# nature portfolio

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# **Reporting Summary**

Nature Portfolio 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 Portfolio policies, see our <u>Editorial Policies</u> and the <u>Editorial Policy Checklist</u>.

Please do not complete any field with "not applicable" or n/a. Refer to the help text for what text to use if an item is not relevant to your study. For final submission: please carefully check your responses for accuracy; you will not be able to make changes later.

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Statistics			
For all statistical an	alyses, confirm that the following items are present in the figure 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	nt on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly		
The statist	ical test(s) used AND whether they are one- or two-sided on tests should be described solely by name; describe more complex techniques in the Methods section.		
X A descript	ion of all covariates tested		
A descript	ion of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons		
A full desc	ription of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) tion (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)		
	pothesis testing, the test statistic (e.g. $F$ , $t$ , $r$ ) with confidence intervals, effect sizes, degrees of freedom and $P$ value noted as as exact values whenever suitable.		
X For Bayesi	$\overline{\mathbf{X}}$ $\square$ For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings		
X For hierard	X For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes		
☐ X Estimates	$\left  \mathbf{X} \right $ Estimates of effect sizes (e.g. Cohen's <i>d</i> , Pearson's <i>r</i> ), indicating how they were calculated		
1	Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.		
Software and	d code		
Policy information a	about <u>availability of computer code</u>		
Data collection	Preprocessing code is available at: https://github.com/neurolabusc/nii_preprocess		

All scripts used for analysis will be made available on Dr Bonilha and Dr Rorden's Github.

For manuscripts utilizing 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 reviewers. We strongly encourage code deposition in a community repository (e.g. GitHub). See the Nature Portfolio guidelines for submitting code & software for further information.

## Data

Data analysis

Policy information about availability of data

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

- Accession codes, unique identifiers, or web links for publicly available datasets
- A description of any restrictions on data availability
- For clinical datasets or third party data, please ensure that the statement adheres to our policy

Source data will be shared upon reasonable request to the corresponding author. Due to the limitations of our ethics, a request is deemed as reasonable if it does not require identifiable information relating to the participants. Data used to create graphs are included in Supplementary Data 1.

Research involving	human participants, their data, or biological material
Policy information about student and sexual orientation and ra	dies with <u>human participants or human data</u> . See also policy information about <u>sex, gender (identity/presentation),</u> ice, ethnicity and racism.
Reporting on sex and gende	Found in Supplementary Table 1.
Reporting on race, ethnicity other socially relevant groupings	y, or Found in Supplementary Table 1.
Population characteristics	Found in Methods and Supplementary Table
Recruitment	Information found in Methods, under sections 'Healthy Controls' and 'Stroke Participants'
Ethics oversight	Found in Methods section
Note that full information on the	approval of the study protocol must also be provided in the manuscript.
Field-specific	
Please select the one below t	that is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.
Life sciences	X Behavioural & social sciences Ecological, evolutionary & environmental sciences
For a reference copy of the documen	t with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>
Life sciences	study design
All studies must disclose on t	hese points even when the disclosure is negative.
Sample size	
Data exclusions	
Replication	
Randomization	
Blinding	
-	
Behavioural 8	& social sciences study design
All studies must disclose on t	hese points even when the disclosure is negative.
Study description	Investigation into regional brain aging in stroke aphasia and how it relates to behavior.
Research sample	232 healthy participants and 89 participants with stroke aphasia
Sampling strategy	Recruited for previous studies at the University of South Carolina
Data collection	Data were collected at the University of South Carolina Research labs.
Limina	Participants were recruited either as part of the ongoing ABC@USC project, or as part of the POLAR clinical trial (NCT0341678) which spanned 5 years.
Data exclusions	lealthy controls were excluded if they were not proficient in English, stroke participants were excluded if they had severely limited verbal utput or comprehension, or contra-indications for MRI.

Data exclusions Non-participation

Randomization

N/A N/A

Ecological, e	volutionary & environmental sciences study design
All studies must disclose on	these points even when the disclosure is negative.
Study description	
Research sample	
Sampling strategy	
Data collection	
Timing and spatial scale	
Data exclusions	
Reproducibility	
Randomization	
Blinding	
Did the study involve field	work? Yes No
Field work, collect	ion and transport
Field conditions	
Location	
Access & import/export	
Disturbance	
We require information from a	r specific materials, systems and methods uthors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, vant 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 & experime	ntal systems Methods
n/a   Involved in the study	n/a Involved in the study
Antibodies	X ChIP-seq
Eukaryotic cell lines	X   Flow cytometry
Palaeontology and a  Animals and other o	— I—
Clinical data	
Dual use research of	concern
<b>X</b> Plants	
Antibodies	
Antibodies used	
Validation	

Eukaryotic cell lin	es
Policy information about ce	ell lines and Sex and Gender in Research
Cell line source(s)	
Authentication	
Mycoplasma contaminati	ion
Commonly misidentified (See <u>ICLAC</u> register)	lines
Palaeontology an	d Archaeology
Specimen provenance	
Specimen deposition	
Dating methods	
Tick this box to confir	m that the raw and calibrated dates are available in the paper or in Supplementary Information.
Ethics oversight	
Note that full information on t	he approval of the study protocol must also be provided in the manuscript.
Animals and othe	r research organisms
	udies involving animals; ARRIVE guidelines recommended for reporting animal research, and Sex and Gender in
Laboratory animals	
Wild animals	
Reporting on sex	
Field-collected samples	
Ethics oversight	
Note that full information on t	he approval of the study protocol must also be provided in the manuscript.
Clinical data	
Policy information about <u>cl</u> All manuscripts should comply	inical studies with the ICMJE guidelines for publication of clinical research and a completed CONSORT checklist must be included with all submissions.
Clinical trial registration	NCT0341678
Study protocol	Described in the Methods
Data collection	Described in the Methods
Outcomes	Data from a clinical trial were used, but this manuscript does not report outcomes of the clinical trial.
Dural management	

#### Dual use research of concern

Policy information about <u>dual use research of concern</u>

### Hazards

Could the accidental, deliberate or reckless misuse of agents or technologies generated in the work, or the application of information presented in the manuscript, pose a threat to:

No Yes  Public health  National security  Crops and/or livest  Ecosystems  Any other significan	
Experiments of concer	n
No Yes  Demonstrate how to the confer resistance to the confer resistan	to render a vaccine ineffective o therapeutically useful antibiotics or antiviral agents nce of a pathogen or render a nonpathogen virulent ibility of a pathogen e of a pathogen diagnostic/detection modalities nization of a biological agent or toxin lly harmful combination of experiments and agents
Plants	
Seed stocks	
Novel plant genotypes	
Authentication	
ChIP-seq	
Confirm that you have	y and final processed data have been deposited in a public database such as <u>GEO</u> . e deposited or provided access to graph files (e.g. BED files) for the called peaks.
Data access links May remain private before public	cation.
Files in database submissi	on (
Genome browser session (e.g. <u>UCSC</u> )	
Methodology	
Replicates	
Sequencing depth	
Antibodies	
Peak calling parameters	
Data quality	

Software	
Flow Cytometry	
The axis scales are clearly vi	rker and fluorochrome used (e.g. CD4-FITC). sible. Include numbers along axes only for bottom left plot of group (a 'group' is an analysis of identical markers). ith outliers or pseudocolor plots.
A numerical value for numb	er of cells or percentage (with statistics) is provided.
Methodology	
Sample preparation	
Instrument	
Software	
Cell population abundance	
Gating strategy	
Tick this box to confirm that	a figure exemplifying the gating strategy is provided in the Supplementary Information.
Magnetic resonance i	maging
Experimental design	
Design type	Cohort study
Design specifications	Cross-sectional cohort
Behavioral performance measu	res Western Aphasia Battery
Imaging type(s)	T1-weighted and T2-weighted MRI
Field strength	3T Siemens Trio
Sequence & imaging parameter	MP-RAGE sequence with 1mm isotropic voxels, a 256 x 256 matrix size, a 9 degree flip angle, and a 92-slice squence with repetition time (TR) = 2250ms, inversion time(TI)=925ms, and echo time (TE) = 4.11ms.
Area of acquisition	Brain
Diffusion MRI Used	$\overline{f X}$ Not used
Preprocessing	
Preprocessing software	Matlab, SPM12, ASLtbx, MRItrix
Normalization	BrainAgeR analysis pipeline - github.com/james-cole/brainageR
Normalization template	BrainAgeR analysis pipeline - github.com/james-cole/brainageR
Noise and artifact removal	BrainAgeR analysis pipeline - github.com/james-cole/brainageR
Volume censoring	BrainAgeR analysis pipeline - github.com/james-cole/brainageR
Statistical modeling & inference	ence
Model type and settings	Multiple linear regression, mixed effects ANCOVA, independent samples t-test, correlations
Effect(s) tested	Brain ages between hemisphere, and controls vs patients, which variables predict behavior

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Specify type of analysis:	ROI-based X Both	
Statistic type for inference Correlati	on, independent samples t-test, mixed effects ANCOVA, multiple linear regression	
(See Eklund et al. 2016)		
Correction Adjusted	alpha levels to account for multiple comparisons	
Models & analysis		
n/a   Involved in the study		
X Functional and/or effective connectivity		
X Graph analysis		
Multivariate modeling or predictive analysis		
Functional and/or effective connectivity		
Graph analysis		
Multivariate modeling and predictive analys	Multiple linear regression to estimate multivariate relationship between gray matter volume and brain age	