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Reporting Summary

Nature Research wishes to improve the reproducibility of the work that we publish. This form provides structure for consistency and transparency in reporting. For further information on Nature Research policies, see our <u>Editorial Policies</u> and the <u>Editorial Policy Checklist</u>.

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FOL	all Statistical af	laryses, commit that the following items are present in the rigure legend, table legend, main text, or Methods Section.			
n/a	Confirmed				
	The exact	sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement			
	X A stateme	ent on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly			
	The statistical test(s) used AND whether they are one- or two-sided Only common tests should be described solely by name; describe more complex techniques in the Methods section.				
	X descript	tion of all covariates tested			
\boxtimes	A description of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons				
	A full description of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) AND variation (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)				
	For null hypothesis testing, the test statistic (e.g. <i>F</i> , <i>t</i> , <i>r</i>) with confidence intervals, effect sizes, degrees of freedom and <i>P</i> value noted Give <i>P</i> values as exact values whenever suitable.				
\boxtimes	For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings				
\boxtimes	For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes				
\square Estimates of effect sizes (e.g. Cohen's d , Pearson's r), indicating how they were calculated					
,		Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.			
Software and code					
Polic	cy information	about <u>availability of computer code</u>			
Da	ta collection	Altoida NMI			
Da	ta analysis	SPSS 22.0 for Mac			
		g custom algorithms or software that are central to the research but not yet described in published literature, software must be made available to editors and			

Data

Policy information about <u>availability of data</u>

All manuscripts must include a data availability statement. This statement should provide the following information, where applicable:

- Accession codes, unique identifiers, or web links for publicly available datasets
- A list of figures that have associated raw data $% \left(1\right) =\left(1\right) \left(1\right) \left($
- A description of any restrictions on data availability

The data that support the findings of this study are available from Altoida Inc. but restrictions apply to the availability of these data, which were used under license for the current study, and so are not publicly available. Data are however available from the authors upon reasonable request and with permission of Altoida Inc.

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Please select the one be	low that is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.
Life sciences	Behavioural & social sciences Ecological, evolutionary & environmental sciences
For a reference copy of the doo	cument with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>
Rehavioura	Il & social sciences study design
	, —
All studies must disclose	e on these points even when the disclosure is negative.
Study description	Quantitative prospective longitudinal study
Research sample	(1) 55-90 years of age, (2) fluency in English, French, Spanish, Greek, German or Italian, and (3) familiarity with digital devices, including currently possessing and actively using an iPad Pro or iPhone with an at-home Wi-Fi network for the remote assessments.
Sampling strategy	The MCI and AD cohorts were included independently on their biomarker status if their diagnosis was consistent with MCI and Alzheimer's dementia diagnosis according to core criteria of NIA-AA revised guidelines (Jack et al., 2011). The participant cohort in Study B (Table 2) is further detailed in Buegler & colleagues 2020. The cohort in Study A of symptomatic AD patients from the Hirslanden Clinic, Zurich, Switzerland was added for comparison (n=29).
Data collection	Using these criteria, we first recruited a control group of 283 cognitively healthy individuals that underwent the same procedure at the Global Brain Health Institute (GBHI) at Trinity College, Dublin. In recruiting participants with cognitive impairments, the biomarkers (CSF, brain MRI and ApoE genotype) were used as a criterion, and cognitive deficits compatible with MCI diagnosis were found in 213 subjects: 170 from the memory clinics and primary care centers in Europe and 43 from the community centers in the USA. Seven participants were excluded from the data analysis due to poor data quality. The Study B cohort consisted of HC (n=283) and patients with MCI who are at high risk of developing AD within 18-40 months (n=213), assessed every 6 months.
Timing	October 17, 2016 recruitment start, February 21, 2020 study completion.
Data exclusions	Seven participants were excluded from the data analysis due to poor data quality.
Non-participation	N/A
Randomization	The MCI and AD cohorts were included independently on their biomarker status if their diagnosis was consistent with MCI and Alzheimer's dementia diagnosis according to core criteria of NIA-AA revised guidelines (Jack et al., 2011).

Reporting for specific materials, systems and methods

We require information from authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, system or method listed is relevant to your study. If you are not sure if a list item applies to your research, read the appropriate section before selecting a response.

Materials & experimental systems	Methods	
n/a Involved in the study	n/a Involved in the study	
Antibodies	ChIP-seq	
Eukaryotic cell lines	Flow cytometry	
Palaeontology and archaeology	MRI-based neuroimaging	
Animals and other organisms		
Human research participants		
Clinical data		
Dual use research of concern		

Corfu, Greece where the study was initiated

Human research participants

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Population characteristics	See above
Recruitment	Semi-naturalistic observational multicenter study performed in ten European memory clinics and primary care centers and two primary care community centers in the USA
Ethics oversight	Both studies were approved by the local institutional review board (IRB), Bioethics committee of the Ionian University in

Note that full information on the approval of the study protocol must also be provided in the manuscript.

Clinical data

Policy information about <u>clinical studies</u>

All manuscripts should comply with the ICMJE guidelines for publication of clinical research and a completed CONSORT checklist must be included with all submissions.

Clinical trial registration NCT02050464; NCT02843529

Study protocol Clinicaltrials.gov

Data collection Recruitment from the memory clinics and primary care centers in Europe and another sample from the community centers in the USA. Locale was English, French, Spanish, Greek, German or Italian. October 17, 2016 recruitment start and February 21, 2020 study

completion.

Outcomes Primary outcome was Sensitivity, specificity and accuracy of models for diagnosis of memory disorders and also change in Diagnostic

Area Under the Receiver Operating Characteristic Curve (ROC-AUC).