## nature research

Corresponding author(s):	Dr Melissa Herbst-Kralovetz
Last updated by author(s):	Apr 2, 2021

## **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.

_				
5	۲a	Ť١	c†	ics

For	all statistical an	alyses, 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 stateme	nt on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly			
	The statis	tical test(s) used AND whether they are one- or two-sided on tests should be described solely by name; describe more complex techniques in the Methods section.			
	A descript	ion of all covariates tested			
	A descript	ion of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons			
	A full desc AND varia	ription of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) tion (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)			
	For null hy Give P valu	pothesis testing, the test statistic (e.g. $F$ , $t$ , $r$ ) with confidence intervals, effect sizes, degrees of freedom and $P$ value noted as as exact values whenever suitable.			
$\boxtimes$	For Bayes	ian analysis, information on the choice of priors and Markov chain Monte Carlo settings			
	For hierar	chical and complex designs, identification of the appropriate level for tests and full reporting of outcomes			
	$\square$ Estimates of effect sizes (e.g. Cohen's $d$ , Pearson's $r$ ), 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 <u>availability of computer code</u>					
Da	ata collection	Bio-Rad Bio-Plex Manager 5.0 software			

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

Data analysis

Policy information about availability of data

All manuscripts must include a data availability statement. This statement should provide the following information, where applicable:

5) Prism v8 Software (GraphPad): (https://www.graphpad.com/scientific-software/prism/)

- Accession codes, unique identifiers, or web links for publicly available datasets

1) Metaboanalyst 4.0: (https://www.metaboanalyst.ca/)
2) Metabolon Inc: (https://www.metabolon.com/)
3) R (version 3.6.3): (https://www.r-project.org/)
4) ClustVis: (https://biit.cs.ut.ee/clustvis/)

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

The authors declare that the data supporting the findings of this study are available within the paper and its supplementary information files. Additional data are available from the corresponding author upon reasonable request.

Field-spe	ecific reporting			
Please select the or	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 t	the document with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>			
Life scier	nces study design			
All studies must dis	sclose on these points even when the disclosure is negative.			
Sample size	Sample size was determined based on previous studies and the number of replicates needed to minimize variation and maximize statistical significance. For the immunoproteomics and metabolomics analysis we performed at least three independent batches of bioreactors run in duplicate for all analyses.			
Data exclusions	No data was excluded from the analysis.			
Replication	The data presented is representative of at least three independent replicates of the experiments.			
Randomization	Cell culture samples were randomly evaluated for downstream analyses. However during the set up of each experiment, cell culture wells and experimental group were not randomized on the plate for the in vitro studies.			
Blinding	The investigators performing the experiments were not blinded as that was not relevant to our study and would be difficult to complete.			
Reporting for specific materials, systems and methods  We require information from authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, system or method listed 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 & exp	perimental systems Methods			
n/a Involved in th	ne study n/a   Involved in the study			
✓ Antibodies     ✓ ChIP-seq				
☐ ☑ Eukaryotic cell lines ☑ ☐ Flow cytometry				
Palaeontology and archaeology MRI-based neuroimaging				
Animals and other organisms				
Human research participants				
Clinical data  Dual use research of concern				
M   Dual disc re	Scalar of concern			
Eukaryotic c	ell lines			
Policy information	about <u>cell lines</u>			
Cell line source(s	Human A2EN cervical epithelial cell line			

STR DNA profiling was performed and these cells are not contaminated with other cell lines found in available databases.

The cell lines were not tested for mycoplasma contamination recently

The cell line used in this study is not commonly misidentified

Authentication

(See <u>ICLAC</u> register)

Mycoplasma contamination

Commonly misidentified lines