nature research

Corresponding author(s):	Catherine A. Wakeman
Last updated by author(s):	Dec 28, 2020

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 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,

_					
C-4	ta:	٠,	\sim t	- 1	~
`	1 4		N.		

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
\boxtimes	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. <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 statistics for biologists contains articles on many of the points above

Software and code

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

Data collection FV Analysis software (Olympus FluoView 3000) (confocal imaging and z-stack); Hitachi PC-SEM software and Quartz PCI software (SEM micrograph); FlexImaging (MALDI IMS)

Data analysis

Rockhopper 2 (RNA-seq analysis); ImageJ (live/dead cell counting for confocal images); GraphPad Prism 8.0 (data analysis for CV, qPCR, and CFUS); FlexAnalysis (MALDI IMS)

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 Research guidelines for submitting code & software for further information.

Data

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

 $All \ manuscripts \ must \ include \ a \ \underline{data \ availability \ statement}. \ This \ statement \ should \ provide \ the \ following \ information, \ where \ applicable:$

- 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

The raw RNA-seq data sets generated during this study are available through NCBI's BioProject database under accession number PRJNA664520. The raw data sets generated by MALDI IMS are available upon request. The authors declare that all other relevant data supporting the claims of the paper are available either in the main text or supplementary files.

-· ı									
-ıel	C	l-SK	oec	ITIC	rei	00	rti	n	g

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.			
Life sciences	Behavioural & social sciences Ecological, evolutionary & environmental sciences			
For a reference copy of	the document with all sections, see nature.com/documents/nr-reporting-summary-flat.pdf			
Life scier	nces study design			
All studies must dis	sclose on these points even when the disclosure is negative.			
Sample size	All data sets were replicated on at least 3 independent days and performed in triplicate on each day. The same day triplicates were averaged and treated as technical replicates.			
Data exclusions	No data exclusions were made.			
Replication	All data sets were replicated on at least 3 independent days and performed in triplicate on each day. The same day triplicates were averaged and treated as technical replicates. By repeating experiments in triplicate both on the same day as well as performing these experiments on multiple days, we are confident in the reproducibility of these data.			
Randomization	This is not relevant to our study.			
Blinding	This was not performed as the same person collecting the data was performing the analysis for the bulk of the data sets. Therefore, blinding was not readily achievable.			
Poportin	σ for specific materials, systems and methods			
<u> </u>	g for specific materials, systems and methods			
	on from authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, 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.			

n/a	Involved in the study
∇	Antibodies

Eukaryotic cell lines

Palaeontology and archaeology
Animals and other organisms

Human research participants

Clinical data

Dual use research of concern

Methods

n/a	Involved in the study
\boxtimes	ChIP-seq
\boxtimes	Flow cytometry