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Armin P. Schoech Corresponding author(s): Alkes L. Price

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 <u>Authors & Referees</u> and the <u>Editorial Policy Checklist</u>.

When statistical analyses are reported, confirm that the following items are present in the relevant location (e.g. figure legend, table legend, main

Statistical parameters

text, or Methods section).					
n/a	Confirmed				
	\square The exact sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement				
	An indication of 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.				
	A description of all covariates tested				
	A description of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons				
	A full description of the statistics including <u>central tendency</u> (e.g. means) or other basic estimates (e.g. regression coefficient) AND <u>variation</u> (e.g. standard deviation) or associated <u>estimates of uncertainty</u> (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 <i>Give P values as exact values whenever suitable.</i>				
	🔀 For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings				
	For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes				
	Estimates of effect sizes (e.g. Cohen's <i>d</i> , Pearson's <i>r</i>), indicating how they were calculated				
	Clearly defined error bars State explicitly what error bars represent (e.g. SD, SE, CI)				

Our web collection on statistics for biologists may be useful.

Software and code

 Policy information about availability of computer code

 Data collection
 We used GCTA (version 1.24.7) to calculate the variance components and likelihood for a given GRM. GRMs were calculated using software designed by the authors, which can be found at "https://github.com/arminschoech/GRM-MAF-LD".

 Data analysis
 Data was also analyzed using R (version 3.4.1).

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/reviewers upon request. We strongly encourage code deposition in a community repository (e.g. GitHub). See the Nature Research guidelines for submitting code & software for further information.

Data

Policy information about availability of data

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
- A description of any restrictions on data availability

The study involved publicly available data sets (UK10K and UK Biobank project data).

Field-specific reporting

Please select the best fit for your research. If you are not sure, read the appropriate sections before making your selection.

Life sciences

s Behavioural & social sciences

Ecological, evolutionary & environmental sciences

For a reference copy of the document with all sections, see <u>nature.com/authors/policies/ReportingSummary-flat.pdf</u>

Life sciences study design

All studies must dis	close on these points even when the disclosure is negative.		
Sample size	We use the UK Biobank phase 1 data release, which comprises of data from 152,729 individuals. We only used data from all 113,851 individuals who have self-reported and confirmed British ancestry and pairwise genetic relatedness < 5%. Reasonable error size in our estimates confirm that the sample size is sufficient.		
Data exclusions	see 'Sample size' answer above		
Replication	NA (no experiments were performed)		
Randomization	NA (we did not allocate participants in experimental groups)		
Blinding	NA (we used existing UK Biobank data)		

Reporting for specific materials, systems and methods

Materials & experimental systems

Unique biological materials

 Methods		
 n/a	Involved in the study	
\boxtimes	ChIP-seq	
\mathbf{X}	Flow cytometry	

\times		Antibodies
\mathbf{X}	\square	Eukaryotic cell lines

Involved in the study

n/a

 \boxtimes

Palaeontology

Animals and other organisms

Human research participants

Human research participants

Policy information about studies involving human research participants

NA

Population characteristics

The study involved publicly available data sets (UK10K and UK Biobank project data).

Recruitment

MRI-based neuroimaging