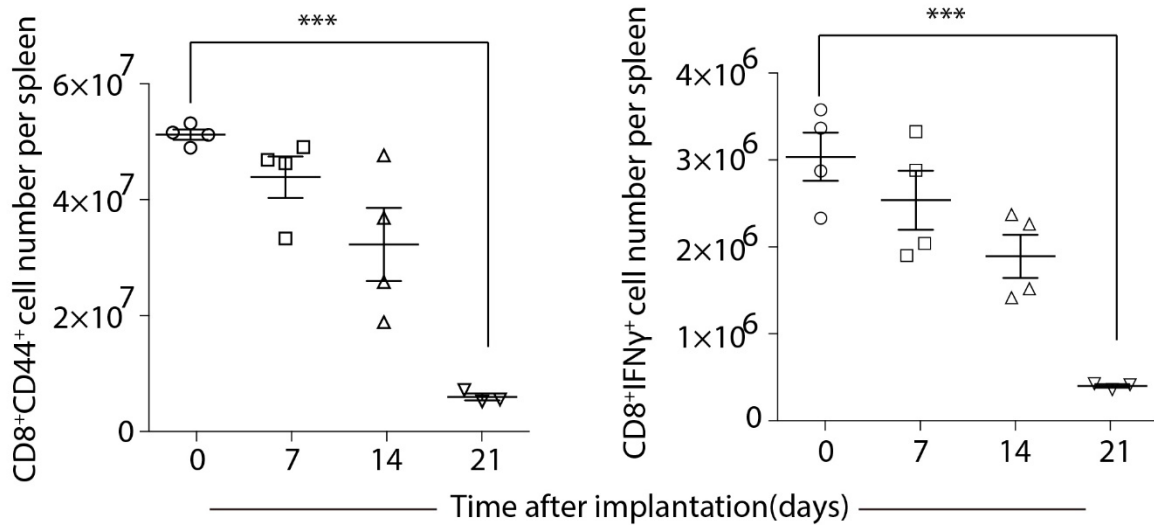
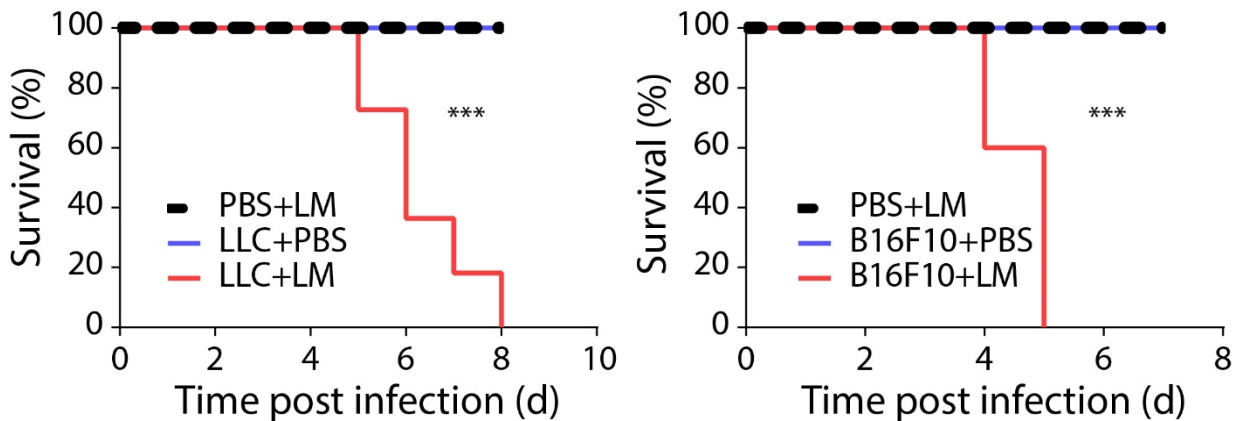


Supplementary Figures and Legends

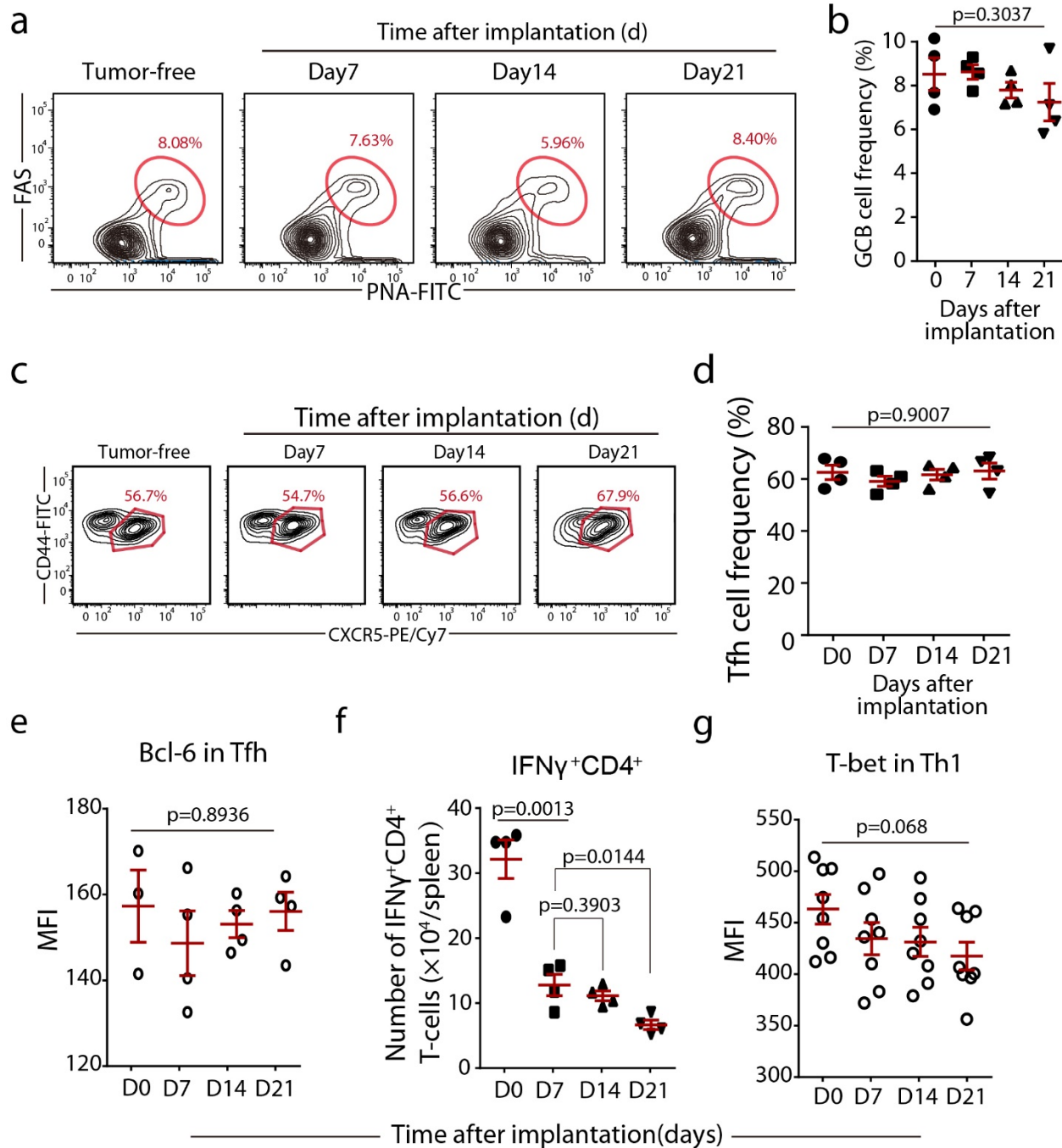
a



b

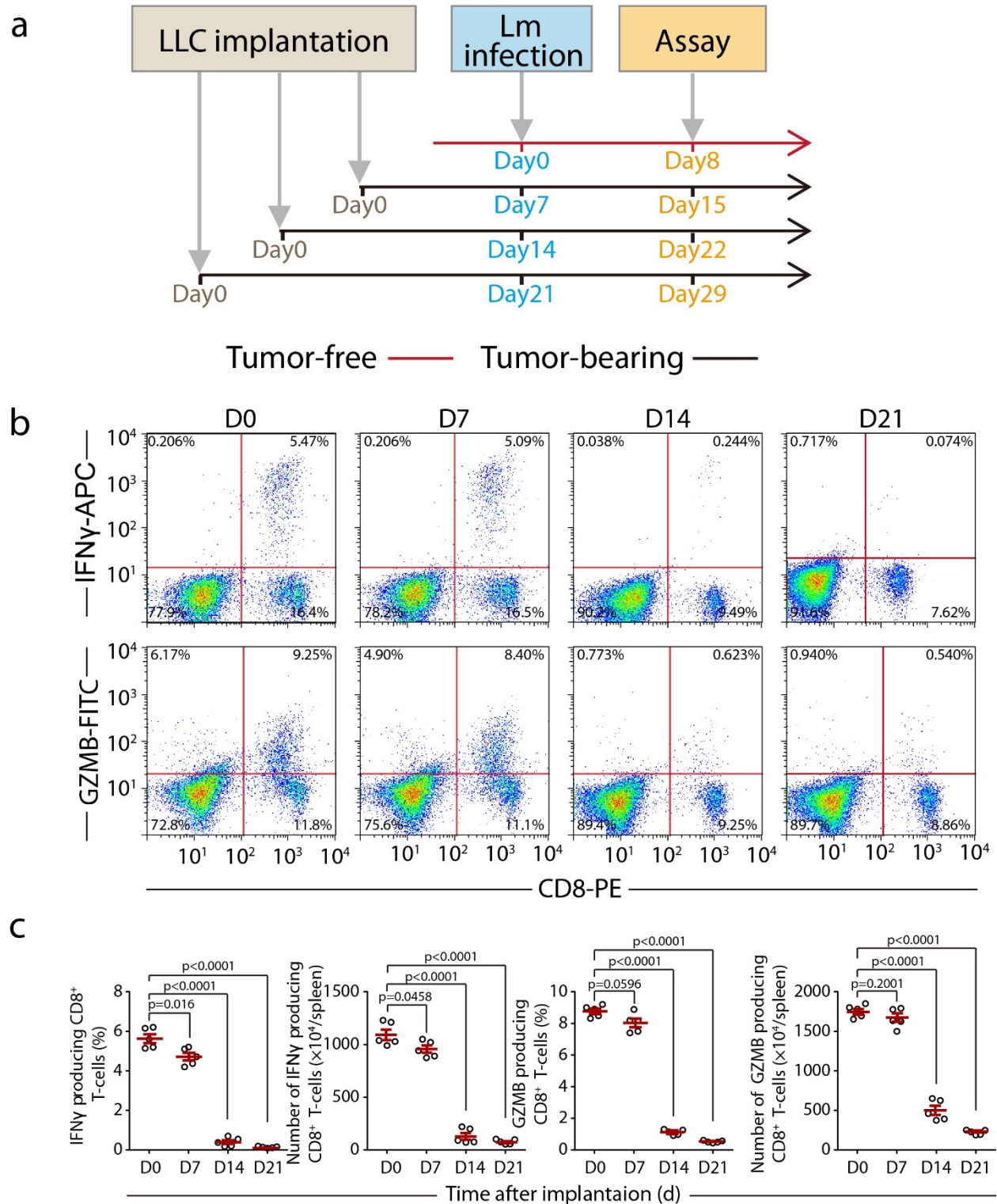


Supplementary Fig. 1. Reduced CD8⁺ T cell response and survival of tumor-bearing mice following viral infection. **a**, Mice were infected with LCMV-Armstrong at different time points following LLC inoculation (0, 7, 14 and 21 days) and sacrificed on day 8 post-infection. The total numbers of CD8⁺CD44⁺ T cells and IFN-γ producing CD8⁺ T cells stimulated with viral antigen in the spleens were analyzed. **b**, 14 days after LLC cell inoculation, or 10 days after B16F10 melanoma cell inoculation, tumor-bearing mice (n=10) were infected with 2 × 10⁵ CFU rLmOVA. Infected, non-tumor-bearing mice (n=10) and uninfected tumor-bearing mice (n=10) were used as controls. Mouse survival time was analyzed and plotted.



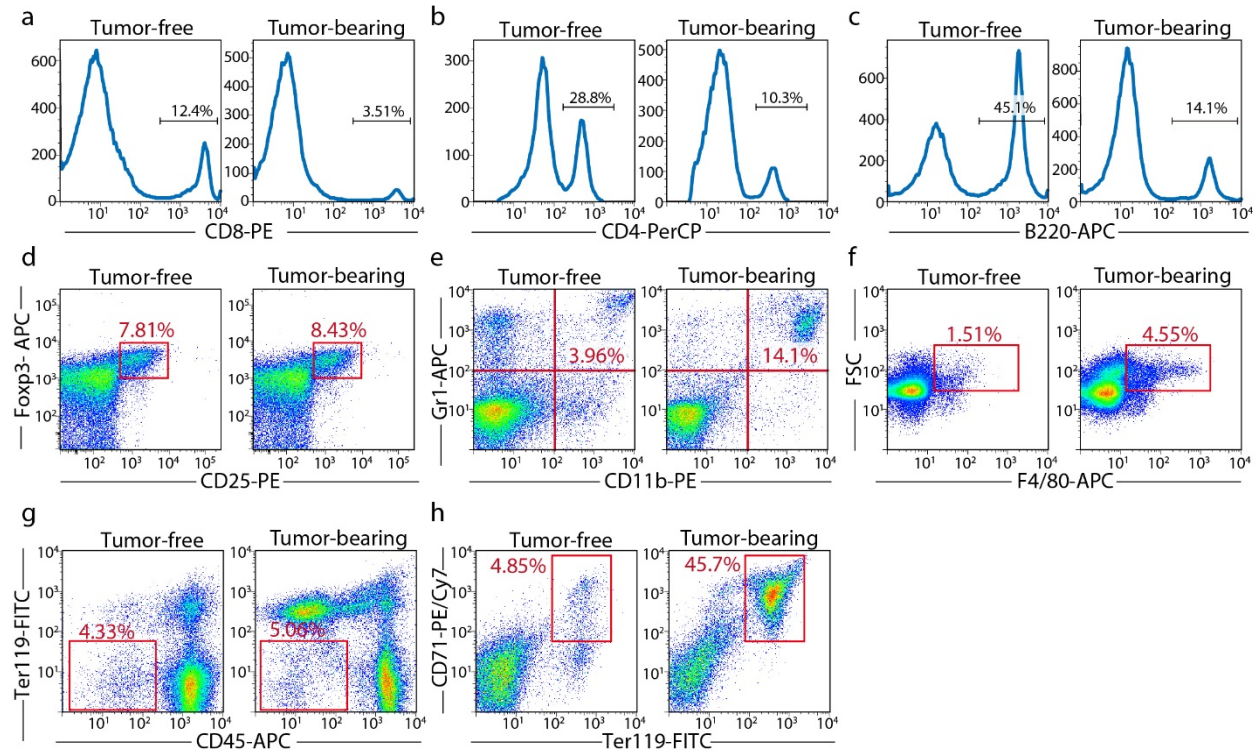
Supplementary Fig. 2. Infection-induced differentiation of germinal center B cells and Tfh cells is not affected in tumor-bearing mice. **a-b**, Mice were infected with LCMV-Cl13 at the indicated times after LLC inoculation and sacrificed on day 8 post-infection. Representative flow cytometry plots (**a**) and cumulative composite data (**b**) show the frequency of germinal center B cells (GCB, gated on B220) in the spleens of LCMV CL13-infected tumor-bearing mice at different time points (0, 7, 14 and 21 days, $n=4$) after LLC inoculation. **c-d**, Representative flow cytometry plots (**c**) and cumulative composite data (**d**) show Tfh cell differentiation cells in spleen ($n=4$). **e**, Tfh cell Bcl-6 expression in the spleens of tumor-bearing mice at the indicated time points ($n=3$ or 4) was detected by flow cytometry. **f**, Splenocytes were collected from mice

at the indicated times (n=4) after tumor inoculation and 8 days after LCMV infection. Cells were stimulated in vitro with GP a.a.61-80 peptide, and the number of antigen-specific CD4⁺ T cells was determined by intracellular IFN- γ staining and flow cytometry. **g**, T-bet expression in Th1 cells in the spleens of tumor-bearing mice at the indicated time points (n=8) was determined by flow cytometry. Data are representative of at least three independent experiments. Two-tailed Student's t-tests were used for all analyses. Error bars represent the means \pm SEM.

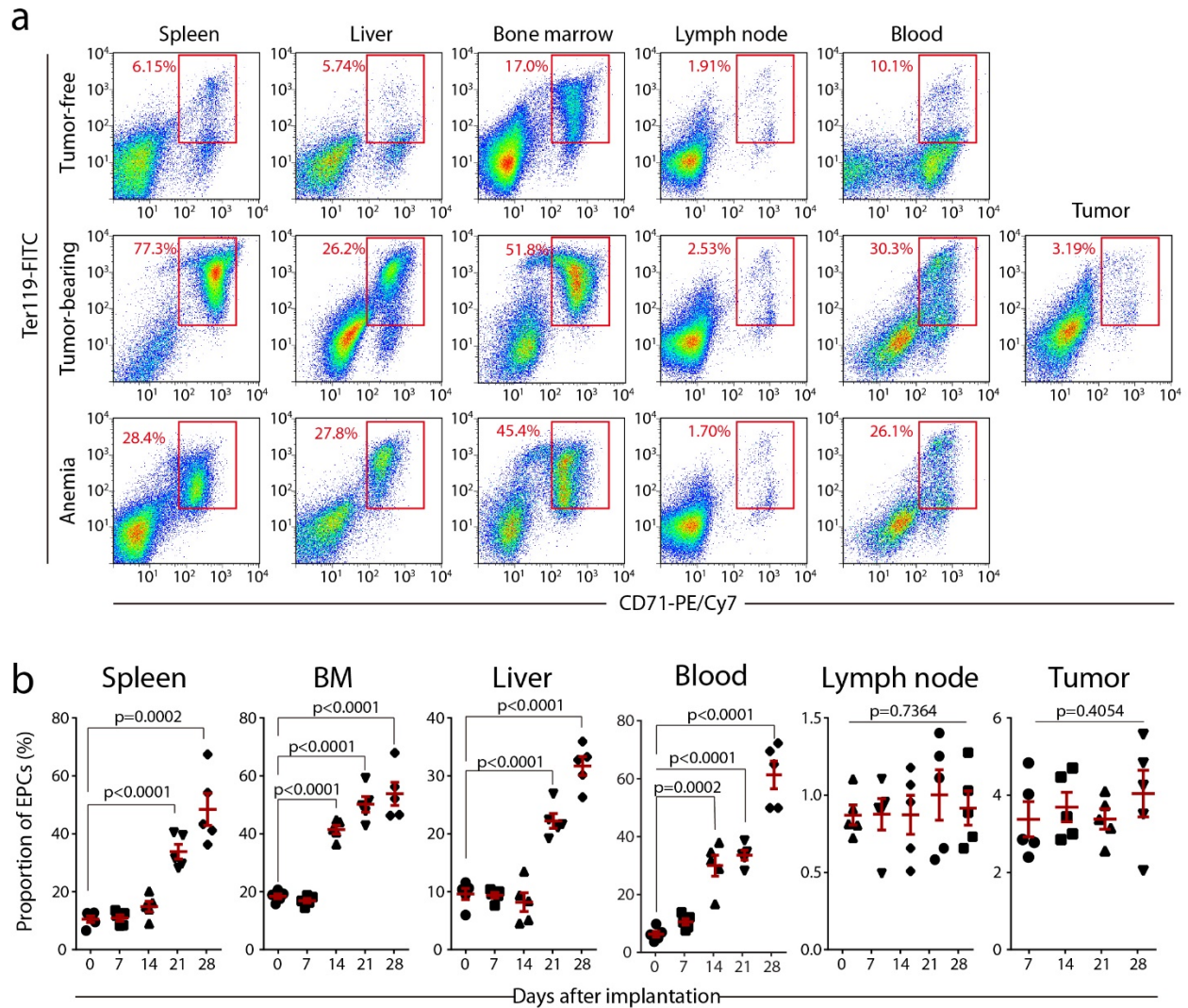


Supplementary Fig. 3. Impaired anti-bacterial response of CD8⁺ T cells in mice with established tumors. **a**, Schematic of experimental procedure. Mice were infected with 4×10^6 CFU Δ actArLmOVA at different time points (0, 7, 14 and 21 days, n=5) after LLC inoculation and sacrificed on day 7 post-infection. **b**, Antigen-specific CD8⁺ T cells were quantified by

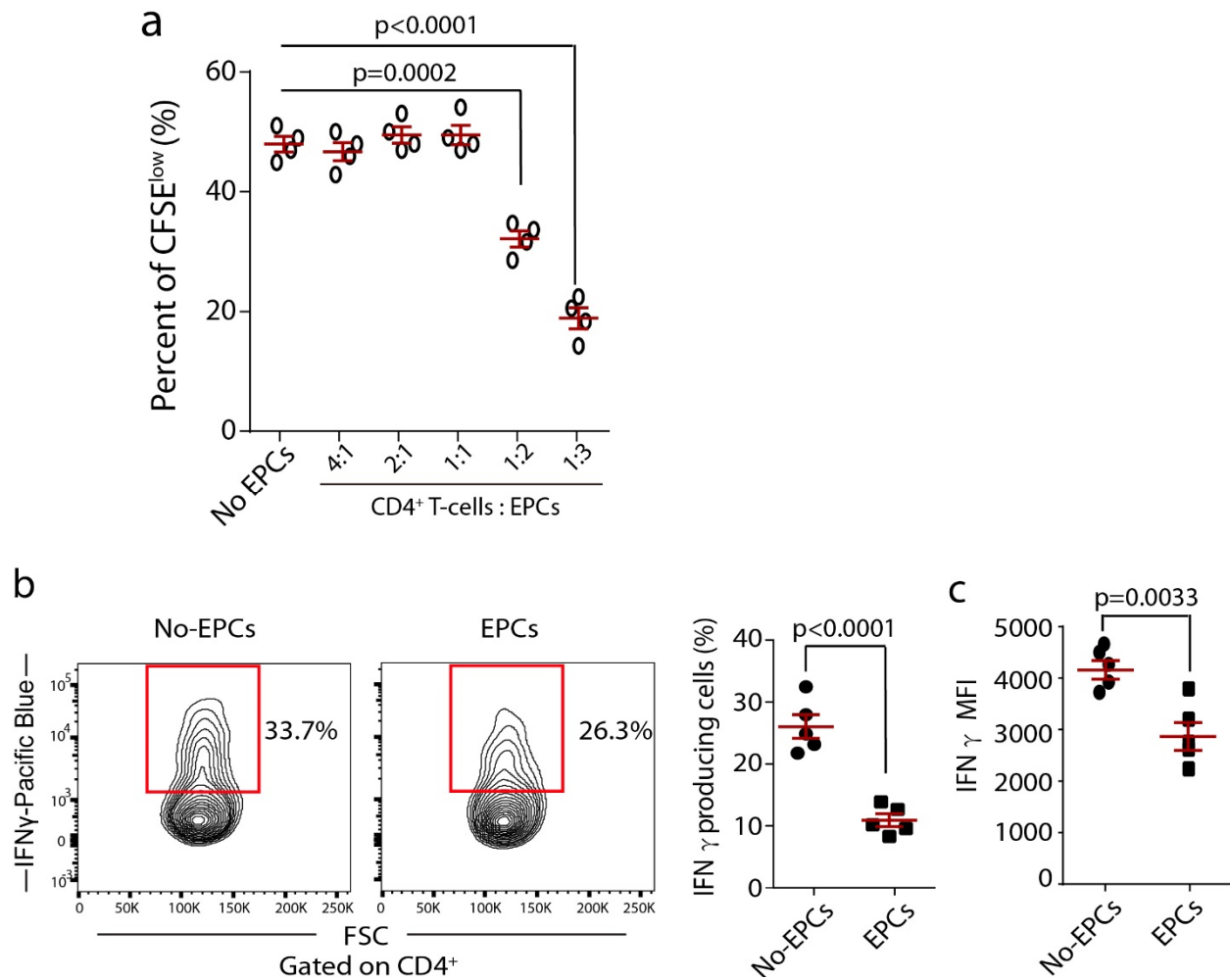
staining for intracellular IFN- γ and GzmB using flow cytometry after stimulating splenocytes with a specific OVA peptide in vitro. **c**, Frequency and absolute number of antigen-specific CD8⁺ T cells in spleens of tumor-bearing mice. Data are representative of three independent experiments. Two-tailed Student's t-tests were used for all analyses, with the exception of survival curves, for which Gehan-Breslow-Wilcoxon tests were used.



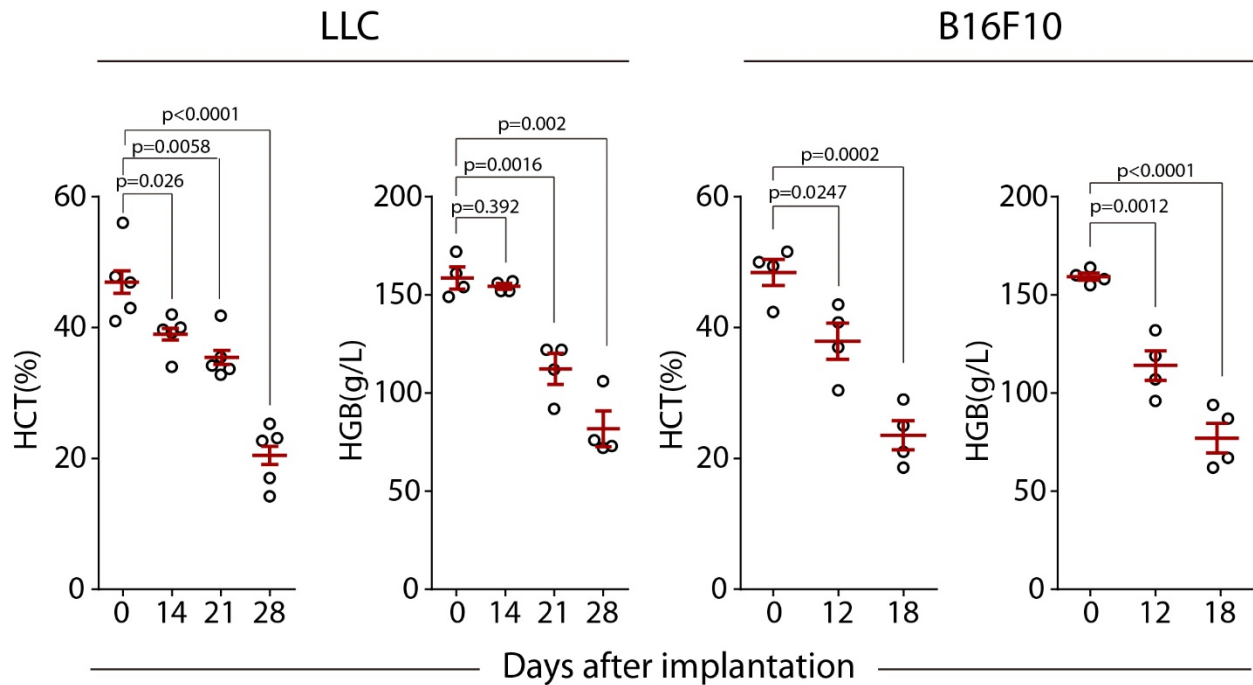
Supplementary Fig. 4. Splenic accumulation of EPCs in tumor-bearing mice. C57BL/6 mice were sacrificed 28 days after LLC cell inoculation, and different cell populations in the spleen were analyzed by flow cytometry. Frequencies of CD8⁺ T (**a**), CD4⁺ T (**b**), B cells (**c**), Tregs (**d**) MDSCs (CD11b⁺Gr1⁺, **e**), macrophages (F4/80⁺, **f**), stromal cells (CD45⁻TER119⁻, **g**), and EPCs (CD71⁺TER119⁺, **h**) are shown. Data are representative of at least three independent experiments.



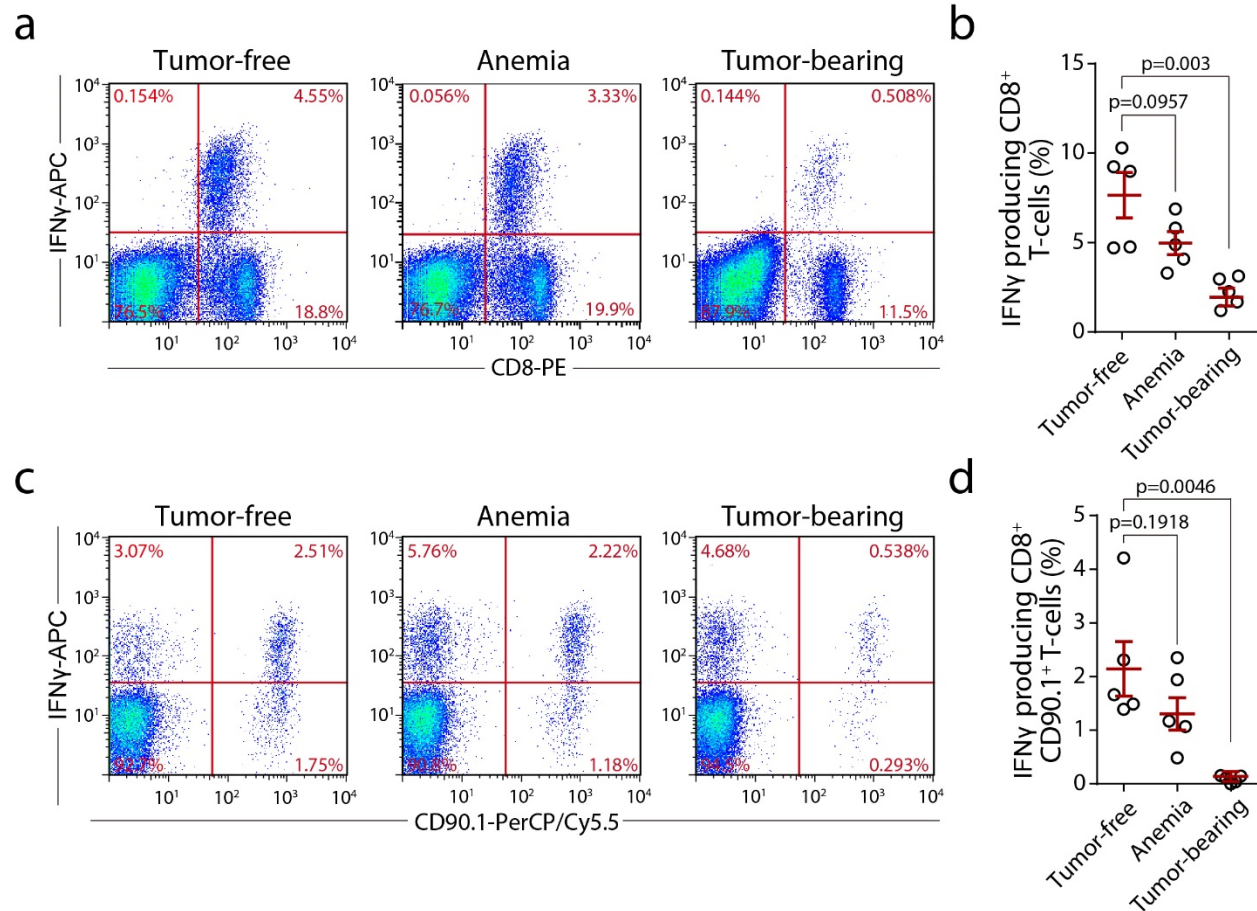
Supplementary Fig. 5. EPC accumulation in tissues of tumor-bearing and anemic mice. a, Representative flow cytometry results showing the frequency of CD71⁺TER119⁺ cells in different tissues from control (n=5), tumor-bearing (n=5) and anemic (n=5) mice. **b,** Cumulative composite flow cytometry data show the frequency of CD71⁺TER119⁺ cells in different tissues of tumor-bearing mice at different time points (n=5) after LLC cell inoculation.



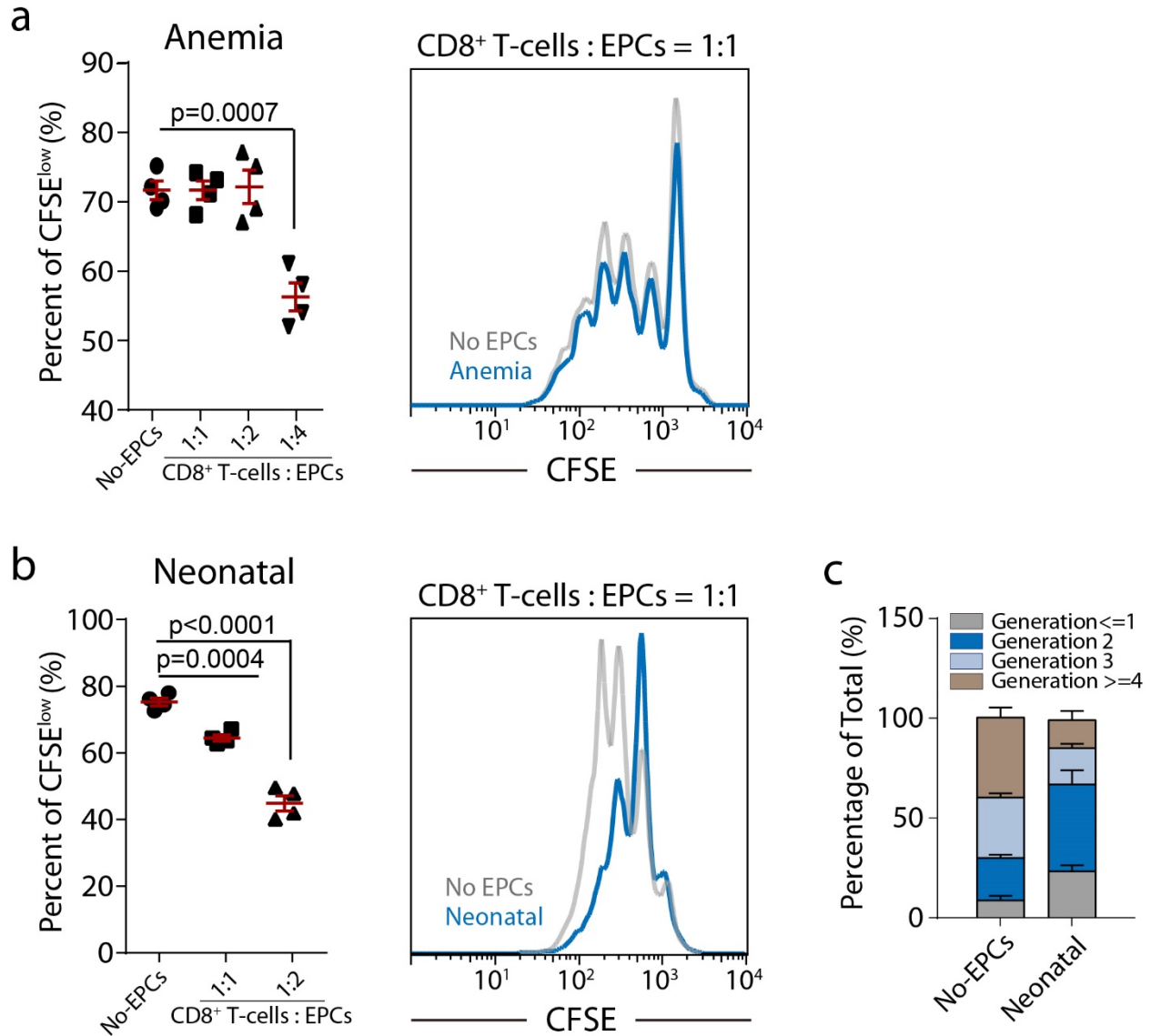
Supplementary Fig. 6. CD71⁺TER119⁺ EPCs inhibit CD4⁺ T cell proliferation and differentiation. **a**, CD71⁺TER119⁺ EPCs isolated from the spleens of tumor-bearing mice were co-cultured with anti-CD3 and anti-CD28 antibodies, and stimulated CFSE-labeled CD4⁺ T cells at different CD4⁺ T cell:EPC ratios. Cumulative composite data show the proliferation of CFSE-labeled CD4⁺ T cells(n=4). **b**, CD4⁺ T cells purified from wild-type mice were cocultured with CD71⁺TER119⁺ cells (CD4⁺ T cell: EPC ratio 1:2) and stimulated with anti-CD3 and anti-CD28 under TH1 differentiation culture conditions. After 4 days, intracellular IFN-γ expression (gated on CD4) was determined by flow cytometry(n=5). **c**, Cumulative composite data shows the MFI of IFN-γ(n=5). Data are representative of three independent experiments and were analyzed by two-tailed unpaired t-test. Error bars denote the SEM.



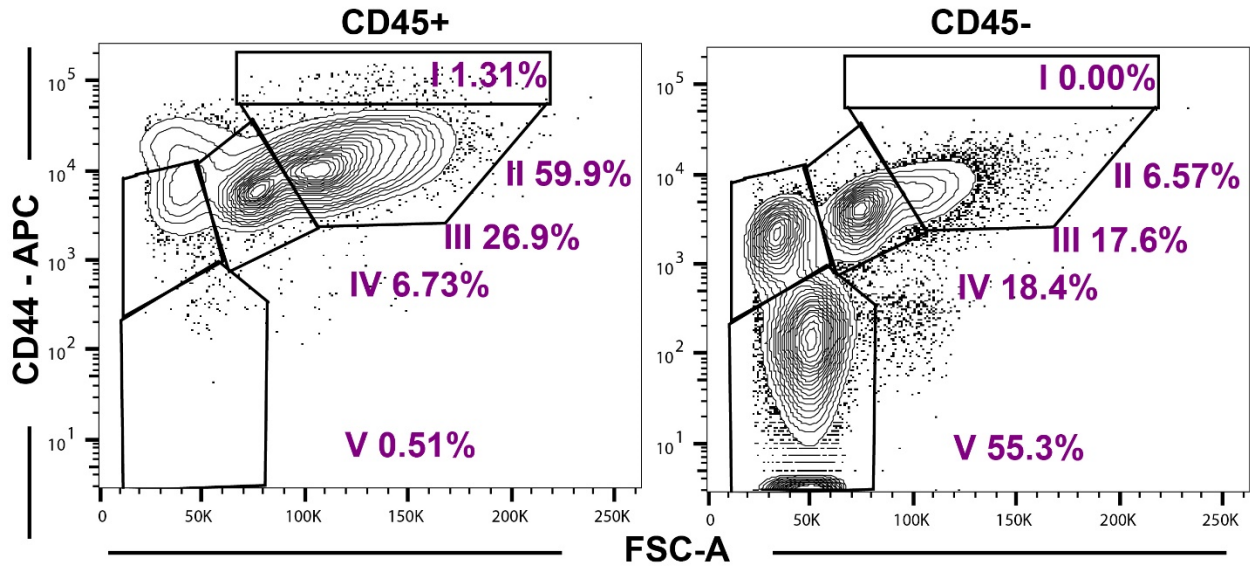
Supplementary Fig. 7. Anemia induced by tumorigenesis. Hematocrit (HCT) and hemoglobin (HGB) were measured in tumor-bearing mice at different time points (n=4 or 5) after LLC or B16F10 inoculation, as indicated. Each point represents data from an individual mouse, and data are representative of at least three independent experiments. Two-tailed Student's t-tests were used for all analyses. Error bars represent the means \pm SEM.



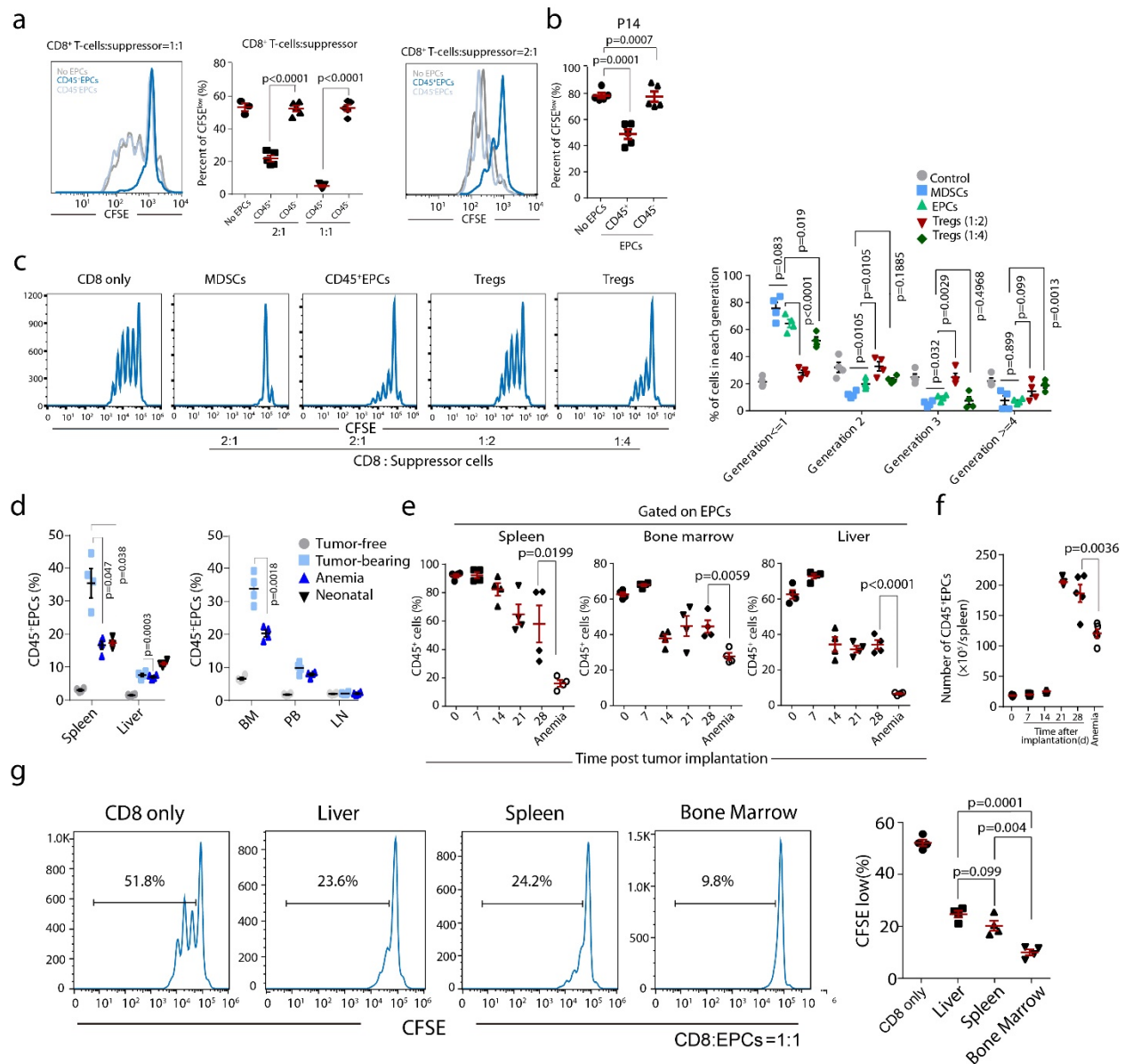
Supplementary Fig. 8. Listeria-specific CD8⁺ T cell responses remain intact in mice with acute anemia. **a-b**, Mice with established tumors (21 days after LLC inoculation, n=5), acute anemia (n=5), or healthy control mice (n=5) were infected with Δ actArLmOVA. Antigen-specific CD8⁺ T cells were measured by flow cytometric analysis of intracellular IFN- γ in spleen cells on day 7 post infection after stimulating splenocytes with OVA peptide *in vitro*. **c-d**, Mice with established tumors (21 days after LLC inoculation, n=5), acute anemia (n=5) or control healthy mice (n=5) were intravenously injected with equal numbers of CD90.1⁺CD8⁺ OT-1 cells. Six hours after OT-1 transfer, mice were infected with Lm. Splenocytes were harvested on day 7 post-infection and stimulated with OVA peptide *in vitro*. IFN- γ -producing CD90.1⁺CD8⁺ OT-1 cells were quantified by flow cytometry. Data are representative of at least three independent experiments. Two-tailed Student's t-tests were used for all analyses. Error bars represent the means \pm SEM.



Supplementary Fig. 9. Differential suppressive capacity of CD71⁺TER119⁺ EPCs isolated from acutely anemic and neonatal mice. a-b, CD71⁺TER119⁺ EPCs isolated from the spleens of anemic (a) and neonatal (b) mice were co-cultured at different CD8⁺T cell:EPC ratios with CFSE-labeled CD8⁺ T cells that were stimulated with CD3 and CD28 antibodies (n=4). Representative flow cytometry results and cumulative composite data show the proliferation of CFSE-labeled CD8⁺ T cells. Data are representative of at least three independent experiments. Two-tailed Student's t-tests were used for all analyses. Error bars represent the means \pm SEM.

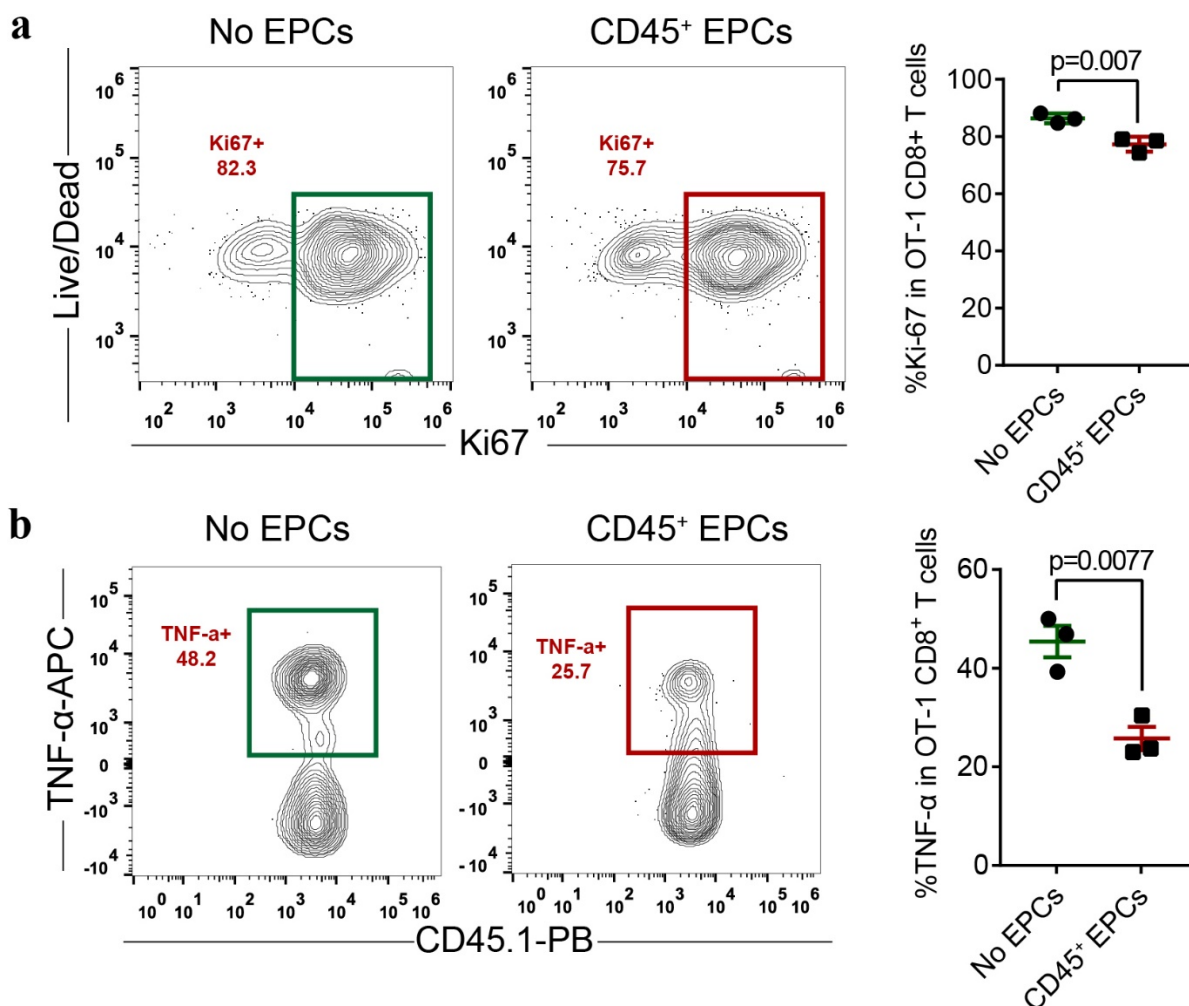


Supplementary Fig. 10. CD45⁺ EPCs are at the earlier developmental stages than CD45⁻ ones. 1×10^6 Lewis lung cancer cells (LLCs) were subcutaneously inoculated into C57BL/6 mice to allow for tumor formation. At day 21 post-inoculation, tumor-bearing mice were sacrificed and spleens were analyzed by FACS. Ter119⁺CD71⁺ cells were separated as CD45⁺ and CD45⁻ subpopulations and plotted as CD44 versus FSC. As defined by Chen et al., the developmental stages of erythroid cells from I (progenitor) to V (fully differentiated red blood cells)¹ were displayed.



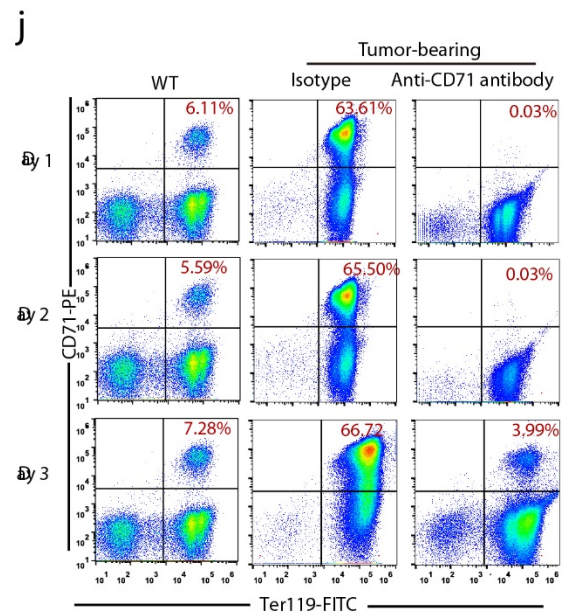
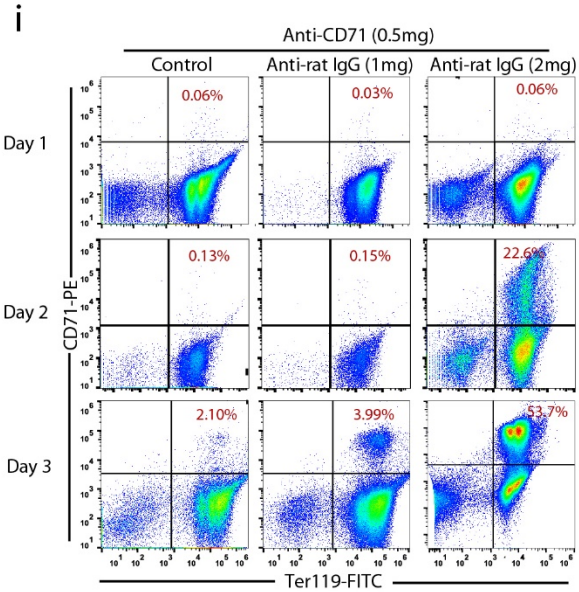
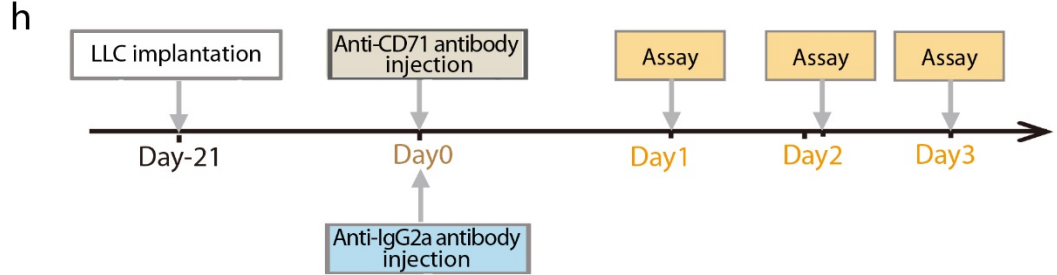
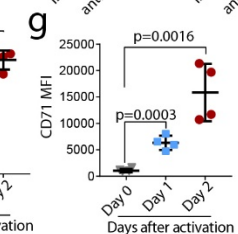
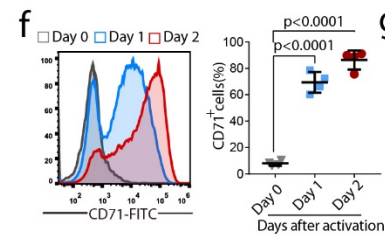
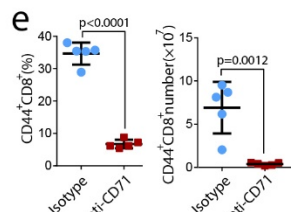
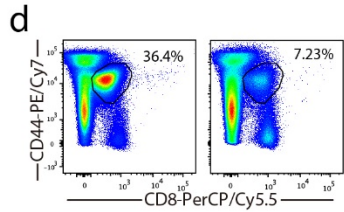
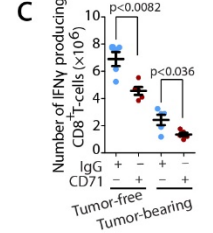
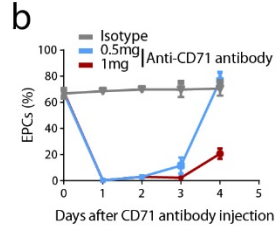
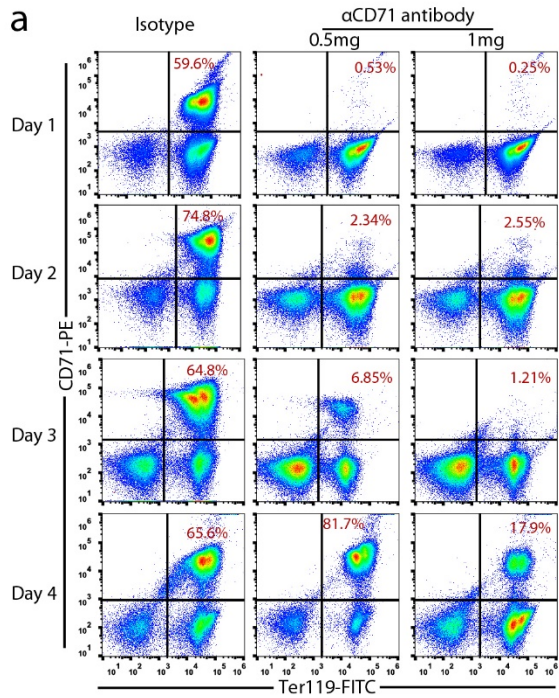
Supplementary Fig. 11. Immunosuppressive CD45⁺ early EPCs uniquely accumulate in tumor-bearing mice. **a**, Proliferation of CFSE-labeled CD8⁺ T cells in response to anti-CD3 and anti-CD28 was analyzed after co-culture with CD45⁺CD71⁺TER119⁺ or CD45⁻CD71⁺TER119⁺ cells isolated from the spleens of tumor-bearing mice at different CD8⁺ T cell:EPC ratios (n=5). **b**, CD45⁺CD71⁺TER119⁺ and CD45⁻CD71⁺TER119⁺ EPCs isolated from the spleens of tumor-bearing mice were co-cultured with CFSE-labeled P14 splenic cells, which were stimulated with a specific peptide, at a ratio of 2:1 (P14 cells:EPCs). Proliferation of CFSE-labeled CD8⁺ T cells (gated on CD8⁺ P14 cells) was then analyzed (n=5). **c**, Proliferative capacity of CFSE-labeled CD8⁺ T cells in response to anti-CD3 and anti-CD28 was analyzed after co-culture with CD45⁺CD71⁺TER119⁺ EPCs, MDSCs or Tregs isolated from the spleen of tumor-bearing mice (day 21) at CD8⁺ T cell:suppressor cell ratios as indicated in the figure (n=4). Two-tailed unpaired t-test of three independent experiments was performed by measuring the distribution of CD8⁺ T cells in each division. **d**, Frequency of CD45⁺CD71⁺TER119⁺ cells in spleen (SPL),

peripheral blood (PB), bone marrow (BM), lymph node (LN) and liver (HEP) for control, tumor-bearing, neonatal and anemic mice (n=4). **e**, Cumulative composite flow cytometry data show the frequencies of CD45⁺ cells within the CD71⁺TER119⁺ cell population in different tissues of anemic and tumor-bearing mice at the indicated times (n=4) after LLC inoculation. **f**, Cumulative composite data show the total number of CD45⁺CD71⁺TER119⁺ cells in the spleens of tumor-bearing and anemic mice(n=5). **g**, Proliferative capacity of CFSE-labeled CD8⁺ T cells in response to anti-CD3 and anti-CD28 antibody stimulation was analyzed after co-culture with CD45⁺CD71⁺TER119⁺ EPCs isolated from spleens of healthy control or tumor-bearing mice (n=5) at days 21 and 28 (CD8⁺ T cell:EPC ratio of 2:1). Data are representative of at least three independent experiments. Two-tailed Student's t-tests were used for all analyses. Error bars represent the means \pm SEM.

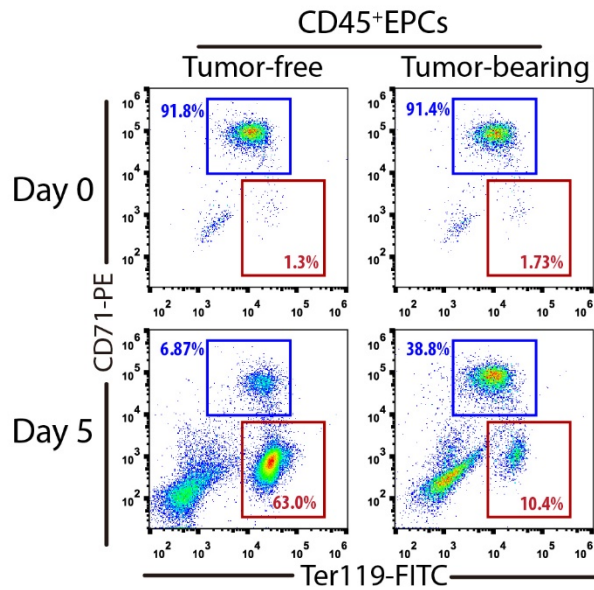
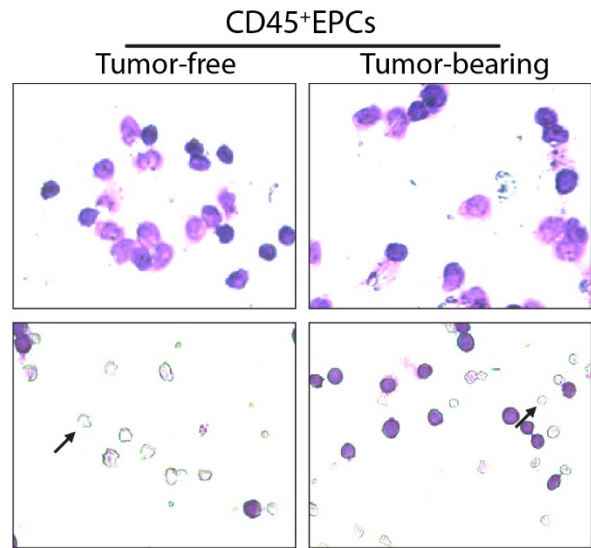


Supplementary Fig. 12. CD45⁺ EPCs suppress tumor-specific CTL responses *in vivo*. A total of 2×10^5 B16-OVA melanoma cells were subcutaneously transplanted into CD45.2⁺ C57BL/6 mice. At day 10 post tumor inoculation, mice were intravenously injected with 2×10^5 CD45.1⁺CD8⁺ OT-I cell mixed with either PBS or 2×10^6 CD45⁺ EPCs purified from other tumor-bearing mice. Mice were sacrificed at day 5 after OT-I transfer and splenocytes were

isolated and restimulated with OVA₂₅₇₋₂₆₄ peptide for 5 hours. The proliferative capacity of transferred OT-I cells was assessed by Ki67 staining (**a**) and their cytotoxicity was measured by intracellular TNF- α staining (**b**). Representative flow cytometry data (**left**) and the statistical analysis of mean frequencies (**right**) are presented. Data are collected from three mice (n=3). Two-tailed Student's t-tests were used for all analyses.

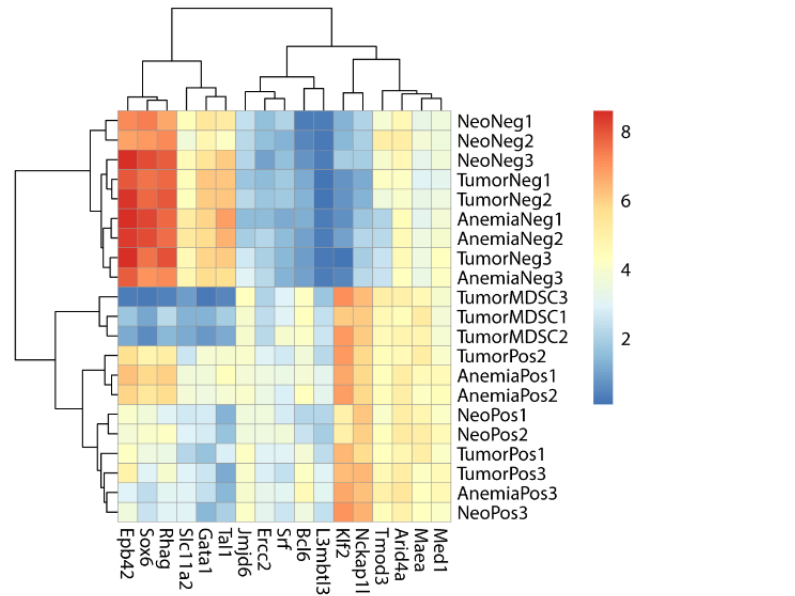


Supplementary Fig. 13. Specific depletion of CD71⁺ EPCs. 1×10^6 LLC cells were subcutaneously injected into C57BL/6 mice at day 0 and different doses of anti-CD71 deletion antibody was injected intravenously at day 21 after tumor cell inoculation (IgG was used as control). Frequencies of CD71⁺TER119⁺ cells in peripheral blood were detected by flow cytometry at different days after antibody administration. Representative flow cytometry **(a)** and cumulative composite data **(b)** show the frequency of CD71⁺TER119⁺ cells in peripheral blood. Mice were infected with LCMV at day 21 following LLC inoculation and anti-CD71 deletion antibody or control IgG was injected intravenously at days 21 and 24 after LLC inoculation. Mice were sacrificed on day 8 post-LCMV infection. Cumulative composite data show the number of CD8⁺IFN γ ⁺ cells in spleen(n=5) **(c)**. C57BL/6 mice infected with LCMV were intravenously injected with anti-CD71 deletion antibody or control IgG on days 0 and 4 after LCMV infection. Mice were sacrificed on day 8 post-LCMV infection(n=5). Representative flow cytometry **(d)** and cumulative composite data **(e, n=5)** show the frequency and number of CD8⁺CD44⁺ cells in spleen. CD8⁺ T cells were isolated and cultured in vitro in the presence of anti-CD3 and anti-CD28. The frequency of CD71⁺ cells among CD8⁺CD44⁺T cells was detected by flow cytometry at the indicates days after entering culture. Representative flow cytometry and cumulative composite data **(f, n=4)** are shown. CD71 MFI of CD8⁺CD44⁺T cells was analyzed **(g, n=4)**. 1×10^6 LLC cells were subcutaneously injected into C57BL/6 mice (PBS was used as control). Anti-CD71 deletion antibody or control IgG was intravenously injected at day 21 after tumor cell inoculation. To attenuate the anti-CD71 deletion antibody, different doses of anti-IgG2a antibody were intravenously injected at the same time **(h)**. The frequency of CD71⁺TER119⁺ cells in peripheral blood was detected by flow cytometry at the indicated times after antibody administration **(i)**. 1×10^6 LLC cells were subcutaneously injected into C57BL/6 mice (PBS was used as control). Anti-CD71 deletion antibody (1 mg/mouse) was intravenously injected at day 21 after tumor cell inoculation (IgG was used as control, 1 mg/mouse). To attenuate the anti-CD71 deletion antibody, anti-IgG2a antibody (3 mg/mouse) was intravenously injected 12h later. Then, we adoptively transferred P14 CD8⁺ T cells (CD90.1, 2×10^6 cells/mouse) and infected mice with LCMV c113 simultaneously 12 h after administration of anti-IgG2a antibody. All mice were sacrificed on day 2 after LCMV infection**(j)**. Representative flow cytometry data shows deletion efficiency of the anti-CD71 antibody **(k)**.

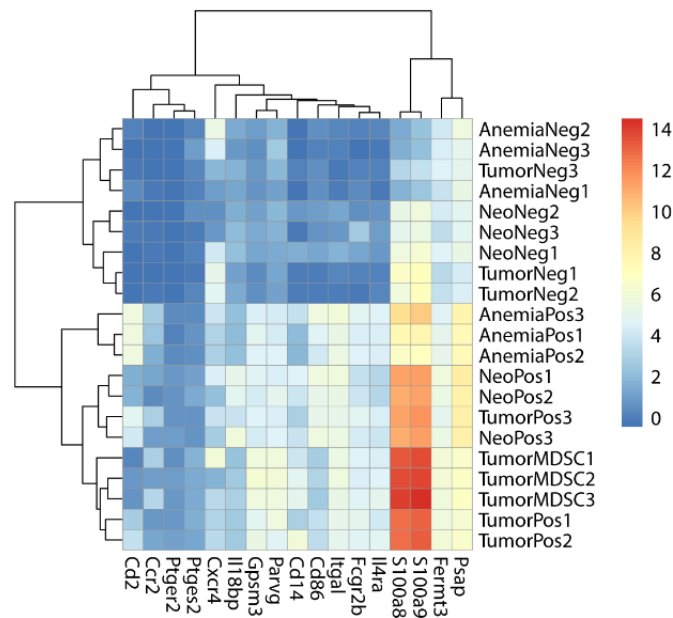
a**b**

Supplementary Fig. 14. CD45⁺CD71⁺TER119⁺ EPCs differentiate into mature RBCs. CD45⁺CD71⁺TER119⁺ EPCs were isolated by flow cytometry from tumor-bearing or healthy control mice and cultured in vitro under conditions that promote erythroid differentiation. Representative flow cytometry (a) and morphology (b) data. Giemsa staining shows erythroid differentiation at days 0 and 5 of culture (b).

a Erythrocyte development pathway (BIOCARTA database)



b Immune signature of MDSC



Supplementary Fig. 15. Transcriptome signatures to characterize EPC physiological and pathological origins. To categorize the differentiation and functional potential of EPCs, signature gene panels for erythroid differentiation (**a**) and MDSC function (**b**) were extracted from the RNAseq data set (Fig. 3a-b). Unsupervised clustering was performed on splenic CD45⁺ and CD45⁻ EPCs purified from neonatal, acutely anemic and tumor-bearing mice. MDSCs purified from spleens of tumor-bearing mice are included as a reference.

Supplementary Tables

Supplementary Table 1. Clinical information for the 167 cancer patients with different types of malignant tumors used in the EBV infection study (Fig. 4e-g).

| No. | ID | Age | Gen der | Diagnosis | Hb (g/L) | EBV (copies/ml) | Stage |
|-----|----------|-----|------------|---|-------------|--------------------|---------|
| 1 | 07895013 | 46 | M | Nasopharyngeal carcinoma | 176 | 851 | III |
| 2 | 07195985 | 68 | M | Small cell lung cancer | 110 | <400 | Limited |
| 3 | 07825720 | 77 | F | Breast cancer | 101 | 643 | III |
| 4 | 05337039 | 62 | F | Small cell lung cancer | 96 | <400 | Limited |
| 5 | 05337039 | 52 | M | Small cell lung cancer | 66 | 6300 | Limited |
| 6 | 05991279 | 60 | F | Breast cancer | 116 | <400 | III |
| 7 | 06003668 | 60 | M | Non-small cell lung cancer | 94 | 434 | III |
| 8 | 00969251 | 62 | M | Lung adenocarcinoma | 130 | <400 | rIV |
| 9 | 07648115 | 52 | F | Rectum cancer | 126 | <400 | rIV |
| 10 | 05732953 | 66 | M | Primary bronchogenic lung squamous cell carcinoma | 71 | 603 | III |
| 11 | 06055174 | 16 | F | Breast cancer | 107 | 500 | III |
| 12 | 05895075 | 13 | M | Pineaal tumor | 113 | <400 | III |
| 13 | 06034647 | 40 | F | Breast cancer | 147 | <400 | IV |
| 14 | 05690867 | 59 | F | Breast cancer | 96 | 463 | rIV |
| 15 | 07659165 | 61 | M | Esophageal squamous cell cancer | 107 | 411 | III |
| 16 | 07853447 | 73 | M | Nasopharyngeal squamous cell cancer | 132 | <400 | IV |
| 17 | 07062881 | 62 | M | Colon cancer | 115 | <400 | III |
| 18 | 07535817 | 41 | F | Colon cancer | 110 | <400 | III |
| 19 | 03415090 | 61 | F | Breast cancer | 102 | <400 | rIV |
| 20 | 07198192 | 69 | M | Laryngeal squamous cell cancer | 127 | 776 | III |
| 21 | 04980245 | 54 | M | Lung squamous cell carcinoma | 135 | 800 | IIIA |
| 22 | 07843380 | 59 | F | Esophageal squamous cell cancer | 115 | <400 | III |
| 23 | 04940642 | 50 | M | Nasopharyngeal squamous cell cancer | 82 | <400 | III |
| 24 | 07821594 | 60 | F | Cervical squamous cell carcinoma | 99 | 615 | III |
| 25 | 07894165 | 58 | F | Rectal caner | 133 | <400 | IV |
| 26 | 05903285 | 36 | F | Ovarian cancer | 129 | <400 | III |
| 27 | 05679620 | 64 | M | Lung squamous cell carcinoma | 104 | 410 | IIIb |
| 28 | 07867140 | 71 | F | Cardia adenocarcinoma | 113 | 550 | III |
| 29 | 05153701 | 42 | F | Lung adenocarcinoma | 108 | <400 | IV |
| 30 | 07811161 | 41 | M | Nasopharyngeal squamous cell | 110 | 496 | IV |
| 31 | 05612175 | 62 | M | Tongue cancer | 103 | 420 | IV |
| 32 | 04929706 | 59 | F | Primary bronchial adenocarcinoma of the lung | 113 | <400 | IV |
| 33 | 07855681 | 33 | M | Primary bronchial adenocarcinoma of the lung | 123 | <400 | III |
| 34 | 07680469 | 67 | M | Tongue cancer | 137 | <400 | III |
| 35 | 07895124 | 64 | M | Lung cancer | 66 | 4370 | III |
| 36 | 06005611 | 24 | M | Tongue squamous cell carcinoma | 120 | <400 | III |
| 37 | 07759139 | 39 | M | Nasopharyngeal squamous cell | 123 | <400 | IIIA |
| 38 | 07903245 | | M | Colon cancer | 128 | <400 | IV |

| | | | | | | | |
|----|----------|----|---|---|-----|------|-----------|
| 39 | 07847305 | 74 | M | Primary bronchial small cell lung cancer | 103 | 1170 | IIIB |
| 40 | 07638833 | 58 | M | Colon cancer | 110 | <400 | III |
| 41 | 07837029 | 47 | M | Primary bronchogenic lung squamous cell carcinoma | 116 | <400 | III |
| 42 | 07900061 | 14 | F | Small cell lung cancer | 132 | 1000 | Extensive |
| 43 | 07579757 | 58 | M | Small cell lung cancer | 112 | <400 | Limited |
| 44 | 05752348 | 56 | F | Cervical squamous cell carcinoma | 132 | 790 | III |
| 45 | 03949479 | 44 | M | Pelvic malignancy | 109 | <400 | IV |
| 46 | 07703736 | 65 | F | Cervical squamous cell carcinoma | 97 | <400 | IV |
| 47 | 07897540 | 48 | F | Cervical squamous cell carcinoma | 97 | <400 | IV |
| 48 | 07709368 | 58 | M | Non-small cell lung cancer | 73 | 548 | IV |
| 49 | 02410882 | 39 | F | Breast cancer | 119 | <400 | III |
| 50 | 05405434 | 73 | F | Non-Hodgkin's lymphoma | 112 | 584 | IV |
| 51 | 05753621 | 72 | M | Small cell lung cancer | 123 | 496 | Extensive |
| 52 | 07846143 | 40 | F | Nasopharyngeal squamous cell | 111 | 424 | III |
| 53 | 07613054 | 63 | M | Non-small cell lung cancer | 84 | 7910 | rIV |
| 54 | 03221140 | 66 | M | Carcinoma of urinary bladder | 132 | 2330 | IV |
| 55 | 04627935 | 53 | M | Lung cancer | 146 | <400 | IV |
| 56 | 07659113 | 55 | M | Non-small cell lung cancer | 98 | 6880 | IV |
| 57 | 03533316 | 62 | F | Non-small cell lung cancer | 128 | <400 | III |
| 58 | 07846569 | 63 | M | Non-small cell lung cancer | 108 | <400 | III |
| 59 | 07839120 | 49 | M | Non-small cell lung cancer | 139 | <400 | IV |
| 60 | 05520961 | 69 | M | Prostatic cancer | 140 | 458 | IV |
| 61 | 07612103 | 53 | M | Small cell lung cancer | 143 | <400 | Extensive |
| 62 | 07063014 | 49 | M | Nasopharyngeal squamous cell | 133 | <400 | III |
| 63 | 07896361 | 76 | F | Metastasis squamous cell | 113 | 790 | III |
| 64 | 07443529 | 29 | M | Rectum cancer | 125 | <400 | IV |
| 65 | 05971281 | 52 | M | Small cell lung cancer | 85 | <400 | Extensive |
| 66 | 07567916 | 60 | M | Nasopharyngeal squamous cell | 122 | 690 | III |
| 67 | 07828686 | 70 | F | Breast cancer | 149 | 1400 | III |
| 68 | 05792179 | 38 | M | Laryngo carcinoma | 108 | <400 | III |
| 69 | 05916328 | 50 | F | Breast cancer | 92 | <400 | rIV |
| 70 | 07895276 | 60 | F | Esophageal squamous cell cancer | 132 | <400 | III |
| 71 | 05734011 | 49 | M | Laryngo carcinoma | 126 | 1180 | III |
| 72 | 07793060 | 53 | M | Non-small cell lung cancer | 129 | <400 | IV |
| 73 | 07895366 | 53 | M | Nasopharyngeal squamous cell | 109 | 1210 | III |
| 74 | 07140879 | 48 | M | Non-small cell lung cancer | 112 | <400 | IV |
| 75 | 07442213 | 52 | M | Rectum cancer | 135 | <400 | IIB |
| 76 | 68103580 | 38 | F | Breast cancer | 118 | <400 | IIIA |
| 77 | 07321276 | 52 | M | Multiple myeloma | 147 | <400 | III |
| 78 | 05124803 | 50 | F | Non-small cell lung cancer | 127 | 497 | IV |
| 79 | 06059938 | 72 | M | Non-Hodgkin's lymphoma | 137 | 1600 | III |
| 80 | 07888019 | 70 | M | Nasopharyngeal squamous cell | 77 | 8400 | IV |
| 81 | 06051555 | 63 | M | Non-small cell lung cancer | 130 | <400 | III |
| 82 | 07841848 | 55 | M | Small cell lung cancer | 143 | <400 | Extensive |
| 83 | 05105660 | 72 | M | Rectum cancer | 125 | 1540 | rIV |
| 84 | 07820551 | 67 | M | Non-Hodgkin's lymphoma | 74 | 1150 | III |
| 85 | 07575124 | 60 | F | Non-small cell lung cancer | 107 | <400 | rIV |

| | | | | | | | |
|-----|----------|----|---|---------------------------------|-----|------|-----------|
| 86 | 07724974 | 30 | F | Carcinoma of submaxillary gland | 98 | 1540 | III |
| 87 | 04650882 | 37 | M | Nasopharyngeal squamous cell | 142 | <400 | III |
| 88 | 05920670 | 51 | M | Laryngo carcinoma | 138 | <400 | III |
| 89 | 06053980 | 66 | M | Rectum cancer | 118 | 1800 | IV |
| 90 | 07185514 | 64 | F | Breast cancer | 96 | 1130 | IV |
| 91 | 07647544 | 29 | M | Osteoblastoma | 134 | <400 | III |
| 92 | 07632546 | 42 | M | Nasopharyngeal squamous cell | 141 | 408 | IV |
| 93 | 02651458 | 67 | M | Esophageal squamous cell cancer | 88 | <400 | III |
| 94 | 06069054 | 34 | M | Tongue cancer | 89 | 928 | III |
| 95 | 07880370 | 57 | M | Non-small cell lung cancer | 144 | 730 | IV |
| 96 | 05989028 | 49 | M | Esophageal squamous cell cancer | 136 | <400 | III |
| 97 | 07860785 | 53 | M | Nasopharyngeal squamous cell | 119 | <400 | III |
| 98 | 07890643 | 49 | M | Nasopharyngeal squamous cell | 150 | <400 | III |
| 99 | 07774183 | 43 | F | Cervical cancer | 87 | 1080 | IV |
| 100 | 07044368 | 70 | F | Melanoma | 94 | 803 | rIV |
| 101 | 05920081 | 81 | F | Ovarian cancer | 112 | 6410 | IIIb |
| 102 | 07869420 | 49 | M | Nasopharyngeal squamous cell | 113 | 1490 | IV |
| 103 | 04366533 | 29 | F | Breast cancer | 103 | <400 | III |
| 104 | 07856083 | 68 | M | Laryngo carcinoma | 107 | <400 | III |
| 105 | 04801797 | 65 | M | Esophageal squamous cell cancer | 98 | <400 | III |
| 106 | 07343965 | 51 | M | Renal carcinoma | 96 | <400 | rIV |
| 107 | 07730104 | 52 | F | Ovarian cancer | 88 | <400 | IIIb |
| 108 | 07181329 | 60 | M | Non-small cell lung cancer | 96 | <400 | IV |
| 109 | 05688509 | 68 | F | Non-small cell lung cancer | 135 | 780 | IV |
| 110 | 07207312 | 41 | M | Small cell lung cancer | 133 | <400 | Extensive |
| 111 | 07815225 | 65 | M | Non-small cell lung | 80 | 1920 | IV |
| 112 | 05689100 | 78 | F | Non-small cell lung cancer | 93 | 910 | IIIA |
| 113 | 07810012 | 64 | M | Non-small cell lung cancer | 126 | <400 | IV |
| 114 | 05935348 | 60 | M | Non-small cell lung cancer | 77 | 930 | IV |
| 115 | 04023326 | 78 | F | Hepatocellular carcinoma | 97 | 2280 | IV |
| 116 | 07379305 | 50 | M | Non-small cell lung cancer | 133 | <400 | IV |
| 117 | 07903046 | 52 | F | Thyroid carcinoma | 136 | <400 | II |
| 118 | 07832602 | 48 | F | Non-small cell lung cancer | 112 | <400 | IV |
| 119 | 70009677 | 52 | M | Sinus cancer | 124 | <400 | IV |
| 120 | 07257155 | 50 | F | Ovarian cancer | 102 | <400 | IV |
| 121 | 07645628 | 8 | M | Glioma | 138 | 620 | III |
| 122 | 07764784 | 59 | M | Non-Hodgkin's lymphoma | 83 | <400 | III |
| 123 | 04997009 | 58 | M | Rectum carcinoma | 123 | 410 | rIV |
| 124 | 07486030 | 63 | M | Non-small cell lung cancer | 129 | 876 | III |
| 125 | 07461780 | 56 | M | Non-small cell lung cancer | 123 | 600 | IV |
| 126 | 07811295 | 64 | M | Non-small cell lung cancer | 104 | 2000 | IV |
| 127 | 07837308 | 52 | M | Non-small cell lung | 142 | <400 | III |
| 128 | 07609270 | 10 | F | Intracranial germinomas | 85 | <400 | rIII |
| 129 | 07778200 | 51 | M | Esophageal squamous cell cancer | 95 | <400 | III |
| 130 | 07316014 | 55 | M | Small cell lung cancer | 89 | <400 | Extensive |
| 131 | 07502501 | 49 | F | Cholangiocellular carcinoma; | 95 | 799 | rIV |
| 132 | 07895716 | 53 | M | Non-small cell lung cancer | 145 | <400 | IV |
| 133 | 07830955 | 75 | F | Peritoneal carcinoma | 82 | 3490 | III |
| 134 | 07895402 | 52 | M | Nasopharyngeal squamous cell | 140 | 520 | IV |

| | | | | | | | |
|-----|----------|----|---|---------------------------------|-----|-------|------|
| 135 | 03320072 | 62 | M | Non-small cell lung cancer | 121 | 458 | rIV |
| 136 | 07577962 | 56 | M | Non-small cell lung cancer | 132 | 554 | IIIb |
| 137 | 02888053 | 72 | M | Non-small cell lung cancer | 91 | <400 | III |
| 138 | 07903283 | 69 | M | Tongue cancer | 137 | 502 | III |
| 139 | 07846087 | 66 | F | Esophageal squamous cell cancer | 110 | 3010 | III |
| 140 | 07896281 | 56 | F | Non-small cell lung cancer | 111 | <400 | III |
| 141 | 04228779 | 63 | F | Nasopharyngeal squamous cell | 126 | 1440 | III |
| 142 | 0160065 | 64 | M | Nasopharyngeal squamous cell | 117 | 1070 | rIV |
| 143 | 07899027 | 63 | M | Non-small cell lung cancer | 112 | <400 | IV |
| 144 | 04462246 | 34 | M | Nasopharyngeal squamous cell | 102 | <400 | IV |
| 145 | 05862096 | 48 | F | Non-small cell lung cancer | 85 | <400 | IV |
| 146 | 07498447 | 58 | M | Non-small cell lung cancer | 79 | 932 | IV |
| 147 | 06029263 | 43 | M | Non-small cell lung cancer | 72 | 4300 | III |
| 148 | 07221093 | 64 | M | Rectum cancer | 55 | 935 | IV |
| 149 | 05679611 | 47 | M | Nasopharyngeal squamous cell | 85 | 7710 | III |
| 150 | 07911662 | 71 | F | Rectum cancer | 77 | 752 | III |
| 151 | 07483019 | 64 | M | Rectum cancer | 65 | <400 | III |
| 152 | 07888019 | 70 | M | Nasopharyngeal squamous cell | 62 | 4020 | III |
| 153 | 07316014 | 55 | M | Non-small cell lung | 54 | <400 | IIIB |
| 154 | 07585360 | 72 | F | Esophageal squamous cell cancer | 68 | 693 | III |
| 155 | 06038755 | 70 | M | Non-small cell lung cancer | 62 | 593 | IV |
| 156 | 04801797 | 65 | M | Esophageal squamous cell cancer | 76 | <400 | IV |
| 157 | 06017371 | 46 | F | Breast cancer | 65 | <400 | III |
| 158 | 06202415 | 51 | M | Non-small cell lung cancer | 51 | 730 | IV |
| 159 | 00031631 | 59 | M | Esophageal squamous cell cancer | 62 | 1710 | III |
| 160 | 03912538 | 54 | M | Non-small cell lung cancer | 65 | 1630 | III |
| 161 | 08221390 | 54 | F | Cervical carcinoma | 67 | 1440 | IV |
| 162 | 00237414 | 53 | M | Esophageal squamous cell cancer | 71 | 1060 | IV |
| 163 | 07418835 | 62 | M | Non-small cell lung cancer | 74 | 3120 | III |
| 164 | 08229641 | 78 | F | Rectum cancer | 74 | 5050 | IV |
| 165 | 04676514 | 66 | F | Non-small cell lung | 77 | 9100 | III |
| 166 | 05984900 | 60 | M | Non-small cell lung cancer | 78 | <400 | III |
| 167 | 05700972 | 51 | M | Hepatocellular carcinoma | 57 | 11500 | IV |

Supplementary Table 2. Clinical information for the 5 healthy donor and 41 patients with different types of malignant tumors used to study EPC frequency in peripheral blood (Fig. 4e-g and i).

| No | ID | Age | Gender | Diagnosis | Hb(g/L) | Stage |
|----|----------|-----|--------|---|---------|-------|
| 1 | 07612059 | 46 | F | Nasopharyngeal carcinoma | 53 | III |
| 1 | 07612059 | 46 | F | Nasopharyngeal carcinoma | 53 | III |
| 2 | 07408347 | 58 | M | Diffuse large B-cell lymphoma | 63 | III |
| 3 | 05108828 | 57 | F | Pleomorphic liposarcoma of the peritoneum | 67 | III |
| 4 | 05994934 | 64 | M | Primary bronchogenic lung cancer | 74 | IV |
| 5 | 07337183 | 69 | M | Lung squamous cell carcinoma | 77 | IV |
| 6 | 07708983 | 20 | F | Mediastinal malignant germ cell tumors | 78 | III |
| 7 | 07737176 | 61 | M | Lung sarcomatoid carcinoma | 81 | III |
| 8 | 07622038 | 27 | F | Tonsillar lymphoma | 85 | III |
| 9 | 07531718 | 48 | F | Primary bronchogenic lung cancer | 88 | III |
| 10 | 07673672 | 69 | M | Diffuse large B-cell lymphoma | 88 | III |
| 11 | 05152855 | 64 | F | Primary bronchial adenocarcinoma of the lung | 88 | IV |
| 12 | 07277665 | 46 | F | Primary bronchogenic lung squamous cell carcinoma | 89 | III |
| 13 | 07468522 | 68 | F | Diffuse large B-cell lymphoma | 89 | III |
| 14 | 05310357 | 71 | M | Lung adenocarcinoma | 90 | III |
| 15 | 01201637 | 70 | F | Lung squamous cell carcinoma | 92 | IV |
| 16 | 05908951 | 53 | M | Left breast infiltrating ductal carcinoma | 93 | IV |
| 17 | 05992241 | 34 | M | Gastric adenocarcinoma | 95 | IV |
| 18 | 07219424 | 52 | M | Lung adenocarcinoma | 98 | III |
| 19 | 07658793 | 55 | F | Gallbladder cancer | 98 | III |
| 20 | 07701980 | 79 | F | Lung adenocarcinoma | 109 | III |
| 21 | 05991279 | 52 | M | Primary bronchial small cell lung cancer | 107 | III |
| 22 | 05689100 | 32 | F | Primary bronchial adenocarcinoma of the lung | 105 | IV |
| 23 | 05898617 | 66 | F | Primary bronchial adenocarcinoma of the lung | 99 | III |
| 24 | 00945960 | 64 | F | Tongue squamous cell carcinoma | 93 | III |
| 25 | 05905574 | 42 | F | Nasopharyngeal squamous cell | 100 | III |
| 26 | 07672747 | 46 | M | Colon cancer | 109 | IV |
| 27 | 07564619 | 63 | M | Primary bronchial small cell lung cancer | 108 | III |
| 28 | 07215539 | 75 | M | Non-Hodgkin's lymphoma | 105 | III |
| 29 | 07626046 | 58 | M | Primary bronchogenic lung squamous cell carcinoma | 103 | III |
| 30 | 07623835 | 60 | F | Small cell lung cancer | 110 | IV |
| 31 | 07679195 | 36 | M | Pelvic malignancy | 150 | IV |
| 32 | 07500671 | 52 | F | Cervical squamous cell carcinoma | 139 | IIIB |
| 33 | 06008744 | 59 | F | Cervical squamous cell carcinoma | 138 | IIIB |
| 34 | 07497250 | 65 | F | Primary bronchogenic lung squamous cell carcinoma | 126 | III |
| 35 | 07188428 | 49 | M | Duodenal adenocarcinoma | 136 | III |
| 36 | 07535269 | 69 | M | Hodgkin 's lymphoma | 127 | III |
| 37 | 04083507 | 70 | M | Left breast infiltrating ductal carcinoma | 133 | III |
| 38 | 05724108 | 49 | F | Primary bronchial adenocarcinoma of the lung | 117 | IV |
| 39 | 07547089 | 54 | F | Nasopharyngeal squamous cell | 119 | IV |
| 40 | 07273674 | 74 | M | Small cell lung cancer | 132 | III |

| | | | | | | |
|----|----------|----|---|------------------------|-----|------|
| 41 | 07587762 | 55 | F | Small cell lung cancer | 125 | IV |
| 42 | None | 31 | M | Health donner | 125 | None |
| 43 | None | 27 | M | Health donner | 133 | None |
| 44 | None | 24 | F | Health donner | 134 | None |
| 45 | None | 28 | F | Health donner | 136 | None |
| 46 | None | 28 | M | Health donner | 127 | None |

Supplementary Table 3. Clinical information for with the 15 cancer patients used to study the inhibitory effects of CD45⁺ and CD45⁻ EPCs on CD8⁺ T cell function (Fig. 4h, j-n).

| No. | ID | Age | Gender | Diagnosis | Hb(g/L) | Stage |
|------------|-----------|------------|---------------|---|----------------|--------------|
| 1 | 05108828 | 57 | F | Pleomorphic liposarcoma of the peritoneum | 67 | III |
| 2 | 05994934 | 64 | M | Primary bronchogenic lung cancer | 74 | IV |
| 3 | 07337183 | 69 | M | Lung squamous cell carcinoma | 77 | IV |
| 4 | 07708983 | 20 | F | Mediastinal malignant germ cell tumors | 78 | III |
| 5 | 07531718 | 48 | F | Primary bronchogenic lung cancer | 88 | III |
| 6 | 05152855 | 64 | F | Primary bronchial adenocarcinoma of the lung | 88 | IV |
| 7 | 07277665 | 46 | F | Primary bronchogenic lung squamous cell carcinoma | 89 | III |
| 8 | 05310357 | 71 | M | Lung adenocarcinoma | 90 | III |
| 9 | 01201637 | 70 | F | Lung squamous cell carcinoma | 92 | IV |
| 10 | 05908951 | 53 | M | Left breast infiltrating ductal carcinoma | 93 | IV |
| 11 | 05992241 | 34 | M | Gastric adenocarcinoma | 95 | IV |
| 12 | 07219424 | 52 | M | Lung adenocarcinoma | 98 | III |
| 13 | 07658793 | 55 | F | Gallbladder cancer | 98 | III |
| 14 | 07701980 | 79 | F | Lung adenocarcinoma | 109 | III |
| 15 | 05991279 | 52 | M | Primary bronchial small cell lung cancer | 107 | III |

Supplementary Table 4. Primer sequences used for RT-qPCR.

| Gene | Forward | Reverse |
|-------------|-------------------------|-------------------------|
| human Fes | AGGACCGTGACAAGGCTAAG | CCTTCAGGATGCAAGCCATCT |
| human Glrx | CCCATCAAACAAGGGCTTCTG | CTGCATCCGCCTATACAATCTT |
| human Gpx3 | AGAGCCGGGGACAAGAGAA | ATTGCCAGCATACTGCTTGA |
| human Gpx4 | GAGGCAAGACCGAAGTAAACTAC | CCGAACTGGTTACACGGGAA |
| human Gsr | CACTTGCGTGAATGTTGGATG | TGGGATCACTCGTGAAGGCT |
| human Mgst1 | ATGACAGAGTAGAACGTGTACGC | TACAGGAGGCCAATTCCAAGA |
| human Mpo | TGCTGCCCTTTGACAACCTG | TGCTCCCGAAGTAAGAGGGT |
| human msra | GGCCATCTACCCGACCTCT | GCCATIGGGGTTCTTGCTCA |
| human prnp | AGTCAGTGGAACAAGCCGAG | CTGCCGAAATGTAIGATGGGC |
| human cybb | CACAGGCCTGAAACAAAAGA | GCTTCAGGTCCACA GAGGAA. |
| mouse cybb | TGTGGTTGGGGCTGAATGTC | CTGAGAAAGGAGAGCAGATTTCG |