## nature research

Corresponding author(s):	Daniel J. Benjamin
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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 Editorial Policies and the Editorial Policy Checklist.

For all statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.

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n/a	Confirmed
	$\square$ The exact sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement
	🔀 A statement 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.
	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 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 <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
$\boxtimes$	For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes
	$\boxtimes$ Estimates of effect sizes (e.g. Cohen's $d$ , Pearson's $r$ ), indicating how they were calculated
	Our web collection on statistics for biologists contains articles on many of the points above.

## Software and code

Policy information about <u>availability of computer code</u>

Data collection

No software was used for data collection.

Data analysis

The following software packages were used for data analysis: Python version 3.7.4 with packages pandas 0.25.1, scipy 1.3.1, numpy 1.17.2, matplotlib 3.1.1 and argparse 1.1 (https://anaconda.org); R version 4.0.3 with packages EasyQC 9.2, plotrix 3.7.8, tidyr 1.1.3 and readstata13 0.9.2, R version 3.6 (https://www.r-project.org); GCTA 1.93.2beta (https://yanglab.westlake.edu.cn/software/gcta/#Overview); GCTB 2.03 (https://cnsgenomics.com/software/gctb/#Overview); Stata 16.1 (https://www.stata.com); PLINK 1.9 (https://www.cog-genomics.org/plink/1.9); PLINK 2 (https://www.cog-genomics.org/plink/2.0); LDpred 1.0.11 (https://github.com/bvilhjal/ldpred); METAL release 2011-03-25 (https://genome.sph.umich.edu/wiki/METAL\_Documentation); BOLT-LMM 2.3 (https://alkesgroup.broadinstitute.org/BOLT-LMM/BOLT-LMM\_manual.html); LDSC 1.0.1 (https://github.com/bulik/ldsc); SNIPar (https://github.com/AlexTISYoung/SNIPar/tree/EA4).

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 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

GWAS summary statistics can be downloaded from http://www.thessgac.org/data subject to a Terms of Use to ensure responsible use of the data. We provide association results for all SNPs that passed quality-control filters in autosomal, X chromosome, and dominance GWAS meta-analyses that excludes the research

participants from 23andMe. SNP-level summary statistics from analyses based entirely or in part on 23andMe data can only be reported for up to 10,000 SNPs. For the complete dominance GWAS meta-analysis, which includes 23andMe, clumped results for the 1,000 SNPs with the smallest P values are provided. For the complete autosomal and X chromosome GWAS meta-analyses, respectively, clumped results for the 8,617 and 143 SNPs with P < 10-5 are provided; this P value threshold was chosen such that the total number of SNPs across the analyses that include data from 23andMe does not exceed 10,000. The full GWAS summary statistics from 23andMe will be made available through 23andMe to qualified researchers under an agreement with 23andMe that protects the privacy of the 23andMe participants. Please visit https://research.23andme.com/collaborate/#dataset-access/ for more information and to apply to access the data.

Field-specific reporting				
Please select the one below	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 docume	ent with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>			
Behavioural	& social sciences study design			
All studies must disclose on	these points even when the disclosure is negative.			
Study description	This is a genome-wide association study (GWAS) meta-analysis of educational attainment (EA) in a sample of ~3 million individuals.  All data used in this study (genetic and phenotype data) are quantitative.			
Research sample	The research sample consists of $^{\sim}3$ million individuals from 71 research cohorts. We meta-analyzed three sets of summary statistics: publicly available results from Lee et al. (2018) that exclude 23andMe and UKB (N = 324,162), new association results from 23andMe (N = 2,272,216), and new association results from a GWAS we conducted in UKB with an improved coding of the EA measure (N = 441,121). The large study sample was required for us to have sufficient statistical power in detecting single nucleotide polymorphisms (SNPs) with small effect sizes and for our follow-up analyses.			
Sampling strategy	We obtained the largest sample we could.			
Data collection	Data collection was performed independently by each participating cohort.			
Timing	Data was collected from multiple cohorts with variable data collection periods.			
Data exclusions	All observations reporting less than seven years of schooling were dropped to exclude outliers (there were fewer than 50 such observations; see Supplementary Note 1.1.4).			
Non-participation	No participants dropped out or declined participation.			
Randomization	Participants were not allocated into experimental groups.			
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 & 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 a	rchaeology MRI-based neuroimaging			
Animals and other organisms				
Human research participants				
Clinical data  Dual use research of concern				
Dual use research of	Concern			
Human research រុ	participants			

Policy information about studies involving human research participants

See above.

Population characteristics

Recruitment

Recruitment strategies were particular to each cohort.

Ethics oversight

All analyses are on anonymized, secondary data. Nonetheless, the analyses reported in the paper fall under National Bureau of Economic Research IRB protocols 19\_434, 19\_465, and 20\_041.

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