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Reporting Summary

X Life sciences

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>.

Statistics	
For all statistical analy	rses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.
n/a Confirmed	
☐ ☐ The exact sa	mple 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 Only common	al test(s) used AND whether they are one- or two-sided tests should be described solely by name; describe more complex techniques in the Methods section.
A description	n of all covariates tested
A description	n of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons
	otion of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) on (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)
	othesis testing, the test statistic (e.g. F , t , r) with confidence intervals, effect sizes, degrees of freedom and P value noted as exact values whenever suitable.
For Bayesian	analysis, information on the choice of priors and Markov chain Monte Carlo settings
For hierarch	ical and complex designs, identification of the appropriate level for tests and full reporting of outcomes
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 abo	out <u>availability of computer code</u>
Data collection	Provide a description of all commercial, open source and custom code used to collect the data in this study, specifying the version used OR state that no software was used.
Data analysis	Provide a description of all commercial, open source and custom code used to analyse the data in this study, specifying the version used OR state that no software was used.
	stom algorithms or software that are central to the research but not yet described in published literature, software must be made available to editors/reviewers. e deposition in a community repository (e.g. GitHub). See the Nature Research guidelines for submitting code & software for further information.
Data	
All manuscripts mus - Accession codes, u - A list of figures tha	out <u>availability of data</u> t include a <u>data availability statement</u> . This statement should provide the following information, where applicable: nique identifiers, or web links for publicly available datasets t have associated raw data y restrictions on data availability
The data that support the	he findings of this study are available from the corresponding author upon reasonable request.
Field-spec	rific reporting

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.

Ecological, evolutionary & environmental sciences

Behavioural & social sciences

Life sciences study design

Materials & experimental systems

Sample size	Sample size of three (n=3, distinct samples) was used for all in-solution, in-vitro and in-vivo toxicity studies. Sample size of five (n=5, distict samples) was used for in-vitro biodistribution and in-vivo tumor regression studies.
Data exclusions	No data excluded
Replication	Describe the measures taken to verify the reproducibility of the experimental findings. If all attempts at replication were successful, confirm this OR if there are any findings that were not replicated or cannot be reproduced, note this and describe why.
Randomization	For each groups, cells/animals were allocated/chosen at random.
Blinding	Describe whether the investigators were blinded to group allocation during data collection and/or analysis. If blinding was not possible, describe why OR explain why blinding was not relevant to your study.

Reporting for specific materials, systems and methods

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.

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 organi	sms
Human research participa	ants
Clinical data	
Antibodies	
Antibodies used	Describe all antibodies used in the study; as applicable, provide supplier name, catalog number, clone name, and lot number.
Validation	Describe the validation of each primary antibody for the species and application, noting any validation statements on the manufacturer's website, relevant citations, antibody profiles in online databases, or data provided in the manuscript.
Eukaryotic cell lines	
Policy information about cell line	<u>es</u>
Cell line source(s)	HeLa and Cal27 cells were obtained from ATCC.
Authentication	The cell lines are authenticated by ATCC and they were used directly.
Mycoplasma contamination	All cells lines were tested for mycoplasma contamination using the mycoAlert, mycoplasma detection kit by Roche. They were confirmed mycoplasma free before use.
Commonly misidentified lines (See ICLAC register)	No misidentified cell lines were used in the study

Palaeontology

Dating methods

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.

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 stu	dies involving animals; ARRIVE guidelines recommended for reporting animal research
Laboratory animals	Balb/C nude mice, aged 5- 6 weeks were used for the study
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.

Singapore and Institutional Animal Care and Use Committee Singapore Health Services Pte Ltd.

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."

Ethical approval and guidance was obtained from Institutional Animal Care and Use Committee (IACUC), National University of

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

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 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.

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

Study protocol

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. <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

Sequencing depth	(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

Briefly, the cells treated with NIR laser were trypsinized and filtered through a 40 µm cell strainer to get a single cell suspension.

The cells were then centrifuged and re-suspended at a density of 1 × 106 cells/mL in the pre-warmed RPMI-1640 medium

containing 2% FBS. The cells were obtained from ATCC as reported in the Methods section.

Instrument Cytoflex LX (Beckman Coulter)

Software Cytexpert

Cell population abundance After gating for viable single cell events, approximately 95-98% of the sub-population was included for fluorescence analysis.

Gating strategy

Initially forward and side scatter were used to gate the viable cell population and doublets were excluded by forward scatter height vs forward scatter area. Finally single parameter histograms were used to measure fluorescence intensity of the target cells

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

Magnetic resonance imaging

Behavioral performance measures

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.

or block (If trials are blocked) and interval between trial

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, MN/305, 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 & inference

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: Whole 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.

 $\textit{Describe the type of correction and how it is obtained for multiple comparisons (e.g. FWE, FDR, permutation or Monte to the type of correction and how it is obtained for multiple comparisons (e.g. FWE, FDR, permutation or Monte to the type of correction and how it is obtained for multiple comparisons (e.g. FWE, FDR, permutation or Monte to the type of correction and how it is obtained for multiple comparisons (e.g. FWE, FDR, permutation or Monte to the type of correction and how it is obtained for multiple comparisons (e.g. FWE, FDR, permutation or Monte to the type of correction and how it is obtained for multiple comparisons (e.g. FWE, FDR, permutation or Monte to the type of correction and type of correction and the type of correctio$

Models & analysis

Correction

n/a	Involved in the study
\boxtimes	Functional and/or effective connectivity
X	Graph analysis
\boxtimes	Multivariate modeling or predictive analysis