

Corresponding author(s):	Matthew Rogers
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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 Authors & Referees and the Editorial Policy Checklist.

Sta	tis	tics	
For a	ll st	atistical ana	alyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.
n/a	Cor	nfirmed	
	\times	The exact s	sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement
	\boxtimes	A stateme	nt on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly
	\boxtimes	The statist	ical 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.
	\times	A descripti	ion of all covariates tested
	\times	A descripti	ion of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons
	\boxtimes	A full desci AND variat	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)
	\boxtimes	For null hy Give P value	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 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 d , Pearson's r), indicating how they were calculated
1			Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.
Sof	tw	are and	d code
Polic	y in	formation a	about <u>availability of computer code</u>
Dat	ta c	ollection	Cycle threshold values (Ct) were determined using the Applied Biosystems SDS2.4 Software. Ct values were input into Biogazelle qBasePLUS software. For normalisation calculations, candidate control genes were tested with geNorm and Normfinder programmes. Luminex microspheres were quantified using a Luminex® 100 System (Luminexcorp).
Dat	ta a	nalysis	Data was analysed with GraphPad Prism software version 5.0. R version 3.4.1 with the Mclust package was used for the Gaussian mixture model-based cluster analysis.
			custom algorithms or software that are central to the research but not yet described in published literature, software must be made available to editors/reviewers. ode deposition in a community repository (e.g. GitHub). See the Nature Research 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 list of figures that have associated raw data
- A description of any restrictions on data availability

All data generated or analysed during this study are included in this published article (and its supplementary information files).

Field-specific reporting				
Please select the or	be 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	ne document with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>			
Life sciences study design				
All studies must dis	close on these points even when the disclosure is negative.			
Sample size	The sample size of analysed bites was based on a previous study (Kimblin et al. 2008, PNAS).			
·	The sample sizes used in most infection experiments will be 6 mice per group, maintained for 80 days. Our previous studies (Rogers et al. 2009, PLoS Pathogens) have shown this is sufficient to show a 30% difference in parasite load with a power of 80% and at 95% confidence limits.			
Data exclusions	No data were excluded from the analyses.			
Replication	Experiments analysing the composition of the infectious dose from individual infected sandflies were obtained by pooling the results from 4 independent replicates. Experiments investigating the immunological consequence of dose composition were obtained from a representative experiment of 3 replicates.			
Randomization	Infected sandflies were randomly selected to bite mice. All sandflies and their bites were analysed.			
Blinding	No prior knowledge of the sandfly infection status was revealed to the researcher performing the RTqPCR.			
Reportin	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, ed 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 & experimental systems Methods				
n/a Involved in th	e study n/a Involved in the study			
Antibodies	ChIP-seq			
Eukaryotic				
Palaeontol				
Animals and other organisms				
Human research participants				
Clinical dat				
Antibodies				
Antibodies used	CCL3 (MIP-1a), CXCL2 (MIP-2) and IL-1b as part of a custom Bio-Plex Pro Mouse Cytokine Luminex assay (BIO-RAD UK)			
Validation	http://www.bio-rad.com/webroot/web/pdf/lsr/literature/Bulletin_3156.pdf http://www.bio-rad.com/webroot/web/pdf/lsr/literature/Bulletin_5297.pdf			
Animals and	other organisms			
Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research				
Laboratory anima	Mouse, BALB/c, Female, Age 6-8 weeks, average weight 21g, supplier: Charles River UK.			

Laboratory animals

Mouse, BALB/c, Female, Age 6-8 weeks, average weight 21g, supplier: Charles River UK.

Wild animals

The study did not involve wild animals.

 $\begin{tabular}{ll} Field-collected samples & The study did not involve samples from the field. \end{tabular}$

Ethics oversight

All animal experiments were carried out in accordance with the UK Animal Scientific Procedure Act (ASPA) 1986, which transposes European Directive 2010/63/EU into UK national law.

The animal studies were approved by the UK home office in granting Project licence 70/8427 under the Animal Scientific Procedure Act and all protocols had undergone appropriate local ethical review procedures by the Animal welfare and Ethical

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

Review Board (AWERB) of The London School of Hygiene and Tropical Medicine.