nature research

Corresponding author(s):	Li Zhao
Last updated by author(s):	Feb 10, 2022

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 Editorial Policies and the Editorial Policy Checklist.

For all statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.

$\overline{}$					
Ç	+~	١+،	ist	10	c
J	LC	ı	IΣL	IL	2

n/a	Confirmed
	$oxed{x}$ 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
X	For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes
	x 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 statistics for biologists contains articles on many of the points above.

Software and code

Policy information about availability of computer code

Data collection

qRT-PCR data was collected by LightCycler 480 real-time PCR system (Roche); Flow cytometry was performed on LSRFortessa analyzer (Becton Dickinson); Fluorescent images were acquired using LeicaDMI3000B fluorescence microscopy.

Data analysis

Cluster 3.0 and DAVID 6.8 (https://david.ncifcrf.gov/tools.jsp) were used for analyses of RNA-seq data; FlowJo V10 was used for flow cytometry data; SPSS 17.0 and GraphPad Prism 8.0 were used for statistical analyses.

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.

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

Jaspar database(http://jaspar.genereg.net).

GEO datasets (GEO Accession Number: GSE126153, GSE126154, GSE161430, GSE75299 and GSE152699) and TCGA-THCA dataset available from tcga.org are used for analyses in this study.

The RNA-seq data generated in this study have been deposited in the Gene Expression Omnibus (GEO) under the accession GSE166513.

All data associated with this study are present in the paper or the Supplementary information. Source data are provided with this paper.

_					• (100	•
H	lel	C	-S	ре	CI1	TIC	re	po	rti	ıng

Please select the or	ne below that is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.					
x Life sciences	Behavioural & social sciences Ecological, evolutionary & environmental sciences					
For a reference copy of t	he document with all sections, see nature.com/documents/nr-reporting-summary-flat.pdf					
Life scier	nces study design					
All studies must dis	close on these points even when the disclosure is negative.					
Sample size	The sample size for each experiment is specified in the Methods section and figure legends. The sample size was determined by relevant published research or previous experience with similar type of experiments (Saqcena, M. et al. Cancer Discov (2020). Li, X. M. et al. Oncogene 37, 2773-2792 (2018).) to confirm a result with statistically significant.					
Data exclusions	For tumor formation experiments and inhibitor treatment in mouse assays (Fig. 5e, Fig.6g-h and Fig. 7i), the highest and lowest tumor of each group were rexcluded from analysis according to the pre-established exclusion criteria.					
Replication	For each experiment, the number of biological replicates is reported in the figure legend.					
Randomization	All cell samples were randomly allocated to experimental and control groups. In each mice experimental series, animals were assigned to various groups in random according to planned age and weight.					
Blinding	In each mice experimental series, the administration of treatments and the subsequent analysis were performed by different operator. The immunohistochemical analysis of the PTC was finished by two pathologists who did not know each other. For the other cell experiments, two investigators performed with different cell lines, respectively. And data analysis was blinded to the experimental conditions.					

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 & experimental systems		Methods		
n/a	Involved in the study		Involved in the study	
	x Antibodies	x	ChIP-seq	
	x Eukaryotic cell lines		x Flow cytometry	
x	Palaeontology and archaeology	x	MRI-based neuroimaging	
	X Animals and other organisms			
	Human research participants			
x	Clinical data			
×	Dual use research of concern			

Antibodies

Antibodies used

All antibodies used in the study are reported in Methods.

TBX3 Polyclonal Antibody Invitrogen Cat# 42-4800

Monoclonal Anti-a-Tubulin Simga Cat# T5168 Clone B-5-1-2

B-Raf (D9T6S) Rabbit mAb CST Cat# 14814 Clone D9T6S

c-Jun (60A8) Rabbit mAb CST Cat# 9165 Clone 60A8

p44/42 MAPK (Erk1/2) (137F5) Rabbit mAb CST Cat# 4695 Clone 137F5

Phospho-p44/42 MAPK (Erk1/2) (Thr202/Tyr204) (D13.14.4E) XP® Rabbit mAb CST Cat# 4370 Clone D13.14.4E

FosB (5G4) Rabbit mAb CST Cat# 2251 Clone 5G4 $\,$

JunB (C37F9) Rabbit mAb CST Cat# 3753 Clone C37F9

c-Fos (9F6) Rabbit mAb CST Cat# 2250 Clone 9F6

GAPDH (D16H11) XP® Rabbit mAb CST Cat# 5174 Clone D16H11

 β -Actin (8H10D10) Mouse mAb CST Cat# 3700 Clone 8H10D10

NF-κB p65 (D14E12) XP® Rabbit mAb CST Cat# 8242 Clone D14E12

Phospho-NF-κB p65 (Ser536) (93H1) Rabbit mAb CST Cat# 3033 Clone 93H1

IKKβ (D30C6) Rabbit mAb CST Cat# 8943 Clone D30C6

Phospho-IKK α/β (Ser176/180) (16A6) Rabbit mAb CST Cat# 2697 Clone 16A6

TLR2 Polyclonal Antibody Bioss Cat# bs-1019R

```
Akt (pan) (C67E7) Rabbit mAb CST Cat# 4691 Clone C67E7
```

Phospho-Akt (Ser473) (D9E) XP® Rabbit mAb CST Cat# 4060 Clone D9E

p38 MAPK (D13E1) XP® Rabbit mAb CST Cat# 8690 Clone D13E1

Phospho-p38 MAPK (Thr180/Tyr182) (D3F9) XP® Rabbit mAb CST Cat# 4511 Clone D3F9

SAPK/JNK Antibody CST Cat# 9252

Phospho-SAPK/JNK (Thr183/Tyr185) (G9) Mouse mAb CST Cat# 9255 Clone G9

Stat3 (124H6) Mouse mAb CST Cat# 9139 Clone 124H6

Phospho-Stat3 (Tyr705) (D3A7) XP® Rabbit mAb CST Cat# 9145 Clone D3A7

Anti-Ki67 antibody Abcam Cat# ab16667 Clone SP6

CXCL1/GRO alpha antibody GeneTex Cat# GTX31184

CXCL2/GRO beta antibody GeneTex Cat# GTX31171

Anti-Thyroglobulin antibody Abcam Cat# ab156008 Clone EPR9730

TBX3 Polyclonal Antibody Proteintech Cat# 16741-1-AP

CD8α (D4W2Z) XP® Rabbit mAb (Mouse Specific) CST Cat# 98941 Clone D4W2Z

Anti-CD4 antibody Abcam Cat# ab183685 Clone EPR19514

Anti- MRP8 antibody Abcam Cat# ab92331 Clone EPR3554

Purified anti-mouse Ly-6G/Ly-6C (Gr-1) Antibody Biolegend Cat# 108401 Clone RB6-8C5

Anti-CD11b antibody Abcam Cat# ab133357 Clone EPR1344

IL-8 Polyclonal Antibody ImmunoWay Cat# T5153

Alexa Fluor® 700 anti-mouse CD45 Antibody Biolegend Cat# 147716 Clone I3/2.3

FITC anti-mouse CD45 Antibody Biolegend Cat# 103108 Clone 30-F11

PE anti-mouse F4/80 Antibody Biolegend Cat# 123110 Clone BM8

Pacific Blue™ anti-mouse/human CD11b Antibody Biolegend Cat# 101224 Clone M1/70

FITC anti-mouse CD3ε Antibody Biolegend Cat# 100306 Clone 145-2C11

PerCP/Cyanine5.5 anti-mouse CD3ɛ Antibody Biolegend Cat# 100328 Clone 145-2C11

APC anti-mouse NK-1.1 Antibody Biolegend Cat# 108709 Clone PK136

Brilliant Violet 605™ anti-mouse CD4 Antibody Biolegend Cat# 100451 Clone GK1.5

Alexa Fluor® 700 anti-mouse CD4 Antibody Biolegend Cat# 100430 Clone GK1.5

Brilliant Violet 510™ anti-mouse CD8a Antibody Biolegend Cat# 100752 Clone 53-6.7

PerCP/Cyanine5.5 anti-mouse Ly-6G/Ly-6C (Gr-1) Antibody Biolegend Cat# 108428 Clone RB6-8C5

Brilliant Violet 605™ anti-mouse Ly-6G Antibody Biolegend Cat# 127639 Clone 1A8

Brilliant Violet 510™ anti-mouse Ly-6C Antibody Biolegend Cat# 128033 Clone HK1.4

PE/Dazzle™ 594 anti-mouse CD11c Antibody Biolegend Cat# 117347 Clone N418

PE/Cyanine7 anti-mouse CD19 Antibody Biolegend Cat# 115519 Clone 6D5

Purified Rat Anti-Mouse CD16/CD32 (Mouse BD Fc Block™) Becton Dickinson Cat# 553141 Clone 2.4G2

PE-Cy7 anti-Ly-6C Antibody Biolegend Cat# 128017 Clone HK1.4

APC anti-CD115 Antibody Biolegend Cat# 135509 Clone AFS98 $\,$

PE anti-CD34 Antibody Biolegend Cat# 152203 Clone SA376A4

PE-CF594 anti-CD135 Antibody Biolegend Cat# 313319 Clone BV10A4H2

BV421 anti-CD117 Antibody Biolegend Cat# 105827 Clone 2B8

FITC anti-CD16/32 Antibody Biolegend Cat# 101305 Clone 93

AF-700 anti-Sca-1 Antibody Biolegend Cat# 108141 Clone D7
Percp-cy5.5 anti-Ter-119 Antibody Biolegend Cat# 116227 Clone TER-119

APC anti-IFN-γ Antibody Biolegend Cat# 505809 Clone XMG1.2

PE anti- Granzyme B Antibody Biolegend Cat# 96405 Clone QA18A28

Anti-Digoxigenin-AP antibody Roche Cat# 11093274910

InVivoMAb rat IgG2a isotype control, anti-trinitrophenol Bio-XCell Cat# BE0089 Clone 2A3

InVivoMAb anti-mouse Ly6G/Ly6C (Gr-1) Bio-XCell Cat# BE0075 Clone RB6-8C5

InVivoMAb anti-mouse CD8 Bio-XCell Cat# BE0061 Clone 2.43

All antibodies were obtained from commercial sources with reported validation, and we confirmed with negative and positive control before the study.

Abbreviation for species cross reactivity: H-human, M-mouse, R-rat, Hm-hamster, Mk-monkey, Vir-virus, Mi-mink, C-chicken, Dm-Drosophila melanogaster, X-xenopus, Z-zebrafish, B-bovine, Dg-dog, PG-pig, Sc-Saccharomyces cerevisiae, Ce-Caenorhabditis elegans, Hr-horse, K-Kangaroo, Su-Sea urchin, Ch-Chlamydomonas, S-Sheep

TBX3 Polyclonal Antibody Invitrogen Cat# 42-4800 H, M, R PMID: 27110270

Monoclonal Anti-a-Tubulin Simga Cat# T5168 Clone B-5-1-2 C, K, R, SU, R, CH, B, H, MK, M PMID: 31523176

B-Raf (D9T6S) Rabbit mAb CST Cat# 14814 Clone D9T6S H, M, R, M PMID: 34992144

c-Jun (60A8) Rabbit mAb CST Cat# 9165 Clone 60A8 H, M, R, M PMID: 35027468

p44/42 MAPK (Erk1/2) (137F5) Rabbit mAb CST Cat# 4695 Clone 137F5 H, M, R, Hm, Mk, Mi, Dm, Z, B, Dg, Pg, Ce PMID: 35028009 Phospho-p44/42 MAPK (Erk1/2) (Thr202/Tyr204) (D13.14.4E) XP® Rabbit mAb CST Cat# 4370 Clone D13.14.4E H, M, R, Hm, Mk, Mi, Dm, Z, B, Dg, Pg, Sc PMID: 34965411

FosB (5G4) Rabbit mAb CST Cat# 2251 Clone 5G4 H, M, R PMID: 34352786

JunB (C37F9) Rabbit mAb CST Cat# 3753 Clone C37F9 H, M, R, M PMID: 34744629

c-Fos (9F6) Rabbit mAb CST Cat# 2250 Clone 9F6 H, M, R PMID: 34673574

GAPDH (D16H11) XP* Rabbit mAb CST Cat# 5174 Clone D16H11 H, M, R, Mk PMID: 34955649

 $\beta\text{-Actin}$ (8H10D10) Mouse mAb CST Cat# 3700 Clone 8H10D10 H, M, R, Hm, Mk, Dg PMID: 34977268

NF-ĸB p65 (D14E12) XP® Rabbit mAb CST Cat# 8242 Clone D14E12 H, M, R, Hm, Mk, Dg PMID: 34558536

Validation

Phospho-NF-кВ p65 (Ser536) (93H1) Rabbit mAb CST Cat# 3033 Clone 93H1 H, M, R, Hm, Mk, Pg PMID: 34901529

IKKβ (D30C6) Rabbit mAb CST Cat# 8943 Clone D30C6 H, M, R, Hm, Mk PMID: 33971703

Phospho-IKKα/β (Ser176/180) (16A6) Rabbit mAb CST Cat# 2697 Clone 16A6 H, M, R, Hm, Mk PMID: 35002723

TLR2 Polyclonal Antibody Bioss Cat# bs-1019R H, M, R, B, Pg, S PMID: 23295061

Akt (pan) (C67E7) Rabbit mAb CST Cat# 4691 Clone C67E7 H, M, R, Mk, Dm, PMID: 35005567

Phospho-Akt (Ser473) (D9E) XP® Rabbit mAb CST Cat# 4060 Clone D9E H, M, R, Hm, Mk, Dm, Z, B PMID: 35024764

p38 MAPK (D13E1) XP® Rabbit mAb CST Cat# 8690 Clone D13E1 H, M, R, Hm, Mk, B, Pg PMID: 34935053

Phospho-p38 MAPK (Thr180/Tyr182) (D3F9) XP® Rabbit mAb CST Cat# 4511 Clone D3F9 H, M, R, Mk, Mi, Pg, Sc PMID: 34935053

SAPK/JNK Antibody CST Cat# 9252 H, M, R, Hm, Mk, Z, B, Sc PMID: 35027468

Phospho-SAPK/JNK (Thr183/Tyr185) (G9) Mouse mAb CST Cat# 9255 Clone G9 H, M, R, Hm, Sc PMID: 35028009

Stat3 (124H6) Mouse mAb CST Cat# 9139 Clone 124H6 H, M, R, Mk PMID: 35020440

Phospho-Stat3 (Tyr705) (D3A7) XP® Rabbit mAb CST Cat# 9145 Clone D3A7 H, M, R, Mk PMID: 34938610

Anti-Ki67 antibody Abcam Cat# ab16667 Clone SP6 H, M, R PMID: 34284046

CXCL1/GRO alpha antibody GeneTex Cat# GTX31184 validated in the manufacturers' websites.

CXCL2/GRO beta antibody GeneTex Cat# GTX31171 validated in the manufacturers' websites.

Anti-Thyroglobulin antibody Abcam Cat# ab156008 Clone EPR9730 H, M, R PMID: 33162555

TBX3 Polyclonal Antibody Proteintech Cat# 16741-1-AP H, M PMID: 29620145

CD8α (D4W2Z) XP® Rabbit mAb (Mouse Specific) CST Cat# 98941 Clone D4W2Z M PMID: 34952899

Anti-CD4 antibody Abcam Cat# ab183685 Clone EPR19514 M PMID: 32401602

Anti- MRP8 antibody Abcam Cat# ab92331 Clone EPR3554 H, M PMID: 33301706

Purified anti-mouse Ly-6G/Ly-6C (Gr-1) Antibody Biolegend Cat# 108401 Clone RB6-8C5 M PMID: 22965162

Anti-CD11b antibody Abcam Cat# ab133357 Clone EPR1344 H, M, R PMID: 33283987

IL-8 Polyclonal Antibody ImmunoWay Cat# T5153 H PMID: 28033248

Anti-Digoxigenin-AP antibody Roche Cat# 11093274910 PMID: 24318810

InVivoMAb rat IgG2a isotype control, anti-trinitrophenol Bio-XCell Cat# BE0089 Clone 2A3 PMID: 30097293

InVivoMAb anti-mouse Ly6G/Ly6C (Gr-1) Bio-XCell Cat# BE0075 Clone RB6-8C5 PMID: 29311363

InVivoMAb anti-mouse CD8 Bio-XCell Cat# BE0061 Clone 2.43 PMID: 27775706

The following antibodies for Flow Cytometry were all validated in the manufacturers' websites with a large number of research.

Alexa Fluor® 700 anti-mouse CD45 Antibody Biolegend Cat# 147716 Clone I3/2.3

FITC anti-mouse CD45 Antibody Biolegend Cat# 103108 Clone 30-F11

PE anti-mouse F4/80 Antibody Biolegend Cat# 123110 Clone BM8

Pacific Blue™ anti-mouse/human CD11b Antibody Biolegend Cat# 101224 Clone M1/70

FITC anti-mouse CD3ε Antibody Biolegend Cat# 100306 Clone 145-2C11

PerCP/Cyanine5.5 anti-mouse CD3 EAntibody Biolegend Cat# 100328 Clone 145-2C11

APC anti-mouse NK-1.1 Antibody Biolegend Cat# 108709 Clone PK136

Brilliant Violet 605™ anti-mouse CD4 Antibody Biolegend Cat# 100451 Clone GK1.5

Alexa Fluor® 700 anti-mouse CD4 Antibody Biolegend Cat# 100430 Clone GK1.5

Brilliant Violet 510™ anti-mouse CD8a Antibody Biolegend Cat# 100752 Clone 53-6.7

PerCP/Cyanine5.5 anti-mouse Ly-6G/Ly-6C (Gr-1) Antibody Biolegend Cat# 108428 Clone RB6-8C5

Brilliant Violet 605™ anti-mouse Ly-6G Antibody Biolegend Cat# 127639 Clone 1A8

Brilliant Violet 510™ anti-mouse Ly-6C Antibody Biolegend Cat# 128033 Clone HK1.4

PE/Dazzle™ 594 anti-mouse CD11c Antibody Biolegend Cat# 117347 Clone N418

PE/Cyanine7 anti-mouse CD19 Antibody Biolegend Cat# 115519 Clone 6D5

Purified Rat Anti-Mouse CD16/CD32 (Mouse BD Fc Block™) Becton Dickinson Cat# 553141 Clone 2.4G2

PE-Cy7 anti-Ly-6C Antibody Biolegend Cat# 128017 Clone HK1.4 $\,$

APC anti-CD115 Antibody Biolegend Cat# 135509 Clone AFS98

PE anti-CD34 Antibody Biolegend Cat# 152203 Clone SA376A4

PE-CF594 anti-CD135 Antibody Biolegend Cat# 313319 Clone BV10A4H2

BV421 anti-CD117 Antibody Biolegend Cat# 105827 Clone 2B8

FITC anti-CD16/32 Antibody Biolegend Cat# 101305 Clone 93

AF-700 anti-Sca-1 Antibody Biolegend Cat# 108141 Clone D7

Percp-cy5.5 anti-Ter-119 Antibody Biolegend Cat# 116227 Clone TER-119

APC anti-IFN-γ Antibody Biolegend Cat# 505809 Clone XMG1.2

PE anti- Granzyme B Antibody Biolegend Cat# 396405 Clone QA18A28.

Eukaryotic cell lines

Policy information about <u>cell lines</u>

HEK293T (ACS-4500) was purchased from the American Type Culture Collection (ATCC). And all cancer cell lines were obtained from Tianjin Medical University Cancer Institute and Hospital with STR profiling.

Authentication

Cell line source(s)

Cell lines have been confirmed with STR Authentication.

Mycoplasma contamination

All cell lines have tested for mycoplasma and found to be negative

Commonly misidentified lines (See ICLAC register)

We did not use commonly misidentified lines.

4

Animals and other organisms

Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research

Laboratory animals

And BALB/c nude mouse of female at 5-6w at the beginning of study. C57BL/6 strain including Tpo-cre, LSL-BrafV600ECA, Tbx3flox/flox mice, Tbx3-GFP, TPO-creER, Rosa-mTmG of both sexes at 4w-12m. All mice were maintained under SPF-condition at normal

room temperature with a 12/12 h light/dark cycle and normal ambient humidity.

Wild animals No wild animals were used.

Field-collected samples The study did not involve field-collected samples.

Ethics oversight

All mouse experiment procedures and protocols were evaluated and authorized by the Regulations of Tianjin Laboratory Animal

Management and strictly followed the guidelines under the Institutional Animal Care and Use Committee of Tianjin Medical

University.

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

Human research participants

Policy information about studies involving human research participants

Population characteristics PTC samples were collected from (1) 10-30 years old including 3 men and 19 women; (2) 30-50 years old including 16 men

and 59 women; and (3) >50 years old including 5 men and 38 women. Fresh samples from adjacent normal tissues, pathological grade I, II, III and IV were frozen in liquid nitrogen immediately after resection or fixed in 4% paraformaldehyde (Sigma Aldrich) at $4^{\circ}C$ overnight before embedded in paraffin. Patients with a past history of radiation therapy or chemotherapy were excluded from this study. The healthy volunteers used in neutrophil chemotaxis assays were 25-30 years

old including 2 men and 2 women.

Recruitment Participants were recruited by the doctors participating in the study at Tianjin cancer institute and hospital

(Tianjin, China). Tumor tissue samples were collected according to standard clinical procedures. The study did not look at the results of individual patients, nor did the doctors or patients involved in the study rule out bias. In addition, the healthy volunteers in this study were recruited with normal certificate of physical examination and no selection bias was observed.

Ethics oversight

All PTC samples were obtained from Tianjin cancer institute and hospital, and the patients signed an informed consent form issued by the 'Ethics Committee of Tianjin Cancer Institute and Hospital'. In addition, healthy volunteers signed informed

consent issued by the 'Ethics Committee of Tianjin Cancer Institute and Hospital'. In addition, healthy volunteers signed inform

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

Flow Cytometry

Plots

Confirm that:

- $m{x}$ 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 Mouse PTC tumors were minced into small fragments on ice and then separated into single cells with tissue dissociation

buffer including 1mg/ml collagenase I (Sigma) and 0.5 mg/ml dispase II (Invitrogen) dissolved in PBS at 37°C for 60-90min. Samples were mashed through 40µm filters into FACS buffer (1% FBS in PBS) and washed for three times. The single cells were stained with Zombie NIR™ Fixable Viability Kit (#423105, Biolegend) for 15min in dark on ice, and then incubated with

anti-CD16/CD32 (#553141, Becton Dickinson) for blocking and other indicated antibodies for 30 min on ice.

Instrument LSRFortessa analyzer (Becton Dickinson).

Software FlowJo V10 Software.

Cell population abundance The purity of the population after sort was more than 95%.

Gating strategy First eliminate duplex, gate on size and granularity to narrow down specific populations using FSC/SSC; and then live/dead

discrimination with NIR signal; finally, set gates with indicated staining.

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