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Last updated by author(s): May 4, 2020

## **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.

For all statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section

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101	an statistical analyses, commit that the following items are present in the figure regend, thair text, or interious 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.
$\boxtimes$	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

### Software and code

Policy information about availability of computer code

Data collection MaxQuant (version 1.5.0.35)

R version 3.6.1 (2019-07-05) Data analysis

pheatmap\_1.0.12 gridSVG 1.7-1 Biostrings\_2.54.0 XVector\_0.26.0 IRanges\_2.20.1

S4Vectors\_0.24.0 RColorBrewer\_1.1-2 hwriter\_1.3.2 RBDmap\_0.0.17 XML\_3.98-1.20

limma\_3.42.0 Biobase\_2.46.0 BiocGenerics\_0.32.0 VennDiagram\_1.6.20 futile.logger\_1.4.3 ggrepel\_0.8.1

ggplot2\_3.2.1

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/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.

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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	fany restrictions on data availability
	oroteomic data is deposited to the ProteomeXchange Consortium via the PRIDE partner repository with the dataset identifier PXD009789. nerated in 15 and was re-analyzed here to illustrate the analytical workflow (Figure 3 and 5).
Field-spe	ecific reporting
Please select the o	ne below 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>
	sclose on these points even when the disclosure is negative.  3 biological replicates with 3 experimental conditions each
·	
Data exclusions	No data excluded.
Replication	Experiments were performed from 3 biological replicates and in the primary papers candidate proteins were functionally tested
Randomization	Not applicable
Blinding	Not applicable

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

### **Antibodies**

Antibodies used

β-Actin (Sigma-Aldrich, A1978)

• Histone H4 (Abcam, ab10158)

• Polypyrimidine tract binding protein (PTBP1) (Sigma, WH0005725M1)

• Cold shock domain-containing protein E1 (CSDE1)/UNR (Proteintech, 13319–1-AP)

• Non-POU domain-containing octamer-binding protein (NonO) (Novus Biologicals, NBP1–95977)

• ELAV-like protein 1 (ELAVL1)/Hu-antigen R (HuR) (Proteintech, 11910–1-AP)

Validation

Antibodies are commercially available and validated by provider

### Eukaryotic cell lines

Policy information about <u>cell lines</u>

Cell line source(s)

HeLa cells (ATCC, cat. no. CCL-2)

Jurkat cells (DSM7. ACC-282)

Jurkat cells (DSMZ, ACC-282) HEK293 (ECACC, #85120602)

Authentication Cell lines authenticated by supplier

Mycoplasma contamination All lines are tested regularly and are negative

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

None