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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 <u>Authors & Referees</u> and the <u>Editorial Policy Checklist</u>.

When statistical analyses are reported, confirm that the following items are present in the relevant location (e.g. figure legend, table legend, main

Statistical parameters

text	text, or Methods section).				
n/a	Confirmed				
	\boxtimes	The $\underline{\text{exact sample size}}$ (n) for each experimental group/condition, given as a discrete number and unit of measurement			
	\boxtimes	An indication of 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.			
	\boxtimes	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 statistics including <u>central tendency</u> (e.g. means) or other basic estimates (e.g. regression coefficient) AND <u>variation</u> (e.g. standard deviation) or associated <u>estimates of uncertainty</u> (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>			
\times		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			
		Clearly defined error bars State explicitly what error bars represent (e.a. SD. SE. CI)			

Our web collection on <u>statistics for biologists</u> may be useful.

Software and code

Policy information about availability of computer code

Data collection

All software used is reported in the methods under the "Flow cytometry," "DNA sequencing and isolation," or "RNA sequencing" sections.

All software used is reported in the methods under the "Flow cytometry," "DNA sequencing and isolation," "RNA sequencing, differential gene expression analysis and enrichment," or "Statistical analysis" sections.

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/reviewers upon request. 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 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 relevant data are available from the authors. DNA sequencing data have been submitted under BioProject ID PRJNA353361. RNA sequencing data have been deposited in the Gene Expression Omnibus (GEO) database (accession number GSE115495).

Field-specific reporting				
Please select 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	he document with all sections, see nature.com/authors/policies/ReportingSummary-flat.pdf			
Life scier	nces study design			
All studies must dis	close on these points even when the disclosure is negative.			
Sample size	Sample sizes were chosen following empirical statistical power analysis based on previous pilot studies (Khader et al. 2007 Nat Immunol, Griffiths et al, 2016 etc).			
Data exclusions	One data point in Figure 1d and Supplementary Figure 1 (same mouse) was excluded based on Graphpad Prism's Grubb's outliers test (p<0.05)			
Replication	Experiments were replicated at least twice to ensure reproducibility. All attempts at replication were successful.			
Randomization	No method of randomization was used.			
Blinding	J.R-M. was blinded during the histological analyses.			
Materials & experimental systems Methods n/a Involved in the study n/a Involved in the study □ Unique biological materials □ ChIP-seq □ Antibodies □ Flow cytometry □ Eukaryotic cell lines □ MRI-based neuroimaging □ Palaeontology □ Animals and other organisms				
Human research participants				
Antibodies				
Antibodies used	All antibodies are described in the methods under "Flow cytometry" or "in vitro Mtb infection" sections.			
Validation	All antibodies were validated for the species and application by the manufacturer.			
Animals and other organisms				
Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research				
Laboratory animals The strain, sex, and age of all mice (Mus musculus) used in the study are reported in the methods in the "mice" section.				

Wild animals

Field-collected samples

The study did not involve wild animals.

The study did not involve field-collected samples.

Flow Cytometry

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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	Sample preparation is described in methods in the "Flow cytometry" section.			
Instrument	Becton Dickinson (BD) Fortessa flow cytometer using FACS Diva software, or the BD FACSJazz flow cytometer using FACS Sortware software (BD).			
Software	Flow cytometry experiments were analyzed using FlowJo (Tree Star Inc) and percentages were calculated using previously published gating strategies (Griffiths et al, Nat Comm 2016; Treerat et al, 2017, Mucosal Immunol)			
Cell population abundance	N/A			
Gating strategy	Reported in the methods in the "Flow cytometry" section. Percentages of different cell types were gated based on published gating strategies (Griffiths et al, 2016, Nature Comm; Treerat Et al, Mucosal Immunol 2017) and total cell numbers/lung were			

back calculated based on counted lung cell numbers.

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