nature portfolio

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Last updated by author(s):	February 1, 2024

Reporting Summary

Nature Portfolio 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 Portfolio policies, see our <u>Editorial Policies</u> and the <u>Editorial Policy Checklist</u>.

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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.
\boxtimes	A description of all covariates tested
\boxtimes	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>
\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
	Estimates of effect sizes (e.g. Cohen's <i>d</i> , Pearson's <i>r</i>), indicating how they were calculated
	Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.
So	ftware and code
Poli	cy information about <u>availability of computer code</u>
Di	ata collection No software was used for data collection

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 Portfolio guidelines for submitting code & software for further information.

Data

Data analysis

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

https://github.com/rayanramin/APOBEC3B-UPDSeq

- A description of any restrictions on data availability
- For clinical datasets or third party data, please ensure that the statement adheres to our policy

The raw sequence files generated in this study are deposited at NCBI-SRA BioProject PRJNA1005650 (https://www.ncbi.nlm.nih.gov/bioproject/?term=1005650). It is publicly available and there are no restrictions on these data.

Research inv	volving hu	man participants, their data, or biological material
		vith human participants or human data. See also policy information about sex, gender (identity/presentation), thnicity and racism.
Reporting on sex	and gender	N/A
Reporting on rac other socially rele groupings	• • • • • • • • • • • • • • • • • • • •	N/A
Population chara	cteristics	N/A
Recruitment		N/A
Ethics oversight		N/A
Note that full informa	ation on the appr	oval of the study protocol must also be provided in the manuscript.
Field spe	oific ro	norting
Field-spe		
Please select the o	ne below that is	s the best fit for your research. If you are not sure, read the appropriate sections before making your selection.
X Life sciences	□ B	ehavioural & social sciences Ecological, evolutionary & environmental sciences
For a reference copy of	the document with	all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>
Life scier	nces stu	udy design
All studies must dis	sclose on these	points even when the disclosure is negative.
Sample size	A minimum of t	three biological replicates were included in each experiment.
Data exclusions	None.	
Replication	All replicates ar	e biological replicates, not technical replicates.
Randomization	N/A	
Blinding	N/A	
Reportin	g for si	pecific materials, systems and methods
We require informati	on from authors	about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, your study. If you are not sure if a list item applies to your research, read the appropriate section before selecting a response.
Materials & ex	perimental s	ystems Methods
n/a Involved in th	•	n/a Involved in the study
Antibodies Sukaryotic		ChIP-seq
	logy and archaeol	logy
	nd other organism	
Clinical dat		
Dual use re	esearch of concer	n
Plants		

Antibodies

Antibodies used

Rabbit anti-APOBEC3B monoclonal antibody, 1:500 or 1:2,000 dilution as indicated, NIH AIDS Reagent Program, catalog number 5210-87-13.

Validation

A3B-CTD expressed in E. coli and purified to homogeneity.

Eukaryotic cell lin	es	
Policy information about <u>ce</u>	Il lines	and Sex and Gender in Research
Cell line source(s)		N/A
Authentication		N/A
Mycoplasma contaminati	on	N/A
Commonly misidentified (See <u>ICLAC</u> register)	lines	N/A
Palaeontology and	d Arc	chaeology
Specimen provenance	N/A	
Specimen deposition	N/A	
Dating methods	N/A	
Tick this box to confirm that the raw and calibrated dates are available in the paper or in Supplementary Information.		
Ethics oversight	sight N/A	
Note that full information on the	ne appro	oval of the study protocol must also be provided in the manuscript.
Animals and othe	r res	earch organisms
Policy information about <u>studies involving animals</u> ; <u>ARRIVE guidelines</u> recommended for reporting animal research, and <u>Sex and Gender in Research</u>		
Laboratory animals	N/A	
Wild animals	N/A	
Reporting on sex	N/A	
Field-collected samples	N/A	
Ethics oversight	N/A	
Note that full information on the approval of the study protocol must also be provided in the manuscript.		
Clinical data		
Policy information about <u>clinical studies</u> All manuscripts should comply with the ICMJE <u>guidelines</u> for <u>publication</u> of <u>clinical research</u> and a completed <u>CONSORT checklist</u> must be included with all submissions.		
Clinical trial registration	N/A	
Study protocol	N/A	
Data collection	N/A	

Dual use research of concern

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

N/A

Hazards

Outcomes

Could the accidental, deliberate or reckless misuse of agents or technologies generated in the work, or the application of information presented in the manuscript, pose a threat to:

No Yes				
Public health				
National security	- al.			
Crops and/or lives	OCK			
Any other significa	nt area			
Experiments of concer				
Does the work involve ar		periments of concern:		
No Yes	y or these exp	Actimients of confection		
Demonstrate how	o render a vac	cine ineffective		
	o therapeutica	lly useful antibiotics or antiviral agents		
Enhance the virule	nce of a patho	gen or render a nonpathogen virulent		
Increase transmiss				
Alter the host rang				
Enable evasion of Enable the weapo		iction modalities Plogical agent or toxin		
1		nbination of experiments and agents		
Z , p	.,			
Plants				
FIGITES				
Seed stocks	N/A			
Novel plant genotypes	N/A			
, , ,				
Authentication	N/A			
-1				
ChIP-seq				
Data deposition				
Confirm that both rav	and final pro	ocessed data have been deposited in a public database such as <u>GEO</u> .		
Confirm that you have	deposited o	r provided access to graph files (e.g. BED files) for the called peaks.		
Data access links May remain private before publi	N/A N/A			
Files in database submiss	on N/A			
Genome browser session	N/A			
Methodology				
Replicates	N/A			
Sequencing depth	N/A			
Antibodies	N/A			
Peak calling parameters	N/A			
Data quality	N/A			
Software	N/A			

Flow Cytometry		
Plots		
Confirm that:		
The axis labels state the mar	ker and fluorochrome used (e.g. CD4-FITC).	
The axis scales are clearly vis	sible. Include numbers along axes only for bottom left plot of group (a 'group' is an analysis of identical markers).	
	ith outliers or pseudocolor plots.	
A numerical value for number	er of cells or percentage (with statistics) is provided.	
Methodology		
Sample preparation	N/A	
Instrument	N/A	
Software	N/A	
Cell population abundance	N/A	
Gating strategy	N/A	
Tick this box to confirm that	a figure exemplifying the gating strategy is provided in the Supplementary Information.	
Magnetic resonance in	maging	
Experimental design		
Design type	N/A	
Design specifications	N/A	
Behavioral performance measur	res N/A	
Acquisition		
Imaging type(s)	N/A	
Field strength	N/A	
Sequence & imaging parameters	s N/A	
Area of acquisition	N/A	
Diffusion MRI Used	Not used	
Preprocessing		
Preprocessing software	N/A	
Normalization	N/A	
Normalization template	N/A	
Noise and artifact removal	N/A	
Volume censoring	N/A	

Statistical modeling & inference

Model type and settings	N/A
Effect(s) tested	N/A
Specify type of analysis: W	hole brain ROI-based Both

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Š	2	
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N		

Statistic type for inference	N/A
(See Eklund et al. 2016)	
Correction	N/A
Models & analysis n/a Involved in the study	