# nature portfolio

Corresponding author(s):	Ishier Raote
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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>.

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
	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
	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

Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.

For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes

For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings

Estimates of effect sizes (e.g. Cohen's d, Pearson's r), indicating how they were calculated

#### Software and code

Policy information about availability of computer code

Data collection

Xcalibur (version 3.1.)

Give P values as exact values whenever suitable.

Data analysis

**Statistics** 

MaxQuant analysis software and the implemented Andromeda software (1.5.3.8.); Perseus (1.6.2.3.); Instant Clue (0.5.1.)

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

Human FASTA database was used for the sample analysis

### Research involving human participants, their data, or biological material

Reporting on sex and gender	Primary cells were derived from female and male donors
Reporting on race, ethnicity, or other socially relevant groupings	N/A
Population characteristics	no such information is collected
Recruitment	Skin biopsies were obtained from female and male healthy individuals or patients with scleroderma (systemic sclerosis) after written consent.
Ethics oversight	Skin biopsies were obtained with approval of the local ethics committee (Ethics committee of the medical faculty of University of Cologne, 21-1072). The study was conducted in accordance with the criteria set by the Declaration of Helsinki.

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Please select the one below that is 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 the document with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>			
Life sciences study design			

### Life sciences study design

All studies must disclose on these points even when the disclosure is negative.

Sample size Sample sizes were decided based on providing sufficient repeats to carry out robust statistical analyses, taking into accoutn resource availablity and ethical considerations for animal experiments.

Data exclusions No data were excluded

Research sample

Sampling strategy

Data collection

**Timing** 

The main phenotype reported here (inhibition of collagen secretion), was independently verified by at least two collaborating groups Replication performing this study, as detailed in the figure legends.

Randomization Samples were randomly allocated

Analyses were performed single or double blinded Blinding

# Behavioural & social sciences study design

All studies must disclose on these points even when the disclosure is negative.

Briefly describe the study type including whether data are quantitative, qualitative, or mixed-methods (e.g., qualitative cross-sectional, Study description quantitative experimental, mixed-methods case study).

> State the research sample (e.g. Harvard university undergraduates, villagers in rural India) and provide relevant demographic information (e.g. age, sex) and indicate whether the sample is representative. Provide a rationale for the study sample chosen. For

studies involving existing datasets, please describe the dataset and source. Describe the sampling procedure (e.g. random, snowball, stratified, convenience). Describe the statistical methods that were 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. For qualitative data, please indicate whether data saturation was considered, and what criteria were used to decide that no further sampling was needed.

Provide details about the data collection procedure, including the instruments or devices used to record the data (e.g. pen and paper, computer, eye tracker, video or audio equipment) whether anyone was present besides the participant(s) and the researcher, and whether the researcher was blind to experimental condition and/or the study hypothesis during data collection.

Indicate the start and stop dates of data collection. If there is a gap between collection periods, state the dates for each sample

Data exclusions If no data were excluded from the analyses, state so OR if data were excluded, provide the exact number of exclusions and the rationale behind them, indicating whether exclusion criteria were pre-established.

Non-participation

State how many participants dropped out/declined participation and the reason(s) given OR provide response rate OR state that no participants dropped out/declined participation.

Randomization

If participants were not allocated into experimental groups, state so OR describe how participants were allocated to groups, and if allocation was not random, describe how covariates were controlled.

# Ecological, evolutionary & environmental sciences study design

All studies must disclose on these points even when the disclosure is negative.

Study description

Briefly describe the study. For quantitative data include treatment factors and interactions, design structure (e.g. factorial, nested, hierarchical), nature and number of experimental units and replicates.

Research sample

Describe the research sample (e.g. a group of tagged Passer domesticus, all Stenocereus thurberi within Organ Pipe Cactus National Monument), and provide a rationale for the sample choice. When relevant, describe the organism taxa, source, sex, age range and any manipulations. State what population the sample is meant to represent when applicable. For studies involving existing datasets,

describe the data and its source.

Sampling strategy

Note the sampling procedure. Describe the statistical methods that were 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.

Timing and spatial scale Indicate the start and stop dates of data collection, noting the frequency and periodicity of sampling and providing a rationale for these choices. If there is a gap between collection periods, state the dates for each sample cohort. Specify the spatial scale from which the data are taken

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.

Reproducibility

Describe the measures taken to verify the reproducibility of experimental findings. For each experiment, note whether any attempts to repeat the experiment failed OR state that all attempts to repeat the experiment were successful.

repeat the experiment Janea OR state that all attempts to repeat the experiment were successful.

Randomization

Describe how samples/organisms/participants were allocated into groups. If allocation was not random, describe how covariates were controlled. If this is not relevant to your study, explain why.

Describe the extent of blinding used during data acquisition and analysis. If blinding was not possible, describe why OR explain why blinding was not relevant to your study.

Did the study involve field work? Yes No

Blinding

### Field work, collection and transport

Field conditions Describe the study conditions for field work, providing relevant parameters (e.g. temperature, rainfall).

Location State the location of the sampling or experiment, providing relevant parameters (e.g. latitude and longitude, elevation, water depth).

Access & import/export

Describe the efforts you have made to access habitats and to collect and import/export your samples in a responsible manner and in compliance with local, national and international laws, noting any permits that were obtained (give the name of the issuing authority,

the date of issue, and any identifying information).

Disturbance Describe any disturbance caused by the study and how it was minimized.

# 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 & experime	ental system	ns Methods	
n/a Involved in the study n/a Involved in the study			
Antibodies	Antibodies ChIP-seq		
Eukaryotic cell lines			
Palaeontology and archaeology MRI-based neuroimaging			
Animals and other of	organisms		
Clinical data			
Dual use research o	f concern		
Plants			
Antibodies			
Antibodies used  Anti-col1 (1:200, ab23730, Abcam, Cambridge, MA, USA), goat anti-rabbit Cy3 (1:750, RRIDAB AB_2534029, cat. no. A10520) secondary antibody, Goat anti-type I collagen (SouthernBiotech, 1310-01; 1:2000; reduced); rabbit anti-fibrillin1 (kind gift from G. Sengle, Cologne; 1:2000 unreduced); rabbit anti-fibronectin (abcam, ab23750; 1:500, unreduced); and rabbit anti-GAPDH (abcam,			
	ab9485; 1:250	00; reduced), GAPDH (abcam, ab9485; 1:2500; reduced),	
Validation	Antibodies we	ere not validated	
Eukaryotic cell lin	ies		
Policy information about <u>ce</u>	ell lines and Se	ex and Gender in Research	
Cell line source(s)		cells were obtained from the host institution's (Centre for Genomic Regulation) biorepository. RDEB/FB/C7 fibroblasts generated and reported in earlier publications (DOI:10.1038/ng1041 and 10.1016/j.cell.2008.12.025	
Authentication	Cell lir	nes were not authenticated	
Mycoplasma contaminat	ion All cel	l lines test negative for mycoplasma with monthly testing	
Commonly misidentified	lines No mi	cidentified lines were used	
Commonly misidentified lines (See ICLAC register)  No misidentified lines were used			
Palaeontology an	d Archae	ology	
	<u> </u>	<u></u>	
Specimen provenance Provide provenance information for specimens and describe permits that were obtained for the work (including the name of the issuing authority, the date of issue, and any identifying information). Permits should encompass collection and, where applicable, export.			
Specimen deposition	Indicate where the specimens have been deposited to permit free access by other researchers.		
Dating methods  If new dates are provided, describe how they were obtained (e.g. collection, storage, sample pretreatment and measurement), where they were obtained (i.e. lab name), the calibration program and the protocol for quality assurance OR state that no new dates are provided.			
Tick this box to confir	m that the rav	w and calibrated dates are available in the paper or in Supplementary Information.	
Ethics oversight   Identify the organization(s) that approved or provided guidance on the study protocol, OR state that no ethical approval or guidance was required and explain why not.			
Note that full information on the approval of the study protocol must also be provided in the manuscript.			
Animals and othe	er researc	ch organisms	
		g animals; ARRIVE guidelines recommended for reporting animal research, and Sex and Gender in	
Laboratory animals	Zebrafish: TL/I	Ekwill wildtype strain were used at 0-4 days for collagen imaging. 8-month fish were used for wounding experiments	
Wild animals	No wild anima	als were used	
Reporting on sex	eporting on sex Sex / gender was not considered in this study		
Field-collected samples	Id-collected samples No field-collected samples were used		

Ethics oversight

AB wildtype zebrafish were used. Fish were maintained at the PRBB according to the standard procedures of the aquatic facility. All the protocols were following the national and European guidelines and were approved by the institutional Animal Care and Use Ethic Committee (PRBB-IACUEC). The principles of the 3Rs were applied in all zebrafish experiments.

The zebrafish wound healing experiments were approved by the local and federal animal care committees (City of Cologne: 8.87-50.10.31.08.129; LANUV Nordrhein-Westfalen: 81-02.04.2018.A097)

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

#### Clinical data

Policy information about <u>clinical studies</u>	olicy	inforr	nation	about	clinical	studies
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All manuscripts should comply with the ICMJE guidelines for publication of clinical research and a completed CONSORT checklist must be included with all submissions.

Clinical trial registration	Provide the trial registration number from ClinicalTrials.gov or an equivalent agency.
Study protocol	Note where the full trial protocol can be accessed OR if not available, explain why.
Data collection	Describe the settings and locales of data collection, noting the time periods of recruitment and data collection.
Outcomes	Describe how you pre-defined primary and secondary outcome measures and how you assessed these measures.

### Dual use research of concern

Policy information about <u>dual use research of concern</u>

#### Hazards

Could the accidental, deliberate or reckless misuse of agents or technologies generated in the work, or the application of information presented in the manuscript, pose a threat to:

No	Yes
$\boxtimes$	Public health
$\boxtimes$	National security
$\boxtimes$	Crops and/or livestock
$\boxtimes$	Ecosystems
$\boxtimes$	Any other significant area

#### Experiments of concern

Does the work involve any of these experiments of concern:

Vo	Yes
$\boxtimes$	Demonstrate how to render a vaccine ineffective
$\boxtimes$	Confer resistance to therapeutically useful antibiotics or antiviral agents
$\boxtimes$	Enhance the virulence of a pathogen or render a nonpathogen virulent
$\boxtimes$	Increase transmissibility of a pathogen
$\boxtimes$	Alter the host range of a pathogen
$\boxtimes$	Enable evasion of diagnostic/detection modalities
$\boxtimes$	Enable the weaponization of a biological agent or toxin
$\boxtimes$	Any other potentially harmful combination of experiments and agents

#### Plants

Seed stocks

Report on the source of all seed stocks or other plant material used. If applicable, state the seed stock centre and catalogue number. If plant specimens were collected from the field, describe the collection location, date and sampling procedures.

Novel plant genotypes

Describe the methods by which all novel plant genotypes were produced. This includes those generated by transgenic approaches, gene editing, chemical/radiation-based mutagenesis and hybridization. For transgenic lines, describe the transformation method, the number of independent lines analyzed and the generation upon which experiments were performed. For gene-edited lines, describe the editor used, the endogenous sequence targeted for editing, the targeting guide RNA sequence (if applicable) and how the editor was applied.

Authentication

Describe any authentication procedures for each seed stock used or novel genotype generated. Describe any experiments used to assess the effect of a mutation and, where applicable, how potential secondary effects (e.g. second site T-DNA insertions, mosiacism, off-target gene editing) were examined.

#### ChIP-sea

#### Data deposition

Confirm that you have deposited or provided access to graph files (e.g. BED files) for the called peaks. Data access links For "Initial submission" or "Revised version" documents, provide reviewer access links. For your "Final submission" document,

May remain private before publication.

provide a link to the deposited data.

Files in database submission

Provide a list of all files available in the database submission.

Confirm that both raw and final processed data have been deposited in a public database such as GEO.

Genome browser session (e.g. UCSC)

Provide a link to an anonymized genome browser session for "Initial submission" and "Revised version" documents only, to enable peer review. Write "no longer applicable" for "Final submission" documents.

#### Methodology

Replicates

Describe the experimental replicates, specifying number, type and replicate agreement.

Sequencing depth

Describe the sequencing depth for each experiment, providing the total number of reads, uniquely mapped reads, length of reads and whether they were paired- or single-end.

**Antibodies** 

Describe the antibodies used for the ChIP-seq experiments; as applicable, provide supplier name, catalog number, clone name, and

Peak calling parameters

Specify the command line program and parameters used for read mapping and peak calling, including the ChIP, control and index files

Data quality

Describe the methods used to ensure data quality in full detail, including how many peaks are at FDR 5% and above 5-fold enrichment.

Software

Describe the software used to collect and analyze the ChIP-seq data. For custom code that has been deposited into a community repository, provide accession details.

### Flow Cytometry

#### **Plots**

Confirm that:

 $\Box$  The axis labels state the marker and fluorochrome used (e.g. CD4-FITC).

The axis scales are clearly visible. Include numbers along axes only for bottom left plot of group (a 'group' is an analysis of identical markers).

All plots are contour plots with outliers or pseudocolor plots.

A numerical value for number of cells or percentage (with statistics) is provided.

#### Methodology

Sample preparation

Describe the sample preparation, detailing the biological source of the cells and any tissue processing steps used.

Instrument

Identify the instrument used for data collection, specifying make and model number.

Software

Describe the software used to collect and analyze the flow cytometry data. For custom code that has been deposited into a community repository, provide accession details.

(			
Cell population abundance  Describe the abundance of the relevant cell populations within post-sort fractions, providing details on the purity of the samples and how it was determined.			
Gating strategy  Describe the gating strategy used for all relevant experiments, specifying the preliminary FSC/SSC gates of the starting cell population, indicating where boundaries between "positive" and "negative" staining cell populations are defined.			
Tick this box to confirm that a figure exemplifying the gating strategy is provided in the Supplementary Information.			
Magnetic resonance in	naging		
Experimental design			
Design type	Indicate task or resting state; event-related or block design.		
Design specifications	Specify the number of blocks, trials or experimental units per session and/or subject, and specify the length of each trial or block (if trials are blocked) and interval between trials.		
Behavioral performance measures  State number and/or type of variables recorded (e.g. correct button press, response time) and what statistics were use to establish that the subjects were performing the task as expected (e.g. mean, range, and/or standard deviation across subjects).			
Acquisition			
Imaging type(s)			
Field strength	Specify in Tesla		
Sequence & imaging parameters  Specify the pulse sequence type (gradient echo, spin echo, etc.), imaging type (EPI, spiral, etc.), field of view, matrix size, slice thickness, orientation and TE/TR/flip angle.			
Area of acquisition State whether a whole brain scan was used OR define the area of acquisition, describing how the region was determined.			
Diffusion MRI Used	☐ Not used		
Preprocessing			
Preprocessing software	Provide detail on software version and revision number and on specific parameters (model/functions, brain extraction, segmentation, smoothing kernel size, etc.).		
Normalization   If data were normalized/standardized, describe the approach(es): specify linear or non-linear and define image types used for transformation OR indicate that data were not normalized and explain rationale for lack of normalization.			
Normalization template  Describe the template used for normalization/transformation, specifying subject space or group standardized space (e.g. original Talairach, MNI305, ICBM152) OR indicate that the data were not normalized.			
Noise and artifact removal  Describe your procedure(s) for artifact and structured noise removal, specifying motion parameters, tissue signals and physiological signals (heart rate, respiration).			
Volume censoring  Define your software and/or method and criteria for volume censoring, and state the extent of such censoring.			
Statistical modeling & infere	nce		
	Specify type (mass univariate, multivariate, RSA, predictive, etc.) and describe essential details of the model at the first and second levels (e.g. fixed, random or mixed effects; drift or auto-correlation).		
Effect(s) tested	Define precise effect in terms of the task or stimulus conditions instead of psychological concepts and indicate whether ANOVA or factorial designs were used.		
Specify type of analysis: Wh	nole brain ROI-based Both		
Statistic type for inference	Specify voxel-wise or cluster-wise and report all relevant parameters for cluster-wise methods.		
(See Eklund et al. 2016)			
Correction	Describe the type of correction and how it is obtained for multiple comparisons (e.g. FWE, FDR, permutation or Monte Carlo).		

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