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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 paramet | ters |
|---------------------|------|
|---------------------|------|

| When statistical analyses are reported, confirm that the following i text, or Methods section).  | tems are present in the relevant location (e.g. figure legend, table legend, main                                     |  |  |  |
|--|---|--|--|--|
| n/a Confirmed  |   |  |  |  |
| The exact sample size (n) for each experimental group/con  | ndition, 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 to Only common tests should be described solely by name; described                   |   |  |  |  |
| A description of all covariates tested   |   |  |  |  |
| A description of any assumptions or corrections, such as t   | ests of normality and adjustment for multiple comparisons   |  |  |  |
| A full description of the statistics including <u>central tenden</u> <u>variation</u> (e.g. standard deviation) or associated <u>estimates</u> | cy (e.g. means) or other basic estimates (e.g. regression coefficient) AND of uncertainty (e.g. confidence intervals) |  |  |  |
| For null hypothesis testing, the test statistic (e.g. <i>F</i> , <i>t</i> , <i>r</i> ) wing Give P values as exact values whenever suitable.   | th confidence intervals, effect sizes, degrees of freedom and $\it P$ value noted                                     |  |  |  |
| 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 <i>d</i> , Pearson's <i>r</i> ), 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 <u>availability of computer code</u>  |   |  |  |  |
| Data collection NA   |   |  |  |  |
| Data analysis GraphPad Prism 7, Geneious 10.0.2., MEGA7  | , Cytation 5  |  |  |  |
|  | th but not yet described in published literature, software must be made available to editors/reviewers                |  |  |  |

## Data

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

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 list of figures that have associated raw data
- A description of any restrictions on data availability

Data deposit code to be provided before publication

| 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. |  |  |  |  |
| Life sciences                            | Behavioural & social sciences Ecological, evolutionary & environmental sciences                             |  |  |  |  |
| For a reference copy of t                | 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                              | NA  |  |  |  |  |
| Data exclusions                          | NA  |  |  |  |  |
| Replication                              | For all results obtained from cell lines are repeated for three independent experiments                     |  |  |  |  |
| Randomization                            | NA  |  |  |  |  |
| Blinding                                 | NA  |  |  |  |  |
|  |   |  |  |  |  |
| Poportin                                 | a for specific materials, systems and methods   |  |  |  |  |
| керогип                                  | g for specific materials, systems and methods   |  |  |  |  |
|  |   |  |  |  |  |
| Materials & experimental systems Methods |   |  |  |  |  |
| n/a Involved in th                       |   |  |  |  |  |
|  | logical materials ChIP-seq  |  |  |  |  |
| Antibodies                               |   |  |  |  |  |
| Eukaryotic Palaeontol                    |   |  |  |  |  |
|  | d other organisms   |  |  |  |  |
|  | earch participants  |  |  |  |  |
| <u> </u>                                 |   |  |  |  |  |
| Antibodies                               |   |  |  |  |  |
| Antibodies used                          | Commercially available the primary and secondary antibodies were used in Western Blot                       |  |  |  |  |
|  | Anti-Niemann Pick C1 antibody   |  |  |  |  |
|  | Cat#: ab108921 Host species: Rabbit   |  |  |  |  |
|  | Supplier: Abcam Dilution: 1:1,000   |  |  |  |  |
|  |   |  |  |  |  |
|  | HRP-goat anti-mouse IgG Cat#:405306   |  |  |  |  |
|  | Host species: Goat Supplier: BioLegend  |  |  |  |  |
|  | Dilution: 1:10,000  |  |  |  |  |
|  | Monoclonal anti-beta-actin antibody   |  |  |  |  |
|  | Cat#:A2228 Host species: Mouse  |  |  |  |  |
|  | Supplier: Sigma   |  |  |  |  |
|  | Dilution: 1:10,000  |  |  |  |  |
|  | Goat anti-rabbit IgG-HRP Cat#:sc-2357   |  |  |  |  |

Host species: Goat Supplier: Santa Cruz biotechnology Dilution: 1:10,000

Validation

Describe the validation of each primary antibody for the species and application, noting any validation statements on the manufacturer's website, relevant citations, antibody profiles in online databases, or data provided in the manuscript.

## Eukaryotic cell lines

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

Cell line source(s)

The primary bat cell lines were derived from Myotis davidii kidney (MdKi), R. leschenaultii kidney (RlKi), E. spelaea kidney (EsKi), E. spelaea lung (EsLu), and E. spelaea intestine (EsIn). Human embryonic kidney fibroblast cells (HEK293, ATCC#CRL-1573), human cervix epithelial cells (HeLa, ATCC#CCL-2), human lung epithelial cells (A549, ATCC#CCL-185), African green monkey kidney cells (Vero, clone E6, ATCC#CRL-1586), Rhesus monkey kidney epithelial cells (LLC-MK2, ATCC#CCL-7), Madin-Darby Canine kidney cells (MDCK, ATCC#CRL-2935), baby hamster kidney cells (BHK21, ATCC#CCL-10), were obtained from ATCC.

Authentication

All monkey and human cells were from ATCC with authentication. Bat cells made by ourselves were from organ. We guarantee they were from the organs described but there was no further authentication.

Mycoplasma contamination

We confirm that all cells were tested as mycoplasma negative.

Commonly misidentified lines (See ICLAC register)

None of the cell lines used are listed in the ICLAC database.