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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 our Editorial Policies Editorial Policy Checklist

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For	all statistical an	alyses, 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			
	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)			
x	For null hypothesis testing, the test statistic (e.g. F t r P ,,) with conGive P values as exact values whenever suitable.			
×	For Bayesi	an analysis, information on the choice of priors and Markov chain Monte Carlo settings		
x	For hierard	chical and complex designs, identification of the appropriate level for tests and full reporting of outcomes		
\blacksquare Estimates of effect sizes (e.g. Cohen's $d r$				
Our web collection on <u>statistics for biologists</u>				
So	ftware and	d code		
Poli	cy information a	about <u>availability of computer code</u>		
D	ata collection	Zen 2.0 (Carl Zeiss) software Multi-user software package (FLUOstar® Omega) software BD CellQuest™ Pro software SATURNO software (Crony instruments)		
D	ata analysis	Fiji (NIH) software FlowJo™ v10.0.8 (Tree Star) software GraphPad Prism 6.0 software Microsoft Office 2010 Photoshop PSCS6		

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

Data

Policy information about availability of data

All manuscripts must include a data availability statement

- 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 data generated or analyzed during this study are included in this published article. All relevant data are also available upon request from the corresponding

authors.		
Field-spe	ecific 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.	
x Life sciences	Behavioural & social sciences Ecological, evolutionary & environmental sciences	
For a reference copy of t	the document with all sections, see nature.com/documents/nr-reporting-summary-flat.pdf	
Life scier	nces study design	
	close on these points even when the disclosure is negative.	
Sample size	The sample size of animal studies was chosen taking into account previous works in this area.	
	Lopez-Gil et al, 2013. doi:10.1371/journal.pntd.0002309 Busquets et al, 2010. DOI: 10.1089=vbz.2009.0205	
	Borel et al, 2005. doi: 10.1007/s11259-005-0844-0.	
Data exclusions	No data have been excluded from the analyses	
Replication	All the analyses have been replicated	
Randomization	Animals were allocated in experimental groups randomly	
Blinding	The scientists were blinded during data collection or analyses	
Behaviou	ıral & social sciences study design	
	close on these points even when the disclosure is negative.	
Study description		
	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.

Data collection

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.

Timing

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

Data exclusions

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.

Non-participation

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.

Randomization

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

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, collect	tion 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.		
· · · · · · · · · · · · · · · · · · ·	r specific materials, systems and methods		
•	authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, evant 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 & experime	ental systems Methods		
n/a Involved in the study Antibodies	n/a Involved in the study 		

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

Antibodies

Antibodies used

Anti sheep IgG Alexa fluor 488 cat number A11015 lot 687630 (Thermofisher) Anti Mouse IgG alexa fluor 596 Cat number A11032 lot 2048170 (Thermofisher)

CD8a-PerCP-Vio700, Clone 53-6.7 mouse. Order no: 130-120-756 lot 5190507445 Miltenyi CD107a (LAMP-1)-FITC, Clone H4A3 mouse. Order no: 130-102-191 lot 5180607338 Miltenyi Anti-IFN-y-PE, mouse. Clone AN.18.17.24. Order no: 130-102-388 lot 5180958661 Miltenyi

Goat anti-mouse immunoglobulins-HRP from Dako (P0447)

Validation

Sheep antisera raised against RVFV from previous work (Lorenzo et al 2018) was used as primary antibody in protein expression analysis by fluorescence and immunohistochemistry

Mouse antisera raised against BTV from previous experiments (Marín-López et al, 2018) used as primary antibody in protein expression analysis by fluorescence

Eukaryotic cell lines

Policy information about cell lines

Cell line source(s) Vero cells (ATCC; catalog no. CCL-81)

BHK-21 cells (ATCC; catalog no. CCL-10)

Chicken embryo fibroblasts (DF-1) (ATCC; catalog no. CRL-12203)

Authentication Authentication of cell lines by STR profiling

Mycoplasma contamination All cell lines tested negative for mycoplasma

Commonly misidentified lines (See ICLAC

No misidentified cell lines have been used

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.

Type I interferon receptor defective mice ,IFNAR (-/-) on a 129 Sv/Ev background, female, 8 weeks old

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

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BALB/c mice, female, 8 weeks old

Sheep (Spanish Churra sheep breed) , female, aged two years

Wild animals

Laboratory 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

Animal experimental protocols were approved by the Ethical Review Committee at the INIA-CISA and Comunidad de Madrid (Permit number: PROEX 037/15) in strict accordance with EU guidelines 2010/63/UE about protection of animals used for experimentation

	and other scientific purposes and Spanish Animal Welfare Act 32/2007.	
Note that full information on tl	he approval of the study protocol must also be provided in the manuscript.	
Human research p	participants	
	udies 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 tl	he approval of the study protocol must also be provided in the manuscript.	
Clinical data		
Policy information about <u>cli</u> All manuscripts should comply	inical studies with the ICMJEguidelines for publication of clinical research CONSORT checklist	
Clinical trial registration	Provide the trial registration number from ClinicalTrials.gov or an equivalent agency.	
Study protocol	Note where the full trial protocol can be accessed OR if not available, explain why.	
Data collection	Describe the settings and locales of data collection, noting the time periods of recruitment and data collection.	
Outcomes	Describe how you pre-defined primary and secondary outcome measures and how you assessed these measures.	
Dual use research	of concern	
Policy information about <u>du</u>	ual use research of concern	
Hazards		
Could the accidental, deli in the manuscript, pose a	berate or reckless misuse of agents or technologies generated in the work, or the application of information presented	
No Yes		
Public health		
National security		
Crops and/or livest	rock	
x Ecosystems		
Any other significant area		

Experiments of concern

Doe	s the work involve any of these experiments of concern:
No	Yes
x	Demonstrate how to render a vaccine ineffective
x	Confer resistance to therapeutically useful antibiotics or antiviral agents
x	Enhance the virulence of a pathogen or render a nonpathogen virulent

Increase transmissibility of a pathogen

Alter the host range of a pathogen

Enable evasion of diagnostic/detection modalities

Enable the weaponization of a biological agent or toxin

Any other potentially harmful combination of experiments and agents

ChIP-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 publication.

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 database submission

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

Describe the software used to collect and analyze the ChIP-seq data. For custom code that has been deposited into a community repository, provide accession details.

Flow Cytometry

Plots

Confirm that:

- $m{x}$ The axis labels state the marker and fluorochrome used (e.g. CD4-FITC).
- The axis scales are clearly visible. 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 outliers or pseudocolor plots.
- A numerical value for number of cells or percentage (with statistics) is provided.

Methodology

Cell population abundance

 Sample preparation
 Splenocytes were collected from mice and ammonium chloride was used to lyse red blood cells before flow cytometry assays

 Instrument
 FACSCalibur (Becton Dickinson)

 Software
 BD CellQuest™ Pro was used to collect data Flow Jo V10 .0.7 was used to analyse data

Live cells were counted after isolation of splenocytes. Abundance of lymphocytes from mouse spleen is usually 75-80%, and percentage of CD8+cells is aproximately 20%

Gating strategy

Analyses were done using FlowJo software. Discrimination of lymphocytes is based on scatter parameters (FSC/SSC). Within scatter parameters, the pulse height versus pulse width plots was used to isolate single cells. After gating lymphocyte population, CD8+ cells were selected by staining with anti-mouse CD8 PerCP-Vio700 antibody. For determination of CD8+IFN-

gamma+ or CD8+CD107a+ using IFN-γ-PE and CD107a/LAMP-1-FITC antibodies.

Tick this box to confirm that a figure exemplifying the gating strategy is provided in the Supplementary Information.

Magnetic resonance imaging

Graph analysis

Magnetic resonance in	TAPILIP				
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 measure	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					
Preprocessing software	Provide detail on software version and revision number and on specific parameters (model/functions, brain extraction, segmentation, smoothing kernel size, etc.).				
Normalization If data were normalized/standardized, describe the approach(es): specify linear or non-linear and define image types used 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.				
Noise and artifact removal	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.				
Statistical modeling & inferen	nce				
	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: Wh	nole brain ROI-based Both				
Statistic type for inference (See <u>Eklund et al. 2016</u>	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 Graph analysis Multivariate modeling or pr					
Functional and/or effective conn	Report the measures of dependence used and the model details (e.g. Pearson correlation, partial correlation,				

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.