nature portfolio

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Last updated by author(s):	20.08.2025

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>.

Statistics				
or all statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.				
y/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 <i>Give P values as exact values whenever suitable.</i>				
For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings				
For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes				
Estimates of effect sizes (e.g. Cohen's <i>d</i> , Pearson's <i>r</i>), 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>				
Data collection no software was used				
Data analysis Python (version 3.12.10), Scanpy (1.11.), Rsubread package (v1.22.2), DESeq2 R package (v1.26.0)				
or 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 eviewers. 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				

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

- For clinical datasets or third party data, please ensure that the statement adheres to our policy

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

Policy information about availability of data

- A description of any restrictions on data availability

Research involving human participants, their data, or biological materia

Clinical data

| Plants

Dual use research of concern

Field-specific re Please select the one below that is Life sciences Be Be Life sciences Stu Life sciences stu Sample size Sample size Sample sizes Sample sizes we statistical sample	Please specify the socially constructed or socially relevant categorization variable(s) used in your manuscript and explain why they were used. Please note that such variables should not be used as proxies for other socially constructed/relevant variables (for example, race or ethnicity should not be used as a proxy for socioeconomic status). Provide clear definitions of the relevant terms used, how they were provided (by the participants/respondents, the researchers, or third particily, and the method(s) used to classify people into the different categories (e.g. self-report, census or administrative data, social media data, etc.) Please provide details about how you controlled for confounding variables in your analyses. Describe the covariate-relevant population characteristics of the human research participants (e.g. age, genotypic information, past and current diagnosis and treatment categories). If you filled out the behavioural & social sciences study design questions and have nothing to add here, write "See above." Describe how participants were recruited. Outline any potential self-selection bias or other biases that may be present and how these are likely to impact results. Identify the organization(s) that approved the study protocol. Deval of the study protocol must also be provided in the manuscript. Porting Step be the study protocol must also be provided in the manuscript. Ecological, evolutionary & environmental sciences Ecological, evolutionary & environmental sciences all sections, see nature.com/documents/nc-reporting-summary-flat.pdf Light design points even when the disclosure is negative.	
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Data exclusions No data were ex	vere determined based on our prior published studies and our previous experience with similar experimental models; no formal ple size calculation was performed.	
	xcluded from the analysis.	
Replication Animal experime	experiment was performed once with four animals per group; results were consistent within each cohort.	
Randomization Randomization	mization was performed by assigning animals to experimental groups using a computer-generated random number list.	
Blinding For lung tumor	tumor studies, the investigators were blinded to allocation during experiments and outcome assessment.	
We require information from authors a	Decific materials, systems and methods about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material syour study. If you are not sure if a list item applies to your research, read the appropriate section before selecting a response.	
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Antibodies	n/a Involved in the study ☐ ChIP-seq	
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Antibodies

Antibodies used

Anti-I-A/I-E (clone M5/114.15.2) from BD Biosciences. Anti-CD4 (clone GK1.5) and anti-CD11b (clone M1/70) from BioLegend.

Validation

According to BD and Biolegend, these antibodies are routinely quality control tested by flow cytometric analysis.

Eukaryotic cell lines

Policy information about cell lines and Sex and Gender in Research

Cell line source(s)

GFP-MC38 were donated by Gerhard Christofor, Department of Biomedicine, University of Basel, Basel, Switzerland (females).

Tumor cell line culture KP1.9 were donated by Mikael Pittet, Ludwig Cancer Research, University of Geneva, Switzerland

(males).

Cell line authentication was performed before and after cell sorting by Short Tandem Repeat (STR) DNA genotype analysis Authentication

(Microsynth, Balgach, Switzerland).

Mycoplasma contamination

All cells used in this study were confirmed to be negative for mycoplasma.

Commonly misidentified lines (See ICLAC register)

Name any commonly misidentified cell lines used in the study and provide a rationale for their use.

Animals and other research organisms

Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research, and Sex and Gender in Research

Laboratory animals

6-8 weeks old C57BL/6J mice were used for the study. 6-8 old weeks Vav-Cre mice tdTomato, Spp1-IRES-tdTomato and Arg1-ires-YFP mice were used for macrophage isolation and OT-II mice for CD4 T cell isolation. Mice of both sex were used.

Wild animals

Provide details on animals observed in or captured in the field; report species and age where possible. Describe how animals were caught and transported and what happened to captive animals after the study (if killed, explain why and describe method; if released, say where and when) OR state that the study did not involve wild animals.

Reporting on sex

Indicate if findings apply to only one sex; describe whether sex was considered in study design, methods used for assigning sex. Provide data disaggregated for sex where this information has been collected in the source data as appropriate; provide overall numbers in this Reporting Summary. Please state if this information has not been collected. Report sex-based analyses where performed, justify reasons for lack of sex-based analysis.

Field-collected samples

For laboratory work with field-collected samples, describe all relevant parameters such as housing, maintenance, temperature, photoperiod and end-of-experiment protocol OR state that the study did not involve samples collected from the field.

Ethics oversight

All animal experiments were performed according to animal experimentation licenses as approved by the Swiss Federal Veterinary

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

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

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.

Flow Cytometry

Gating strategy

Plots

11000	
Confirm that:	
The axis labels state the m	narker 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 num	aber of cells or percentage (with statistics) is provided.
Methodology	
Sample preparation	BM cells were washed with PBS. Cells were preincubated with Mouse BD Fc Block™ (≤ 1 µg/million cells in 100 µl, BD Biosciences) at 4°C for 10 min before AB staining.
Instrument	Analysis was performed with a LSR II Fortessa 4L with HTS.
Software	The data were analyzed using FlowJo software (version 10.7.1).
Cell population abundance	no sort has been performed.

Cells were first gated based on forward scatter (FSC) and side scatter (SSC) parameters to exclude debris and aggregates and

to select the main cell population. Singlets were discriminated by FSC-A/FSC-H gating, and viability dye exclusion was applied where appropriate. Within this population, macrophages were identified as CD11b* cells, and expression of MHC-II was analyzed. The boundary between positive and negative staining populations was set using fluorescence minus one (FMO) controls and unstained samples. For proliferation assays, responder T cells were identified by CFSE staining, and successive

cell divisions were quantified as a progressive dilution of CFSE intensity.

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