# 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 <u>Editorial Policies</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
	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.
X	A description of all covariates tested
X	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>
$\times$	For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings
X	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 <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 availability of computer code

Data collection

FluorEssence V3.5, Light Cycler 480 II (Roche), PatchControl, Patch Master, NAMD 2.12, SoftMax Pro (Molecular Devices), Zetasizer Nano-ZS (Malvern Instruments), Gaussian 16, Gaussview 6.0

Data analysis

OriginPro 2016, ImageJ, ChemDraw 20.1.1, Clampfit 10.6.1, VMD (Visual Molecular Dynamics) 1.9.4, MestReNova V9.0.1

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 Portfolio 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 description of any restrictions on data availability
- For clinical datasets or third party data, please ensure that the statement adheres to our policy

All the databases/datasets used in the study, along with appropriately accessible links/accession codes are mentioned in the manuscript under the "Data availability" section as well as in this reporting summary. For example: PDB code 1XAV, optimized structures of TBP2 and TBP2 along with cations and anions, initial and final configurations of MD simulations etc.

Policy information ab and sexual orientatio		vith human participants or human data. See also policy information about sex, gender (identity/presentation), thnicity and racism.		
Reporting on sex ar	nd gender	Not Applicable		
Reporting on race, ethnicity, or other socially relevant groupings		Not Applicable		
Population characteristics		Not Applicable		
Recruitment		Not Applicable		
Ethics oversight		Not Applicable		
Note that full information	on on the appro	oval of the study protocol must also be provided in the manuscript.		
Field-spec	cific re	porting		
Please select the one	below that is	s the best fit for your research. If you are not sure, read the appropriate sections before making your selection.		
Life sciences	Ве	ehavioural & social sciences		
For a reference copy of the	e document with a	all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>		
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## **Antibodies**

Antibodies used

Plants

Clinical data

Dual use research of concern

Anti-DNA G-quadruplex Antibody, clone 1H6 mouse monoclonal [(Merck, USA, Catalog no. MABE1126)(1:200 dilution in 1X PBS)], and Alexafluor 647-conjugated secondary antibody [(Invitrogen, Catalog no. A56576) (1:1000 dilution in 1X PBS)]

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 <u>cell lines and Sex and Gender in Research</u>

Cell line source(s)

HeLa, K562, A549, NKE and HEK293T cell lines were procured from NCCS, Pune, India

Authentication

Cell lines were authenticated by NCCS, Pune, India. We have not performed further authentication of the commercially available cell lines.

Mycoplasma contamination

Cell lines were certified as testing negative for mycoplasma contamination by the supplier. We have not performed further mycoplasma testing.

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

No commonly misidentified cell lines were used in the study.

## 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,

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

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 research organisms

Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research, and Sex and Gender in Research

Laboratory animals

For laboratory animals, report species, strain 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 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.

Reporting on sex

Indicate if findings apply to only one sex; describe whether sex was considered in study design, methods used for assigning sex. Provide data disaggregated for sex where this information has been collected in the source data as appropriate; provide overall numbers in this Reporting Summary. Please state if this information has not been collected. Report sex-based analyses where performed, justify reasons for lack of sex-based analysis.

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.

#### 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	Ollection 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 researd	h of concern		
Policy information about <u>c</u>	dual use research of concern		
Hazards			
Could the accidental, de in the manuscript, pose	liberate or reckless misuse of agents or technologies generated in the work, or the application of information presented a threat to:		
No Yes			
Public health			
National security  Crops and/or live:	stack		
Ecosystems	NOCK		
Any other signific	ant area		
Experiments of conce	ern		
Does the work involve a	ny of these experiments of concern:		
No Yes			
Demonstrate how	v to render a vaccine ineffective		
	to therapeutically useful antibiotics or antiviral agents		
- -	ence of a pathogen or render a nonpathogen virulent		
	rsibility of a pathogen		
	diagnostic/detection modalities		
	onization of a biological agent or toxin		
	ially harmful combination of experiments and agents		
Plants			
Fidills			
Seed stocks	Report on the source of all seed stocks or other plant material used. If applicable, state the seed stock centre and catalogue number. If plant specimens were collected from the field, describe the collection location, date and sampling procedures.		
Novel plant genotypes	Describe the methods by which all novel plant genotypes were produced. This includes those generated by transgenic approaches, gene editing, chemical/radiation-based mutagenesis and hybridization. For transgenic lines, describe the transformation method, the number of independent lines analyzed and the generation upon which experiments were performed. For gene-edited lines, describe the editor used, the endogenous sequence targeted for editing, the targeting guide RNA sequence (if applicable) and how the editor		
Authentication	was applied.  Describe any authentication procedures for each seed stock used or novel genotype generated. Describe any experiments used to assess the effect of a mutation and, where applicable, how potential secondary effects (e.g. second site T-DNA insertions, mosiacism, off-target gene editing) were examined.		
ChIP-seq			
'			
Data deposition  Confirm that both ra	w and final processed data have been deposited in a public database such as <u>GEO</u> .		
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 pub	For "Initial submission" or "Revised version" documents, provide reviewer access links. For your "Final submission" document, provide a link to the deposited data.		

Provide a list of all files available in the database submission.

Files in 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 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 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

## Flow Cytometry

#### **Plots**

Confirm that:	
The axis labels state the m	arker 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 num	ber 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 that	at 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.

repository, provide accession details.

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 (ij thats are blocked) and interval between thats

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.		nctional, structural, diffusion, perfusion.
Field strength Specify in		Tesla
		e pulse sequence type (gradient echo, spin echo, etc.), imaging type (EPI, spiral, etc.), field of view, matrix size, ness, orientation and TE/TR/flip angle.
Area of acquisition	State when	ther a whole brain scan was used OR define the area of acquisition, describing how the region was determined.
Diffusion MRI Used	☐ Not u	sed
Preprocessing		
Preprocessing software		n software version and revision number and on specific parameters (model/functions, brain extraction, 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		mplate used for normalization/transformation, specifying subject space or group standardized space (e.g. ch, 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 soft	ware 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: W	hole brain	ROI-based Both
Statistic type for inference	Specify voxel-wi	se or cluster-wise and report all relevant parameters for cluster-wise methods.
(See Eklund et al. 2016)		
Correction	Describe the typ	be 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 conn	ectivity	Report the measures of dependence used and the model details (e.g. Pearson correlation, partial correlation, mutual information).
Graph 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.).
Multivariate modeling and predi	ctive analysis	Specify independent variables, features extraction and dimension reduction, model, training and evaluation metrics.