

Reporting Summary

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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 Confirmed

- 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. F , t , r) with confidence intervals, effect sizes, degrees of freedom and P value noted
Give P values as exact values whenever suitable.
- 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 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 [availability of computer code](#)

Data collection The genome-wide association summary-level data was obtained via publicly available links online. The individual phenotypic data was obtained by application to UK Biobank.

Data analysis The GSMR analysis was performed using GCTA 1.93.1beta: <https://cnsgenomics.com/software/gcta/>
All results was analyzed using R 4.0.2: <https://www.r-project.org/>.
The bivariate genomic scan was performed using R package "MultiABEL": <https://github.com/xiashen/MultiABEL>
The pathway enrichment analysis for GWAS summary statistics was performed on FUMA platform: <https://fuma.ctglab.nl/>
The genetic correlation analysis was performed using: LDSC v1.0.1: <https://github.com/bulik/ldsc> and HDL v1.3.8: <https://github.com/zhenin/HDL>

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 used in this study are publicly available (for URLs, see Table S1). The individual-level phenotype data are available in the UK Biobank (<http://www.ukbiobank.ac.uk/>) upon application and with permission of UKBB's Research Ethics Committee. The source data for the main figures can be accessed

as Supplementary Data 2-6. The bivariate GWAS summary statistics of aging-related COVID-19 generated in this study are available on Figshare (https://figshare.com/articles/dataset/combined_ukbbCOVID_meta_txt/16416822).

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 document with all sections, see nature.com/documents/nr-reporting-summary-flat.pdf

Life sciences study design

All studies must disclose on these points even when the disclosure is negative.

Sample size	The work described in this manuscript uses summary statistics from previously conducted genome-wide association studies and the phenotype data from the UK Biobank cohort. Therefore, we do not control the sample size for these studies. Sample sizes for all summary statistics included are given in Supplementary Tables S1 and S3. Sample sizes for the UK Biobank traits are shown in the Figure 2 and S2.
Data exclusions	Duplicated markers were excluded from the summary statistics prior to the analysis. We didn't exclude any sample from the UK Biobank dataset.
Replication	The results are reproducible by re-running the analysis pipeline. No additional replication studies was applicable.
Randomization	The work used publicly available summary statistics and UK Biobank observational data, therefore no randomization was applied.
Blinding	The investigators had no involvement in data collection. No group allocation was applicable.

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 & experimental systems

n/a	Involved in the study
<input checked="" type="checkbox"/>	<input type="checkbox"/> Antibodies
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<input checked="" type="checkbox"/>	<input type="checkbox"/> Palaeontology and archaeology
<input checked="" type="checkbox"/>	<input type="checkbox"/> Animals and other organisms
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<input checked="" type="checkbox"/>	<input type="checkbox"/> Clinical data
<input checked="" type="checkbox"/>	<input type="checkbox"/> Dual use research of concern

Methods

n/a	Involved in the study
<input checked="" type="checkbox"/>	<input type="checkbox"/> ChIP-seq
<input checked="" type="checkbox"/>	<input type="checkbox"/> Flow cytometry
<input checked="" type="checkbox"/>	<input type="checkbox"/> MRI-based neuroimaging

Human research participants

Policy information about [studies involving human research participants](#)

Population characteristics	The data used in this work are summary statistics from 46 genome wide association studies and phenotypic information from the UK Biobank cohort. These studies were all conducted in adults. The detailed ancestry information are provided in Supplementary Table S1.
Recruitment	We did not recruit the study participants.
Ethics oversight	Besides the publicly available summary-level data, ethical approval was provided by the UK Biobank cohort.

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