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
	\square 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.
\boxtimes	A description of all covariates tested
	A description of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons
\boxtimes	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 <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.
So	ftware and code

Software and code

Policy information about <u>availability of computer code</u>

Data collection No software was used for data collection

Data analysis All in silico analyses were performed in R Statistical Software (v4.0.2; R Core Team 2020) using in-house codes.

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

RNA-seq data files from the present study have been deposited into the Gene Expression Omnibus database (www.ncbi.nlm.nih.gov/geo/) with accession number GSE230639. Other datasets used in the study can be found in their respective original publications as listed in Table 1 [44,66-71]. The list of 252 statistically significant (FDR < 0.05) lipids and metabolites can be found in Supplementary Data 1. Source data for all figures/graphs are available in Figshare (DOI: 10.6084/

m9.figshare.251344	70). Relevant R s	cripts and all other data is available upon reasonable request from the corresponding author.		
Human rese	arch part	cipants		
		nvolving human research participants and Sex and Gender in Research.		
Reporting on sex and gender		NA		
Population characteristics		NA NA		
Recruitment		NA		
Ethics oversight		NA NA		
Note that full informa	ation on the appi	oval of the study protocol must also be provided in the manuscript.		
Field-spe		eporting s the best fit for your research. If you are not sure, read the appropriate sections before making your selection.		
Life sciences For a reference copy of t	E the document with	Behavioural & social sciences		
		points even when the disclosure is negative.		
Sample size	Describe how sample size was determined, detailing any statistical methods used to predetermine sample size OR if no sample-size calculation was performed, describe how sample sizes were chosen and provide a rationale for why these sample sizes are sufficient.			
Data exclusions	Describe any data exclusions. If no data were excluded from the analyses, state so OR if data were excluded, describe the exclusions and the rationale behind them, indicating whether exclusion criteria were pre-established.			
Replication	Describe the measures taken to verify the reproducibility of the experimental findings. If all attempts at replication were successful, confirm this OR if there are any findings that were not replicated or cannot be reproduced, note this and describe why.			
Randomization	Describe how samples/organisms/participants were allocated into experimental groups. If allocation was not random, describe how cover were controlled OR if this is not relevant to your study, explain why.			
Blinding Describe whether the investigators were blinded to group all describe why OR explain why blinding was not relevant to yo		ner the investigators were blinded to group allocation during data collection and/or analysis. If blinding was not possible, R explain why blinding was not relevant to your study.		
<u> </u>		pecific materials, systems and methods about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material,		
system or method list	ted 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.		
Materials & ex	·	· · · · · · · · · · · · · · · · · · ·		
n/a Involved in the study				
Antibodies Lukaryotic cell lines		Flow cytometry		
Animals and other organisms				
Clinical data				
Dual use research of concern				

Animals and other research organisms

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

Laboratory animals	C57BI6/J mice
Wild animals	(NA
Reporting on sex	male mice
Field-collected samples	The details of the animal protocol is attached
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Ethics oversight	All experiments were conducted according to the regulations of the Bern Animal Welfare Committee, Canton of Bern, Switzerland (BE42/19).

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