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# **FY 2021 Financial Results & Business Highlights**

March 28, 2022

# 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, Acumen's ability to achieve its strategic and financial goals, including its projected use of cash, cash equivalents and marketable securities and the sufficiency of its cash resources, and the therapeutic potential of Acumen's product candidate, ACU193, including its potential for improved safety and efficacy as compared to other monoclonal antibodies in development, as well as the expectations concerning the INTERCEPT-AD trial and Acumen's planned Phase 2/3 clinical trial, including the expected timing of initiation, enrollment and reporting data, and risks and uncertainties relating to the progression and duration of the COVID-19 pandemic and responsive measures thereto and related effects on Acumen. 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 Quarterly Report on Form 10-Q for the quarter ended September 30, 2021, and future filings and reports by Acumen, including Acumen's Annual Report on Form 10-K for the year ended December 31, 2021. 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 Potential Best-/First-In-Class Antibody for Early Alzheimer's disease (Early AD)



Alzheimer's Represents an Enormous Market Driven by High Unmet Need and Recent Scientific and Regulatory Momentum



Scientific Consensus Supports **Amyloid-Beta Oligomers (A $\beta$ O<sub>s</sub>)** as the Most toxic form of A $\beta$  and a Novel Target for Effective AD Treatment



**ACU193**: First, Clinical-Stage Monoclonal Antibody (mAb) to Selectively Target A $\beta$ O<sub>s</sub> and has Promising Pre-Clinical Evidence Supporting its Differentiation



**Experienced Leadership** Comprised of Industry Leaders with AD Drug Discovery, Development, and Regulatory Expertise from Eli Lilly & Co.



Strong Balance Sheet:  
2021 Series B Tranche \$30M  
2021 IPO  
~\$184M Gross  
~\$225M in Cash and Marketable Securities at 12/31/21



Phase 1 Clinical Trial in Early AD Patients Ongoing  
Proof of Mechanism / Target Engagement / Safety Data  
**Topline Results Expected 1H 2023**

We believe Acumen has the organizational expertise and fiscal resources to advance ACU193 through multiple anticipated clinical development milestones during 2022 through 2025

# Strong Momentum Heading into 2022

- **2021: Transformational year for Acumen**

- ✓ Series B Tranche gross proceeds of \$30M and IPO with gross proceeds of \$184M
- ✓ INTERCEPT-AD trial launched; first patient dosed in October
- ✓ Trial design and program presented at CTAD in November
- ✓ December 31, 2021: Cash, cash equivalents and marketable securities of ~\$225M representing expected cash runway through 2025









- **Expanded team to enhance depth and breadth of expertise of the Company**

- ✓ Appointed three new senior team members in 4Q 2021
  - Head of Clinical Operations
  - Head of HR
  - Corporate Controller and Treasurer
- ✓ Kim Drapkin, CPA, appointed to the Board and to serve as Chair of Audit Committee effective April 1, 2022

# INTERCEPT-AD Trial Update

- **INTERCEPT-AD: Phase 1 clinical trial of ACU193 in patients with early AD (RCT)**
  - Topline results under full database lock expected in 1H 2023
    - Safety / ARIA-E
    - PK
    - Target engagement
  - Trial timeline also adjusted for COVID-related impact
  - Trial enrollment on-going at 8 active sites, 6 additional sites selected for potential activation
  - Strong cash position has provided us the ability to expand study footprint to support recruitment and capture complete follow-up period (Cohort 7 Day 168) prior to read out
  - Complete trial results anticipated for presentation at major Alzheimer's meeting mid-2023
- **Phase 2/3 'Ready' Activities**
  - ✓ Chronic GLP toxicity testing initiated
  - ✓ New drug substance production process and drug product formulation being finalized
  - ✓ Developing Phase 2/3 study design and planning for FDA End of Phase 2 meeting

# ACU193's High Selectivity for toxic A $\beta$ O<sub>2</sub>s, Combined with its Expected Lack of ARIA-related Safety Concerns, Is Anticipated to Provide Superior Efficacy Compared to Peers

Company	Asset	TARGET SELECTIVITY <sup>+</sup>				SAFETY PROFILE
		Amyloid plaque	A $\beta$ fibrils	A $\beta$ monomers	A $\beta$ oligomers	Lack of ARIA
 ACUMEN	ACU193	✗	untested	✗	✓	✓
 Biogen	Aduhelm™	✓	✓	✗	✓	✗
 Eisai	lecanemab	✓	✓	✗	✓	✗
 Roche	gantenerumab	✓	✓	✗	✓	✗
 Lilly	donanemab	✓	untested	✗	✗	✗
 Lilly	solanezumab*	✗	✗	✓	✗	✓
 Genentech	crenezumab*	✓	✓	✓	✓	✓
 Pfizer Janssen	bapineuzumab*	✓	✓	✓	✓	✗

<sup>+</sup> There have been no head-to-head trials between any of the product candidates listed above. Study designs and protocols for each product candidate were different, and results may not be comparable between product candidates.

\*Phase 3 discontinued for primary AD indication

# (ACU-001) INTERCEPT-AD trial: Phase 1 Overview

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## **TRIAL DESIGN:**

### Randomized Placebo Controlled Phase 1

- Part A : Single-Ascending Doses
- Part B : Multiple-Ascending Doses

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## **ENROLLMENT CRITERIA:**

### Early AD

- Mild Cognitive Impairment and Mild Dementia due to AD (amyloid positive by PET)

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## **TRIAL OBJECTIVES:**

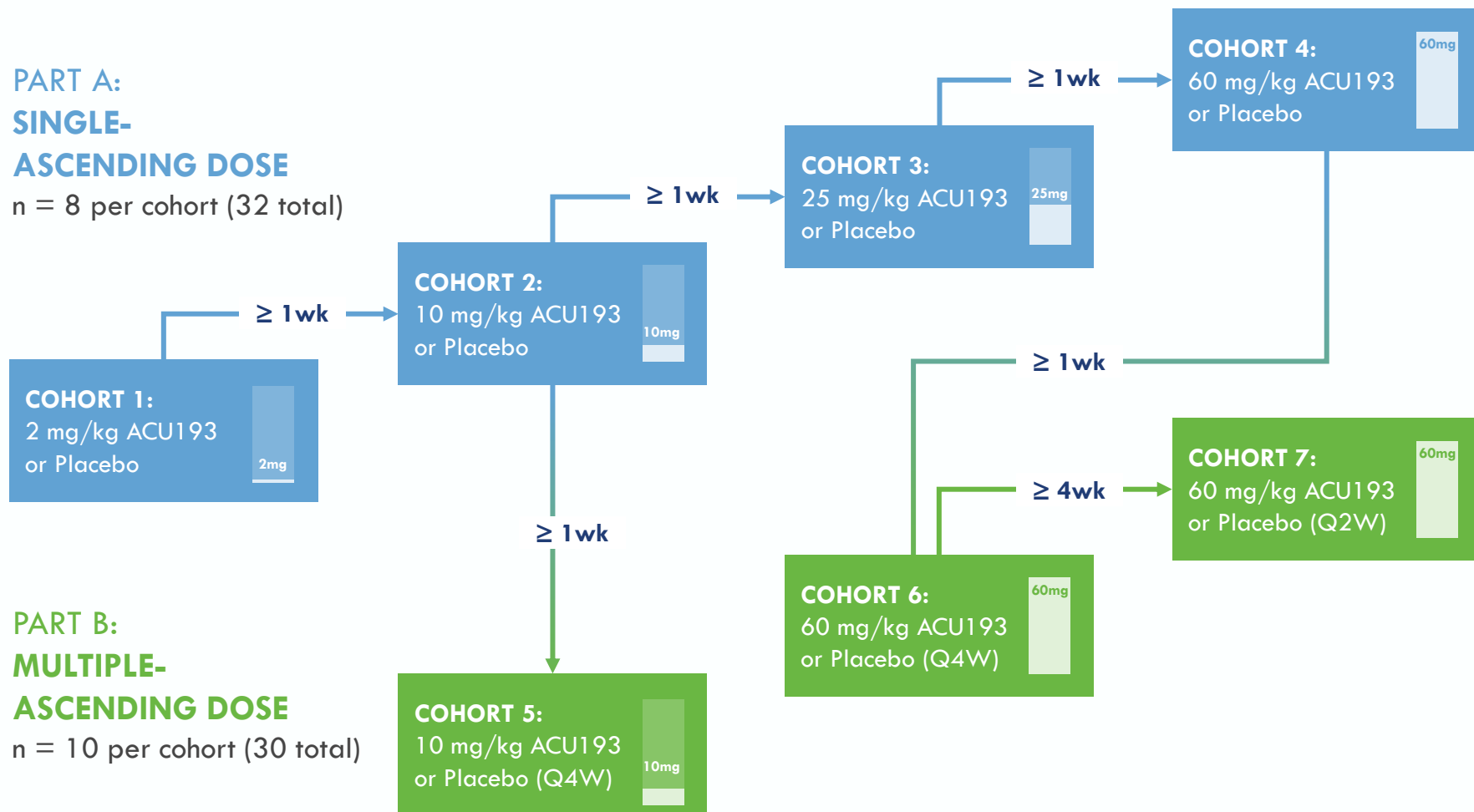
### Proof of Mechanism (PoM)

- Safety and tolerability
  - Pharmacokinetics
  - Target Engagement
  - Biomarkers; cognition
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# INTERCEPT-AD a Randomized Placebo Controlled Phase 1 in Early AD patients

## PART A: SINGLE- ASCENDING DOSE

n = 8 per cohort (32 total)



## PART B: MULTIPLE- ASCENDING DOSE

n = 10 per cohort (30 total)



# Phase 1 Objectives: Proof of Mechanism

## 1. SAFETY AND TOLERABILITY

- Assessment of ARIA-E
- Absence of problematic immunogenicity

## 2. PHARMACOKINETICS

- Peripheral and Central

## 3. EVIDENCE OF TARGET ENGAGEMENT

- CSF level of ACU193:A $\beta$ O complexes (bound)

## 4. FLUID BIOMARKER EFFECTS

- Phospho-tau, Neurofilament light, et. al.

## 5. CLINICAL MEASURES

- Assessment of clinical cognitive measures, computerized tests (Cogstate Ltd.)

## 6. MRI EFFECTS

- Potential improvements in cerebral blood flow shown with MRI ASL pulse sequence



## PROOF OF MECHANISM

### Requirements for Phase 2/3

- ✓ Acceptable safety and tolerability
- ✓ Show ACU193 gets into central compartment
- ✓ Target engagement
- ✓ Other indicators of target mechanism of action

Topline Results anticipated in 1H 2023: primary outcomes Safety / ARIA-E, PK and Target Engagement. Detailed study results anticipated to be presented at major Alzheimer's meeting

# Acumen is Well Capitalized, with Expected Cash Runway through 2025 to Achieve Multiple Anticipated Clinical Milestones

MILESTONES	STATUS/EXPECTED TIMING
Initiated Ph1 clinical trial INTERCEPT-AD	✓
INTERCEPT-AD trial updates	2022
Proof of Mechanism Topline results	1H 2023

**~\$225M**  
Cash and marketable securities  
as of December 31, 2021

## Key FY 2021 Financial Results:

- R&D expenses of \$12.3M
- G&A expenses of \$7.3M
- Loss from operations of \$19.6M

FY 2021 Net Loss of \$101M  
includes non-cash expense of  
more than \$81 million

We believe Acumen has the organizational expertise and cash and marketable securities on hand to advance ACU193 through multiple anticipated clinical milestones 2022 through 2025

Thank you - Q&A