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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 <u>Authors & Referees</u> and the <u>Editorial Policy Checklist</u>.

When statistical analyses are reported, confirm that the following items are present in the relevant location (e.g. figure legend, table legend, main

Statistical parameters

text	, or I	Methods section).
n/a	Cor	nfirmed
		The $\underline{\text{exact sample size}}(n)$ for each experimental group/condition, given as a discrete number and unit of measurement
		An indication of 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 statistics including <u>central tendency</u> (e.g. means) or other basic estimates (e.g. regression coefficient) AND <u>variation</u> (e.g. standard deviation) or associated <u>estimates of uncertainty</u> (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
		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 d, Pearson's r), indicating how they were calculated
		Clearly defined error bars State explicitly what error bars represent (e.g. SD SE CI)

Our web collection on <u>statistics for biologists</u> may be useful.

Software and code

Policy information about availability of computer code

Data collection

The data were collected at a core facility using the standard Illumina Hi-seq software and pipeline.

Standard bioinformatics software was used to analyze the next-gen sequencing data. The details are provided in the methods section. A supplementary file is included with the submission that contains custom code for estimating cell doublet removal.

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 upon request. 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

Next-generation sequencing data are available at the Gene Expression Omnibus under accession number GSE125971.

Field-specific reporting						
Please select 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	he document with all sections, see <u>nature.com/authors/policies/ReportingSummary-flat.pdf</u>					
Life scier	nces study design					
All studies must dis	close on these points even when the disclosure is negative.					
Sample size	Cell numbers selected for the bulk- and single-cell analyses are typical for the field.					
Data exclusions	Some libraries were filtered from the single-cell data to remove cell doublets. This procedure is described in the Methods section of the manuscript.					
Replication	All bulk-cell samples were performed in biological replicate, and all results were highly consistent.					
Randomization	Randomization was not necessary for testing the ACT-seq method.					
Blinding	Blinding was not necessary for testing the ACT-seq method.					
Materials & experimental systems n/a Involved in the study Unique biological materials Antibodies Palaeontology Animals and other organisms Human research participants Methods n/a Involved in the study ChIP-seq Flow cytometry MRI-based neuroimaging						
Unique biological materials						
Policy information about <u>availability of materials</u>						
Obtaining unique materials The Tn5-PA vector was deposited to Addgene and is freely available.						
Antibodies						
Antibodies used	Validated antibodies were used from commercial sources. The details are in the Methods section of the manuscript.					
Validation	Commercial validation was performed by the respective companies. See the antibody catalog numbers in the Methods section.					
Eukaryotic cell lines						
Policy information	about <u>cell lines</u>					
Cell line source(s)	HEK293T cells were ordered from ATCC.					
Authentication	Cells have been authenticated in many experiments by transcript analysis and genome sequencing.					

The cell line was not tested for mycoplasma contamination.

No such cell lines were used.

Mycoplasma contamination

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