nature portfolio

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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 Editorial Policies and the Editorial Policy Checklist.

Please do not complete any field with "not applicable" or n/a. Refer to the help text for what text to use if an item is not relevant to your study. For final submission: please carefully check your responses for accuracy; you will not be able to make changes later.

For all statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.

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n/a	Confirmed		
	The exact	sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement	
	X A stateme	ent on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly	
	The statist	tical test(s) used AND whether they are one- or two-sided on tests should be described solely by name; describe more complex techniques in the Methods section.	
\times	A descript	ion of all covariates tested	
	A descript	ion 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 Give <i>P</i> values as exact values whenever suitable.		
\times	For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings		
\times	For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes		
\times	\square Estimates of effect sizes (e.g. Cohen's d , Pearson's r), indicating how they were calculated		
	Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.		
Software and code			
Poli	cy information a	about availability of computer code	
Da	ita collection	We used CRISPR direct [https://crispr.dbcls.jp], PhosphoSitePlus (ver. 6.5.9.3) [https://www.phosphosite.org/homeAction], RegPhos2.0 [http://140.138.144.141/~RegPhos/index.php], and the Universal Protein Resource (Uniprot) [https://www.uniprot.org].	
Da	ita analysis	We used BIO_CYCLE (ver. 0.9.3), ShinyGO v0.61 [http://bioinformatics.sdstate.edu/go61/], Proteome Discoverer 2.2, and MATHEMATICA 12.0.	
For m	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		

Data

Policy information about availability of data

All manuscripts must include a data availability statement. This statement should provide the following information, where applicable:

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.

- Accession codes, unique identifiers, or web links for publicly available datasets
- 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 RNA-seq data obtained in this study have been deposited in the Gene Expression Omnibus (GEO) under accession code GSE199061 [https://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSE199061]. The proteomics data obtained in this study have been deposited in PRIDE with the dataset identifier PXD035414 [http://www.ebi.ac.uk/pride/archive/projects/PXD035414]. Also, the processed RNA-seq data and proteomics data are provided in the Supplementary Data file. The codes and readme file for simulation of the mathematical models are available in Zenodo database under accession code [https://doi.org/10.5281/zenodo.6512360]. The codes are implemented by MATHEMATICA 12.0.

Field-specific reporting

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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 <u>nature.com/documents/nr-reporting-summary-flat.pd</u> f		
Life scier	nces study design	
All studies must dis	close on these points even when the disclosure is negative.	
Sample size	We did not perform sample size test. We determined sample sizes based on related studies: Hirano et al., Cell, 2013, Terajima et al., Nat. Genet., 2017, and Imamura et al., Proc. Natl. Acad. Sci. U. S. A., 2018.	
Data exclusions	No data were excluded from analyses.	
Replication	The experiments were performed at least three times.	
Randomization	For animal and cellular experiments, experimental groups were based on genotypes. For individual experiments, animals and cell plates were randomly selected for each time point.	

Behavioural & social sciences study design

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

Study description

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

this study. Analyses of temporal profiles of cells and mouse tissues were performed blindly.

quantitative experimental, mixed-methods case study).

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.

Investigators were not blinded during experiments, because no subjective process is included in all the analyses of the experimental data in

cradical in a second distribution of the desired and second distribution of the desired and second distribution of the desired and second distribution of the desired dist

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 Indicat

Blinding

Research sample

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

Data exclusions If no data were exclude

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

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.

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

Briefly describe the study. For quantitative data include t.

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

Randomization

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 fie	ld work? Yes No	
Field work collec	etion and transport	
Field work, conec	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	5 & import/export Describe the efforts you have made to access habitats and to collect and import/export your samples in a responsible manner a compliance with local, national and international laws, noting any permits that were obtained (give the name of the issuing autitied date of issue, and any identifying information).	
Disturbance	urbance Describe any disturbance caused by the study and how it was minimized.	
Ve require information from	or 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. ental systems Methods	
n/a Involved in the study	<u> </u>	
Antibodies	ChIP-seq	
Eukaryotic cell lines		
Palaeontology and		
☐ ☐ Animals and other ☐ Human research pa		
Clinical data		
Dual use research o	of concern	
Antibodies		
	anti CLOCKY2 (CLCD2, D222, 2). Madical 9. Dialogical Laboratorias), anti ADNIT 22 (D10112, D22F, 2). Madical 9. Dialogical Laboratorias)	
Antibodies used	anti-CLOCK23 (CLSP3; D333-3; Medical & Biological Laboratories), anti-ARNTL23 (B1BH2; D335-3; Medical & Biological Laboratories), anti-PER1 (PM091; Medical & Biological Laboratories), anti-PER2 (PM083; Medical & Biological Laboratories), anti-CRY1 (PM081; Medical & Biological Laboratories), anti-DBP (PM079; Medical & Biological Laboratories), anti-NR1D1 (PM092; Medical & Biological Laboratories), and anti-ATF2 (C-19; sc-187; Santa Cruz Biotechnology).	
All the antibodies used were validated by the manufacturers and checked by us by Western blot. Validation statements on the manufacturer's website for all primary antibodies used in this study were as follows: anti-CLOCK (CLSP3; D333-3; Medical & Biological Laboratories): mouse (+) and human (+) anti-ARNTL (B1BH2; D335-3; Medical & Biological Laboratories): mouse (+) and human (+) anti-PER1 (PM091; Medical & Biological Laboratories): mouse (+) and human (-) anti-CRY1 (PM081; Medical & Biological Laboratories): mouse (+) and human (+)		

anti-CRY2 (PM082; Medical & Biological Laboratories): mouse (+) and human (+) anti-DBP (PM079; Medical & Biological Laboratories): mouse (+) anti-NR1D1 (PM092; Medical & Biological Laboratories): mouse (+), rat (+) and human (+) and anti-ATF2 (C-19; sc-187; Santa Cruz Biotechnology): Xenopus (+), mouse (+), rat (+), dog (+), human (+)

Eukaryotic cell lines

Policy information about cell lines

Cell line source(s)

NIH3T3 cells were obtained from RIKEN Cell Bank.

Authentication

The cell line was not authenticated.

Mycoplasma contamination

We confirmed that the cell line was negative for mycoplasma.

Commonly misidentified lines (See ICLAC register)

No commonly misidentified cell lines were 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). Permits should encompass collection and, where applicable, export.

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

Male C57BL/6J mice aged 5-24 weeks were used. Bmal1- Δ RRE mice were generated in this study. WT mice were obtained from Kiwa Laboratory Animals (Wakayama, Japan). PER2::LUC mice originate from mice generated by Yoo et al. (Proc Natl Acad Sci U S A., 2004,101(15):5339-46).

The mice were maintained in a light-tight chamber at a constant temperature (23 \pm 1 °C), humidity (55 \pm 10%), and 12-hr light/12-hr dark (LD) cycles.

Wild animals

No wild animals were used in this study.

Field-collected samples

No field-collected samples were used in this study.

Ethics oversight

All the animal experiments were conducted according to the animal ethics guideline of the University of Tokyo.

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 <u>cli</u> All manuscripts should comply	nical studies 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	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 du	ual use research of concern	
Hazards		
in the manuscript, pose a No Yes Public health National security Crops and/or livest Ecosystems Any other significa Experiments of concer Does the work involve an No Yes Demonstrate how	ock nt area n y of these experiments of concern: to render a vaccine ineffective	
Confer resistance to therapeutically useful antibiotics or antiviral agents 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		
	and final processed data have been deposited in a public database such as <u>GEO</u> . 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 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.

Describe the sequencing depth for each experiment, providing the total number of reads, uniquely mapped reads, length of reads and Sequencing depth

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Sequencing depth	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	scribe the software used to collect and analyze the ChIP-seq data. For custom code that has been deposited into a community ository, provide accession details.	
Flow Cytometry		
Plots		
Confirm that:		
The axis labels state the	ne marker and fluorochrome used (e.g. CD4-FITC).	
The axis scales are cle	arly visible. Include numbers along axes only for bottom left plot of group (a 'group' is an analysis of identical markers).	
All plots are contour p	olots with outliers or pseudocolor plots.	
A numerical value for	number of cells or percentage (with statistics) is provided.	
Methodology		
Sample preparation	Describe the sample preparation, detailing the biological source of the cells and any tissue processing steps used.	
Instrument	Identify the instrument used for data collection, specifying make and model number.	
Software	Describe the software used to collect and analyze the flow cytometry data. For custom code that has been deposited into a community repository, provide accession details.	
Cell population abundance	Describe the abundance of the relevant cell populations within post-sort fractions, providing details on the purity of the samples and how it was determined.	
Gating strategy	Describe the gating strategy used for all relevant experiments, specifying the preliminary FSC/SSC gates of the starting cell population, indicating where boundaries between "positive" and "negative" staining cell populations are defined.	
Tick this box to confirm	m that a figure exemplifying the gating strategy is provided in the Supplementary Information.	
Magnetic resonar	nce imaging	
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 r	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 para	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 for transformation OR indicate that data were not normalized and explain rationale for lack of normalization.
Normalization template	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

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

physiological signals (heart rate, respiration).

Volume censoring

Graph analysis

statistical modeling & inferen	nce	
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	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: WI	nole 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 connectivity Graph analysis Multivariate modeling or predictive analysis		
Functional and/or effective conne	Report the measures of dependence used and the model details (e.g. Pearson correlation, partial correlation, mutual information).	

Multivariate modeling and predictive analysis

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

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,

