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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>.

Statistical parameters

When statistical analyses are reported	, confirm that the following items are	present in the relevant	location (e.g. figur	re legend, table	legend, mair
text, or Methods section).					

n/a	Confirmed
	\times The 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.
\boxtimes	A description of all covariates tested
\boxtimes	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 Give P values as exact values whenever suitable.
\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
	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

Confocal images were acquired using Zeiss Axio Observer Z1 microscope (equipped with an Andor camera and a spinning-disk confocal scan head, FV10-ASW (version 2.0, for Olympus FV1200) and Zen (version 11.0.0.190, for Zeiss LSM700 and LSM880).

3D-SIM images were acquired on the DeltaVision OMX V3 imaging system (Applied Precision, GE). SIM image stacks were reconstructed using softWoRx (version 5.0, Applied Precision).

Some images in Figure S5 and S7 were acquired with Axio Imager M2 microscope (Carl Zeiss).

Data analysis

Microsoft Excel and GraphPad Prism (version: 5.0) were used for analyzing and graphing.

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 <u>availability of data</u>

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

The quantification analyses in all figures have associated raw data. The raw data are available from the corresponding author upon reasonable request.		
Field-spe	cific reporting	
Please select the b	est fit for your research. If you are not sure, read the appropriate sections before making your selection.	
☐ Behavioural & social sciences ☐ Ecological, evolutionary & environmental sciences		
For a reference copy of	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	The sample size and the statistical method are indicated in the figure legends. There are no specific criteria for sample size in this study.	
Data exclusions	No data were excluded from the analysis.	
Replication	All attempts at replication were successful.	
Randomization	This is not relevant to our study.	
Blinding	The investigators were blinded to group allocations.	
Reporting for specific materials, systems and methods		
Materials & expe	erimental systems Methods	
n/a Involved in th		
Unique biological materials ChIP-seq		
Antibodies Flow cytometry		
Eukaryotic cell lines MRI-based neuroimaging Palaeontology		
Animals and other organisms		
Human res	earch participants	
Antibodies		
Antibodies used	Mouse Anti-HA mAb(Zsbio #TA-04),rabbit anti-calreticulin (Abcam #ab2907)	
Validation Mouse Anti-HA mAb(Zsbio #TA-04) http://www.zsbio.com/product/TA-04 rabbit anti-calreticulin (Abcam #ab2907) https://www.abcam.com/calreticulin-antibody-ab2907-references.html		

Eukaryotic cell lines

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

Cell line source(s)

COS-7 cells were obtained from ATCC (Cat#ATCC CRL-1651, RRID: CVCL_0224).

Authentication	None of the cell lines were authenticated.
Mycoplasma contamination	All cell lines were tested negative for mycoplasma contamination.
Commonly misidentified lines (See <u>ICLAC</u> register)	None

Animals and other organisms

Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research		
Laboratory animals	C. elegans	
Wild animals	We do not use wild animals in our study.	
Field-collected samples	We do not use Field-collected samples in our study.	