## nature portfolio

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

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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
$\boxtimes$	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.
$\boxtimes$	A description of all covariates tested
$\times$	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)
$\boxtimes$	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 <i>Give P values as exact values whenever suitable.</i>
$\boxtimes$	For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings
$\boxtimes$	For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes
$\boxtimes$	Estimates of 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.

## Software and code

Policy information about availability of computer code

Data collection

Illumina Miseq, QX200 Droplet Digital PCR System.

Data analysis

Perl (version 5.22.1), R (version 3.5.1 or 4.1.1), JupyterLab (version 0.34.9), nucleotide-sequence-clusterizer (version 0.0.7), RDPTools (https://github.com/rdpstaff/RDPTools). The pipeline (v1.2.0) for BarBIQ is available at github (https://github.com/Shiroguchi-Lab/BarBIQ\_Pipeline\_V1\_2\_0).

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 data availability statement. 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
- For clinical datasets or third party data, please ensure that the statement adheres to our policy

The sequencing data generated using the protocol in the relevant study are available from the NCBI Sequence Read Archive under accessions PRJNA639639, PRJNA639647, PRJNA780331, and PRJNA780361. A demonstration dataset is available at github (https://github.com/Shiroguchi-Lab/BarBIQ\_Pipeline\_V1\_2\_0).

Human rese	arch part	icipants			
Policy information	about <u>studies</u>	involving human research participants and Sex and Gender in Research.			
Reporting on sex	and gender	NA			
Population chara	acteristics	NA NA			
Recruitment		NA NA			
Ethics oversight		NA NA			
	ation on the app	roval of the study protocol must also be provided in the manuscript.			
Field-spe	ecific re	eporting			
Please select the o	ne below that	is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.			
X Life sciences		Behavioural & social sciences			
For a reference copy of	the document with	all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>			
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Life scier	nces st	udy design			
All studies must dis	sclose on these	points even when the disclosure is negative.			
Sample size	was not deteri	n absolute minimum based on ethical guidelines, but to statistically compare individuals for our research purpose (sample size mined by a statistic method since the sample size was not large), we used 19 mice in total (6 mice were used in each of two vitamin A study, others were used for technical evaluation).			
Data exclusions	No data were	excluded.			
Replication	All experiment	s were repeated twice or more (exact number of replications for each experiment is shown in the manuscript).			
Randomization	We used rando	om allocation of mice for the vitamin A-dependent experiments which is described in the Methods.			
Blinding	The random al	location in our experiment was not blinded since all mice were apparently indistinguishable by human eyes.			
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Reportin	g for s	pecific materials, systems and methods			
		about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, a your study. If you are not sure if a list item applies to your research, read the appropriate section before selecting a response.			
Materials & ex	perimental :	systems Methods			
	n/a Involved in the study n/a Involved in the study				
Antibodies		ChIP-seq			
	Eukaryotic cell lines    Flow cytometry   Palaeontology and archaeology   MRI-based neuroimaging				
	nd other organisi				
Clinical da	_				
Animals and	l other re:	search organisms			

Policy information about <u>studies involving animals</u>; <u>ARRIVE guidelines</u> recommended for reporting animal research, and <u>Sex and Gender in Research</u>

Laboratory animals

Six-week-old, eight-week-old, or nine-week-old C57BL/6J male or C57BL/6N female mice.

Wild animals	The study did not involve wild animals.
Reporting on sex	We used both male and female mice and reported on sex in the methods section.
Field-collected samples	The study did not involve samples collected from field.
Ethics oversight	All mouse procedures were performed in accordance with the ethical guidelines of the institute under the protocols approved by the Institutional Animal Care and Use Committee of RIKEN or the Animal Experimentation Committee of the Institute for Frontier Life and Medical Sciences. Kvoto University.

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