



Guggenheim Emerging Outlook: Biotech Summit

February 2026



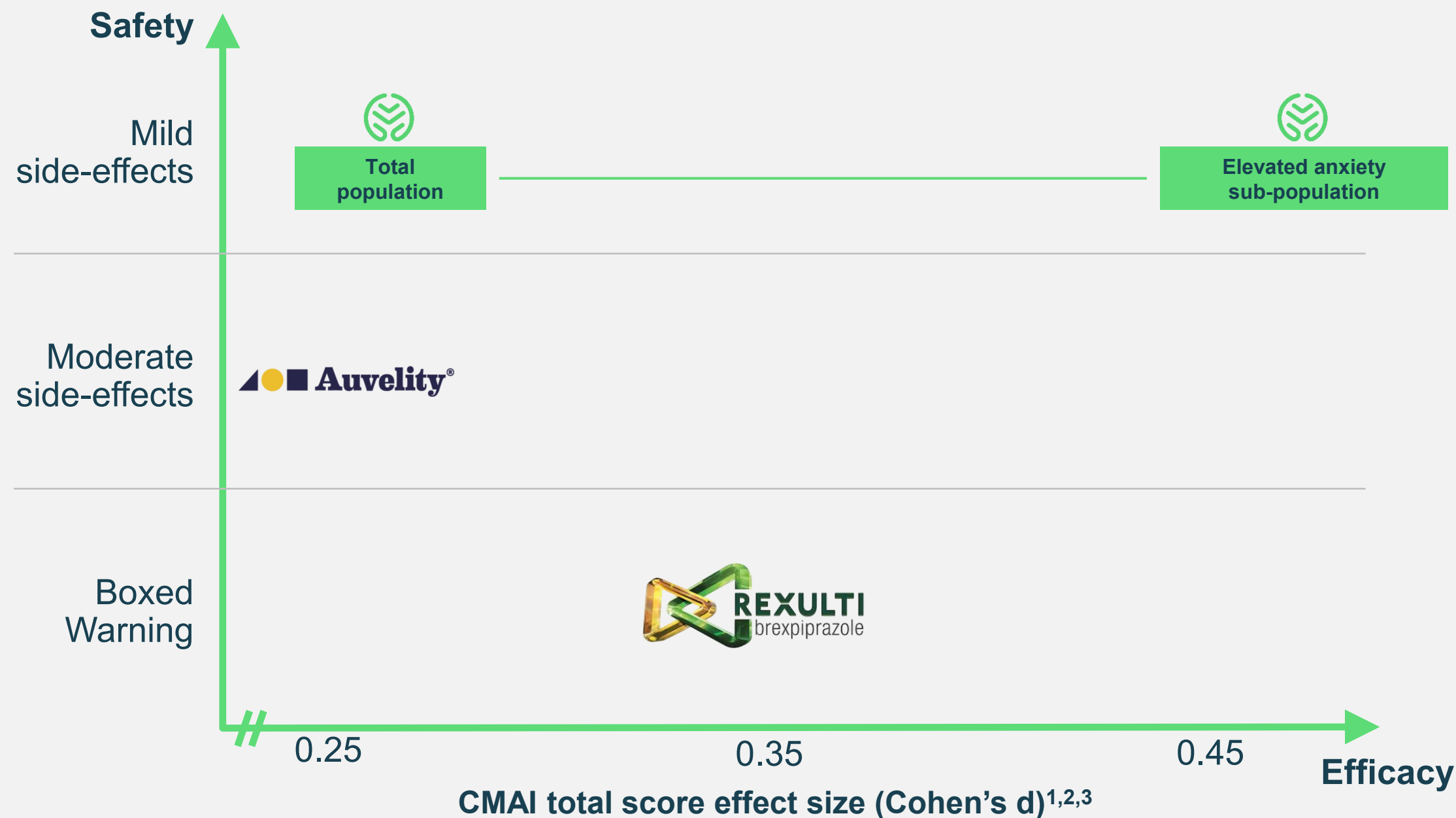
Important Disclosures

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NMRA-511 demonstrated unsurpassed efficacy in patients with AD agitation

Simplified market segmentation and opportunities



NMRA-511 Phase 1b key takeaways

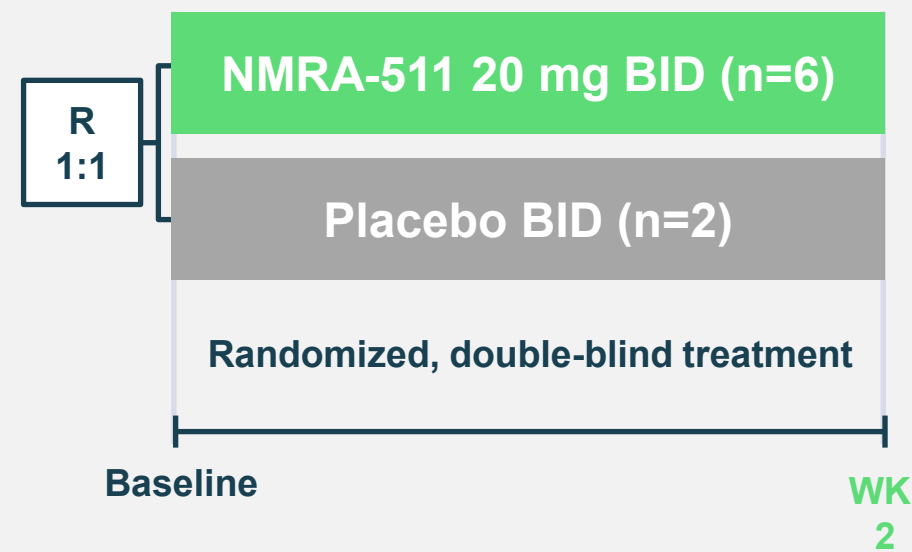
- Well tolerated, with potential for higher dosing
- CMAI effect size similar to Auvelity in total population
- Unsurpassed CMAI effect size in patients with elevated anxiety

For illustrative purposes only. NMRA-511 has not been studied in head-to-head trials against Auvelity or Rexulti, and there are differences in compounds, trial designs and other factors which must be considered.

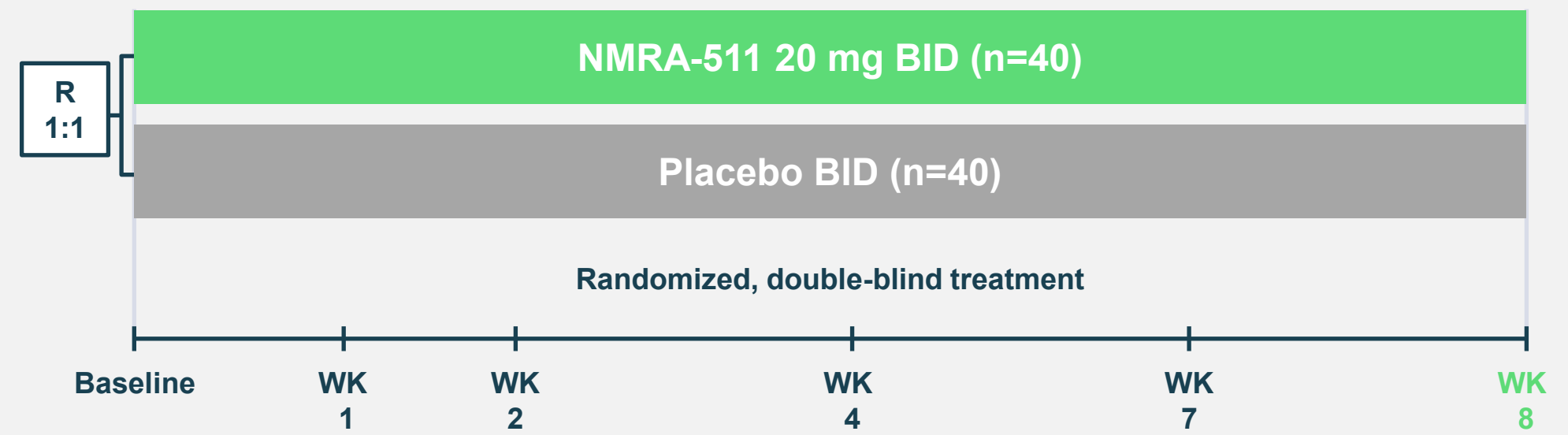
¹Calculated from data: Addressing Dementia Via Agitation-Centered Evaluation (ADVANCE). <https://clinicaltrials.gov/study/NCT03226522?intr=AXS-05&page=1&rank=9&tab=results>. ²Lee D, Slomkowski M, Hefting N, et al. Brexpiprazole for the Treatment of Agitation in Alzheimer Dementia: A Randomized Clinical Trial. JAMA Neurol. 2023;80(12):1307–1316. doi:10.1001/jamaneurol.2023.3810. ³NMRA data on file. CMAI = Cohen-Mansfield Agitation Inventory. Rexulti and Auvelity studies were enriched with an NPI-AA domain cutoff ≥ 4 at baseline. NMRA-511 Phase 1b study included no enrichment. Baseline NPI-AA scores; NMRA-511: 5.1; Rexulti: 7.7 (Study 2); Auvelity: 7.2 (Advance-1)

Study to evaluate the effects of NMRA-511 among healthy elderly and adults with agitation associated with dementia due to Alzheimer's disease

Part A: 2-Week Evaluation Period Enrolling Healthy Elderly Participants



Part B: 8-Week Evaluation Period Enrolling People with Alzheimer's Disease Agitation (ADA)



NMRA-511 Phase 1b Study

- Part A Inclusion Criteria:**
 - Healthy elderly adult participants aged 65-80 years
- Part B Inclusion Criteria:**
 - Adults aged 55-90 years with mild-severe dementia (MMSE score of 5-24) and clinically significant agitation (CMAI total score 45-100)
- Part B Primary Endpoint:**
 - Δ from baseline to Week 8 in CMAI total score
- Part B Other Endpoints Include*:**
 - Δ from baseline to Week 8 in:
 - CGI-S
 - NPI total score
- Prespecified Sub-Populations:**
 - Elevated anxiety (RAID)
- Statistics:**
 - **Study not powered to demonstrate statistical significance**
 - Designed as a signal-seeking study; effect size will inform the potential future development of NMRA-511 in ADA

*Safety Assessments include adverse events, clinical laboratory, vital signs, physical examination, 12-lead electrocardiogram (ECG), Columbia-Suicide Severity Rating Scale (C-SSRS). Δ = Change; BID = twice daily; CMAI = Cohen-Mansfield Agitation Inventory; MMSE = Mini-Mental State Examinations; CGI = Clinical Global Impression of Change for Agitation; NPI = Neuropsychiatric Inventory.

Demographics and baseline characteristics

	Total Population		Pre-specified elevated anxiety population	
	NMRA-511 n=40	Placebo n=40	NMRA-511 n=16	Placebo n=21
Mean age	71.8	72.7	66.8	71.6
Sex, n (%)				
Male	18 (45.0%)	15 (37.5%)	7 (43.8%)	9 (42.9%)
Female	22 (55.0%)	25 (62.5%)	9 (56.3%)	12 (57.1%)
Race, n (%)				
White	27 (67.5%)	30 (75.0%)	11 (68.8%)	14 (66.7%)
Black	10 (25.0%)	9 (22.5%)	3 (18.8%)	6 (28.6%)
Asian	2 (5.0%)	0	1 (6.3%)	0
Other	1 (2.5%)	1 (2.5%)	1 (6.3%)	1 (4.8%)
CMAI Total Score Mean (SD)	68.2 (14.7)	68 (14.3)	69.3 (15.6)	67.7 (14.9)
CGI-S (Agitation) Mean (SD)	4.3 (0.7)	4.2 (0.6)	4.4 (0.8)	4.3 (0.6)
NPI-AA Mean (SD)	5.1 (2.5)	5.9 (2.6)	4.8 (2.7)	5.8 (2.8)
MMSE Mean (SD)	19.0 (3.2)	19.5 (2.8)	19.2 (2.9)	19.4 (2.9)
Baseline anxiety as measured by RAID score (SD)	11.8 (6.4)	14.3 (8.6)	18.3 (4.2)	18.7 (6.5)
Protocol-Defined Medication Non-Adherence ¹	7 (17.5%)	0	N/A	
Modified Analysis Set (n) ²	33	38		

¹70% medication compliance required per protocol

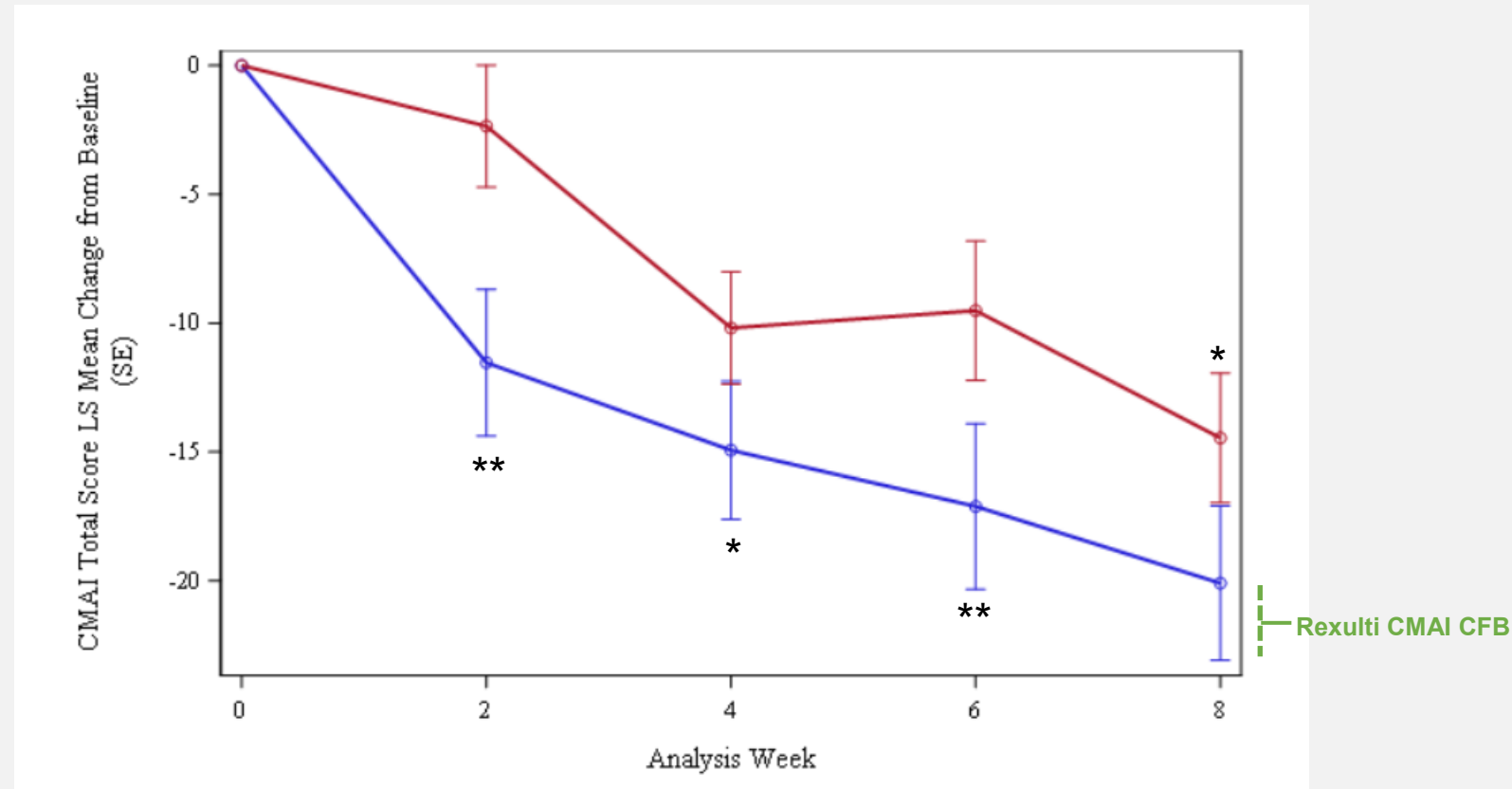
²2 placebo patients excluded based on rater change driving outlier data (>3 standard deviations from the mean)

³Defined as Rating Anxiety In Dementia (RAID) score ≥12



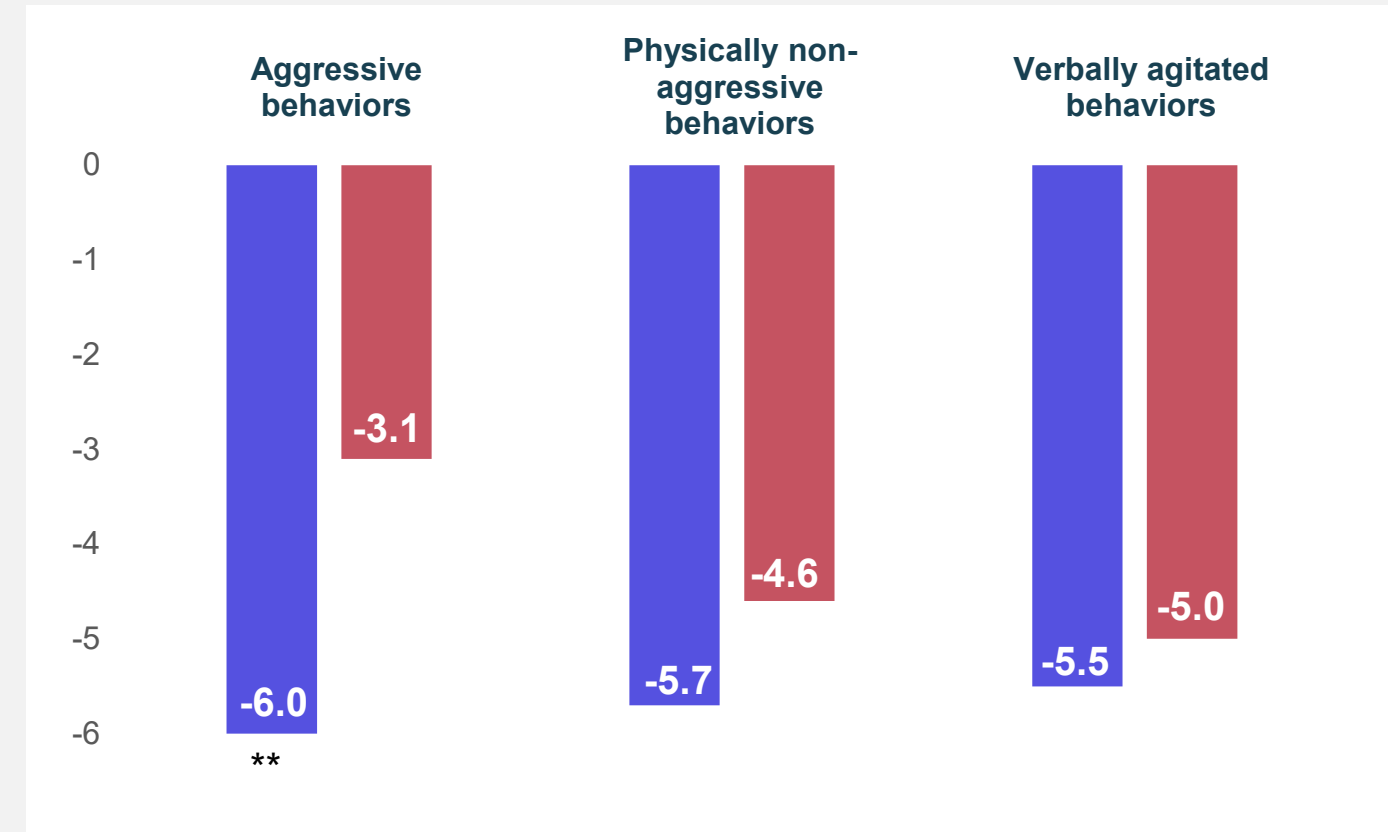
NMRA-511 demonstrated unsurpassed clinical effect size on CMAI total score in patients with elevated anxiety

CMAI Total Score Change from Baseline
(pre-specified elevated anxiety sub-population)



— NMRA-511 — Placebo

Mean Change in CMAI Sub-Scores at Week 8
(pre-specified elevated anxiety sub-population)



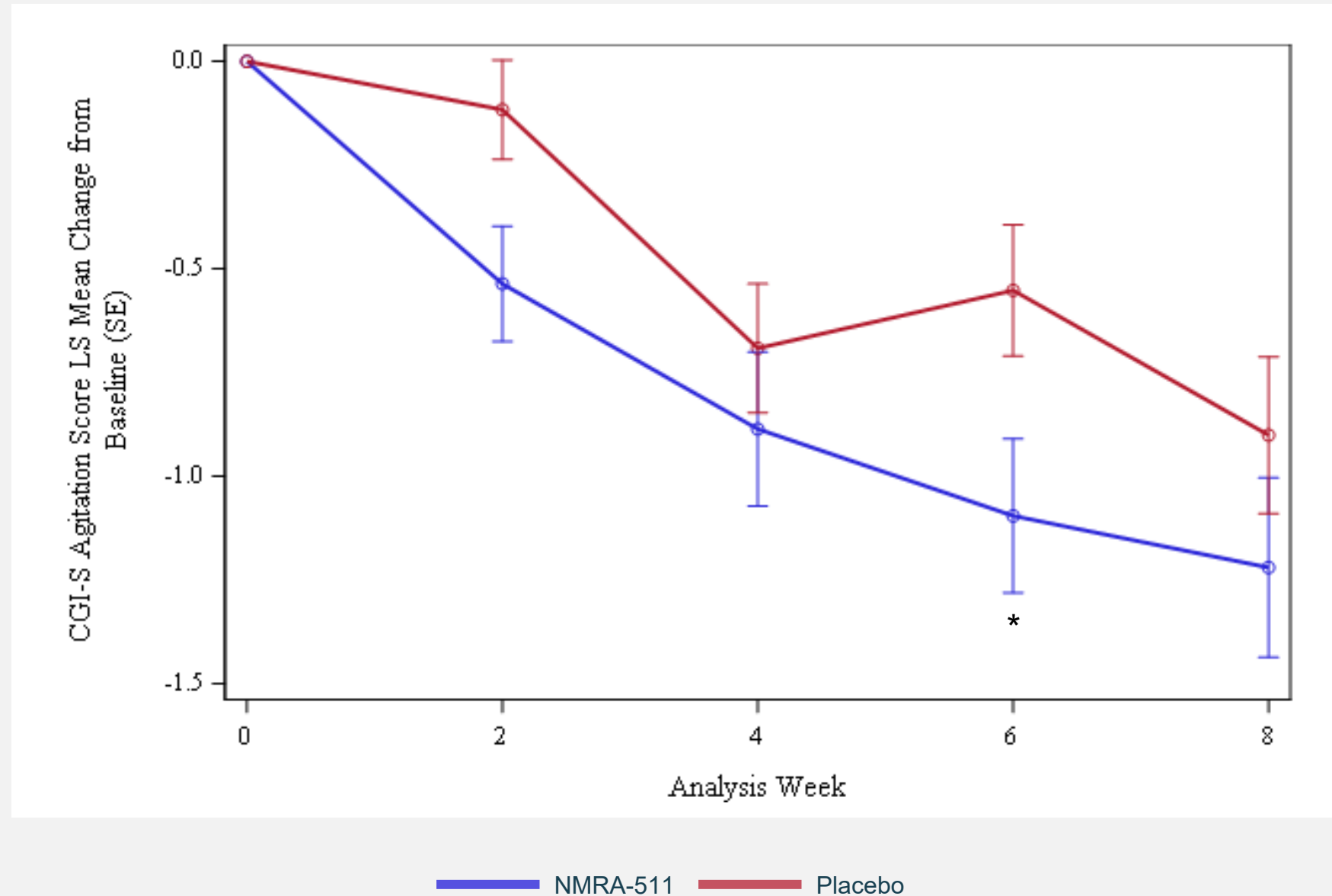
	Week 6	Week 8
LSMD (SE)	-7.6 (4.1)	-5.6 (3.8)
Effect size range (Cohen's d)	0.64	0.51

	Aggressive behaviors	Physically non-aggressive behaviors	Verbally agitated behaviors
Effect size (Cohen's d)	0.82	0.37	0.12

NMRA-511 n=16, placebo n=21
Nominal p-values: **p<0.05, *p<0.1
Cohen's d effect size range for patients with RAID ≥11: 0.45 – 0.54

NMRA-511 drove unsurpassed reductions in CGI-S agitation scores in patients with elevated anxiety at baseline

CGI-S Agitation Change from Baseline
(pre-specified elevated anxiety sub-population)

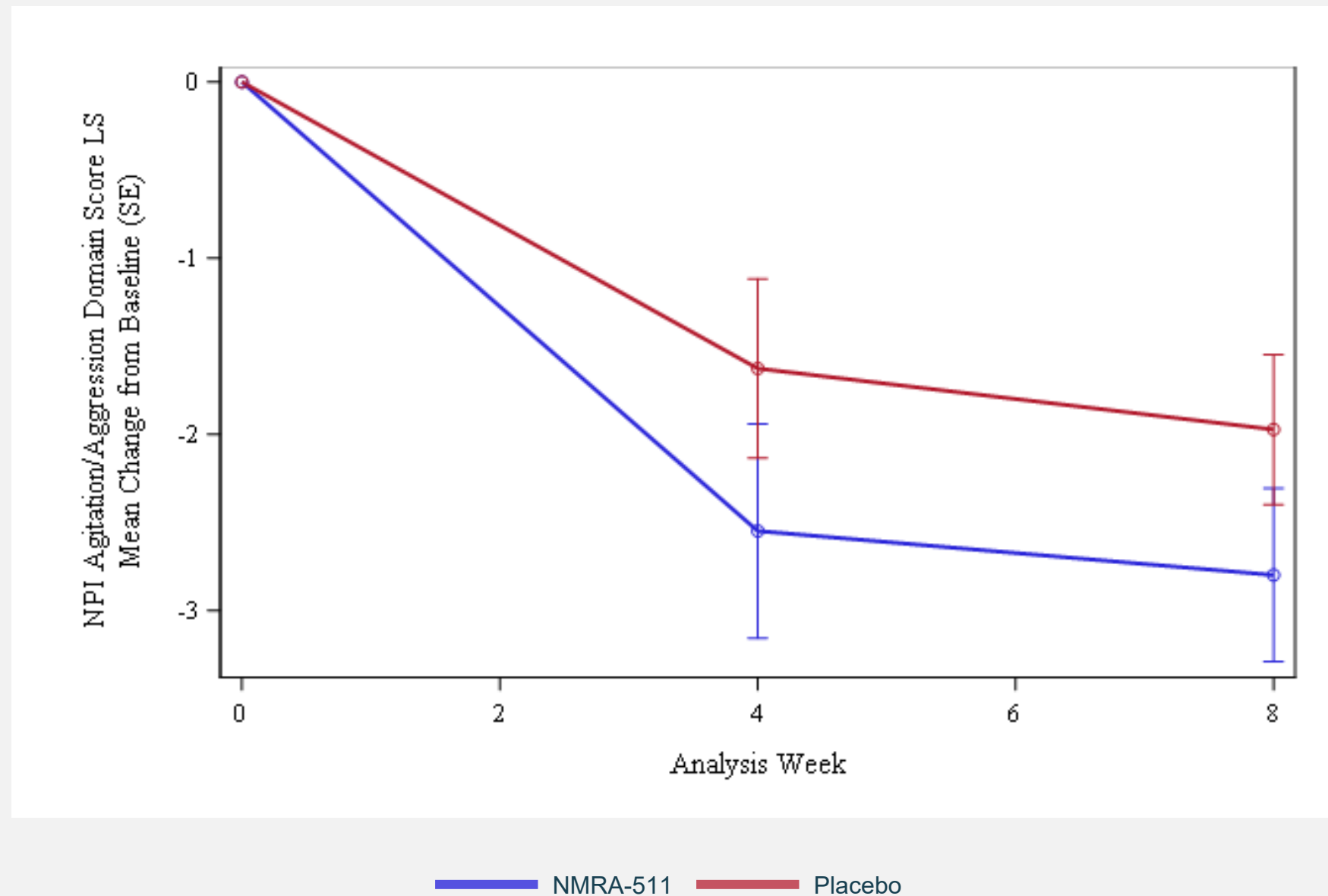


	Week 6	Week 8
LSMD (SE)	-0.5 (0.2)	-0.3 (0.3)
Effect size range (Cohen's d)	0.78	0.38



Strong clinical effect demonstrated on NPI agitation/aggression domain in patients with elevated anxiety at baseline

NPI Agitation/Aggression (NPI-AA) Change from Baseline
(pre-specified elevated anxiety sub-population)



	Week 4	Week 8
LSMD (SE)	-0.9 (0.8)	-0.8 (0.6)
Effect size range (Cohen's d)	0.42	0.46

Favorable tolerability and safety profile demonstrated

NMRA-511 was safe and generally well tolerated



TEAEs Incidence (≥5% in either treatment group)	Placebo n=40	NMRA-511 n=40
Preferred Terms	n (%)	n (%)
Nasopharyngitis	3 (7.5%)	4 (10.0%)
Urinary tract infection	1 (2.5%)	4 (10.0%)
Anemia	1 (2.5%)	2 (5.0%)
Arthralgia	0	2 (5.0%)
Diarrhea	4 (10.0%)	2 (5.0%)
Dizziness	2 (5.0%)	2 (5.0%)
Headache	5 (12.5%)	2 (5.0%)
Hyponatremia	0	2 (5.0%)
Myalgia	1 (2.5%)	2 (5.0%)
Nausea	1 (2.5%)	2 (5.0%)
Vomiting	1 (2.5%)	2 (5.0%)
Abdominal pain	2 (5.0%)	1 (2.5%)

- TEAEs were typically mild to moderate in severity
- Low treatment discontinuations due to TEAEs (2.5%)
- Opportunity to evaluate higher doses of NMRA-511 based on tolerability



One serious adverse event of asthenia (general weakness) reported; resolved by time of discharge from an overnight hospitalization and resolution was maintained at study follow up after treatment discontinuation

NMRA-511 demonstrated consistent unsurpassed efficacy across measures

	CMAI Total Score	CMAI Aggressive Behaviors Score	CGI-S Agitation	NPI
NMRA-511 elevated anxiety population	0.51 – 0.64	0.82 – 1.1	0.38 – 0.78	0.42 – 0.46*
NMRA-511 total population	0.20 – 0.23	0.31 – 0.33	0.25 – 0.35	0.09 – 0.20*
 REXULTI brexpiprazole	0.35	0.33	0.31	0.39^
 Auvelity [®]	~0.2 – 0.25	Not Reported	Not Reported	Not Reported

NMRA-511 in AD agitation

- Well tolerated safety-profile, with potential for higher dosing
- Unsurpassed and consistent treatment effect across a range of measures
- Opportunity for convenient dosing with XR formulation
- Opportunity to enrich future studies for elevated anxiety (Phase 1b not enriched)

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Data presented as Cohen's d effect size; elevated anxiety = RAID ≥12. *NPI-AA ^NPI-nursing home

1. Neumora data on file. 2. Lee D, Slomkowski M, Hefting N, et al. Brexpiprazole for the Treatment of Agitation in Alzheimer Dementia: A Randomized Clinical Trial. JAMA Neurol. 2023;80(12):1307–1316. doi:10.1001/jamaneurol.2023.3810.

3. Axsome Therapeutics corporate materials.

Rexulti and Auvelity studies were enriched with an NPI-AA domain cutoff ≥4 at baseline. NMRA-511 Phase 1b study included no enrichment. Baseline NPI-AA scores; NMRA-511: 5.1; Rexulti: 7.7 (Study 2); Auvelity: 7.2 (Advance-1)

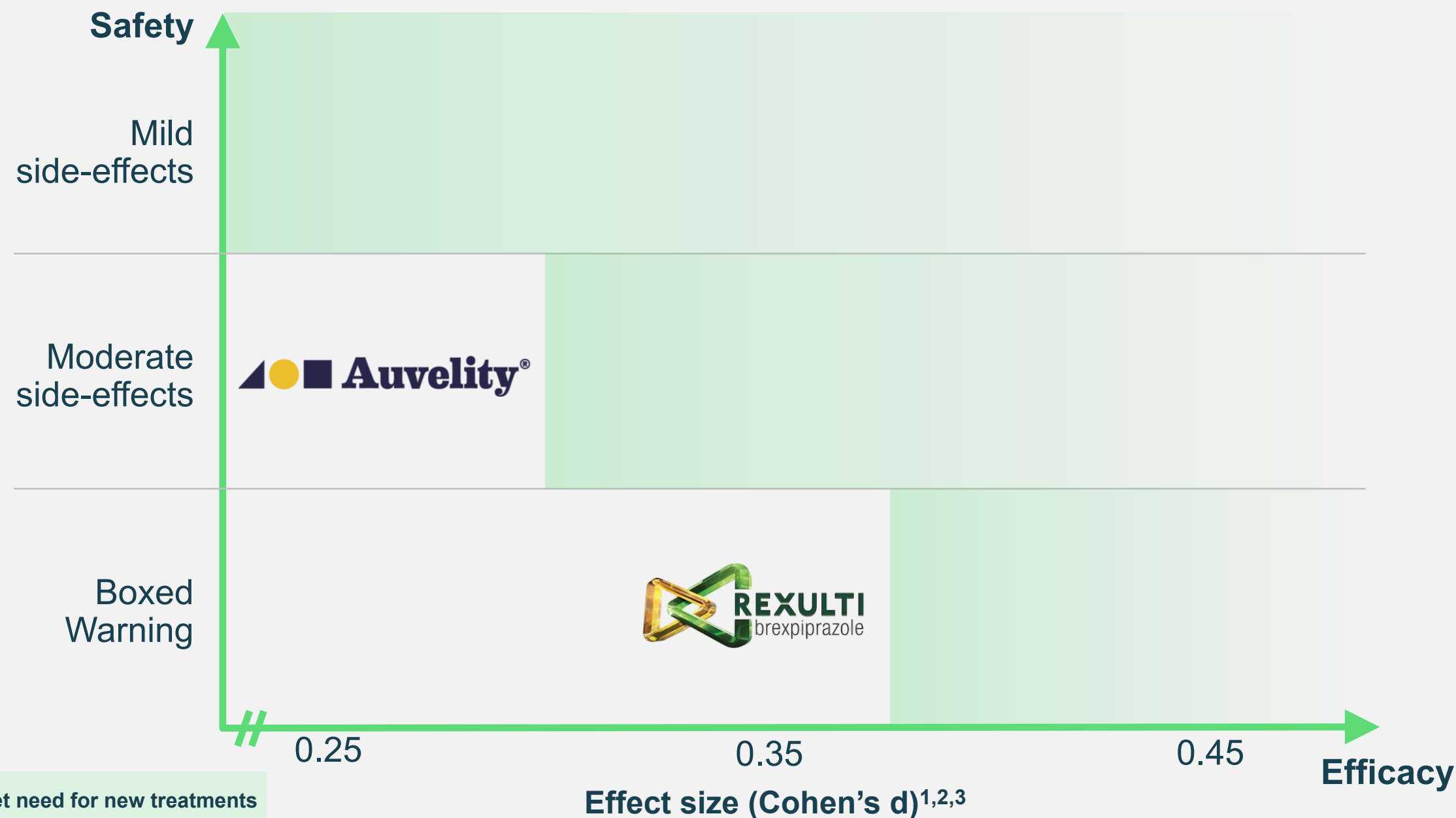


Appendix



Significant opportunity for a product with a differentiated benefit/risk profile

Simplified market segmentation and opportunities



There is an unmet medical need for therapies that reduce agitation with improved tolerability and safety profiles^{3,4}

AD agitation associated with:



Increased morbidity and mortality



Earlier placement in long-term care facilities



Reduced quality of life for patients and caregivers



Inability to maintain independence

Standard-of-care treatment options are insufficient: The only currently approved therapy carries a boxed warning for mortality in elderly people with dementia-related psychosis.

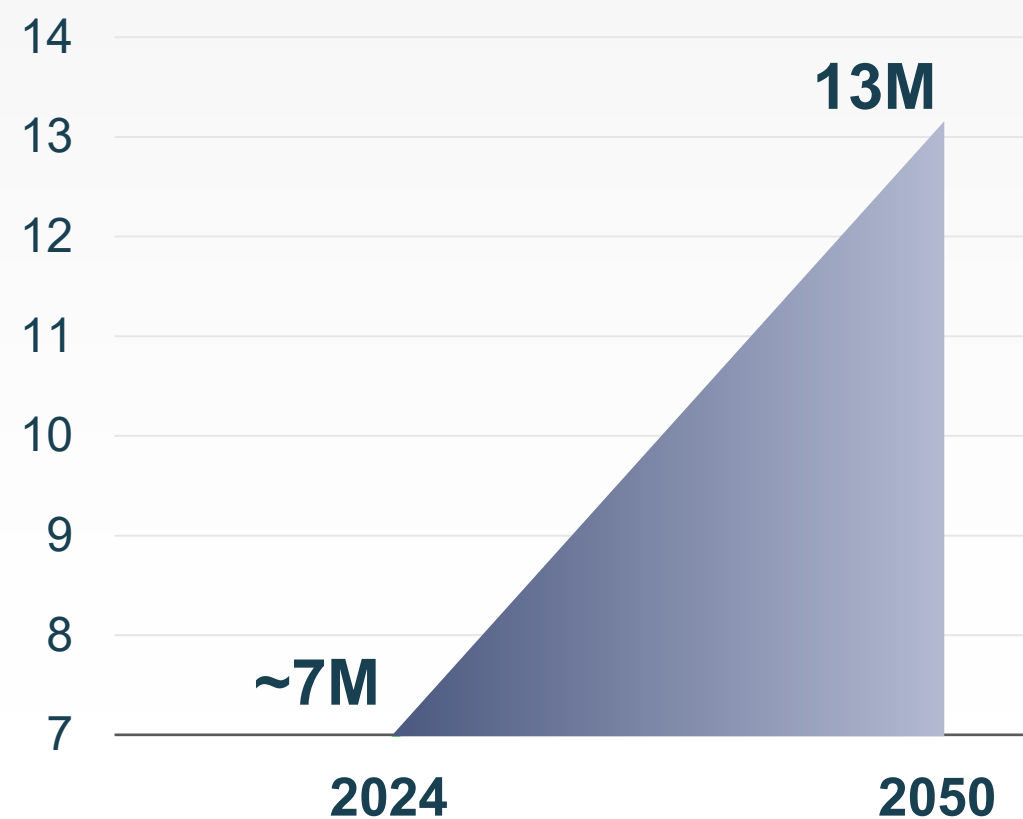
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Alzheimer's disease agitation represents large market opportunity with significant unmet need

Alzheimer's disease agitation is a large and growing health burden

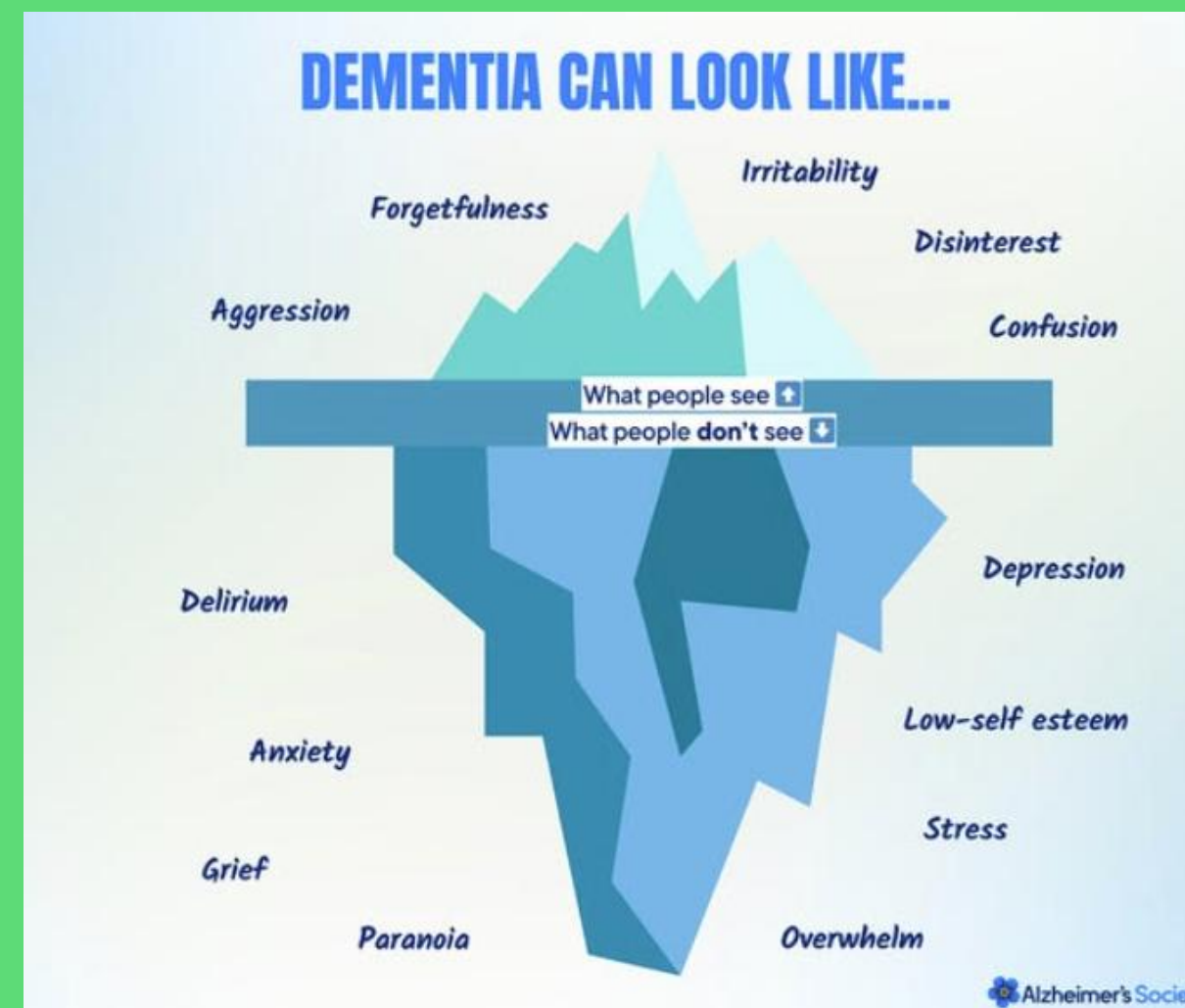
Millions currently living with AD; prevalence expected to increase as the population ages¹

U.S. Adults with Alzheimer's Disease (M)¹



>70%
of people with AD experience agitation at some point in their disease²

Anxiety is a key underlying driver of aggression and irritability in dementia³

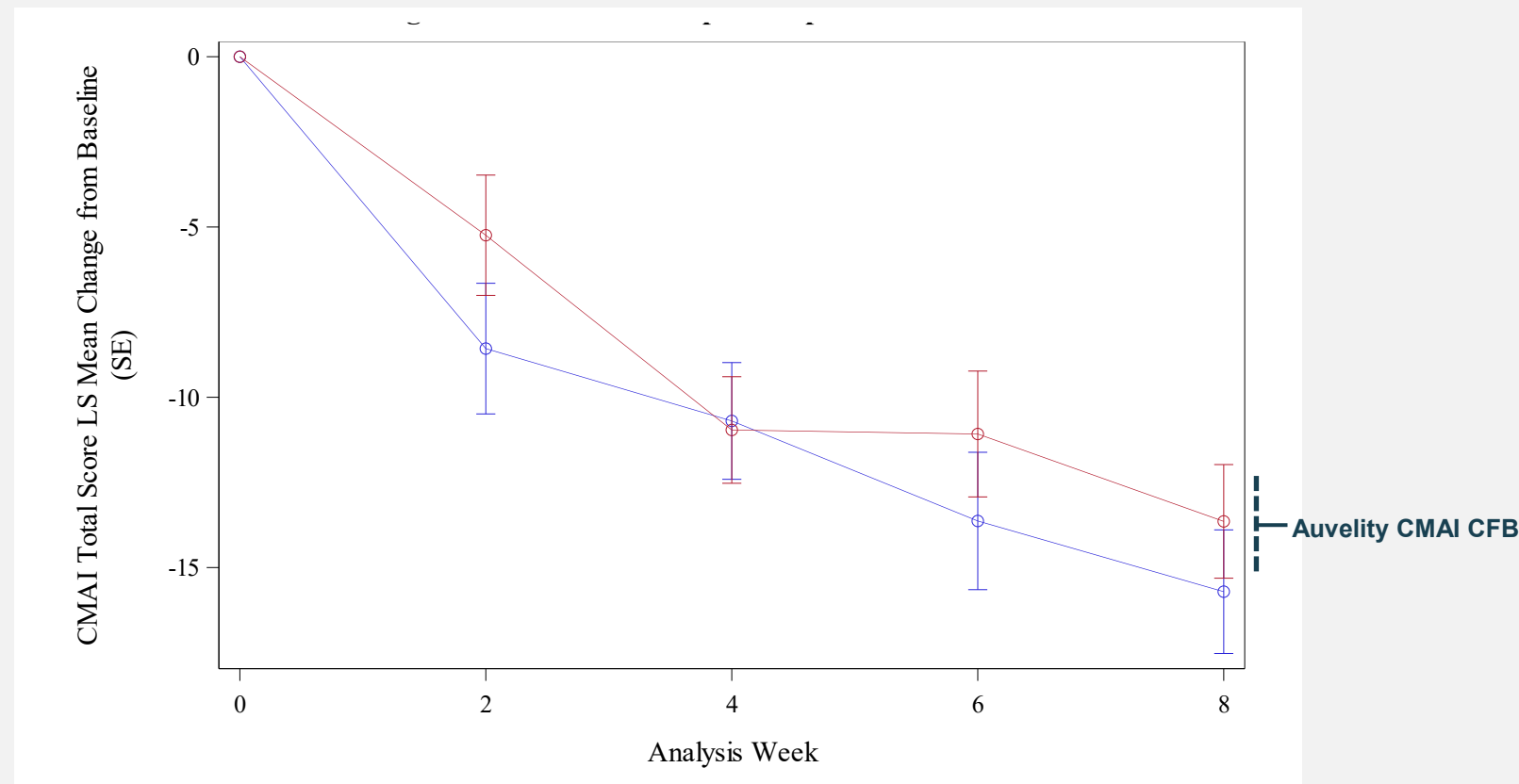


¹Alzheimer's Association. 2025 Alzheimer's Disease Facts and Figures. Alzheimer's Dementia 2025;21(5). ²Van der Musselle S, et al. Aging Ment Health 2015;19(3):247-257. ³Image from Alzheimer's Society

NMRA-511 demonstrated clinically meaningful reduction in CMAI total score and CMAI aggression sub-score

NMRA-511 demonstrated a 15.7-point reduction in CMAI total score at Week 8

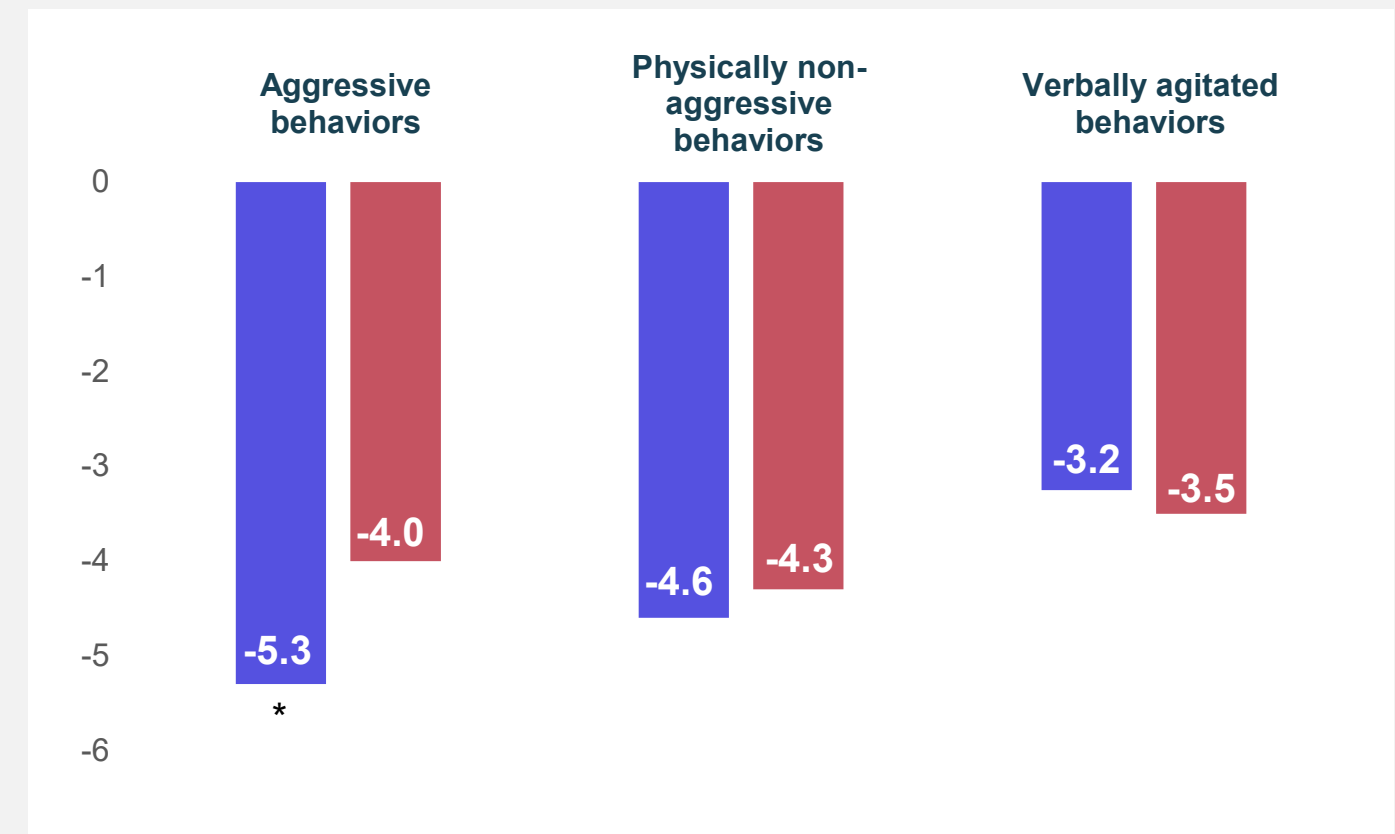
CMAI Total Score Change from Baseline (Modified Analysis Set)



— NMRA-511 — Placebo

CMAI aggression sub-score results suggest improvement on clinically relevant symptoms of AD agitation

Mean Change in CMAI Sub-Scores at Week 8 (Modified Analysis Set)



	Week 6	Week 8
LSMD (SE)	-2.6 (2.7)	-2.1 (2.5)
Effect size range (Cohen's d)	0.23	0.20



