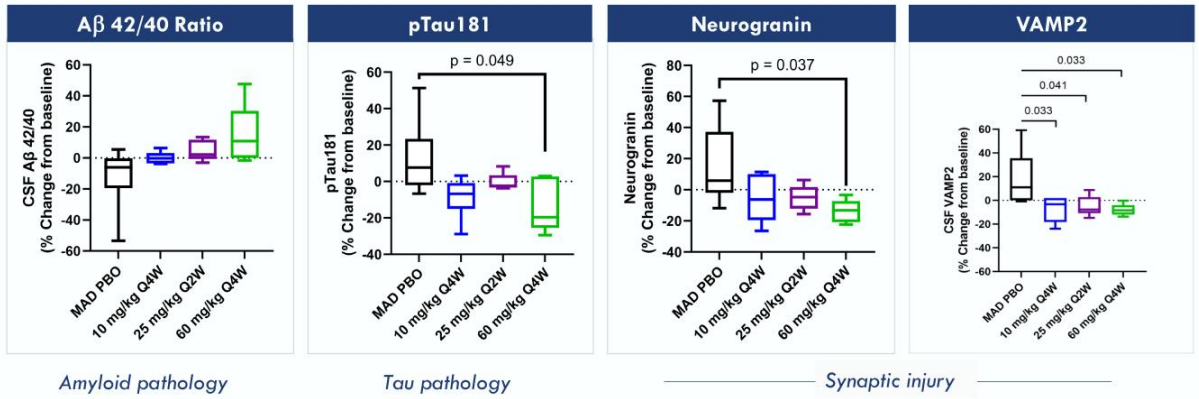
- Biomarkers from cerebrospinal fluid and plasma capture neuronal, synaptic, and axonal injury and reflect the cumulative outcome of different pathological substrates in AD<sup>1</sup>
- Evidence suggests that biomarkers are likely to be better predictors of the underlying pathology of AD than imaging alone<sup>2</sup>

• **After just three administrations of sabirnetug, patients with early AD demonstrated improvements in biomarkers associated with AD pathology**



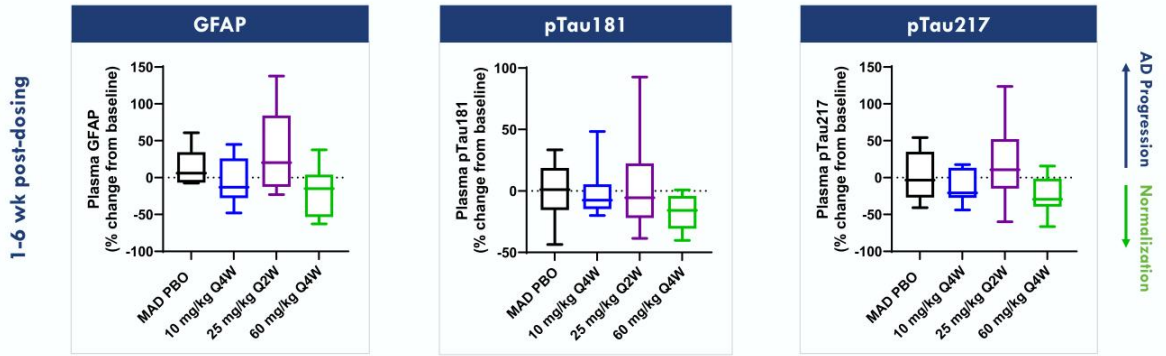
1. Tarawneh, R. Biomarkers: Our Path Towards a Cure for Alzheimer Disease. Biomarker Insights Volume 15: 1–15. 2020; 2. Blennow K, Zetterberg H. The Past and the Future of Alzheimer's Disease Fluid Biomarkers. J Alzheimers Dis. 2018;62(3):1125-1140.

## Consistent Improvement in CSF Amyloid, Tau and Synaptic Biomarkers Indicate Downstream Pharmacology of Sabirnetug After Only Three Doses



n = 8 subjects/treated group; 6 subjects in pooled placebo (PBO); p-values from unpaired, 2-sided Student's t test

## Trend Toward Normalizing Plasma Biomarkers with 10 mg/kg and 60 mg/kg Q4W



- Plasma measurements of glial fibrillary acidic protein (GFAP), pTau181, and pTau217 in 10 mg/kg Q4W & 60 mg/kg Q4W groups were lower than placebo
- More impact to fluid biomarkers was observed with longer dosing duration
  - The 25 mg/kg Q2W cohort differed in dose and sample timing, with drug on board for less time than the 10 mg/kg & 60 mg/kg Q4W cohorts

*n* = 8 subjects/treated group; 6 subjects in pooled placebo (PBO); *p*-values from unpaired, 2-sided Student's *t* test



## Sabirnetug Demonstrates Potential for Best-in-Class Safety

Compelling Overall Safety Profile, with Low Incidence of ARIA-E

### INTERCEPT-AD Phase 1 Safety Data

**5** Total ARIA-E cases,  
or ~10%

**0** Cases of ARIA-E in  
ApoE4 homozygotes  
N=6

**0** Deaths, SAEs Related  
to Study Drug

- ✓ **Limited incidence of ARIA-E**
  - 10 mg/kg Q4W: 1 asymptomatic case
  - 25 mg/kg Q2W: 1 asymptomatic case
  - 60 mg/kg Q4W: 2 asymptomatic cases; 1 symptomatic case
- ✓ **No ARIA-E observed in ApoE4 homozygotes (n=6), despite comprising 13% of study**
  - Differentiated from other antibodies that have ARIA-E rates ~30% to ~40% in participants who are E4-homozygotes
- ✓ **Broad therapeutic index** with convenient monthly dosing
  - Safety profile may support attractive benefit/risk option for large portion of patients

## INTERCEPT-AD Phase 1 Data Support Potential for Sabinetug to Offer Best-in-Class Efficacy and Safety

### Key Takeaways from INTERCEPT-AD

#### Potential for Differentiated Efficacy

- ✓ First mAb to demonstrate selective target engagement of A $\beta$ O $_2$ s (most toxic form of A $\beta$ )
- ✓ Rapid, significant plaque reduction comparable to the current market front-runners at similar timepoints
- ✓ Improvement of AD biomarkers in CSF and plasma are a strong indication of downstream effects

#### Potential for Differentiated Safety

- ✓ Compelling safety profile with low incidence of ARIA-E
- ✓ Absence of ARIA-E observed in ApoE4 homozygotes
- ✓ Broad therapeutic index with convenient monthly dosing

# Clinical Development Plans & Strategic Considerations



## ALTITUDE-AD Study

Currently Enrolling

**Objective:** To evaluate the clinical efficacy, safety and tolerability of sabirnetug

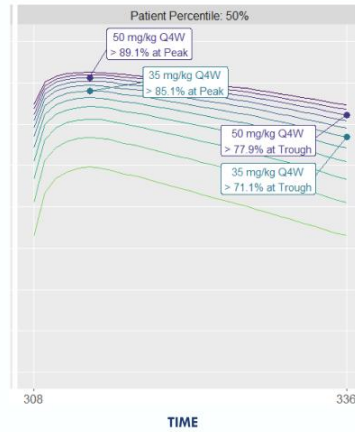
**Patient population:** Patients with early AD (MCI or mild dementia due to early AD)



1. iADRS: Integrated Alzheimer's Disease Rating Scale; CDR-SB: Clinical Dementia Rating – Sum of Boxes; ADAS-cog: Alzheimer's Disease Assessment Scale – Cognitive Subscale; ADCS-ADL: Alzheimer's Disease Cooperative Study – Activities of Daily Living

## Simulated CSF Target Engagement at Steady-State for ALTITUDE-AD Doses

CSF target engagement was simulated at a candidate list of doses given Q4W at steady-state



### Ph2 Dosing Strategy (ALTITUDE-AD)

lower dose: 35 mg/kg Q4W

upper dose: 50 mg/kg Q4W

- Notable **diminishing differentiation** as dose increases
- Doses were selected with **peak-trough** variation in mind: select doses based on trough (end of dosing interval) CSF engagement

## Sabirnetug Subcutaneous Formulation Under Development in Collaboration with Halozyme

*Potential to Broaden Patient Access and Increase Treatment Convenience*

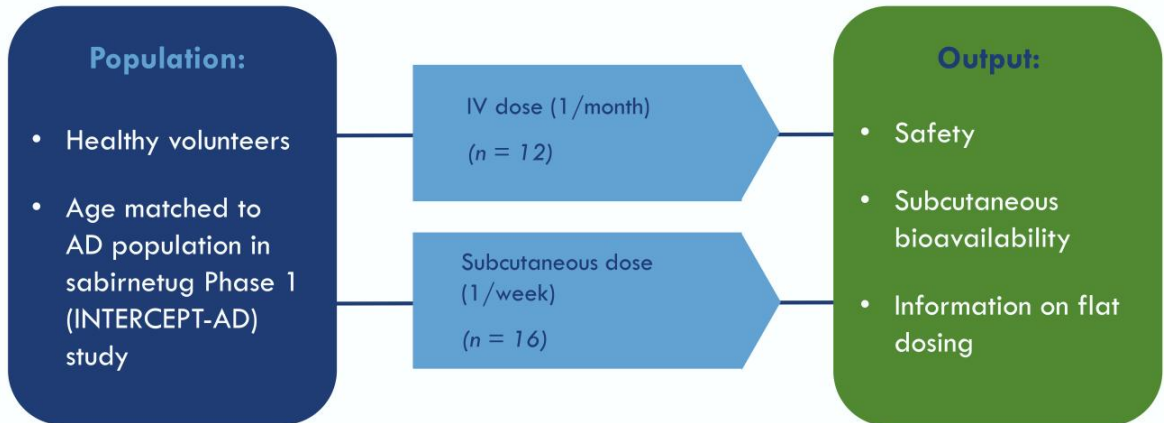


- Announced partnership with Halozyme in November 2023 to develop subcutaneous dosing option for sabirnetug
- Halozyme's drug delivery technology, ENHANZE®, is commercially validated in eight approved therapies available in 100+ countries, with >800,000 patients treated
- Current sabirnetug potential target product profile inclusive of no more than single weekly injection

Phase 1 bioavailability study ongoing to compare the pharmacokinetics of subcutaneous form of sabirnetug to the IV form

## Ongoing Phase 1 Subcutaneous Healthy Volunteer Study

Topline Results Expected in Q1 2025



# Acumen Leadership Team

Experienced in AD/Neuro Drug Development



**DANIEL O'CONNELL**  
Chief Executive Officer  
ACUMEN  
neuroVentures



**JAMES DOHERTY, PHD**  
President & Chief Development Officer  
ACUMEN  
Sage Therapeutics AstraZeneca



**ERIC SIEMERS, MD**  
Chief Medical Officer  
ACUMEN  
Lilly



**MATT ZUGA**  
Chief Financial Officer & Chief Business Officer  
ACUMEN  
HIGHCAPE PARTNERS



**RUSSELL BARTON**  
Chief Operating Officer  
ACUMEN  
Lilly



**JANICE HITCHCOCK, PHD**  
VP, Regulatory Affairs  
ACUMEN  
Lilly



**LIEAN SCHENK**  
VP, Head of CMC  
ACUMEN  
Lilly LONZA NOVAVAX



**SIEW TIN GAN**  
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**JASNA JERICIC, PHD**  
Analytical Methods Leader, Research Scientist  
ACUMEN



**DEREK MEISNER, JD**  
Chief Legal Officer  
ACUMEN  
X4



**JULIE BOCKENSTETTE**  
Executive Vice President, Head of HR  
ACUMEN  
Roche Lilly

Acumen team has decades of experience in Alzheimer's drug discovery and development



## Sabirnetug IP & Market Exclusivity

- Exclusive, perpetual, irrevocable, worldwide, royalty-free license from Merck to its Amyloid Derived Diffusile Ligand (ADDL) IP including issued sabirnetug patents
- Sabirnetug Global IP estate:
  - ✓ Issued patents in 19 countries
  - ✓ Composition of matter patents and methods of use run into July 2031
  - ✓ Patent term extensions may be available, 3-5 years depending on jurisdiction
- Biologics market exclusivity is expected for sabirnetug as a novel biologic drug
  - ✓ US provides 12 years market exclusivity for novel biologics
  - ✓ Europe provides 10 years of market exclusivity for novel biologics

## Milestones Achieved in 2024 and Anticipated in 2025

MILESTONES	STATUS/ EXPECTED TIMING
Initiation of ALTITUDE-AD Phase 2 trial	✓
Initiation of Phase 1 subcutaneous trial	✓
Expected Phase 1 subcutaneous topline results	1Q25
Expected completion of enrollment of ALTITUDE-AD	TBD

~\$281M

Cash, cash equivalents and marketable securities as of June. 30, 2024

We believe that Acumen has the expertise and resources to advance sabirnetug into the first half of 2027

## Summary

### Key Takeaways

- ✓ Significant and growing Alzheimer's population in need of additional treatment options
- ✓ Sabirnetug demonstrates high selectivity for toxic A $\beta$ O<sub>s</sub> in AD patients
- ✓ Positive Phase 1 data strengthen potential for sabirnetug to offer best-in-class efficacy and safety
- ✓ Phase 2 IV study and Phase 1 subcutaneous study ongoing

### Next Steps

- ➔ Anticipate Phase 1 subcutaneous healthy volunteer topline results in Q1 2025
- ➔ Currently enrolling Phase 2 ALTITUDE-AD study

# Appendix

[www.acumenpharm.com](http://www.acumenpharm.com)



# Preclinical Data



# Sabirnetug: Extensive Data Package Supporting Development

## SELECTIVITY

- Nanomolar affinity for A $\beta$ O $_2$ , >500-fold greater selectivity for A $\beta$ O $_2$  over A $\beta$  monomer, with limited or no discernable binding to vascular amyloid or dense core amyloid plaques
- Binds broad range of endogenous A $\beta$ , from dimers to high molecular weight A $\beta$ O $_2$

## PHARMACOLOGY

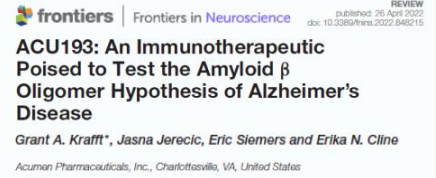
- Dose-dependent effects in multiple in vitro neuroprotection assays
- Positive memory and behavioral effects in multiple in vivo transgenic mouse models for AD

## PK/PD

- Brain penetration and biodistribution demonstrated in multiple species
- Performs like other peripherally administered CNS mAbs

## SAFETY

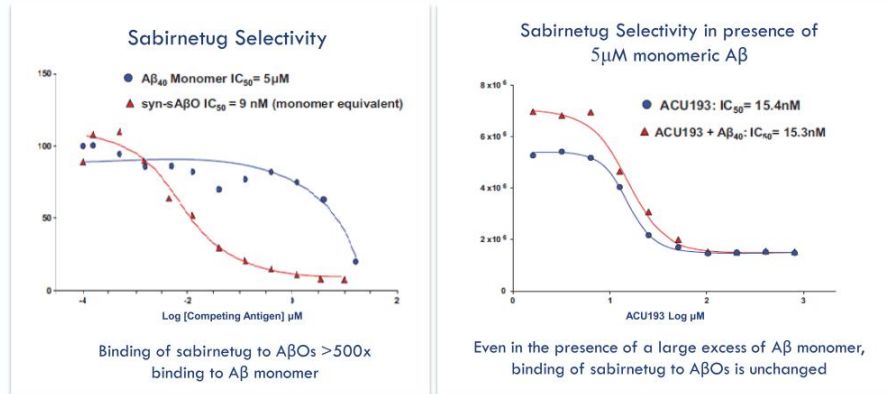
- IgG2 subclass lacks inflammatory effector function signaling (Fc $\gamma$ R binding)
- Nonclinical microhemorrhage studies show no increased risk of microhemorrhage
- GLP studies demonstrated acceptable safety supporting clinical dosing plans including Ph 2



Sabirnetug is a promising immunotherapy for early AD expected to provide meaningful cognitive and functional benefits, slow disease progression, and offer an attractive safety profile.

## Sabirnetug is the First mAb Developed to Selectively Target A $\beta$ O<sub>s</sub>

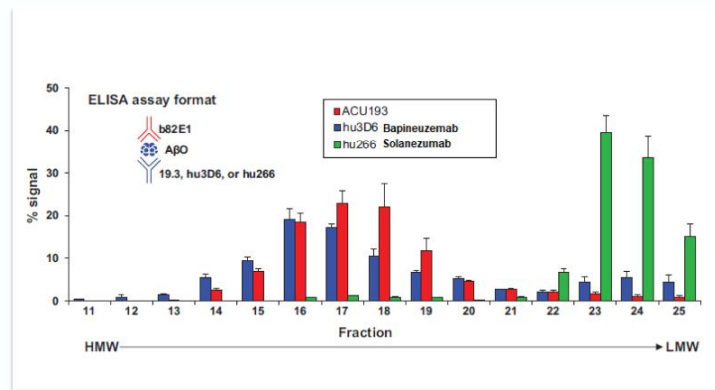
Highly selective for A $\beta$  oligomers versus A $\beta$  monomers



Sabirnetug selective for binding to A $\beta$ O<sub>s</sub> is preserved even in the presence of a large excess of A $\beta$  monomers – such as what is present in the brain, thus limiting ‘target distraction’

## Sabirnetug Binds to a Wide Range of Oligomeric Species of A $\beta$

Comparison of A $\beta$  species-mAb complex signals across SEC fractions

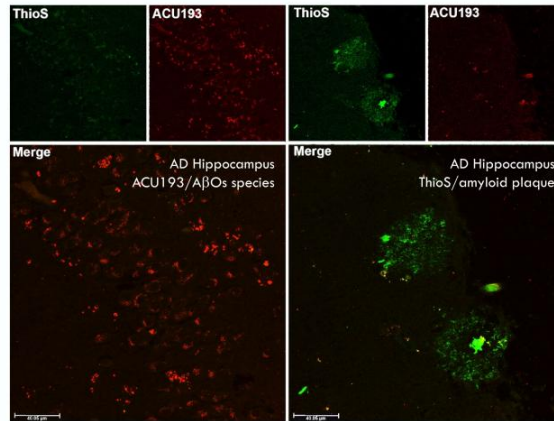


Sabirnetug binds to oligomeric species of A $\beta$  that are differentiated from those bound by hu266 (solanezumab) or hu3D6 (bapineuzumab)



## Sabirnetug is Highly Selective for A $\beta$ O<sub>s</sub> Versus A $\beta$ Plaques

Sabirnetug staining in human AD brain slices sabirnetug (red) binds non-Thioflavin S positive A $\beta$  (green)

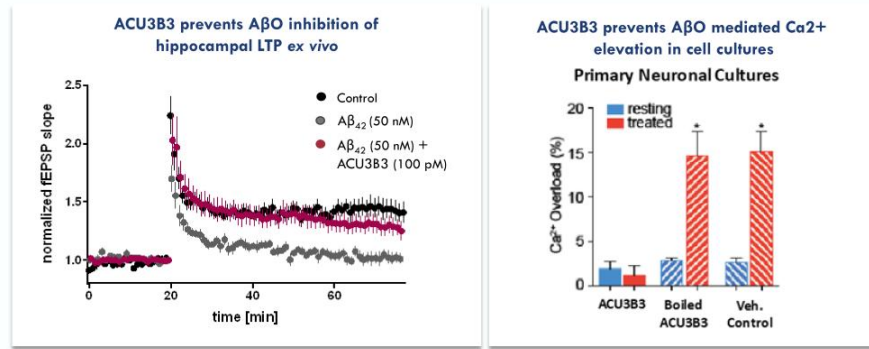


Sabirnetug has little or no binding to thioflavin S positive fibrillar A $\beta$  plaque in human AD brain tissue

Sources: E. Cline et al. CTAD 2019.

## AβOs Bind to Neurons and are Toxic; Mouse Analogue of Sabirnetug Prevents Toxicity

After binding to neurons, AβOs disrupt Long Term Potentiation (LTP) and cause pathologic increases in intracellular calcium that is destructive to cells.

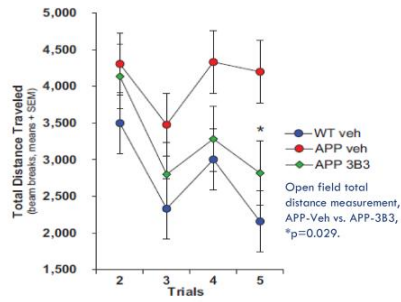


Note: (1) ACU3B3 is the mouse monoclonal antibody precursor to and equivalent of humanized sabirnetug (ACU193)

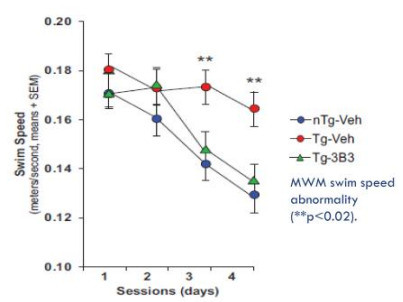
ACU3B3 prevents changes in aberrant neuronal activity thought to underlie memory loss in AD and prevents AβO mediated disruption of calcium homeostasis in neuronal cultures

## Treatment of a Transgenic Mouse Model of AD Results in Behavioral Improvements

Murine parent version of sabirnetug (ACU3B3) was used to treat younger mice with depositing plaque or older mice with abundant plaque

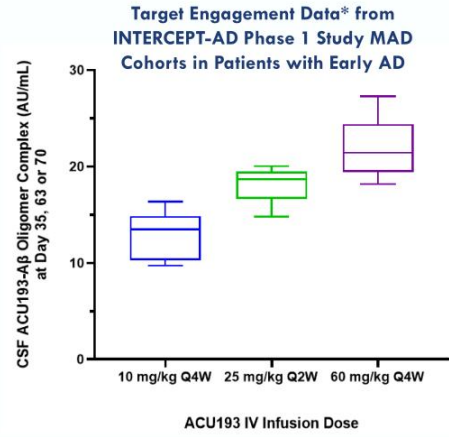
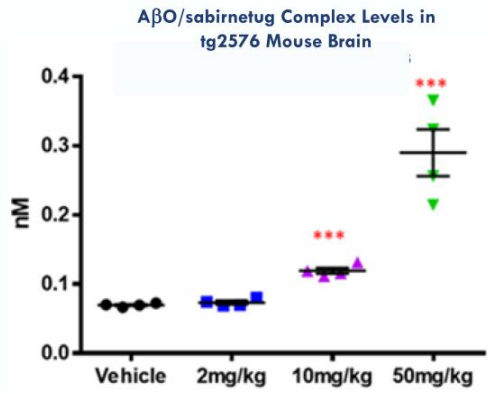


Deficits in younger (5-7 months) transgenic mice are markedly reduced with treatment



Deficits in older (9-10 months) transgenic mice are markedly reduced with treatment

## Sabirnetug Enters the CNS and Binds to A $\beta$ O<sub>s</sub> in Transgenic Mice and Patients with Early AD in a Dose Dependent Manner



Sabirnetug engages target A $\beta$ O<sub>s</sub> in transgenic mouse brain (tg2576) and is found in CSF of patients with early AD

\*One patient from Cohort 5 (10 mg/kg Q4W) excluded because only received one administration of drug (study drug discontinued after lacunar infarct).

