

## Supplementary Online Content

Cummings JL, Lyketsos CG, Peskind ER, et al. Effect of dextromethorphan-quinidine on agitation in patients with alzheimer disease dementia: a randomized clinical trial.

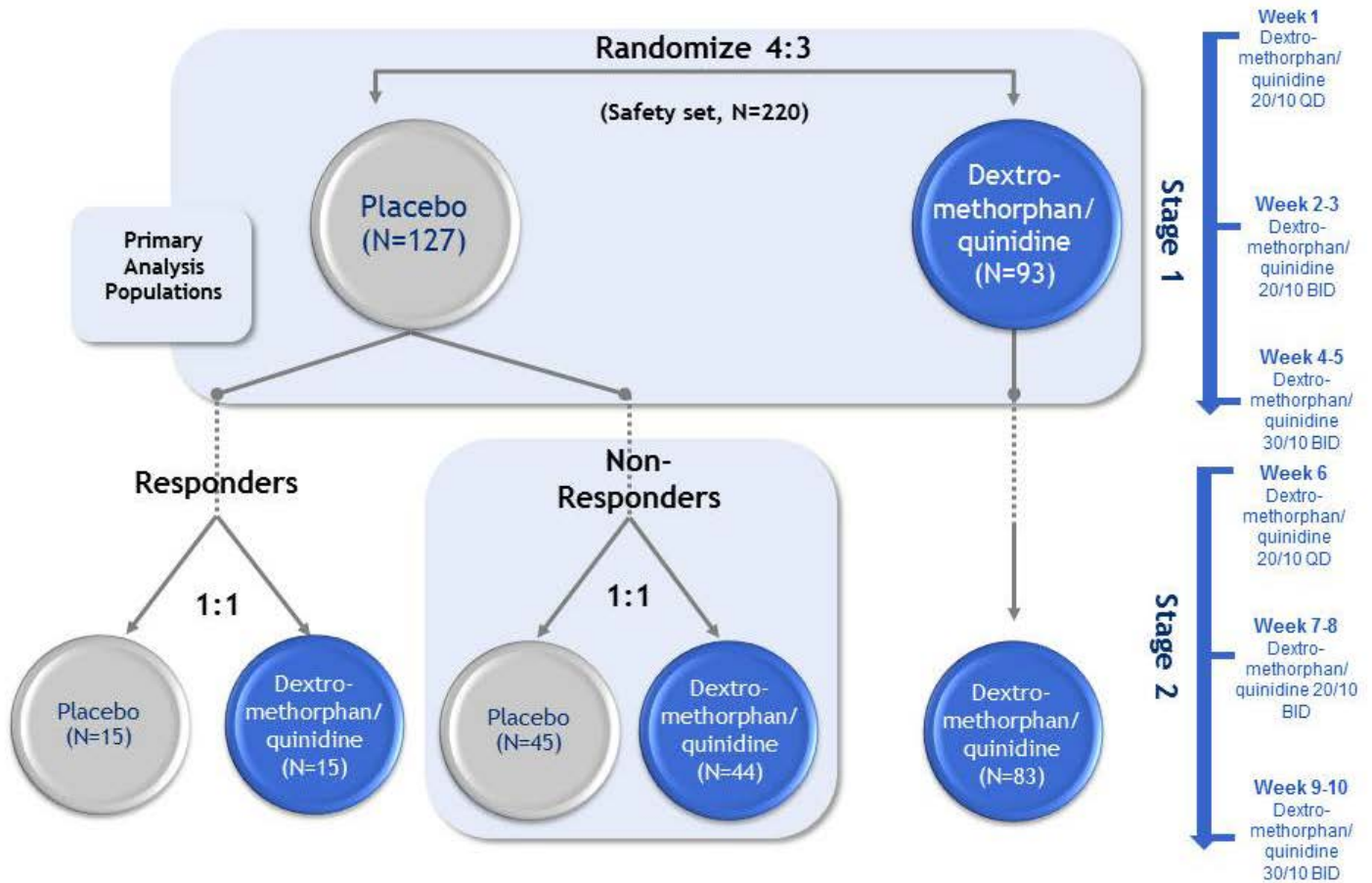
doi:10.1001/jama.2015.10214

**eFigure.** Study Design

**eTable.** Results of Repeated Measures Mixed Model (MMRM) and Seemingly Unrelated Regression (SUR) Sensitivity Analyses of the Primary Endpoint, Modified Intention-to-Treat Population

This supplementary material has been provided by the authors to give readers additional information about their work.

**eFigure. Study Design**



Shading shows the patient groups included in the primary SPCD analysis (all patients randomized in Stage 1 (MITT population) and placebo nonresponders from Stage 1 that were rerandomized in Stage 2, MITT population stage 2). A “response” in Stage 1 was defined as a CGIS score of 1-3 (inclusive) at Visit 4 (week 5/end of Stage 1) and an NPI Agitation/Aggression domain score decrease of  $\geq 25\%$ ; patients not meeting these criteria were considered “nonresponders”. BID = twice per day; QD = once daily.

**eTable.** Results of Repeated Measures Mixed Model (MMRM) and Seemingly Unrelated Regression (SUR) Sensitivity Analyses of the Primary Endpoint, Modified Intention-to-Treat Population

	Stage	N Dextromethorphan/ quinidine	N Placebo	Dextromethorphan/ quinidine, Mean (SD) Change from Baseline	Placebo, Mean (SD) Change from Baseline	Estimated Treatment Effect (95% CI)	Test Statistic <sup>b</sup>	P Value
<b>MMRM Analysis</b>								
NPI–Agitation/ Aggression <sup>a</sup>	1	93	125			-0.74 (-1.17, -0.31)	-3.30	.001
	Wk 1			-2.2 (2.9)	-1.4 (2.7)			
	Wk 3			-2.7 (3.2)	-1.6 (2.9)			
	Wk 5	-3.3 (3.0)	-1.7 (3.1)					
	2	142	60					
	Wk 6			-0.2 (2.8)	-0.9 (2.8)			
Wk 8	-0.6 (3.1)			-0.7 (2.9)				
Wk 10		-0.9 (3.4)	-0.5 (3.4)					
<b>SUR Analysis</b>								
NPI–Agitation/ Aggression <sup>a</sup>	1	93	125	-3.3 (3.0)	-1.7 (3.1)		15.5	<0.0001
	2	44	45	-2.0 (3.2)	-0.8 (3.6)			

<sup>a</sup>Assessed at baseline, weeks 1, 3, 5, 6, 8, and 10.