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Last updated by author(s):	Dec 16, 2020

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.

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1 01	ali statisticai ai	laryses, commit that the following items are present in the righter legend, table legend, main text, or interious section.				
n/a	Confirmed					
	The exact	$oxed{\boxtimes}$ The exact sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement				
\boxtimes	A stateme	ent on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly				
	The statis Only comm	itical test(s) used AND whether they are one- or two-sided non tests should be described solely by name; describe more complex techniques in the Methods section.				
	A descript	tion of all covariates tested				
\boxtimes	A descript	tion of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons				
	A full deso	cription of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) ation (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)				
	For null h	ypothesis testing, the test statistic (e.g. F , t , r) with confidence intervals, effect sizes, degrees of freedom and P value noted uses as exact values whenever suitable.				
\boxtimes	For Bayes	sian analysis, information on the choice of priors and Markov chain Monte Carlo settings				
\boxtimes	For hierar	rchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes				
	Estimates	s of effect sizes (e.g. Cohen's d, Pearson's r), indicating how they were calculated				
	I	Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.				
So	ftware an	d code				
Poli	cy information	about availability of computer code				
Dá	ata collection	R statistical language was used to pull the data from institutional repository approved by IRB.				
Da	Data analysis all analyses were performed using R statistical language.					
		g 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 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 <u>availability of data</u>

All manuscripts must include a <u>data availability statement</u>. 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 $% \left(1\right) =\left(1\right) \left(1\right) \left($
- A description of any restrictions on data availability

Protected Health Information restrictions apply to the availability of the clinical data here, which were used under IRB approval for use only in the current study. As a result, this dataset is not publicly available.

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Ticia spe	terre reporting
Please select the o	ne 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 scier	nces study design
All studies must dis	sclose on these points even when the disclosure is negative.
Sample size	data from the entire COVID-19 population in the institutional repository was used. Iterative random sampling was performed to divided data into train (80%) and test (20%) set.
Data exclusions	For model training, we under-sampled the negative cases due to highly unbalanced labels. To do so, we assumed a patient alive if she did not die within 7-28 days after the COVID-19 PCR test. Patients were initially limited to those who had at least 1 year of medical history (i.e., 1 year time difference between the first and the last medical record before COVID-19 infection date) with Mass General Brigham (MGB).
Replication	exact replication is not possible due to fact that the data cannot be shared publicly.
Randomization	Describe how samples/organisms/participants were allocated into experimental groups. If allocation was not random, describe how covariates were controlled OR if this is not relevant to your study, explain why.
Blinding	Describe whether the investigators were blinded to group allocation during data collection and/or analysis. If blinding was not possible, describe why OR explain why blinding was not relevant to your study.
Reportin	g for specific materials, systems and methods
	on from authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, ted 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.

iviateriais & experimental systems	IVIE	etnoas
n/a Involved in the study	n/a	Involved in the study
Antibodies		ChIP-seq
Eukaryotic cell lines		Flow cytometry
Palaeontology and archaeology		MRI-based neuroimaging
Animals and other organisms		•
Human research participants		
Clinical data		
Dual use research of concern		
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Antibodies

Antibodies used

Describe all antibodies used in the study; as applicable, provide supplier name, catalog number, clone name, and lot number.

Validation

Describe the validation of each primary antibody for the species and application, noting any validation statements on the manufacturer's website, relevant citations, antibody profiles in online databases, or data provided in the manuscript.

Eukaryotic cell lines

Policy information about <u>cell lines</u>	
Cell line source(s)	State the source of each cell line used.
Authentication	Describe the authentication procedures for each cell line used OR declare that none of the cell lines used were authenticated.
Mycoplasma contamination	Confirm that all cell lines tested negative for mycoplasma contamination OR describe the results of the testing for mycoplasma contamination OR declare that the cell lines were not tested for mycoplasma contamination.
Commonly misidentified lines (See <u>ICLAC</u> register)	Name any commonly misidentified cell lines used in the study and provide a rationale for their use.

Palaeontology and Archaeology

Specimen provenance

Provide provenance information for specimens and describe permits that were obtained for the work (including the name of the issuing authority, the date of issue, and any identifying information).

Specimen deposition

Indicate where the specimens have been deposited to permit free access by other researchers.

Dating methods

If new dates are provided, describe how they were obtained (e.g. collection, storage, sample pretreatment and measurement), where they were obtained (i.e. lab name), the calibration program and the protocol for quality assurance OR state that no new dates are provided.

Tick this box to confirm that the raw and calibrated dates are available in the paper or in Supplementary Information.

Ethics oversight

Identify the organization(s) that approved or provided guidance on the study protocol, OR state that no ethical approval or guidance was required and explain why not.

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

Animals and other organisms

Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research

Laboratory animals

For laboratory animals, report species, strain, sex and age OR state that the study did not involve laboratory animals.

Wild animals

Provide details on animals observed in or captured in the field; report species, sex and age where possible. Describe how animals were caught and transported and what happened to captive animals after the study (if killed, explain why and describe method; if released, say where and when) OR state that the study did not involve wild animals.

Field-collected samples

For laboratory work with field-collected samples, describe all relevant parameters such as housing, maintenance, temperature, photoperiod and end-of-experiment protocol OR state that the study did not involve samples collected from the field.

Ethics oversight

Identify the organization(s) that approved or provided guidance on the study protocol, OR state that no ethical approval or guidance was required and explain why not.

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

Human research participants

Policy information about studies involving human research participants

Population characteristics

Describe the covariate-relevant population characteristics of the human research participants (e.g. age, gender, genotypic information, past and current diagnosis and treatment categories). If you filled out the behavioural & social sciences study design questions and have nothing to add here, write "See above."

Recruitment

Describe how participants were recruited. Outline any potential self-selection bias or other biases that may be present and how these are likely to impact results.

Ethics oversight

Identify the organization(s) that approved the study protocol.

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

Clinical data

Policy information about clinical studies

All manuscripts should comply with the ICMJE guidelines for publication of clinical research and a completed CONSORT checklist must be included with all submissions.

Clinical trial registration

Provide the trial registration number from ClinicalTrials.gov or an equivalent agency.

Study protocol

2020P001063

Data collection

16,709 patients with a confirmed case for COVID-19 (confirmed polymerase chain reaction test) between 03/03/2020 and 10/11/2020 with Mass General Brigham (MGB).

Outcomes

death events recorded in the electronic medical records.

Dual use research of concern

Policy information about <u>dual use research of concern</u>

Hazards

Could the accidental, deli in the manuscript, pose a	perate or reckless misuse of agents or technologies generated in the work, or the application of information presented threat to:		
No Yes			
Public health			
National security			
Crops and/or livest	ock		
Ecosystems			
Any other signification	nt area		
Experiments of concer	n		
Does the work involve an	y of these experiments of concern:		
No Yes			
Demonstrate how	to render a vaccine ineffective		
Confer resistance t	o therapeutically useful antibiotics or antiviral agents		
Enhance the virule	nce of a pathogen or render a nonpathogen virulent		
Increase transmissi	bility of a pathogen		
Alter the host rang	e of a pathogen		
Enable evasion of o	liagnostic/detection modalities		
Enable the weapor	ization of a biological agent or toxin		
Any other potentia	lly harmful combination of experiments and agents		
ChIP-seg			
CHIF-SEQ			
Data deposition			
Confirm that both raw	and final processed data have been deposited in a public database such as GEO.		
Confirm that you have	deposited or provided access to graph files (e.g. BED files) for the called peaks.		
Data access links May remain private before public	For "Initial submission" or "Revised version" documents, provide reviewer access links. For your "Final submission" document, provide a link to the deposited data.		
Files in detaless submissi	on Provide a list of all files available in the database submission.		
Files in database submiss	on Provide a list of all files available in the database submission.		
Genome browser session (e.g. <u>UCSC</u>)	Provide a link to an anonymized genome browser session for "Initial submission" and "Revised version" documents only, to enable peer review. Write "no longer applicable" for "Final submission" documents.		
Methodology			
Replicates	Describe the experimental replicates, specifying number, type and replicate agreement.		
Sequencing depth	Describe the sequencing depth for each experiment, providing the total number of reads, uniquely mapped reads, length of reads and whether they were paired- or single-end.		
Antibodies	Describe the antibodies used for the ChIP-seq experiments; as applicable, provide supplier name, catalog number, clone name, and lot number.		
Peak calling parameters	Specify the command line program and parameters used for read mapping and peak calling, including the ChIP, control and index files used.		
Data quality	Describe the methods used to ensure data quality in full detail, including how many peaks are at FDR 5% and above 5-fold enrichment.		
Software			

Flow Cytometry				
Plots Confirm that:				
	and fluorochrome used (e.g. CD4-FITC).			
	. Include numbers along axes only for bottom left plot of group (a 'group' is an analysis of identical markers).			
All plots are contour plots with c	outliers or pseudocolor plots.			
A numerical value for number of	f cells or percentage (with statistics) is provided.			
Methodology				
Sample preparation De	scribe the sample preparation, detailing the biological source of the cells and any tissue processing steps used.			
Instrument	entify the instrument used for data collection, specifying make and model number.			
	scribe the software used to collect and analyze the flow cytometry data. For custom code that has been deposited into a mmunity repository, provide accession details.			
	scribe the abundance of the relevant cell populations within post-sort fractions, providing details on the purity of the mples and how it was determined.			
0 07	cribe the gating strategy used for all relevant experiments, specifying the preliminary FSC/SSC gates of the starting cell ulation, indicating where boundaries between "positive" and "negative" staining cell populations are defined.			
Tick this box to confirm that a fig	gure exemplifying the gating strategy is provided in the Supplementary Information.			
Magnetic resonance ima	aging			
Experimental design				
Design type	Indicate task or resting state; event-related or block design.			
Design specifications	Specify the number of blocks, trials or experimental units per session and/or subject, and specify the length of each trial or block (if trials are blocked) and interval between trials.			
Behavioral performance measures	State number and/or type of variables recorded (e.g. correct button press, response time) and what statistics were used to establish that the subjects were performing the task as expected (e.g. mean, range, and/or standard deviation across subjects).			
Acquisition				
Imaging type(s)	Specify: functional, structural, diffusion, perfusion.			
Field strength	Specify in Tesla			
Sequence & imaging parameters	Specify the pulse sequence type (gradient echo, spin echo, etc.), imaging type (EPI, spiral, etc.), field of view, matrix size, slice thickness, orientation and TE/TR/flip angle.			
Area of acquisition	State whether a whole brain scan was used OR define the area of acquisition, describing how the region was determined.			
Diffusion MRI Used	☐ Not used			

Preprocessing

Normalization

Normalization template

Noise and artifact removal

Preprocessing software Provide detail on software version and revision number and on specific parameters (model/functions, brain extraction, segmentation, smoothing kernel size, etc.).

If data were normalized/standardized, describe the approach(es): specify linear or non-linear and define image types used for transformation OR indicate that data were not normalized and explain rationale for lack of normalization.

Describe the template used for normalization/transformation, specifying subject space or group standardized space (e.g. original Talairach, MNI305, ICBM152) OR indicate that the data were not normalized.

Congress of the second second

Describe your procedure(s) for artifact and structured noise removal, specifying motion parameters, tissue signals and physiological signals (heart rate, respiration).

Volume censoring

Define your software and/or method and criteria for volume censoring, and state the extent of such censoring.

Statistica	I mode	ling 8	& in	ference
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datistical modeling & intere	nec .					
Model type and settings	Specify type (mass univariate, multivariate, RSA, predictive, etc.) and describe essential details of the model at the first and second levels (e.g. fixed, random or mixed effects; drift or auto-correlation).					
Effect(s) tested	Define precise effect in terms of the task or stimulus conditions instead of psychological concepts and indicate whether ANOVA or factorial designs were used.					
Specify type of analysis: W	hole brain ROI-based Both					
Statistic type for inference (See Eklund et al. 2016)	Specify voxel-wise or cluster-wise and report all relevant parameters for cluster-wise methods.					
Correction	Describe the type of correction and how it is obtained for multiple comparisons (e.g. FWE, FDR, permutation or Monte Carlo).					
Models & analysis n/a Involved in the study						
Functional and/or effective conn	Report the measures of dependence used and the model details (e.g. Pearson correlation, partial correlation, mutual information).					
Graph analysis	Report the dependent variable and connectivity measure, specifying weighted graph or binarized graph, subject- or group-level, and the global and/or node summaries used (e.g. clustering coefficient, efficiency, etc.).					

Multivariate modeling and predictive analysis

Specify independent variables, features extraction and dimension reduction, model, training and evaluation metrics.