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Reporting Summary

Nature Portfolio 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 Portfolio policies, see our Editorial Policies 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.		
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.

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. *F*, *t*, *r*) with confidence intervals, effect sizes, degrees of freedom and *P* value noted *Give P 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

 \square Estimates of effect sizes (e.g. Cohen's d, Pearson's r), indicating how they were calculated

 $\textit{Our web collection on } \underline{\textit{statistics for biologists}} \ \textit{contains articles on many of the points above}.$

Software and code

Policy information about availability of computer code

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 OR state that no software was used.

Data analysis

Provide a description of all commercial, open source and custom code used to analyse the data in this study, specifying the version used OR state that no software was used.

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 Portfolio guidelines for submitting code & software for further information.

Data

Policy information about availability of data

All manuscripts must include a <u>data availability statement</u>. This statement should provide the following information, where applicable:

- Accession codes, unique identifiers, or web links for publicly available datasets
- A description of any restrictions on data availability
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Provide your data availability statement here.

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.			
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For a reference copy of the document with all sections, see nature.com/documents/nr-reporting-summary-flat.pdf			
Life sciences study design			
All studies must disclose on these points even when the disclosure is negative.			
Sample size	Sample size is indicated in figure legends. In all experiments we used n=8-11 age and sex-matched mice per group.		
Data exclusions	No data were excluded		
Replication	Replication To increase reproducibility, we analyzed a relatively high sample size in each experiment.		
Randomization	Describe how samples/organisms/participants were allocated into experimental groups. If allocation was not random, describe how covariates were controlled OR if this is not relevant to your study, explain why.		
Blinding	Histological and proteomics analysis of all samples was blind.		
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Field-collected sa	For laboratory work with field-collected samples, describe all relevant parameters such as housing, maintenance, temperature, photoperiod and end-of-experiment protocol OR state that the study did not involve samples collected from the field.		

Experimental procedures with mice were approved by the Animal Care Committee of McGill University (protocol 4966).

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

Ethics oversight