

# Identification of mild cognitive impairment (MCI) with digital biomarkers in individuals presenting with cognitive complaints

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## BACKGROUND

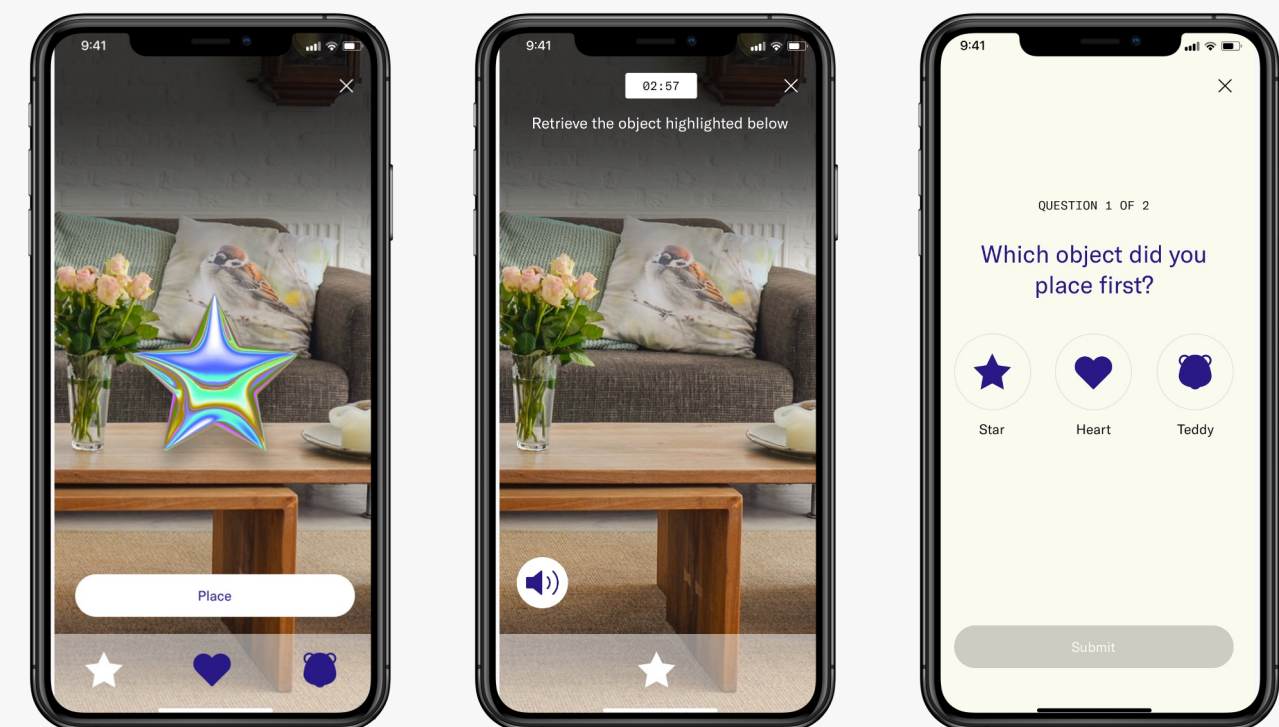
- A major challenge in Alzheimer's disease (AD) is the identification of individuals at the earliest stages, who might benefit from disease-modifying therapies.
- People presenting with subjective cognitive decline (SCD), or mild cognitive impairment (MCI) face a high risk of cognitive worsening over time.
- Digital biomarkers could enable early diagnoses, and thus, streamline the patient journey to specialized care.

## OBJECTIVE

- To compare scores of a digital cognitive assessment (Altoida Digital Neuro Signature; DNS) in SCD vs. MCI.
- To explore correlations between Altoida DNS scores and core AD biomarkers assessed in CSF.

## DIGITAL BIOMARKER ASSESSMENT

- Altoida DNS is a research device based on machine learning (ML) that simulates conducting activities of daily living, providing an objective measure of cognition.
- DNS-MCI is an ML model that can identify MCI/AD with 84% accuracy (Pipeline Version v1.56.0; 13Jun2023).



**Figure 1.** The Altoida DNS assessment evaluates cognitive and functional impairment based on a series of motoric and augmented reality tasks that simulate activities of daily living. The tasks in the Altoida assessment evaluate multi-modal features, including micro-movements, micro-errors, speed, reaction times, or navigation trajectories, which are used to train specific machine learning models. The test can be conducted on a smart device (iOS) and lasts approximately 10 minutes.

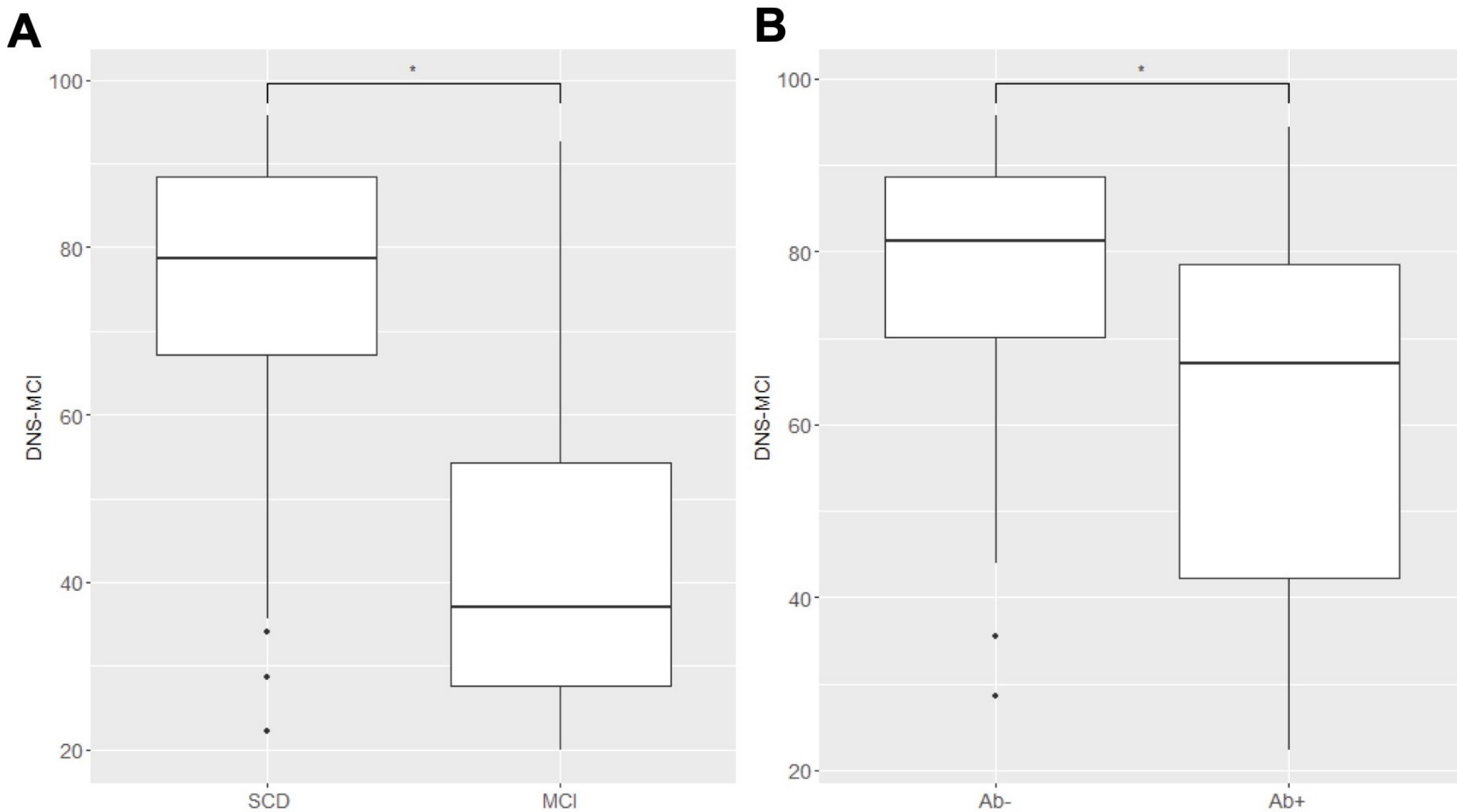
## STUDY POPULATION

Included 102 individuals with cognitive complaints seeking medical advice participating in the  $\beta$ -AARC study (Alzheimer's At Risk Cohort) at BBRC

	SCD (N=94)	MCI (N=8)	Total (N=102)	P-value
<b>Sex</b>				
female	52 (55.3%)	3 (37.5%)	55 (53.9%)	0.548
male	42 (44.7%)	5 (62.5%)	47 (46.1%)	
<b>Age (years)</b>				
Mean (SD)	66.4 (6.19)	70.1 (4.63)	66.7 (6.14)	0.0645
Median [Min, Max]	66.5 [55.6, 80.6]	70.4 [64.4, 76.1]	66.5 [55.6, 80.6]	
<b>Education years</b>				
Mean (SD)	14.8 (3.47)	12.0 (3.66)	14.6 (3.54)	0.0714
Median [Min, Max]	15.0 [8.00, 20.0]	11.0 [8.00, 18.0]	15.0 [8.00, 20.0]	
<b>DNS-MCI</b>				
Mean (SD)	73.9 (17.7)	46.3 (27.7)	71.8 (19.9)	0.0257
Median [Min, Max]	78.7 [22.3, 95.8]	37.0 [19.9, 92.7]	78.0 [19.9, 95.8]	
<b>MMSE</b>				
Mean (SD)	28.5 (1.33)	26.3 (2.76)	28.3 (1.58)	0.058
Median [Min, Max]	29.0 [24.0, 30.0]	27.0 [21.0, 29.0]	29.0 [21.0, 30.0]	
<b>Amyloid status</b>				
Ab-	77 (81.9%)	5 (62.5%)	82 (80.4%)	0.388
Ab+	17 (18.1%)	3 (37.5%)	20 (19.6%)	

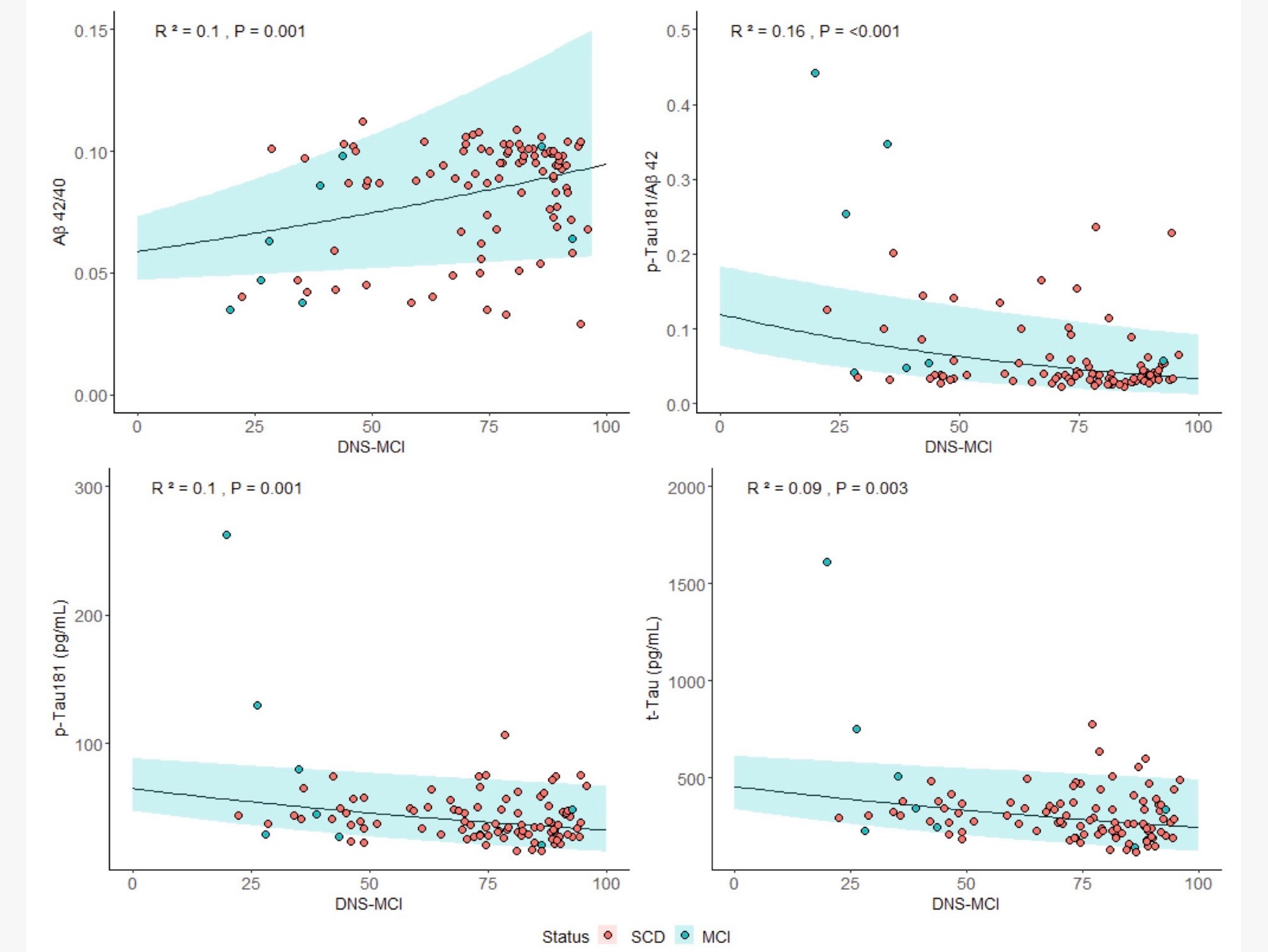
**Table 1.**  $A\beta+$  = amyloid- $\beta$  positive (based on the  $A\beta_{42/40}$  ratio cut-off of  $\leq 0.062$ );  $A\beta-$  = amyloid- $\beta$  negative; MCI: single or multidomain cognitive deficits with preservation of activities of daily living; MMSE = Mini-Mental State Examination; SCD: subjective perception of cognitive decline in the absence of cognitive impairment determined by a formal neuropsychological assessment\*) SD = standard deviation.

## GROUP DIFFERENCES IN DNS ASSESSMENT



**Figure 2.** (A) Comparison of DNS-MCI scores between individuals presenting with SCD (n=94) and MCI (n=8) and (B) between those identified as  $A\beta+$  (n=17) versus  $A\beta-$  (n=77) based on the  $A\beta_{42/40}$  ratio in the SCD subgroup only; \* $p < 0.05$ ; \*\*\* $p < 0.001$ , unpaired t-test.

## CORRELATION BETWEEN DNS & AD BIOMARKERS



**Figure 3.** Correlations between DNS-MCI scores and core CSF AD biomarkers. Beta regression with two-tailed Wald's test on the coefficients was used for the  $A\beta_{42/40}$  ratio and linear regression on the log-transformed values was used for the p-tau181 and t-tau biomarkers.

## CONCLUSIONS

- Altoida DNS-MCI identified differences in cognitive performance between MCI/SCD and  $A\beta+$ / $A\beta-$  in a cohort with cognitive complaints seeking medical advice.
- DNS-MCI scores correlated with core AD biomarkers.

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