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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 our <u>Editorial Policies</u> and the <u>Editorial Policy Checklist</u>.

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For all statistical an	alyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.				
n/a Confirmed					
☐ ☐ The exact	The exact sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement				
A stateme	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.				
A descript	A description of all covariates tested				
A descript	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 Give <i>P</i> values as exact values whenever suitable.					
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				
·	Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.				
Software an	d code				
Policy information	about <u>availability of computer code</u>				
Data collection	NMR data collection: Topspin 4 Xray data collection: PXII beamline software at the Swiss Light Source				
Data analysis	NMR data processing: Topspin 4 and NMRPipe/ NMRDraw version 9.6 NMR data analysis: NMRFAM-SPARKY 1.2 powered by Sparky 3.115, CARA 1.9.1.8a2 and NMRViewJ 9.2.0-b24 NMR structure calculations: Cyana 3.98.13 X-ray structure calculations: XDS 2021, CCP4 7.1, Phaser 2.8.3, Coot 0.9.6 and Phenix 1.18.2-3874 Data fitting: Matlab R2021a Visualization: ChimeraX 1.1 Sequence analysis: ClustalX 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 and 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.

Data

Policy information about <u>availability of data</u>

Dual use research of concern

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 structure factors and coordinates for the Xrn2 core protein have been deposited in the Protein Data Bank under accession code 70PK. The assignments for the Xrn2 zinc finger have been deposited in the Biological Magnetic Resonance Bank under accession number 50997. The atomic coordinates for the Xrn2 zinc finger have been deposited in the Protein Data Bank under accession code 7PVM. All other relevant data are available in the source data provided with this manuscript and are also available upon request to the corresponding author.

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Field-spe	ecific reporting			
Please select the o	ne below that is the best fit for yo	our research. If you are not sure, read the appropriate sections before making your selection.		
Life sciences	Behavioural & socia	al sciences Ecological, evolutionary & environmental sciences		
For a reference copy of t	the document with all sections, see <u>nature</u>	.com/documents/nr-reporting-summary-flat.pdf		
Life sciences study design				
All studies must dis	sclose on these points even when	the disclosure is negative.		
Sample size	The exact sample size and the number of replicate experiments have been described in the figure legends and methods.			
Data exclusions	No data (points) have been omitted.			
Replication	All data were recorded as at least fully independent duplicates. All attempts at replication were successful.			
Randomization	Randomization has no relevance in this study.			
Blinding	Blinding has no relevance in this study.			
Papartin	a for specific m	vatorials systems and mothods		
	-	aterials, 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 th	ne study	n/a Involved in the study		
Antibodies		ChIP-seq		
Eukaryotic		Flow cytometry		
Palaeontology and archaeology		MRI-based neuroimaging		
	nd other organisms			
Human res	search participants			