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Last updated by author(s): Apr 28, 2019

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 <u>Authors & Referees</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							
$oxed{\boxtimes}$ 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.							
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)							
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.							
For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings							
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.							
Software and code							
Policy information about <u>availability of computer code</u>							
Data collection The softwares to collect the data: Bio-rad CFX Manager 3.1; SoftMax Pro6.5.1; Zeiss ZEN 2012.							
Data analysis The softwares to analyse the data: SPSS23.0; Graphpad Prism 6 v6; Image J v18.0; Image Pro Plus 6.0;							
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/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 data availability statement. 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

Gene expression RNA-seq data has been deposited at the NCBI Gene Expression Omnibus (GEO). The accession number is GSE130147 [https://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSE130147]. Data supporting the findings of this study are available within the article and its Supplementary Information files. All relevant data are available from the authors on reasonable request.

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Blinding

Randomization

Please select the one 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 t	For a reference copy of the document with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>						
Life scier	ices study design						
All studies must dis	close on these points even when the disclosure is negative.						
Sample size	We used standard sample sizes reported in the literature previously in mouse studies. The sample size of animal experiments according to similar studies in the field was chose and at least 3 mice per group was used. The number of the independent experiments for cell and biological experiments was indicated in each figure legend.						
Data exclusions	No data were excluded from the analyses.						
Replication	All experiments were performed independently multiple times using biologically independent replicates.						
Randomization	Mice were randomly assigned to groups.						

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, quantitative experimental, mixed-methods case study).

State the research sample (e.g. Harvard university undergraduates, villagers in rural India) and provide relevant demographic information Research sample (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.

The investigators were blinded to group allocation during data collection and analysis. We collected and analyzed the compared samples

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, Data collection 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 cohort.

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.

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

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

Describe the research sample (e.g. a group of tagged Passer domesticus, all Stenocereus thurberi within Organ Pipe Cactus National Research sample 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	ndicate the start and stop dates of data collection, noting the frequency and periodicity of sampling and providing a rationale for hese 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	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 and import/expor	Describe the efforts you have made to access habitats and to collect and import/export your samples in a responsible manner 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.

Ma	terials & experimental systems	Methods			
n/a	Involved in the study	n/a	Involved in the study		
	Antibodies	\boxtimes	ChIP-seq		
	Eukaryotic cell lines	\boxtimes	Flow cytometry		
\boxtimes	Palaeontology	\boxtimes	MRI-based neuroimaging		
	Animals and other organisms				
	Human research participants				
\boxtimes	Clinical data				

Antibodies

Antibodies used

HuR, Millipore, #07-468, rabbit polyclonal antibody, KLH-conjugated, synthetic peptide corresponding to amino acids 1-13 of human HuR, LOT2880766;

ATGL, Abcam, ab109251, EPR3444(2), LOT GR234927-10;

Perilipin-1, Abcam, ab172907, EPR3753(2), LOT GR145311-7;

Akt, cell signaling technology, catalog number 4691, C67E7, LOT 20;

p-Akt(ser473), cell signaling technology, catalog number 4060, D9E, LOT 23;

F4/80, Abcam, ab16911, BM8, Rat monoclonal antibody, LOT GR3243707-3;

HSL,cell signaling technology, catalog number 4107, LOT 3;

ADRB1, Abcam, ab3442, rabbit polyclonal antibody, LOT GR3216445-4;

ADRB2, Abcam, ab182136,EPR707(N) rabbit monoclonal antibody, LOT GR302897-12;

ADRB3, Abclonal, A8607, rabbit polyclonal antibody, LOT1152520201;

β-actin, Proteintech, 66009-1, 2D4H5, LOT 10004156;

GAPDH, Proteintech, 60004-1,1E6D9, LOT 10004129.

Validation HuR for western blot, immunohistochemistry, immunofluroscence, RNA-immnoprecipitation. F4/80 for immunofluroscence of mice adipose tissue.

ATGL, HSL, Perilipin-1, Akt, p-Akt(ser473), ADRB1, ADRB2, ADRB3, β-actin, GAPDH for western blot.

The antibodies are from commercial sources and have been validated by the vendors.

Eukaryotic cell lines

Policy information about cell lines

Cell line source(s) The 3T3-L1 MBX cell line (ATCC CRL-3242) was purchased form ATCC.

Authentication All of the cell lines were authenticated by short tandem repeat (STR)-profiling.

Mycoplasma contamination All cell lines tested negative for mycoplasma contamination.

Commonly misidentified lines (See <u>ICLAC</u> register)

No commonly misidentified cell lines are used in the study.

Palaeontology

Laboratory animals

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

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

Animals and other organisms

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

The animals we used were all male mice of eight weeks. The strains included HuR flox/flox:AdipoQ-Cre-, HuR flox/flox:AdipoQ-Cre+,Ob/ob and C57BL/6J. Mice were housed on a 12-h light/dark cycle and given ad libitum access to food and water. All animal protocols were approved by the Institutional Animal Care and Use Committee of Cheeloo College of Medicine, Shandong

Offiversit

Wild animals The study did not involve wild animals.

Field-collected samples The study did not involve field-collected samples.

Ethics oversight Institutional Animal Care and Use Committee of Cheeloo College of Medicine, Shandong University

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

Human research participants

Population characteristics

Recruitment

Policy information about studies involving human research participants

oney information about <u>studies involving number research participants</u>

Human tissue biopsies from subcutaneous adipose tissue were obtained from the Qilu Hospital of Shandong University during surgery. All subjects provided their written informed consent. All procedures that involved human samples were approved by the Ethics Committee of Qilu Hospital of Shandong University.

Ethics Committee of Qilu Hospital of Shandong University

These patients were recruited while in hospital awaiting surgery. Obese individuals were defined as those with a BMI ≥30 kg m-2. The gender and age was not considered in the analysis and only BMI. There were no obvious study selection biases that

would have influenced the results except for the sample size, thereby increasing variability.

Ethics oversight Ethics Committee of Qilu Hospital of Shandong University

Note that full information on the approval of the study protocol must also be provided in the manuscript. $\frac{1}{2} \int_{\mathbb{R}^{n}} \left(\frac{1}{2} \int_{\mathbb{R}^{$

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 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.

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., UCSC)

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 experimen

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:

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

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 the	nat a figure exemplifying the gating strategy is provided in the Supplementary Information.							
Magnetic resonance	e 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 mea	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 parameter	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 Use	d Not used							
reprocessing								
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 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 & infe	erence							
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:								
Specify type of analysis: Statistic type for inference (See Eklund et al. 2016)	ANOVA or factorial designs were used.							

Models & analysis

*10 G	cis a ariarysis	
n/a	Involved in the study	
	Functional and/or effective connectivity	
	Graph analysis	
	Multivariate modeling or predictive analysis	
Functional and/or effective connectivity		Report the measures of dependence used and the model details (e.g. Pearson correlation, partial correlation, mutual information).
Gra	oh 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.).
Mul	tivariate modeling and predictive analysis	Specify independent variables, features extraction and dimension reduction, model, training and evaluation metrics.