

eLife's transparent reporting form

We encourage authors to provide detailed information within their submission to facilitate the interpretation and replication of experiments. Authors can upload supporting documentation to indicate the use of appropriate reporting guidelines for health-related research (see EQUATOR
Network), life science research (see the BioSharing Information Resource), or the ARRIVE
guidelines for reporting work involving animal research. Where applicable, authors should refer to any relevant reporting standards documents in this form.

If you have any questions, please consult our Journal Policies and/or contact us: editorial@elifesciences.org.

Sample-size estimation

- You should state whether an appropriate sample size was computed when the study was being designed
- You should state the statistical method of sample size computation and any required assumptions
- If no explicit power analysis was used, you should describe how you decided what sample (replicate) size (number) to use

Please outline where this information can be found within the submission (e.g., sections or figure legends), or explain why this information doesn't apply to your submission:

No explicit power analysis was used. Sample sizes were decided based on experience with the techniques performed and practical considerations.

All details pertaining to sample size, number of experiments performed and statistical tests are included in the figure legends, the methods section, or in graphs.

Replicates

- You should report how often each experiment was performed
- You should include a definition of biological versus technical replication
- The data obtained should be provided and sufficient information should be provided to indicate the number of independent biological and/or technical replicates
- If you encountered any outliers, you should describe how these were handled
- Criteria for exclusion/inclusion of data should be clearly stated
- High-throughput sequence data should be uploaded before submission, with a private link for reviewers provided (these are available from both GEO and ArrayExpress)

Please outline where this information can be found within the submission (e.g., sections or figure legends), or explain why this information doesn't apply to your submission:

All experiments were repeated at least twice.

Biological replicates are defined as independent experiments performed on different animals, treated the same way, but at a different time. Technical replicates are defined as individuals treated in the same way, at the same time, within one independent experiment. Biological replicates are clearly distinguished on each graph.

The Source Data File specifies the number of independent experiments performed (biological replicates) and the number of individuals analyzed per experiment (technical replicates) for each figure panel. No outliers or individuals were removed from the data.

Statistical reporting

- Statistical analysis methods should be described and justified
- Raw data should be presented in figures whenever informative to do so (typically when N per group is less than 10)
- For each experiment, you should identify the statistical tests used, exact values of N, definitions of center, methods of multiple test correction, and dispersion and precision measures (e.g., mean, median, SD, SEM, confidence intervals; and, for the major substantive results, a measure of effect size (e.g., Pearson's r, Cohen's d)
- Report exact p-values wherever possible alongside the summary statistics and 95% confidence intervals. These should be reported for all key questions and not only when the p-value is less than 0.05.

Please outline where this information can be found within the submission (e.g., sections or figure legends), or explain why this information doesn't apply to your submission:

Detailed information about imaging, data analysis and statistical tests is provided in the figure legends and/or methods section. Raw data is provided in graphical representations of means where appropriate.

(For large

Graphical numerical data presented as a mean has error bars of +/- SD and includes individual data points for each animal. Results of statistical analysis are superimposed on the graph. Statistical analysis of data represented as means used unpaired t-tests for single comparisons or one-way ANOVA with Tukey Test for multiple comparisons.

Graphical numerical data presented as a proportion has error bars representing +/-95% confidence intervals. Information on sample size and biological replicates are detailed in the figure legend. Statistical analysis of proportional data used Fisher's Exact Test.

Proportions, means and dispersion/precision values were derived from all biological and technical replicates of a given sample.

datasets, or papers with a very large number of statistical tests, you may upload a single table file with tests, Ns, etc., with reference to sections in the manuscript.)

Group allocation

- Indicate how samples were allocated into experimental groups (in the case of clinical studies, please specify allocation to treatment method); if randomization was used, please also state if restricted randomization was applied
- Indicate if masking was used during group allocation, data collection and/or data analysis

Please outline where this information can be found within the submission (e.g., sections or figure legends), or explain why this information doesn't apply to your submission:

Samples were allocated into groups based on RNAi knockdown or treatment as indicated in the figure legends.

Additional data files ("source data")

- We encourage you to upload relevant additional data files, such as numerical data that are represented as a graph in a figure, or as a summary table
- Where provided, these should be in the most useful format, and they can be uploaded as "Source data" files linked to a main figure or table
- Include model definition files including the full list of parameters used
- Include code used for data analysis (e.g., R, MatLab)
- Avoid stating that data files are "available upon request"

Please indicate the figures or tables for which source data files have been provided:

The Source Data File includes raw data for every figure panel in the manuscript, on separate worksheets.