



Zervimesine: a Once-daily Oral Therapeutic Advancing Toward Phase 3

January 2026

Forward-looking Statements

FORWARD-LOOKING STATEMENTS

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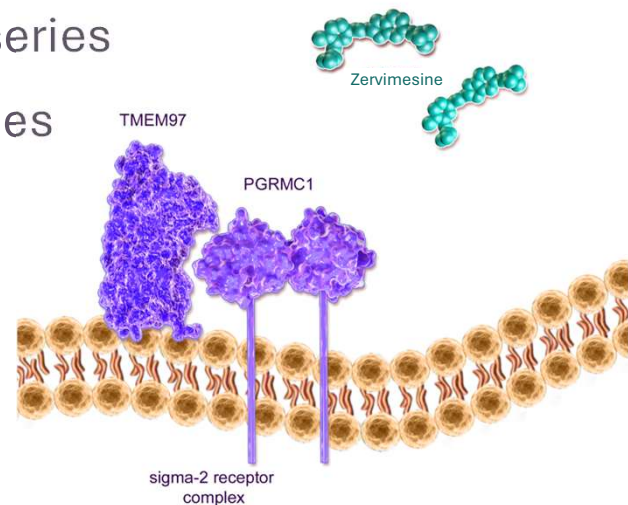
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Projections, estimates, industry data and information contained in this presentation, including the size of and growth in key end markets, are based on information from third-party sources and management estimates. Although we believe that these third party-sources are reliable, we cannot guarantee the accuracy or completeness of these sources. Our management’s estimates are derived from third-party sources, publicly available information, our knowledge of our industry and assumptions based on such information and knowledge. Our management’s estimates have not been verified by any independent source. All of the projections, estimates, market data and industry information used in this presentation involve a number of assumptions and limitations, and you are cautioned not to give undue weight to such information. In addition, projections, estimates and assumptions relating to us and our industry’s future performance are necessarily subject to a high degree of uncertainty and risk due to a variety of factors, including, but not limited to, those described above, that could cause future performance to differ materially from our expressed projections, estimates and assumptions or those provided by third parties.

Novel MoA Discovered Through Founder's Screening Assay

Zervimesine Interrupts Binding of Toxic Oligomers

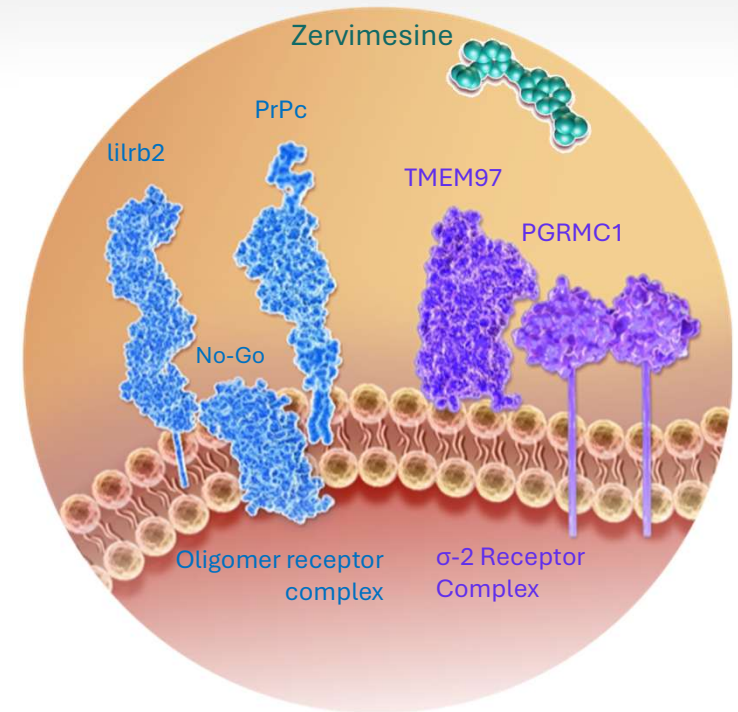
- Phenotypic *in vitro* screen to select molecules that protect neurons from toxic oligomers
- Screened 10,000 compounds, identified 5 unique chemical series
- Zervimesine readily crossed BBB with good drug-like properties
- \$171M in NIH grants funded development through Phase 2
- Zervimesine's MoA - protecting neurons from toxic oligomers – unique, potentially complementary to mAbs



Zervimesine (CT1812) – Lead Product Candidate

Extensive composition of matter IP on zervimesine

- BBB-penetrant small molecule oligomer antagonist
- Target: TMEM97 (sigma-2) receptor
- Oral, once-daily dosing, favorable safety data
- Fast Track granted for Alzheimer's disease
- Potential first-to-market for dementia with Lewy bodies (DLB)



Findings from Completed Studies Support Phase 3 Plans

Patient Population	Program	Treatment period
Alzheimer's disease <i>MCI and Early</i>	Phase 2 COG0203 • START	18 months
Dementia with Lewy bodies <i>Mild-to-moderate</i>	Expanded Access Program	12 months
Completed Studies		
Alzheimer's disease <i>Mild-to-moderate</i>	Phase 2 COG0201 • SHINE	6 months
Dementia with Lewy bodies <i>Mild-to-moderate</i>	Phase 2 COG1201 • SHIMMER	6 months
Dry age-related macular degeneration <i>GA secondary to dry AMD</i>	Phase 2 COG2201 • MAGNIFY	18 months

Takeaways from completed studies

- Phase 2 SHINE Study: efficacy across cognitive measures in mild-to-moderate Alzheimer's disease
- Phase 2 SHIMMER Study: behavioral, functional, cognitive, movement benefit in mild-to-moderate DLB
- Phase 2 MAGNIFY Study: slower lesion growth in geographic atrophy secondary to dry AMD

Regulatory Catalysts for Zervimesine

Anticipate EMA feedback 1Q 2026 for Alzheimer's disease

Completed: AD End-of-Phase 2 Meeting

- Aligned with FDA on following:
 - Disease stage: mild-to-moderate AD
 - Enrichment: p-tau217 at screening $\leq 1.0\text{pg/mL}$
 - Treatment period: 6 months
 - Randomization: 1:1 zervimesine (100mg) vs pbo
 - Endpoints: composite cognitive and functional
 - Open-label extension to follow

Dementia with Lewy Bodies

- FDA Type C meeting scheduled for January 2026
- Will seek alignment with FDA on clinical outcomes that encompass the complex symptomatology of DLB:
 - Neuropsych / behavioral
 - Motor
 - Fluctuations
 - Cognition

Executive Program Summary

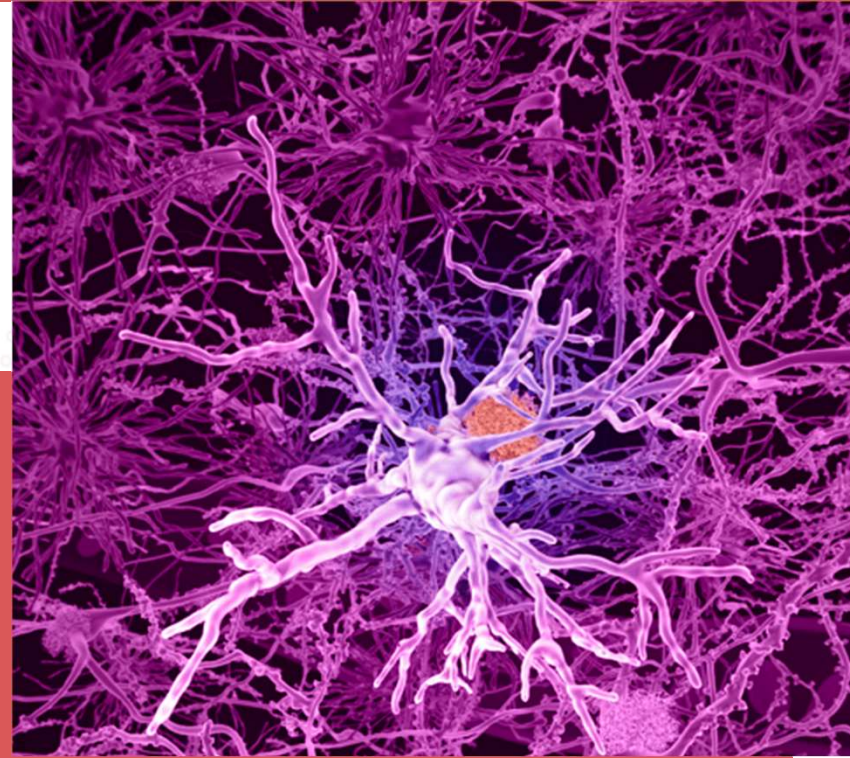
Compelling data with first-in-class candidate supports registrational plan

- **Consistent efficacy** signals in Alzheimer's, DLB and dry AMD
 - Effective in *both* mild & moderate Alzheimer's disease
 - GA lesion growth reduced with zervimesine treatment
- **Well tolerated safety** profile in over 450 people treated to date
 - **ARIA unexpected** based on MoA
 - Modest side effect profile for use in aging population
- **Oral QD** administration
 - Reduced burden compared to infusions for early AD and intravitreal injections for dry AMD; no required imaging surveillance
- **Potential first-to-market for dementia with Lewy bodies (DLB)**
 - Strong responses across **behavioral, functional, cognitive, and movement** measures in Phase 2 SHIMMER study
- **Robust intellectual property** covering platform & compounds, including zervimesine (CT1812) through 2040 with PTE



Alzheimer's Disease

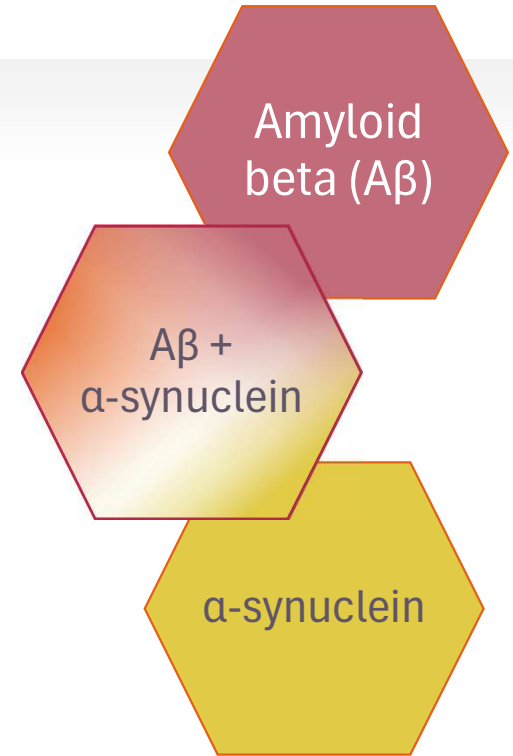
95% slowing of cognitive decline in lower-p-tau217 'SHINE' participants



AD and DLB: Two Diseases with Overlapping Pathology

Primary treatment goal is to slow the progression of disease

- A β : associated with Alzheimer's pathogenesis
- α -synuclein: associated with Lewy body dementias
- Co-pathology is common
 - Up to 80% of DLB patients have BOTH α -synuclein and Amyloid beta (A β)¹
 - Appx 50% of Alzheimer's patients have BOTH A β and α -synuclein²
- Zervimesine has shown protective function against α -synuclein and A β



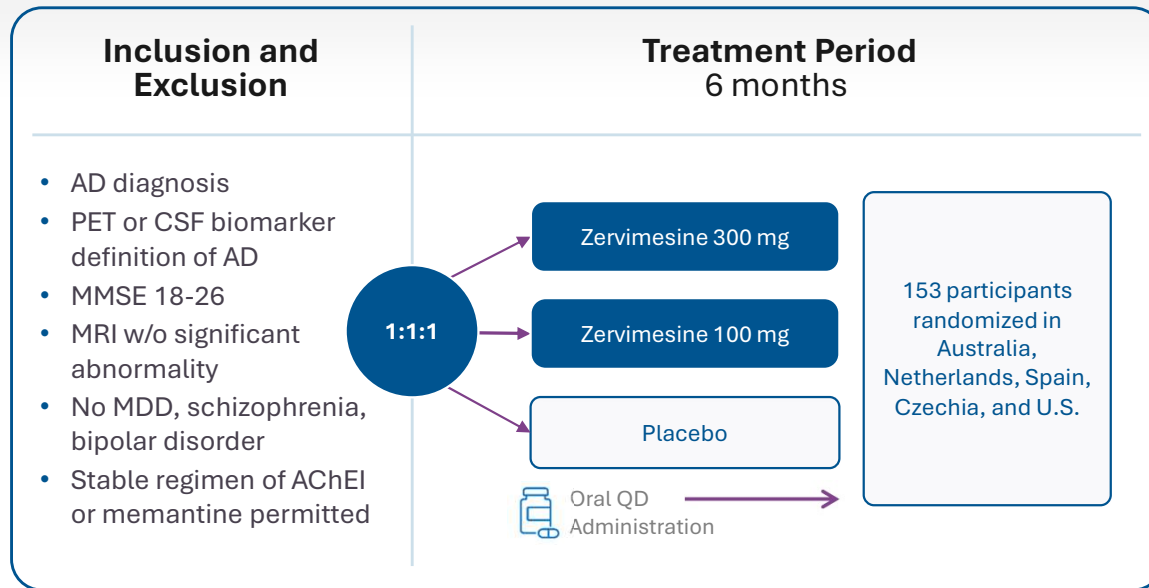
Phase 2 PoC in Mild-to-Moderate Alzheimer's Disease



Well-executed, over-enrolled study, supports advancing clinical development

Enrolled Population:

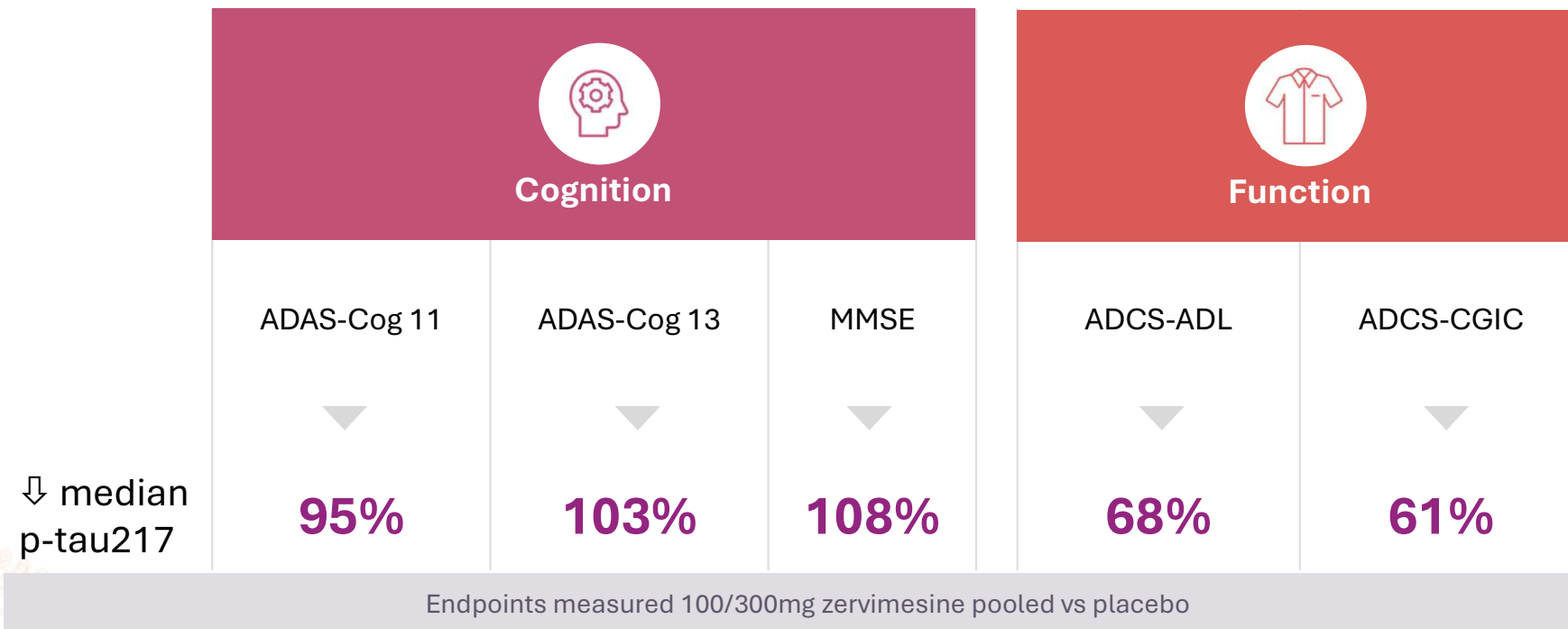
- PET- or biomarker-confirmed AD
- Majority of participants were female (60%), Caucasian (96%), ~ 72 yo
- Mean MMSE score upon entry: 21.37
- ~60% of patients carry the ApoE4 gene
- Characteristics well-balanced between all 3 arms



SHINE COG0201 study (NCT03507790) partially funded by \$31M NIA grant R01AG058660

Up to 108% Slowing on Assessments in Lower p-Tau Population

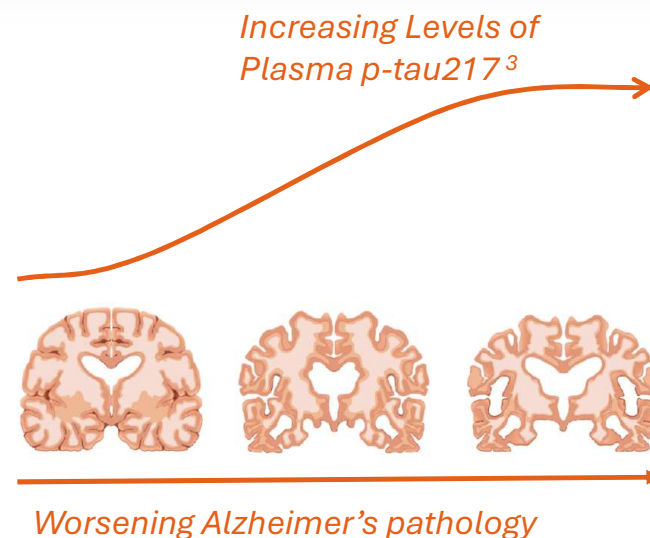
Response of 100 and 300 mg dose groups are similar



Plasma p-Tau217: a Predictive Biomarker of Treatment Response

Plasma p-tau217 level correlates to degree of Alzheimer's pathology

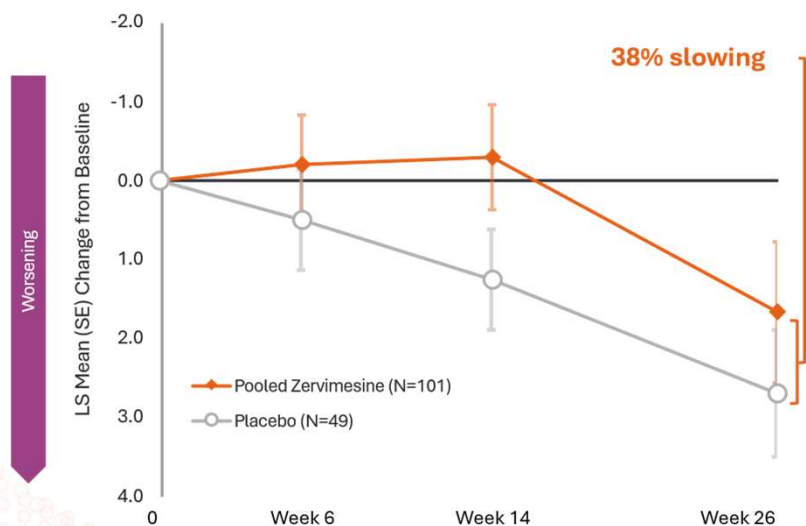
- Individuals with lower levels of plasma p-tau217 at baseline have lower AD pathology. They show a greater response to amyloid-based therapies:
 - Donanemab TRAILBLAZER 2¹
 - Lecanemab CLARITY AD tau sub study²
- Given zervimesine's MoA of displacing A β oligomers, we hypothesized that larger treatment effect may be observed in participants with lower plasma p-tau217
- Prespecified subgroup analysis defined by median baseline plasma p-tau217 within study population



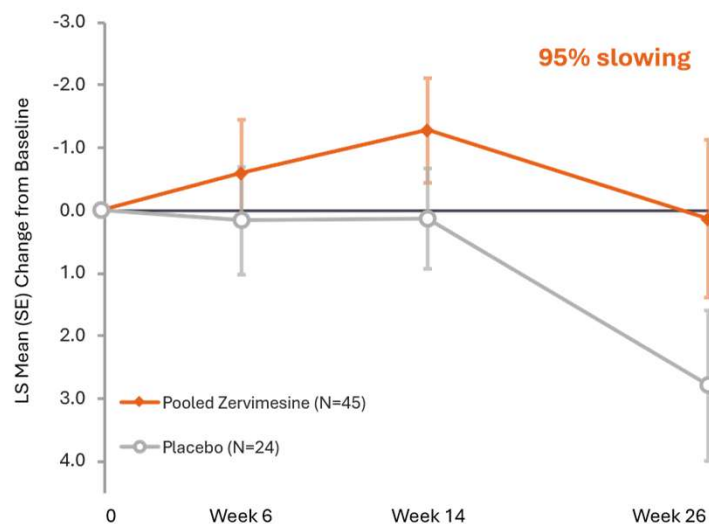
Participants with Below Median p-Tau217 Experienced Profound Treatment Effect

Successful end-of-Phase 2 meeting was based on these results

ADAS-Cog 11* mITT population (n=150)



Below median p-tau217 (n=69)

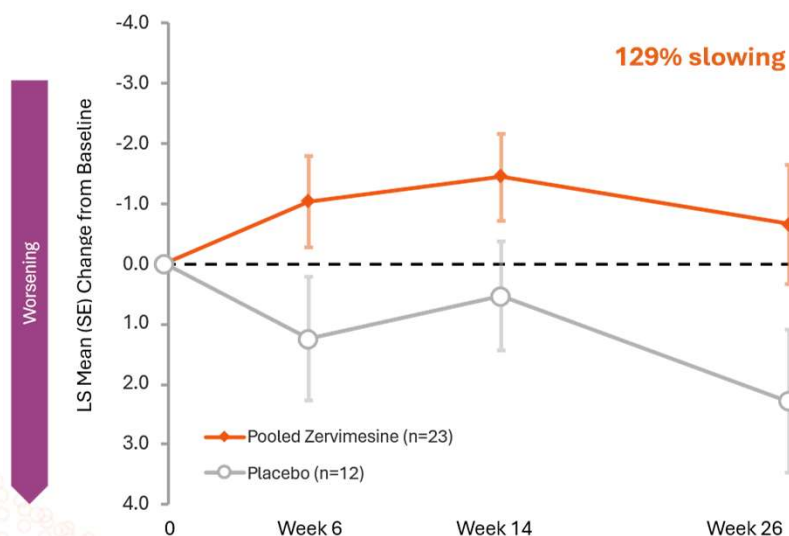


Population selected for Phase 3

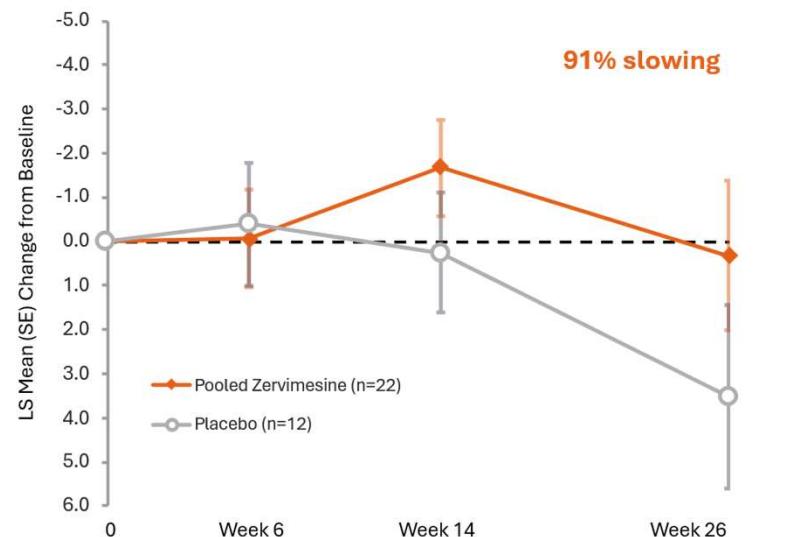
Consistent Treatment Impact in Participants with Lower p-Tau217 Across Baseline MMSE scores

Confirmed enrichment strategy and severity range at end-of-Phase 2 FDA meeting

Zervimesine-treated Mild (MMSE 22-26) Participants



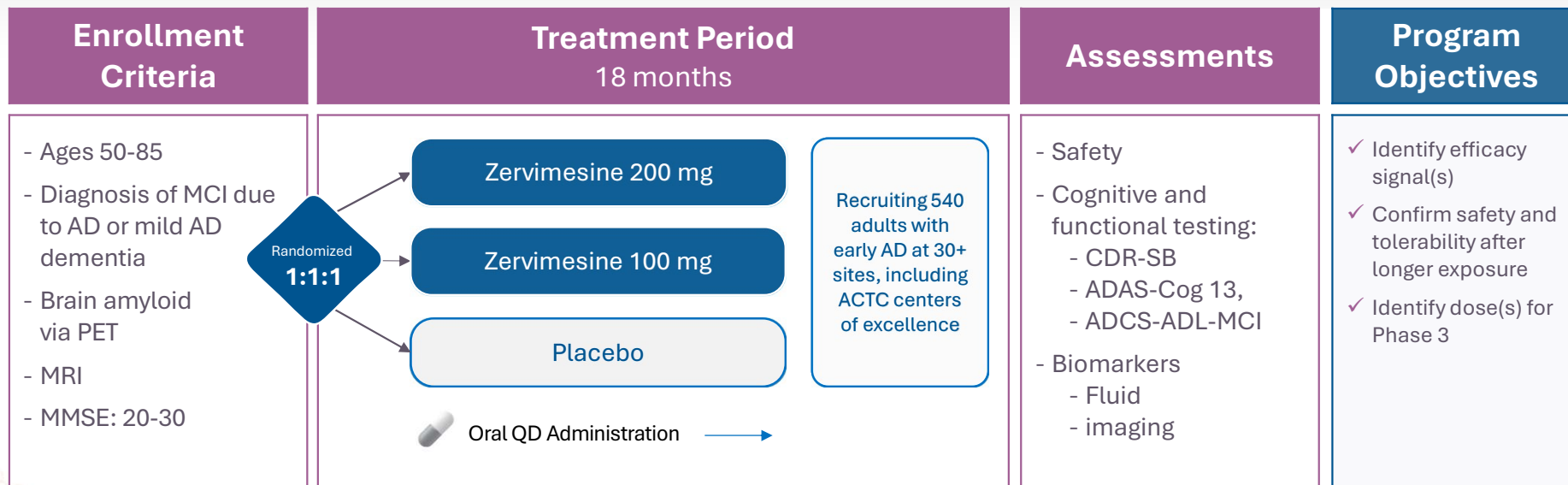
Zervimesine-treated Moderate (MMSE 18-21) Participants



Phase 2 MCI and Early Alzheimer's Study



First study to allow approved mAbs as background therapy in combination with zervimesine



START COG0203 study (NCT05531656) partially funded by \$81M NIA grant R01AG065248

START Study Completed Enrollment 4Q 2025

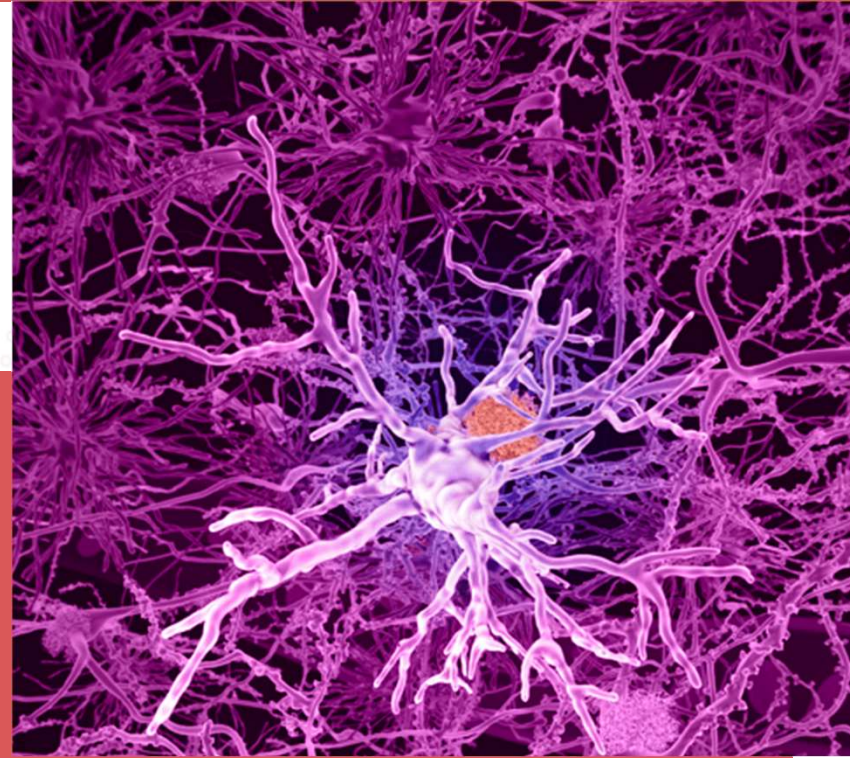


Trial duration: 18 months

- Enrollment complete (n=545)
 - First participant enrolled February 2024
 - Last participant enrolled in December 2025
- Topline results anticipated 2H 2027
- 15-20% of participants enrolled on lecanemab or donanemab
- Sub-studies: MRI, biomarkers, PET

Dementia with Lewy Bodies (DLB)

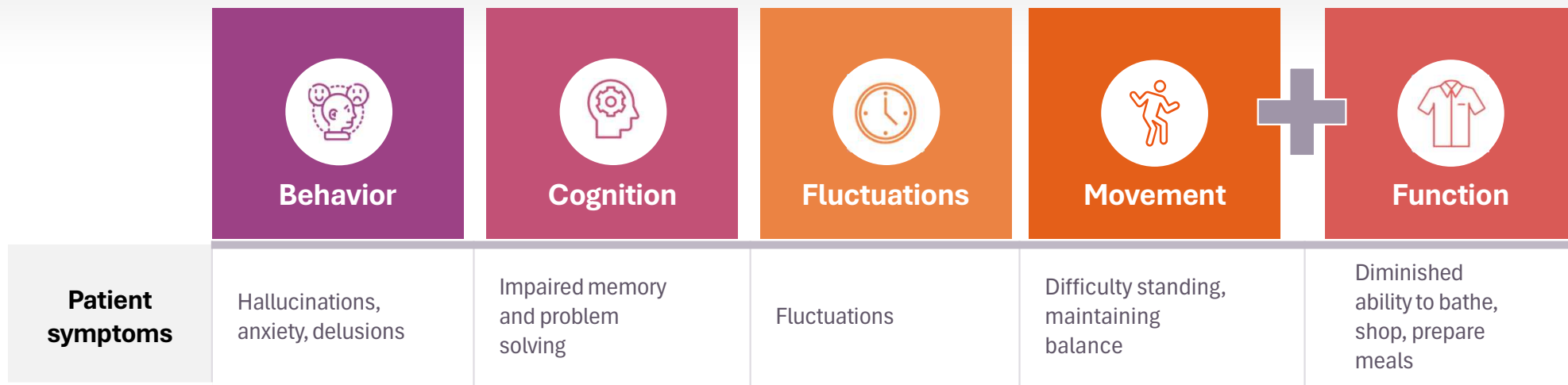
Strong clinical signals across four
major symptom domains in Phase 2
SHIMMER Study



Four Core Domains + Functional Impairment Drive DLB Burden

“A multifactorial disease with a buffet of symptoms”







James E. Galvin, MD, MPH, Univ Miami Miller School of Medicine



- 2nd most common cause of dementia after Alzheimer’s disease
- Correct diagnosis often requires multiple specialist visits over 18 months
- Faster decline than Alzheimer’s
- More common in men

Complex Symptomology Requires Array of Assessments

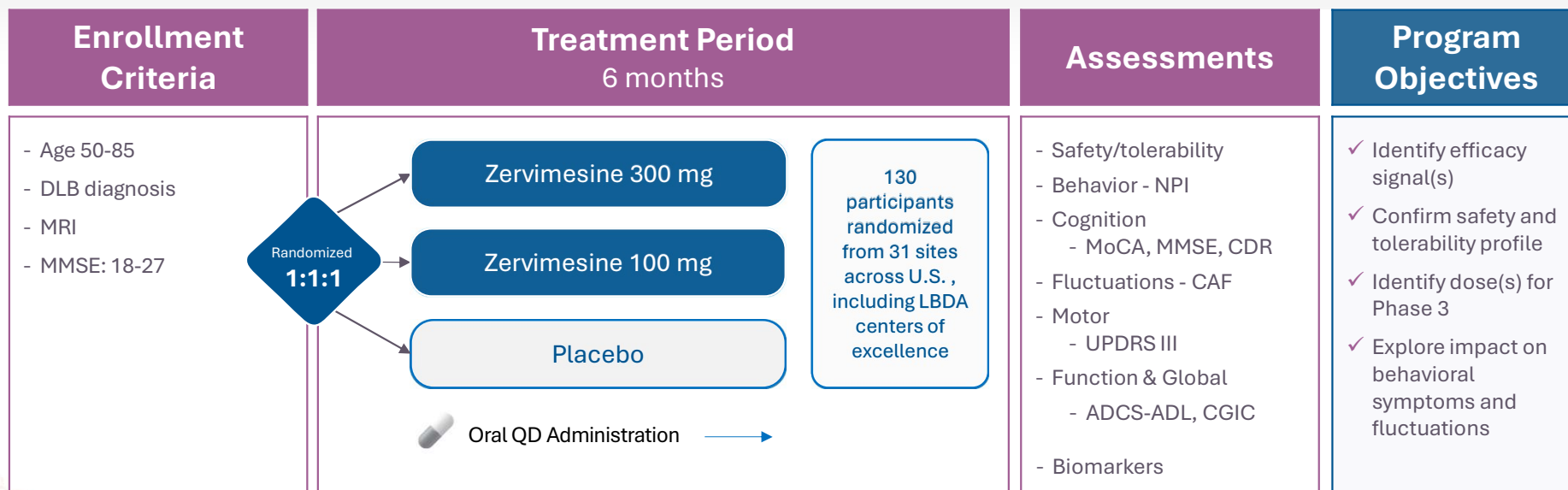
Assessment tools measure a variety of symptoms

	 Behavior	 Cognition	 Fluctuations	 	 Function
Assessment tools	Neuropsychiatric Inventory (NPI)	Cognitive Drug Research (CDR) System	Clinician Assessment of Fluctuation (CAF)	MDS-Unified PD Rating Scale Part III (UPDRS III)	ADCS-Activities of Daily Living (ADL)
	Care Partner's NPI of Distress	Montreal Cognitive Assessment (MoCA)			

Phase 2 Study in Dementia with Lewy Bodies

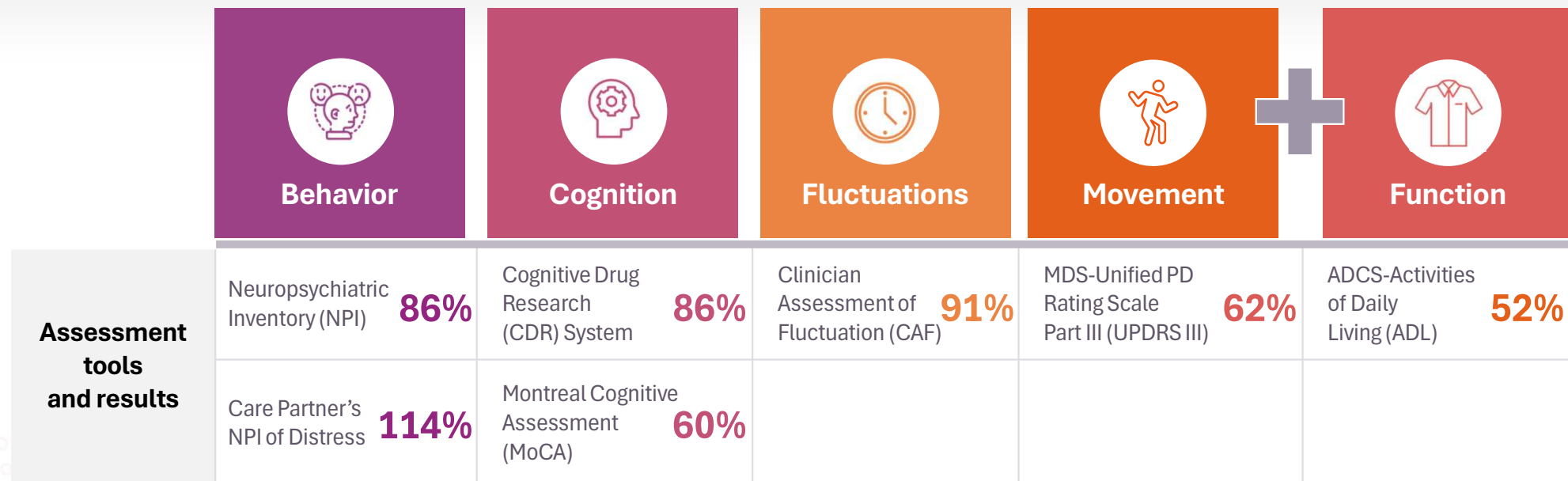


Partially funded by \$30M NIA grant (R01AG071643)



Up to 91% Percent Slowing on Assessments

Strong clinical signals across major DLB symptoms relative to placebo



Endpoints measured 100/300mg zervimesine pooled vs placebo

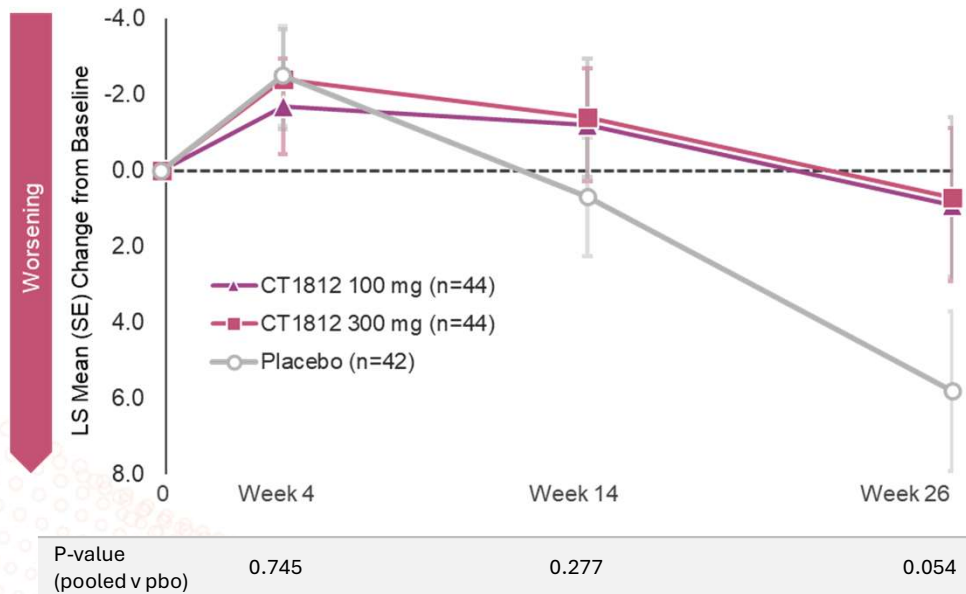
Zervimesine Showed Dramatic 86% Impact on Neuropsychiatric Measures

NPI captures a variety of patient disturbances, including hallucinations, anxiety, and delusions

NPI

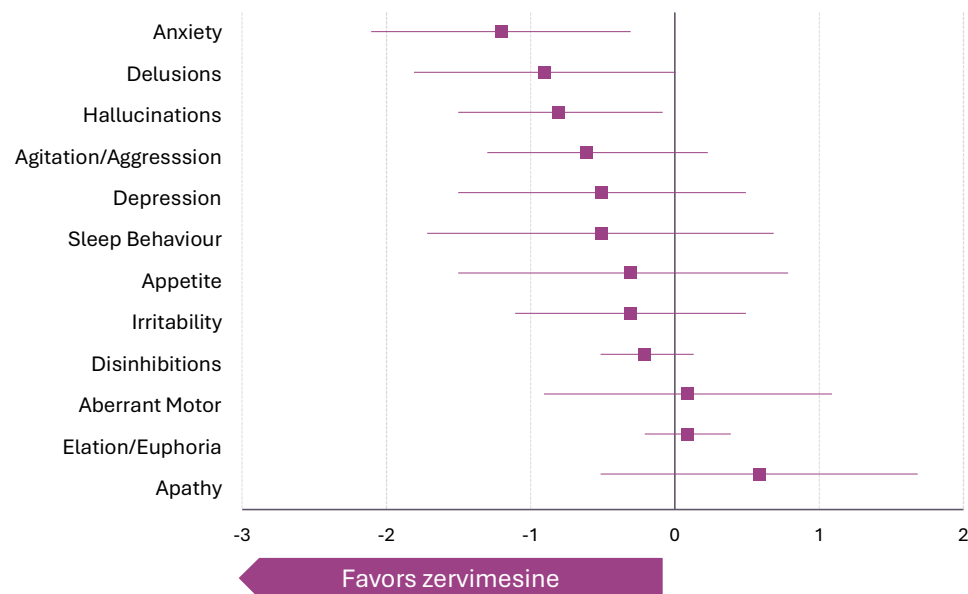
NPI Total Score (A-L)

86% Slowing



NPI favor Treatment with Zervimesine

LS Mean Difference from Placebo 95% CI



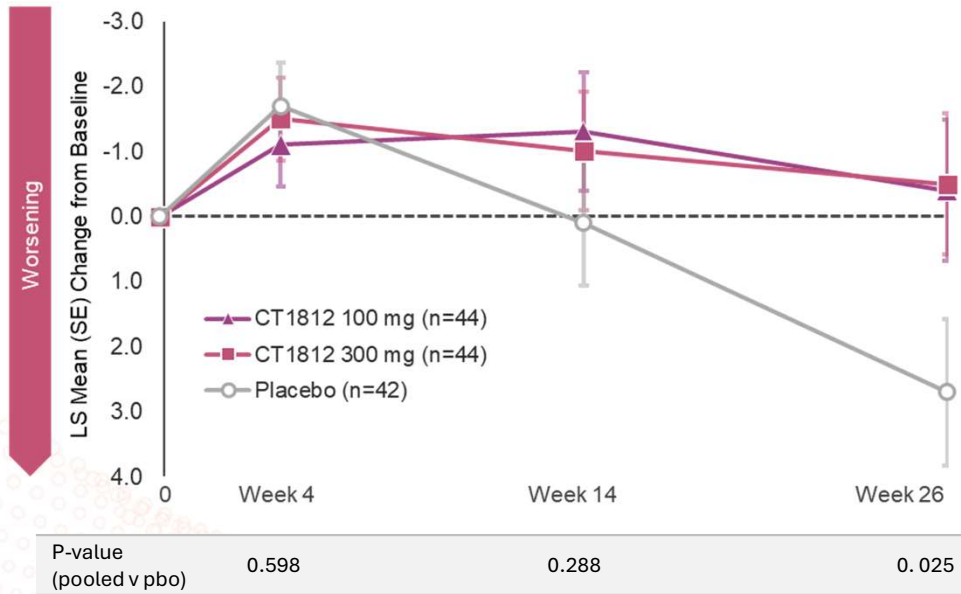
Because Participants Improved in Hallucinations, Delusions & Anxiety, Caregivers Reported Less Distress

Companion tool to measure caregiver distress based on neuropsychiatric symptoms

NPI Distress

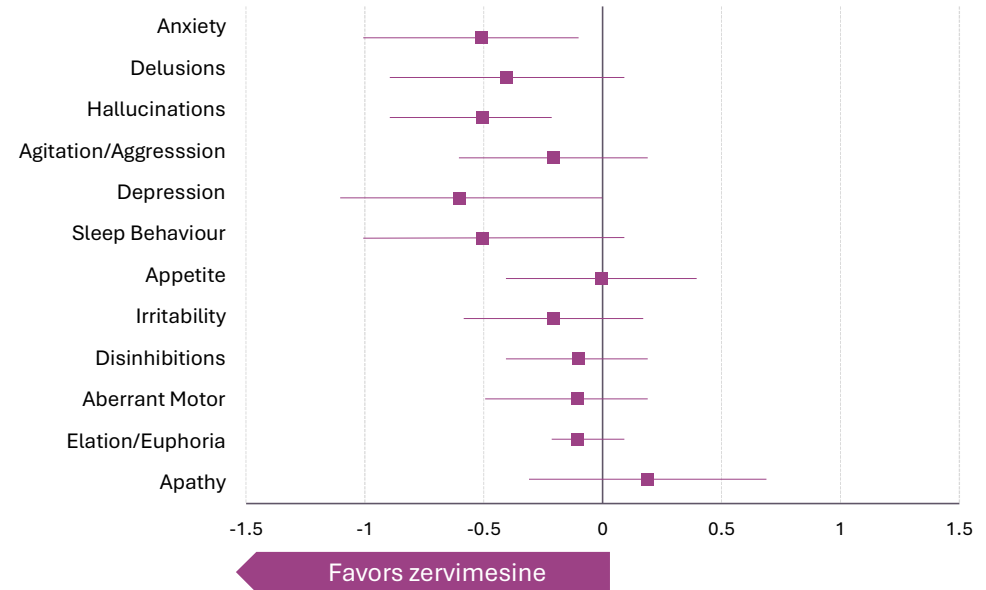
NPI Total Score (A-L) Caregiver Distress

114% Slowing



NPI Distress favors Treatment with Zervimesine

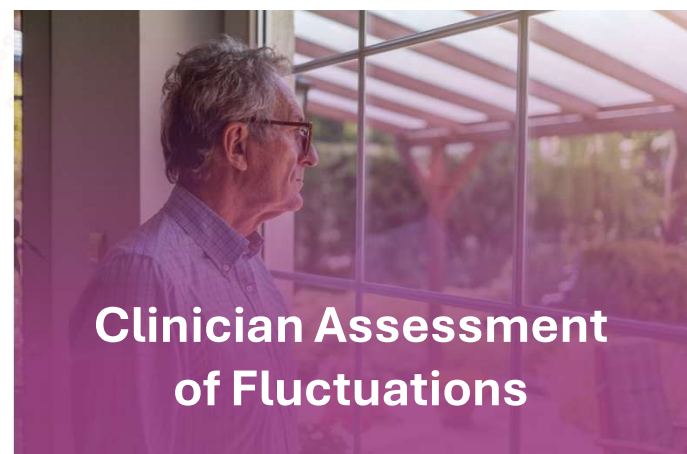
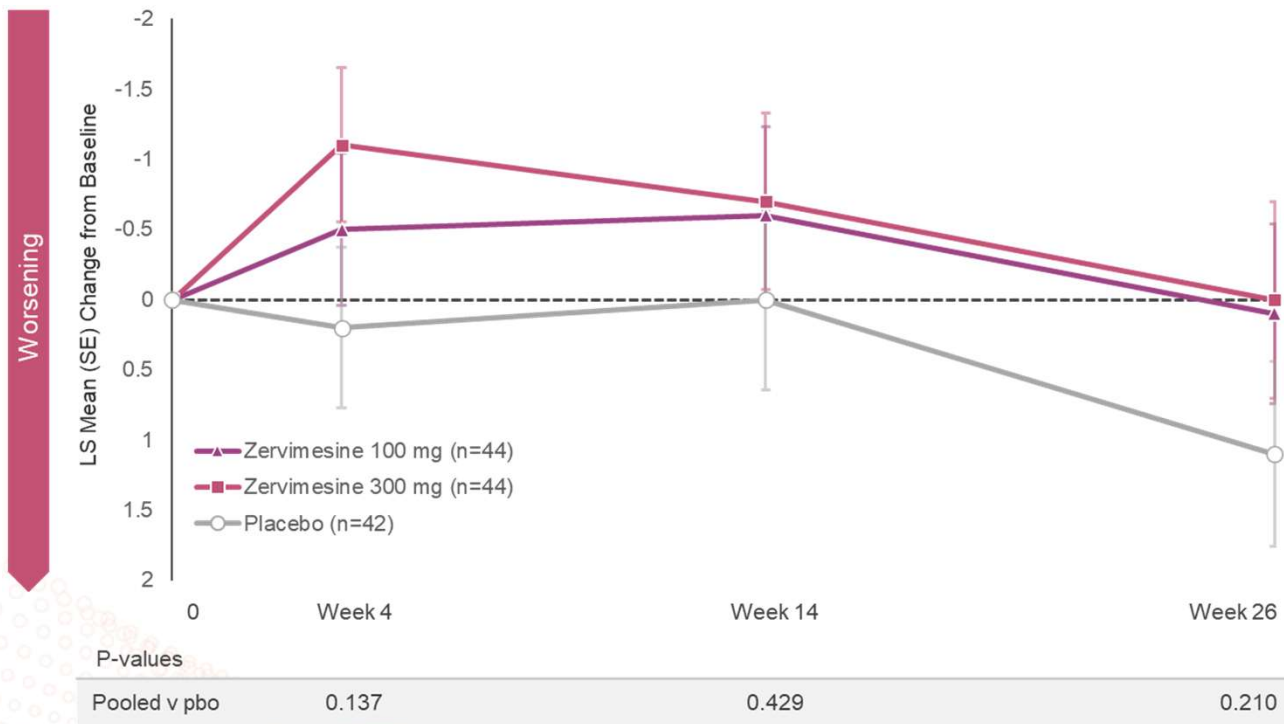
LS Mean Difference from Placebo 95% CI



Fewer & Less Severe Fluctuations After 6 Months of Zervimesine Treatment

CAF

91% slowing



Clinician Assessment of Fluctuations

Unpredictable lapses in attention or consciousness that can last minutes, hours or days

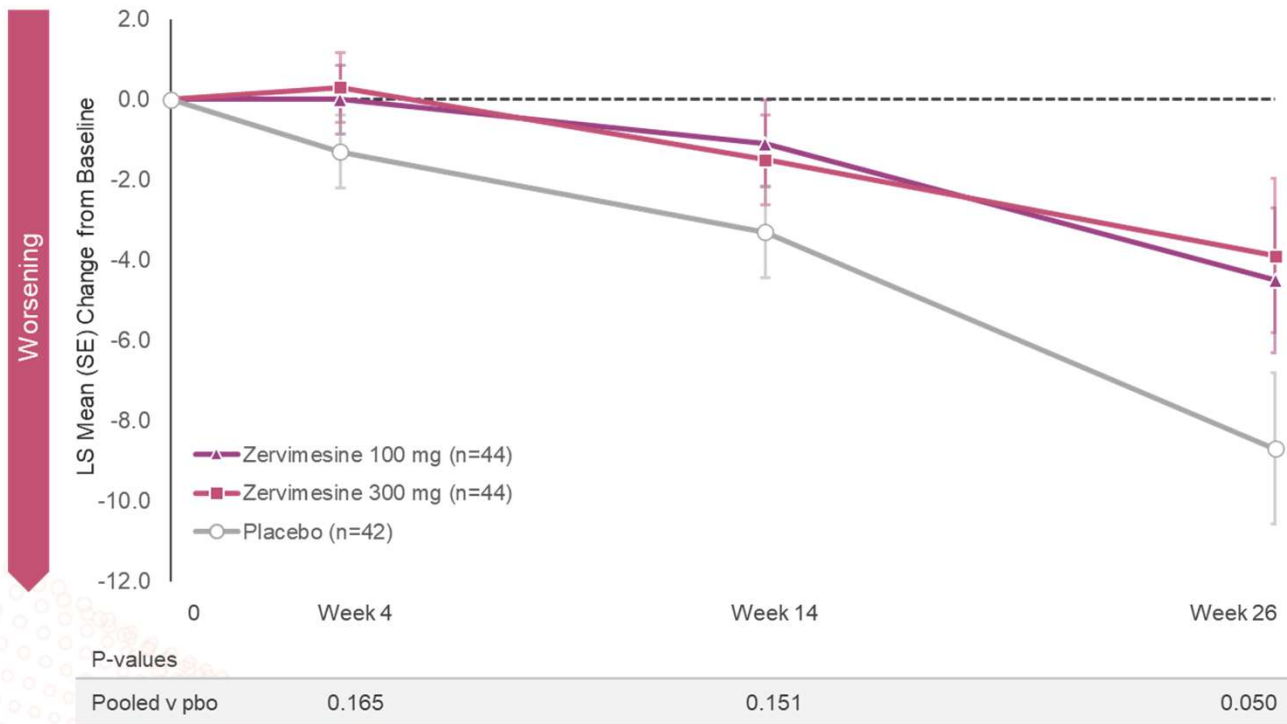


Maintenance of Self-care

52% preservation in activities of daily living (ADL) measures

ADCS-ADL

52% slowing



Components of ADL Score



Bathing



Dressing



Grooming



Feeding



Toileting



Conversing



Shopping



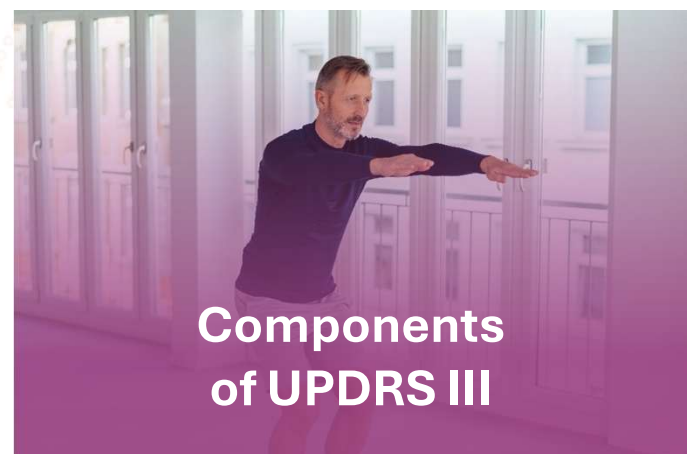
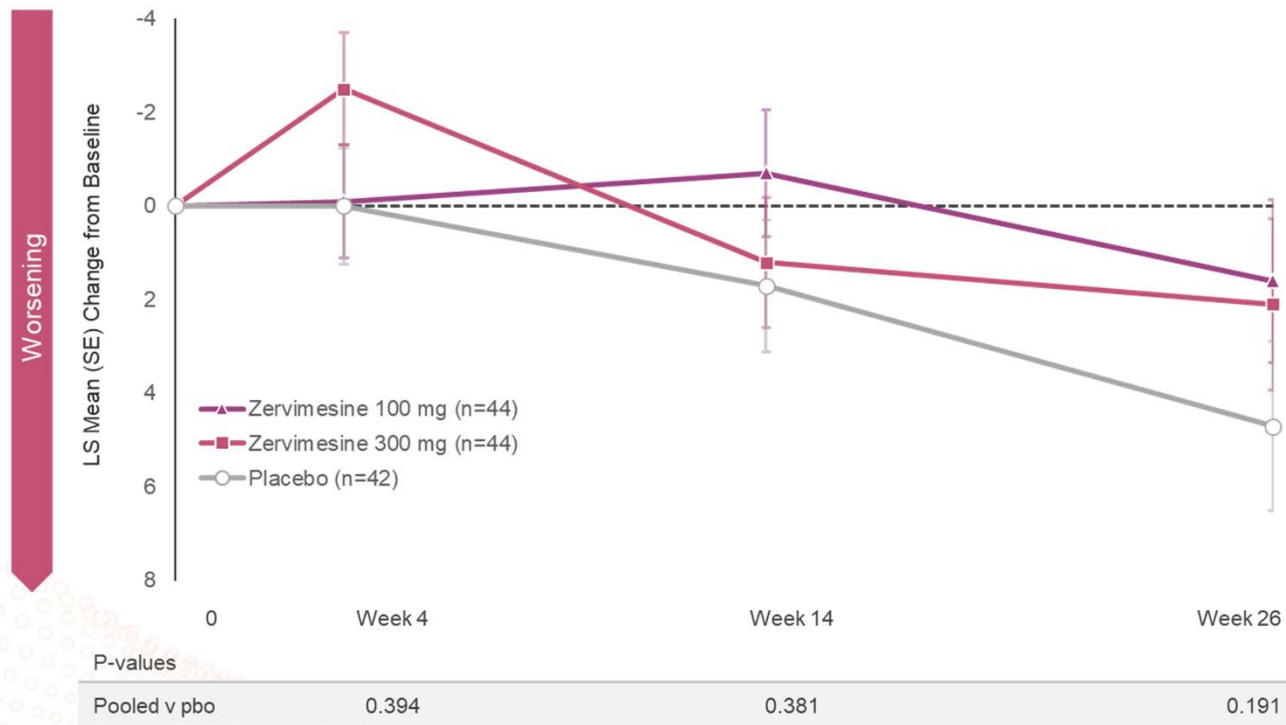
Writing

Maintenance of Motor Function

62% preservation in measures of movement

MDS-UPDRS III

62% slowing



Components of UPDRS III



Balance



Gait



Speech



Facial expression



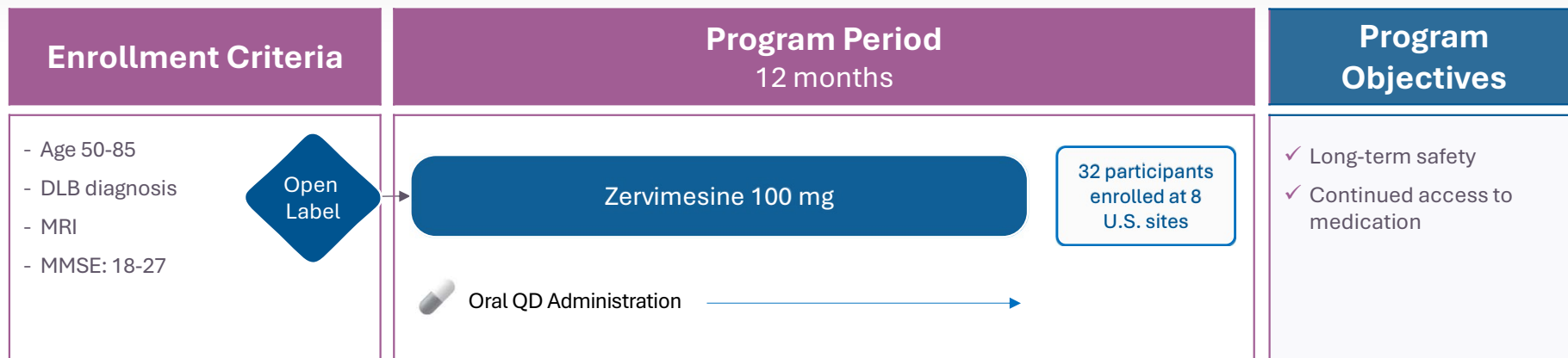
Rigidity



Tremor

DLB Expanded Access Program (COG1202)

Funded by an anonymous donation from SHIMMER family



- Expanded access program (EAP) opened in July 2025
- Majority of enrollees are former Phase 2 participants
- Initial enrollment completed in December 2025

Zervimesine Making an Impact for DLB Patients

Participant from Phase 2 and EAP comments on his experience with zervimesine in SHIMMER

“For years, Susan would order for me at restaurants because I couldn't put sentences together. Now, I can order on my own. I make complete sentences, my speech is different, and my actions are different. It's amazing.”

*SHIMMER and EAP participant
at Banner Sun Health*

“In the mornings he would stand there and forget what to do. Now he gets out of bed and knows exactly what he needs to do. Our lives are absolutely more enriched.”

Care partner



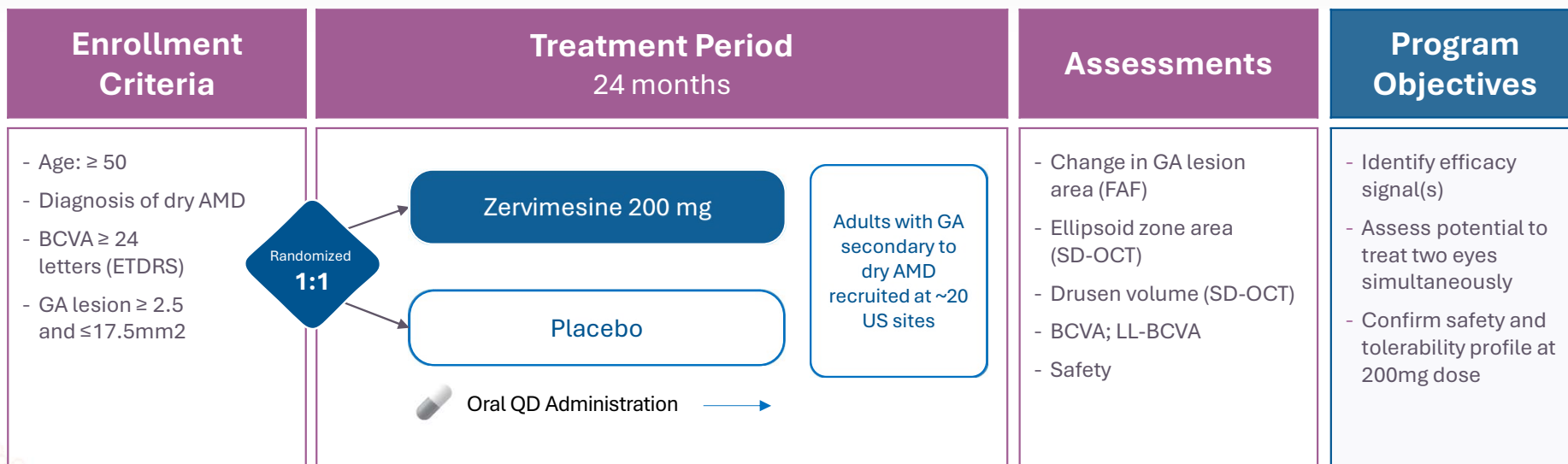
MAGNIFY Topline Results

GA lesion growth slower with
zervimesine treatment



MAGNIFY Trial in Dry AMD/GA

Zervimesine: oral drug development candidate for geographic atrophy



BCVA, best corrected visual acuity; FAF, fundus autofluorescence; SD-OCT, spectral domain optical coherence tomography

Zervimesine Treatment Slowed GA Lesion Growth



Effect size increases with exposure

- 29% mean rate of change (slope) in GA lesion area vs placebo (p=0.0538)
- Change in GA progression rate favored zervimesine vs placebo at each timepoint
 - 6-months: -11.79%
 - 12-months: -15.83%
 - 18-months: -28.19% (p=0.0074)
- Effect size increases with longer study duration
- Safety profile consistent with AD/DLB studies

Zervimesine Slowed GA Lesion Growth Rate (slope) and Area (observed)



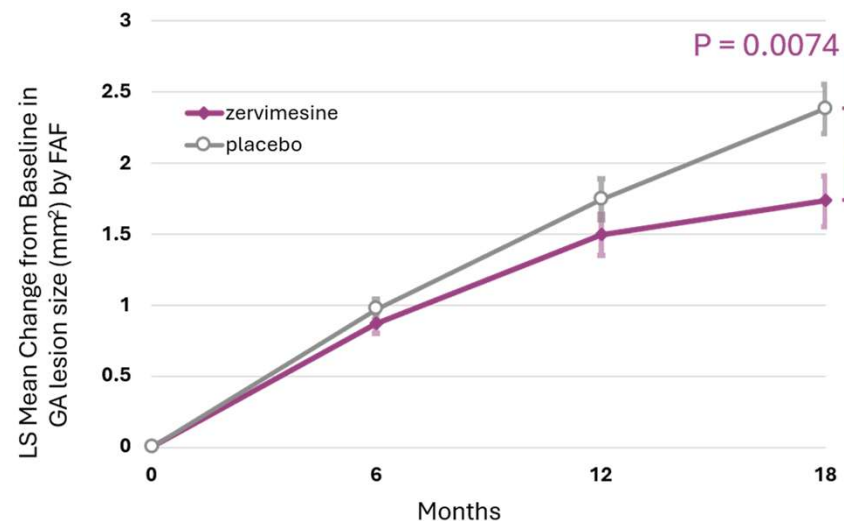
Slope Analysis¹

	Zervimesine	Placebo	Diff
Growth rate (mm ² / month)	0.10 ± 0.015	0.14 ± 0.014	- 0.04
Annualized Growth rate (mm ² / year)	1.23	1.73	- 0.50

Percent Difference from Placebo

29%
(P=0.054)

Mean Area by Time¹



% Difference treated vs placebo	6 Months	12 Months	18 Months
	-11.79	-15.83	-28.19
Zervimesine n's	44	31	16
Placebo n's	43	33	16

Consistent Safety Profile Across Completed Phase 2 Studies

Well tolerated in 271 participants with AD, DLB, dry AMD

- Average age was ~75 years
- Adverse events (AEs) were well balanced between treatment and placebo arms
 - Serious AEs occurred at comparable or higher rates in placebo- compared to zervimesine-treated arms
 - Among zervimesine-treated, there were 25 LFT elevations greater than 3x ULN (9.2%)
 - Majority were in 200mg or 300mg dose groups; only three in 100mg dose groups
 - Elevated liver enzymes normalized after cessation of drug
 - No signs or symptoms of permanent liver injury
- Discontinuations due to AEs are similar between 100mg dose groups (5.8%) and placebo groups (5%)
 - Overall, 12% zervimesine-treated discontinued due to AEs (100, 200 and 300mg)

3 Major Diseases Addressed with Once-Daily Oral Pill

Collective Phase 2 results support advancing zervimesine (CT1812) to registrational studies



Dementia with Lewy Bodies

Marked slowing of progression across multiple domains



Alzheimer's Disease

Slowing of progression; robust response in lower tau cohort



Geographic Atrophy

Slowing of GA growth rate and area

Current Financial Position

Cash runway into 2Q 2027

As of quarter ended September 30, 2025

Cash and cash equivalents \$ 39.8 M

Grant funding for zervimesine studies

Preclinical through Phase 2 ~\$171 M

Approximate funding used (\$135 M)

Remaining grant funding \$36 M





Thank You

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 **COGNITION**™
Therapeutics