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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 Authors & Referees and the Editorial Policy Checklist.

Statistics				
For all statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.				
n/a	Cor	nfirmed		
		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)		
	\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>		
X		For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings		

Our web collection on statistics for biologists contains articles on many of the points above.

Software and code

Policy information about availability of computer code

Data collection

The genotype and death registry information are available through the UK Biobank application 33672.

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 d, Pearson's r), indicating how they were calculated

Software used: Plink2

Data analysis

Analytical results and scripts are available on https://github.com/AprilWei001/CCR5-delta32

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. 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 genotype and death registry information are available with the permission of the UK Biobank. Analytical results and scripts are accessible through (https:// github.com/AprilWei001/CCR5-delta32). In addition, a detailed experimental notebook covering the entire development of this project is available at depository (https://xinzhuaprilwei.weebly.com/download/ccr5-delta32).

Field-specific reporting				
<u> </u>	ne below that is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.			
X Life sciences	Behavioural & social sciences			
For a reference copy of the document with all sections, see nature.com/documents/nr-reporting-summary-flat.pdf				
Life sciences study design				
All studies must dis	close on these points even when the disclosure is negative.			
Sample size	le size 409,693. This is the number of British ancestry volunteer that are genotyped in the UK Biobank. The sample size is sufficient because the delta32 has a relatively high MAF (0.1159).			
Data exclusions	Exclude non-British ancestry volunteers from the UK Biobank to control for the genetic background and for the purpose of calculating Hardy-Weinberg proportion. Exclude samples whose estimated age from year/month of birth do not agree with self-reported age.			
Replication	plication Each reported result is confirmed by several statistical approaches, in addition to two lines of independent evidences confirming each other.			
Randomization	Randomization were not employed.			
Blinding Yes, data is collected and de-identified by UK Biobank.				
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 & exp	perimental systems Methods			
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Antibodies ChIP-seq				
Eukaryotic cell lines Flow cytometry				
Palaeontology MRI-based neuroimaging				
Animals and other organisms				
Human research participants Clinical data				
Human research participants				
Policy information about studies involving human research participants				
Population chara	cteristics British ancestry. Age 40-69 at recruitment. Genotype information from blood-derived DNA.			

Recruitment The UK Biobank recruited volunteers by sending out invitation letters to homes of people aged 40-69. Volunteers then signed up at assessment centers. There can be "healthy volunteer effect" such that people who volunteer are likely healthier than the

general population.

Ethics oversight UC Berkeley, UK Biobank

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