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Last updated by author(s):	YYYY-MM-DD

## **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>.

Stat	tistics					
For al	l statistical analys	es, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.				
n/a	Confirmed					
	The exact sam	nple size $(n)$ for each experimental group/condition, given as a discrete number and unit of measurement				
	A statement of	on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly				
	The statistical  Only common t	test(s) used AND whether they are one- or two-sided ests 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>					
	For Bayesian	analysis, information on the choice of priors and Markov chain Monte Carlo settings				
	For hierarchic	al and complex designs, identification of the appropriate level for tests and full reporting of outcomes				
	Estimates of 6	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 <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 OF state that no software was used.				
Data analysis		Data analysis was performed with R version 3.4.0 (2017-04-21) using base functions and the ggplot package. Perl scripts were run using Perl v5.18.2. The perl script for predicting off target sites is provided in the supplementary information associated with the paper.				
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.						
Dat	а					
All m - 4 - 4	nanuscripts must Accession codes, un A list of figures that	ut <u>availability of data</u> include a <u>data availability statement</u> . This statement should provide the following information, where applicable: ique identifiers, or web links for publicly available datasets have associated raw data restrictions on data availability				
All data generated and analyzed during this study are included in the published article or provided in the supplementary information.						
Fie	eld-speci	fic 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

all studies must disclose on these points even when the disclosure is negative.		
Sample size	No sample-size calculations were performed. The sample sizes were judged to be sufficient based on the reproducibility of measurements between groups.	
Data exclusions	There were no data exclusions.	
Replication	Multiple replicate experiments were performed to verify the conclusions. Typically, a minimum of 3 replicate experiments were performed.	
Randomization	Randomization was not part of the study.	
Blinding	Blinding was not part of the study.	

## 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		
$\boxtimes$	Antibodies	$\boxtimes$	ChIP-seq		
$\boxtimes$	Eukaryotic cell lines	$\boxtimes$	Flow cytometry		
$\boxtimes$	Palaeontology	$\boxtimes$	MRI-based neuroimaging		
$\boxtimes$	Animals and other organisms				
$\boxtimes$	Human research participants				
$\boxtimes$	Clinical data				