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

Statistical parameters

When statistical analyses are reported,	confirm that the following items are pres	sent in the relevant location	(e.g. figure legend	, table legend, main
text, or Methods section).				

n/a	Cor	nfirmed
	\boxtimes	The exact sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement
\boxtimes		An indication of whether measurements were taken from distinct samples or whether the same sample was measured repeatedly
	\boxtimes	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.
	\boxtimes	A description of all covariates tested
	\times	A description of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons
	\boxtimes	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)
	\boxtimes	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>
\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
	\times	Estimates of effect sizes (e.g. Cohen's d, Pearson's r), indicating how they were calculated
	\boxtimes	Clearly defined error bars State explicitly what error bars represent (e.g. SD, SE, CI)

Our web collection on <u>statistics for biologists</u> may be useful.

Software and code

Policy information about availability of computer code

Data collection

No software was used to collect data, as no data were generated.

Data analysis

We used the SLDP package, available on GitHub. We also ran Basset and Plink2, using the 2017 versions of each tool.

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

No data were generated in this study.

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	Life sciences 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	sclose on these points even when the disclosure is negative.			
Sample size	We analyzed available GWAS data, and did not do a new experiment in which we determined the sample size.			
Data exclusions	We excluded the HLA from all analyses and analyzed only autosomes. We excluded signed functional annotations for which Basset AUPRC was <0.3 or for which <5000 reference panel SNPs had non-zero effects. Otherwise, no data were excluded.			
Replication	There were no experimental findings. Where possible, we validated computational results using gene expression data, orthogonal GWAS signals, and gene sets from MSigDB.			

Reporting for specific materials, systems and methods

Materials & experimental systems			Methods		
n/a	Involved in the study		Involved in the study		
\boxtimes	Unique biological materials	\boxtimes	ChIP-seq		
\boxtimes	Antibodies	\boxtimes	Flow cytometry		
\boxtimes	Eukaryotic cell lines	\boxtimes	MRI-based neuroimaging		
\boxtimes	Palaeontology				
\boxtimes	Animals and other organisms				
\boxtimes	Human research participants				

We did not allocate samples into experimental groups.

There was no group allocation.

Randomization

Blinding