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## **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 <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.

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n/a	Confirmed			
	The exact	sample size $(n)$ for each experimental group/condition, given as a discrete number and unit of measurement		
	A stateme	ent on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly		
	The statis Only comm	tical test(s) used AND whether they are one- or two-sided non tests should be described solely by name; describe more complex techniques in the Methods section.		
$\boxtimes$	A descript	tion of all covariates tested		
$\boxtimes$	A descript	tion of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons		
	A full deso	cription of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) ation (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)		
	For null h	ypothesis testing, the test statistic (e.g. $F$ , $t$ , $r$ ) with confidence intervals, effect sizes, degrees of freedom and $P$ value noted use as exact values whenever suitable.		
$\boxtimes$	For Bayes	ian 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 $d$ , Pearson's $r$ ), 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>				
Da	ata collection	CytExpert (v2.4)was used for flow cytometry data collection		

YMAP (v1.0), Phyre2 (v2.0), YASARA (v20,12,24), Autodock Vina (v1.1.2), PyMOL (v2.5.0), Prism 6, Bio-Rad CFX Manager (v3.1), MuTect

### Data

Data analysis

Policy information about availability of data

(v1.1.4.)

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

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.

- 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

Source data are provided with this paper. All additional data, including raw data and images associated with all figures, is available from the corresponding author, L.E.C, upon reasonable request.

Field-spe	cific re	porting				
Please select the or	ne below that is	s the best fit for your research. If you are not sure, read the appropriate sections before making your selection.				
X Life sciences	□В	ehavioural & social sciences Ecological, evolutionary & environmental sciences				
For a reference copy of t	he document with	all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>				
Life scier	ices stu	udy design				
All studies must dis	close on these	points even when the disclosure is negative.				
Sample size		der to use statistical analysis, sample size was always n=3 or greater. Murine explant experiments used 3-5 mice per group based on rical variability data within groups. All experiments were performed in biological duplicate or greater with little deviation between cates.				
Data exclusions	scatter parame	netry experiments, events were excluded from calculation of median fluorescence intensity (MFI) by gating on forward and side neters to eliminate debris and multi-cell clumps that would skew data. Gating removed less than 15% of all acquired events. No ere excluded from analysis.				
Replication	All attempts at	at replication were successful. Unless otherwise noted, all experiments are representative of at least two biological replicates.				
Randomization	For all animal st	or all animal studies, animals were randomly assigned to experimental groups.				
Blinding	Blinding was not relevant to this study as this was not an observational study with no opportunity for bias to factor into quantitative results.  All assays had a quantitative output, rather than qualitative, and therefore, blinding was not required to eliminate user bias.					
Reporting	g for sp	pecific 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 s	ystems Methods				
n/a Involved in th	e study	n/a Involved in the study				
Antibodies		ChIP-seq				
Eukaryotic		Flow cytometry				
	ogy and archaeo					
Animals and other organisms						
Human research participants  Clinical data						
Dual use research of concern						
-,-						
Eukaryotic co	ell lines					
Policy information a	about <u>cell lines</u>					
Cell line source(s)	1	HepG2 cells were provided by ATCC. (ATCC HB-8065).				
Authentication	Cell line was not authenticated as a specific tissue of origin was non-critical to the validity of the results reported.					
Mycoplasma cont	tamination	mination All cell lines tested negative for Mycoplasma contamination.				
Commonly miside (See <u>ICLAC</u> register)		No commonly misidentified cell lines were used.				

## Animals and other organisms

 $Policy\ information\ about\ \underline{studies\ involving\ animals;}\ \underline{ARRIVE\ guidelines}\ recommended\ for\ reporting\ animal\ research$ 

Laboratory animals

C3H/HeN (purchased from Charles River), female mice, 6-weeks old for mouse vaginal explants and Spraw Daley (purchased from Envigo), male rats, 10 weeks old for the rat catheter experiments.

Wild animals

This study did not involve wild animals.

Field-collected samples

The study did not involve samples collected from the field.

Ethics oversight

Murine mouse tissues for the vaginal explant were acquired and analyzed using protocols approved by the Institutional Animal Care and Use Committee (IACUC) of Louisiana State University Health Sciences Center (Protocol 3663). Rat catheter biofilm animal procedures were approved by the Institutional Animal Care and Use Committee at the University of Wisconsin-Madison according to the guidelines of the Animal Welfare Act, The Institute of Laboratory Animals Resources Guide for the Care and Use of Laboratory Animals, and Public Health Service Policy. The approved animal protocol number is DA0031.

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

## Flow Cytometry

#### Plots

Confirm that:

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 Cells used in the flow cytometry experiments include the HWP1p-GFP C. albicans strain. Supplementary Table 1 for this study

includes more information on the strain. C. albicans cells were sub-cultured from a saturated overnight culture to an OD600 of 0.1 in YPD medium and grown for 6 hours at  $42^{\circ}$ C in the absence or presence of 1-ABC or Lactobacillus-conditioned medium. Cells were then pelleted, washed once with PBS, resuspended in PBS, and diluted 1:10 in 1 mL PBS. 250  $\mu$ L of each

sample were added to a flat-bottom transparent 96-well plate (Beckman Coulter).

Instrument CytoFLEX S (Beckman Coulter).

Software CytExpert Software (v2.4).

Cell population abundance The cell population post-sort was approximately 80% of the population.

Gating strategy

The purpose of the gating strategy was to capture the C. albicans population undergoing the morphogenesis process. The gating strategy captured cells with a FSC-A between 40x10^4 and 280x10^4 and SSC-A between 20x10^4 and 190x10^4.

Tick this box to confirm that a figure exemplifying the gating strategy is provided in the Supplementary Information.