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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, seeAuthors & Referees and theEditorial Policy Checklist.

Sta	tistics					
For all statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.						
n/a	a Confirmed					
	<b>x</b> The exact sam	nple size $(n)$ for each experimental group/condition, given as a discrete number and unit of measurement				
	X A statement of	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.					
x	A description of all covariates tested					
x	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.					
x	For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings					
x	For hierarchic	al and complex designs, identification of the appropriate level for tests and full reporting of outcomes				
x	Estimates of e	effect sizes (e.g. Cohen's <i>d</i> , Pearson's <i>r</i> ), indicating how they were calculated				
,		Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.				
Sof	tware and c	code				
Polic	y information abo	ut availability of computer code				
	ta collection	No software was used				
Dat	ta analysis	Graphpad Prism version 5.0				
For ma	anuscripts utilizing custo	om 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 str	ongly encourage code	deposition in a community repository (e.g. GitHub). See the Nature Research guidelines for submitting code & software for further information.				
Dat	:a					
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						
Data available on request from the authors						
Field-specific 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.						
<b>x</b> Li	Life sciences Behavioural & social sciences Ecological, evolutionary & environmental sciences					
For a r	eference copy of the do	ocument with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>				

### Life sciences study design

All studies must disclose on these points even when the disclosure is negative.				
Sample size	Sample size were the maximum chimaeric mice we could obtain with no statistical method used to determine sample size.			
Data exclusions	No data were excluded			
Replication	All attempts to replication were successfull except long live experiments which have not being replicated			
Randomization	There are two groups in this study, one derived from chimeric mice and one as controls.			
Blinding	Blinding was not possible since we know which mice are chimeric and which are not			

## 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.

iviateriais & experimental systems			IVIETNOGS		
n/a	a Involved in the study		Involved in the study		
	<b>✗</b> Antibodies	×	ChIP-seq		
×	Eukaryotic cell lines	×	Flow cytometry		
×	Palaeontology	×	MRI-based neuroimaging		
	🗶 Animals and other organisms				
×	Human research participants				
×	Clinical data				

#### **Antibodies**

Antibodies used

anti-Rap1 (BL735, Bethyl), anti-53BP1 (Novus Biologicals) anti-TRF1 (BED5, Cell Signaling), anti-SMC-1 (Bethyl), GFP (Cell Signaling) and p21 (291H/B5, homemade)

Validation

Validation of the antibodies are available in the website of each company main page or in the CNIO Histopathology unit repository.

#### Animals and other organisms

Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research

Laboratory animals

(12951 female mice (Mus musculus) during their whole life were used.

Wild animals

This study does not involve wild animals

Field-collected samples

The study did not involve samples collected from the field

Ethics oversight

All animal experiments were approved by our Institutional Animal Care and Use Committee (IACUC) and by the Ethical Committee for animal experimentation (CEIyBA) (PROEX 133/15)

Note that full information on the approval of the study protocol must also be provided in the manuscript.