nature research

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Last updated by author(s):	Sep 21, 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 ali	aryses, commit that the following items are present in the right e 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 stateme	ent on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly		
	The statis Only comm	tical 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	cion of all covariates tested		
	🗶 A descript	cion of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons		
	A full desc AND varia	cription of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) tion (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 es as exact values whenever suitable.		
×	For Bayes	ian analysis, information on the choice of priors and Markov chain Monte Carlo settings		
X	For hierar	chical 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 an	d code		
Policy information about <u>availability of computer code</u>				
Da	ata collection	RNA-Seq analysis was performed after ribodepletion and standard library construction using Illumina HiSeq2500 V4 2x100 PE - performed by Genewiz, South Plainfield, NJ.		

Data analysis

MetaCore (v20.2) was used for functional enrichment analysis after RNA-Seq.

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 <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
- A description of any restrictions on data availability

All relevant data are available from the authors. The source data underlying Figs 1i, 3a–f, 4d-f, 5f, and Supplementary Figs 4h, 5b-c, g, h, 6f, and 7e are provided as a Source Data file with this paper. The RNA seq data is accessible at: https://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSE155842

Field-specific reporting

Please select the one below	that is the best fit for	your research. If you a	re not sure, read the	appropriate sections b	efore 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

Describe how sample size was determined, detailing any statistical methods used to predetermine sample size OR if no sample-size calculation was performed, describe how sample sizes were chosen and provide a rationale for why these sample sizes are sufficient.

Data exclusions

Describe any data exclusions. If no data were excluded from the analyses, state so OR if data were excluded, describe the exclusions and the rationale behind them, indicating whether exclusion criteria were pre-established.

Replication

Describe the measures taken to verify the reproducibility of the experimental findings. If all attempts at replication were successful, confirm this OR if there are any findings that were not replicated or cannot be reproduced, note this and describe why.

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.

Behavioural & social sciences study design

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

Briefly describe the study type including whether data are quantitative, qualitative, or mixed-methods (e.g. qualitative cross-sectional, quantitative experimental, mixed-methods case study).

Research sample

State the research sample (e.g. Harvard university undergraduates, villagers in rural India) and provide relevant demographic information (e.g. age, sex) and indicate whether the sample is representative. Provide a rationale for the study sample chosen. For studies involving existing datasets, please describe the dataset and source.

Sampling strategy

Describe the sampling procedure (e.g. random, snowball, stratified, convenience). Describe the statistical methods that were used to predetermine sample size OR if no sample-size calculation was performed, describe how sample sizes were chosen and provide a rationale for why these sample sizes are sufficient. For qualitative data, please indicate whether data saturation was considered, and what criteria were used to decide that no further sampling was needed.

Provide details about the data collection procedure, including the instruments or devices used to record the data (e.g. pen and paper, computer, eye tracker, video or audio equipment) whether anyone was present besides the participant(s) and the researcher, and whether the researcher was blind to experimental condition and/or the study hypothesis during data collection.

Indicate the start and stop dates of data collection. If there is a gap between collection periods, state the dates for each sample cohort.

If no data were excluded from the analyses, state so OR if data were excluded, provide the exact number of exclusions and the rationale behind them, indicating whether exclusion criteria were pre-established.

State how many participants dropped out/declined participation and the reason(s) given OR provide response rate OR state that no participants dropped out/declined participation.

If participants were not allocated into experimental groups, state so OR describe how participants were allocated to groups, and if allocation was not random, describe how covariates were controlled.

Ecological, evolutionary & environmental sciences study design

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

Study description

Study description

Data collection

Data exclusions

Non-participation

Randomization

Timing

Briefly describe the study. For quantitative data include treatment factors and interactions, design structure (e.g. factorial, nested, hierarchical), nature and number of experimental units and replicates.

Research sample	Describe the research sample (e.g. a group of tagged Passer domesticus, all Stenocereus thurberi within Organ Pipe Cactus National Monument), and provide a rationale for the sample choice. When relevant, describe the organism taxa, source, sex, age range and any manipulations. State what population the sample is meant to represent when applicable. For studies involving existing datasets, describe the data and its source.				
Sampling strategy	Note the sampling procedure. Describe the statistical methods that were used to predetermine sample size OR if no sample-size calculation was performed, describe how sample sizes were chosen and provide a rationale for why these sample sizes are sufficient.				
Data collection	Describe the data collection procedure, including who recorded the data and how.				
Timing and spatial scale	Indicate the start and stop dates of data collection, noting the frequency and periodicity of sampling and providing a rationale for these choices. If there is a gap between collection periods, state the dates for each sample cohort. Specify the spatial scale from which the data are taken				
Data exclusions	If no data were excluded from the analyses, state so OR if data were excluded, describe the exclusions and the rationale behind them, indicating whether exclusion criteria were pre-established.				
Reproducibility	Describe the measures taken to verify the reproducibility of experimental findings. For each experiment, note whether any attempts to repeat the experiment failed OR state that all attempts to repeat the experiment were successful.				
Randomization	Describe how samples/organisms/participants were allocated into groups. If allocation was not random, describe how covariates were controlled. If this is not relevant to your study, explain why.				
Blinding	Describe the extent of blinding used during data acquisition and analysis. If blinding was not possible, describe why OR explain why blinding was not relevant to your study.				
Did the study involve field work? Yes No Field work, collection and transport					
Field conditions	Describe the study conditions for field work, providing relevant parameters (e.g. temperature, rainfall).				
Location	State the location of the sampling or experiment, providing relevant parameters (e.g. latitude and longitude, elevation, water depth).				
Access & import/export	Describe the efforts you have made to access habitats and to collect and import/export your samples in a responsible manner and in compliance with local, national and international laws, noting any permits that were obtained (give the name of the issuing authority, the date of issue, and any identifying information).				
Disturbance	Describe any disturbance caused by the study and how it was minimized.				

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		Methods		
Involved in the study	n/a	Involved in the study		
x Antibodies	x	ChIP-seq		
x Eukaryotic cell lines	×	Flow cytometry		
Palaeontology and archaeology	x	MRI-based neuroimaging		
🗶 Animals and other organisms				
Human research participants				
Clinical data				
Dual use research of concern				
	Involved in the study X Antibodies Eukaryotic cell lines Palaeontology and archaeology Animals and other organisms Human research participants Clinical data	Involved in the study X Antibodies X Eukaryotic cell lines Palaeontology and archaeology X Animals and other organisms Human research participants Clinical data		

Antibodies

Antibodies used

rat anti-Mac2, (Cedarlane, CL8942AP, 1:100), anti-Mac3 (BD Pharmingen, 553322, 1:900), rabbit anti-cleaved caspase-3 (Cell Signaling, 9664, 1:400) rabbit anti-HuR/ELAVL1 (12582S Cell Signalling and AB242410, Abcam), biotin goat anti-MerTK (BAF591, R&D Technologies); antibodies were also used against Flag Tag (Cell Signaling, 2368, 1:1000), GAPDH (Cell Signaling, 2118, 1:4000), β-actin (Cell Signaling, #4970, 1:3000), cleaved Caspase-3 (Cell Signalling, 1:1000), cleaved Caspase-8 (Cell Signalling, 1:2000), HuR (12582S, Cell Signalling, 1:3000),

Validation

All antibodies were used per commercial sources and chosen based upon available validation studies for immunohistochemistry or Western analyses.

Eukaryotic cell lines

Policy information about cell lines

Cell line source(s)

Primary bone marrow cells were used from mice and RAW264.7 macrophages were obtained from commercial sources (ATCC)

Authentication Per ATCC for RAW264.7 macrophages.

Mycoplasma contamination Cell lins were free of mycoplasma.

Commonly misidentified lines (See ICLAC register)

n/a

Animals and other organisms

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

Laboratory animals C57BL/6 and LDLR-/- mice(Jackson Laboratory)

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

All protocols concerning animal use were approved by the Institutional Animal Care and Use Committee at Brigham and Women's Hospital and Harvard Medical School, Boston, MA and conducted in accordance with the National Institutes of Health Guide for the Care and Use of Laboratory Animals.

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