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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 Authors & Referees and the Editorial Policy Checklist.

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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.
	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 <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
\boxtimes	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

Policy information about availability of computer code

Data collection

No computer code were used to collect the data.

Data analysis

Reads that mapped to rRNA fasta sequences from the UCSC gene annotation (mm10) using bowtie2 (v.2.2.6) were discarded, and remaining reads were aligned to mm10 using hisat2-align-s (v.2.0.5). Unique reads with high mapping quality were retained using Picard (http://broadinstitute.github.io/picard, v.2.16.0) and SAMtools (v.0.1.09). Differential gene expression between WT and METTL14 R255K were analyzed by DESeq2 using default parameters. For mRNA lifetime RNA-seq data analysis, FPKM of gene were calculated, and the lifetime was calculated as reported in Wang, X. et. al, Nature, 2014. Regions of significant meRIP enrichment relative to input control ("peaks") were identified using MACS2. The parameter '--nomodel' was used for macs2 peak calling. Macs2 bdgdiff was used to identify differential peaks. The enrichment of biological process was conducted using Kobas. Student's t test was carried out using GraphPad Prism 8 version 8.3.0.

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
- A description of any restrictions on data availability

MeRIP—seq data have been deposited in GEO under accession numbers GSE164047. The mass spectrometry data has been deposited in integrated proteome resources (iProX) under accession number IPX0002127000 and ProteomeXchange Consortium under ID PXD018458. All data supporting the findings of this study are available from the corresponding author on reasonable request.

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Please select the one below	v 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.pdf</u>

Life sciences study design

Sample size

Study description

Sampling strategy

Data collection

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

No sample size calculation was performed. Key experiments were repeated in another cell line or another knockout or knock-in clone. Triplicates were performed for key experiments. Number of independent experiments was indicated in the figure legend. Three independent

results were used to perform statistical analysis. If less, no statistical analysis were performed.

Data exclusions No data were excluded from the analyses.

The replicate times of western blots, micrographs, qPCR or LC-MS/MS were shown in the figure legends. For western blots and micrographs, Replication similar results were obtained and a representative figure were shown. The replicates of qPCR and LC/MS-MS were shown in the graph. All

attempts at replication were successful.

Randomization No randomization techniques was used, because the study was based on cellular biology techniques, and no organisms were involved.

Blinding The investigators were not blinded to allocation during experiments and outcome assessment, because the cells were processed in the same way.

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.

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.

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 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 Randomization 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

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	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	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/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
n/a Involved in the study	n/a Involved in the study
Antibodies	ChIP-seq
Eukaryotic cell lines	Flow cytometry
Palaeontology	MRI-based neuroimaging
Animals and other organisms	,
Human research participants	
Clinical data	

Antibodies

Antibodies used

METTL14 (Sigma, HPA038002), WB 1:1000, IP 1:50 PRMT1 (CST, 2449 (A33)), WB 1:1000, IP 1:50 rabbit IgG (Santa, SC-2027), IP 1:100

m6A antibody (Synaptic Systems, 202003), dot blot 1:3000

m6A antibodies (Abcam, ab151230), 1.5 μg per MeRIP-qpcr210 μg per MeRIP-seq

WTAP (Proteintech, 60188-1-Ig), WB 1:1000 METTL3 (Bethyl, A301-567A), WB 1:3000

mono and dimethyl arginine (Abcam, ab412), WB 1:1000

mono-methyl arginine (CST, 8015S), WB 1:1000 GAPDH (Proteintech, 60004-1-lg), WB 1:10000

FLAG (MBL, M185-3L), WB 1:3000

HA (MBL, M180-3), WB 1:3000 MYC (MBL, M192-3), WB 1:1000

Anti-Mouse IgG (H+L) Antibody, Human Serum Adsorbed and Peroxidase-Labeled (KPL, 5220-0341), WB 1;5000

Anti-Rabbit IgG (H+L) Antibody, Peroxidase-Labeled (KPL, 5220-0336), WB 1:5000

VeriBlot for IP Detection Reagent (HRP) (Abcam, ab131366), WB 1:200 Dylight 800 Goat Anti-Rabbit IgG (A23920, Abbkine), WB 1:10000 Dylight 800 and Goat Anti-Mouse IgG (A23910, Abbkine), WB 1:10000

WTAP(Abcam, ab195380 (EPR18744)), WB 1:1000

Validation

All of the antibodies used in this study were commercial. The antibodies were validated based on the information from the manufacturer's instructions and were supported by multiple publications.

Eukaryotic cell lines

Policy information about cell lines

Cell line source(s)

The E14TG2a cells and HEK293T cells were from the National Collection of Authenticated Cell Cultures. The HeLa cells were from Dr. Qinmiao Sun (Institute of Zoology, Chinese Academy of Sciences), and the CGR8 cell were from Dr. Shoujun Huang (Anhui Agriculture University), and published before.

Authentication

The cell line was authenticated by Short Tandem Repeat (STR) profiling analysis.

Mycoplasma contamination

Mycoplasma contamination was negative by routine check.

Commonly misidentified lines (See ICLAC register)

No commonly misidentified cell lines were used.

Palaeontology

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 dates are provided.

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

Laboratory animals

For laboratory animals, report species, strain, sex and age OR state that the study did not involve laboratory animals.

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

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.

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.

olicy information about <u>clinic</u> Il manuscripts should comply wit	al studies h the ICMJE <u>guidelines for publication of clinical research</u> and a completed <u>CONSORT checklist</u> must be included with all submissior		
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.		
	Describe how you pre-defined primary and secondary outcome measures and how you assessed these measures.		
Outcomes ChIP-seq ata deposition	bescribe now you pre-defined primary and secondary outcome measures and now you assessed these measures.		
hIP-seq ata deposition Confirm that both raw ar	nd final processed data have been deposited in a public database such as GEO.		
hIP-seq ata deposition Confirm that both raw ar	nd final processed data have been deposited in a public database such as GEO. eposited or provided access to graph files (e.g. BED files) for the called peaks. For "Initial submission" or "Revised version" documents, provide reviewer access links. For your "Final submission" documents		
hIP-seq ata deposition Confirm that both raw ar Confirm that you have de	nd final processed data have been deposited in a public database such as GEO. eposited or provided access to graph files (e.g. BED files) for the called peaks. For "Initial submission" or "Revised version" documents, provide reviewer access links. For your "Final submission" documents		

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: 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

Software

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.

> 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.

	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.		
	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 that	a figure exemplifying the gating strategy is provided in the Supplementary Information.		
Magnetic resonance in	maging		
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 measur	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 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 & infere	ence		
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: W	hole 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).		

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