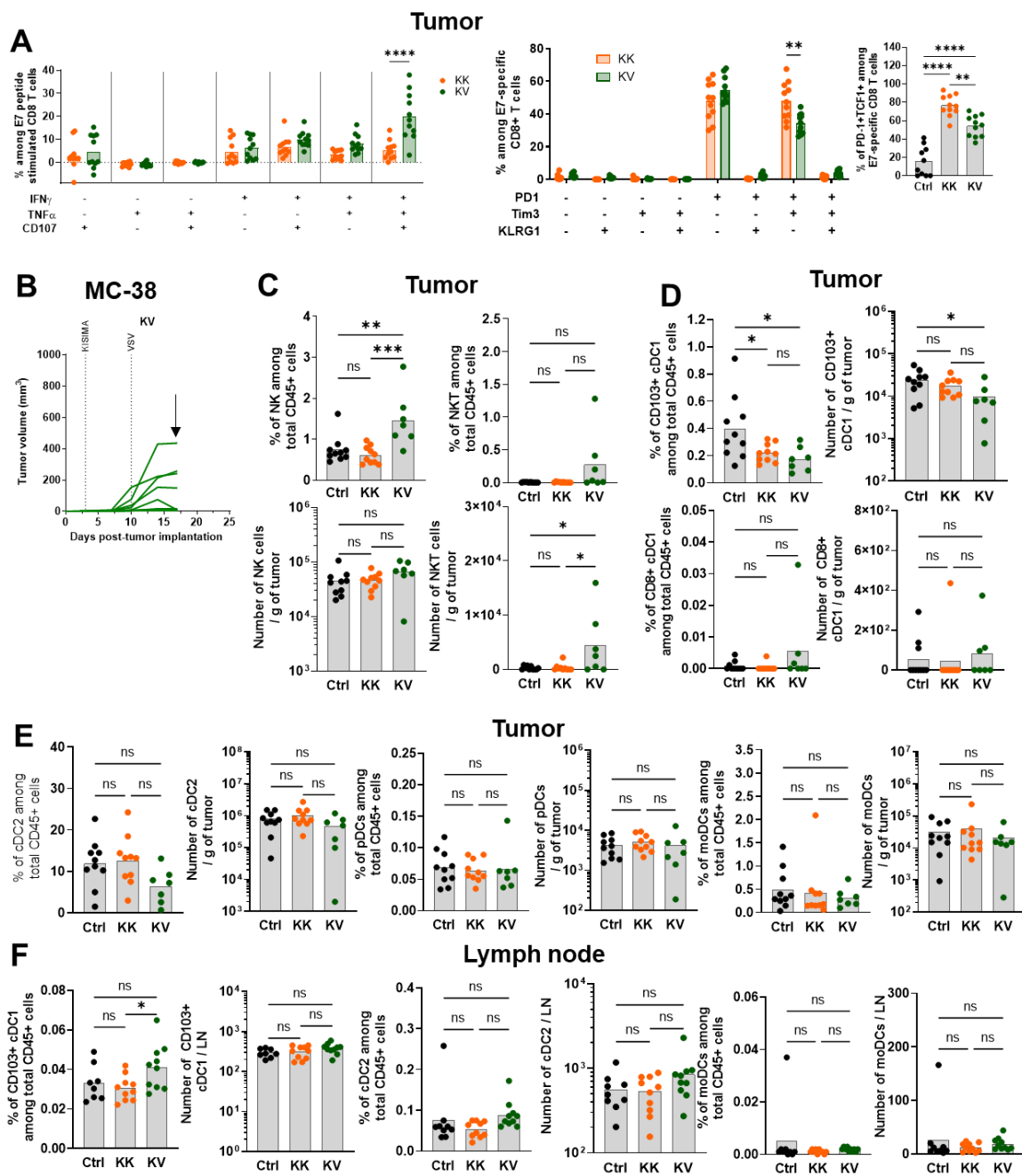
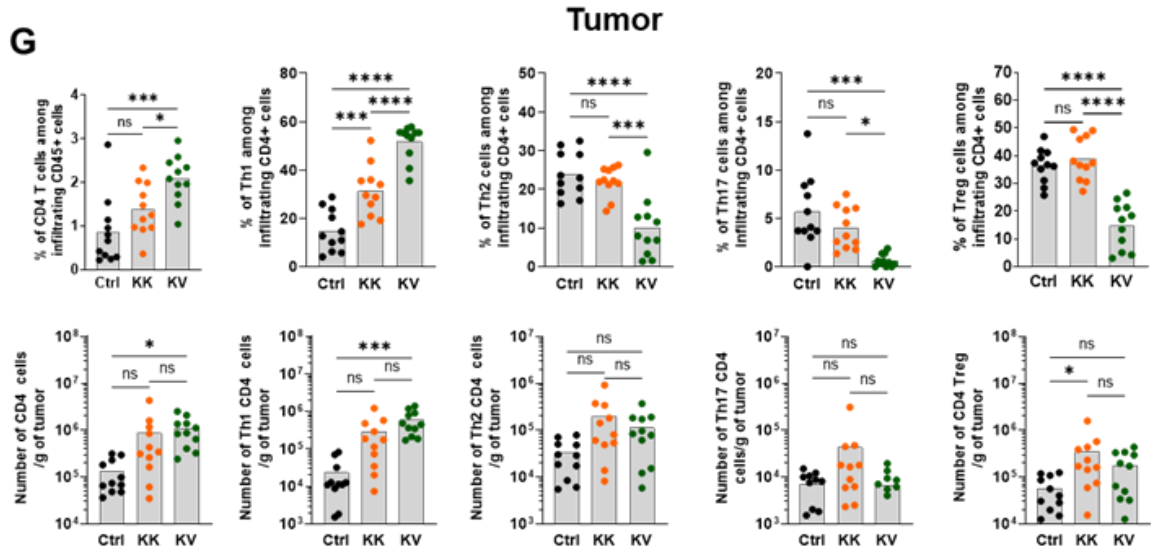
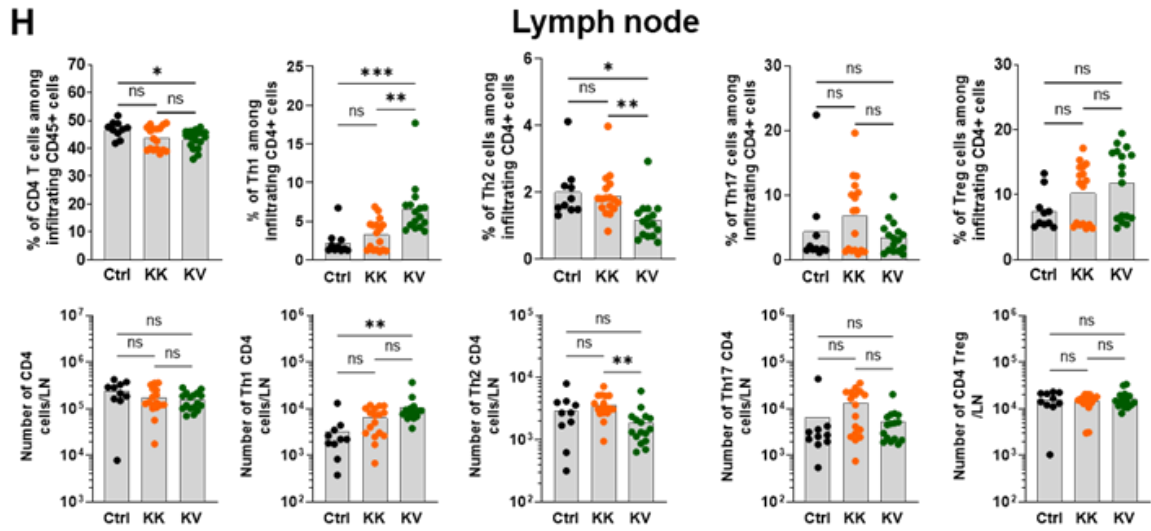
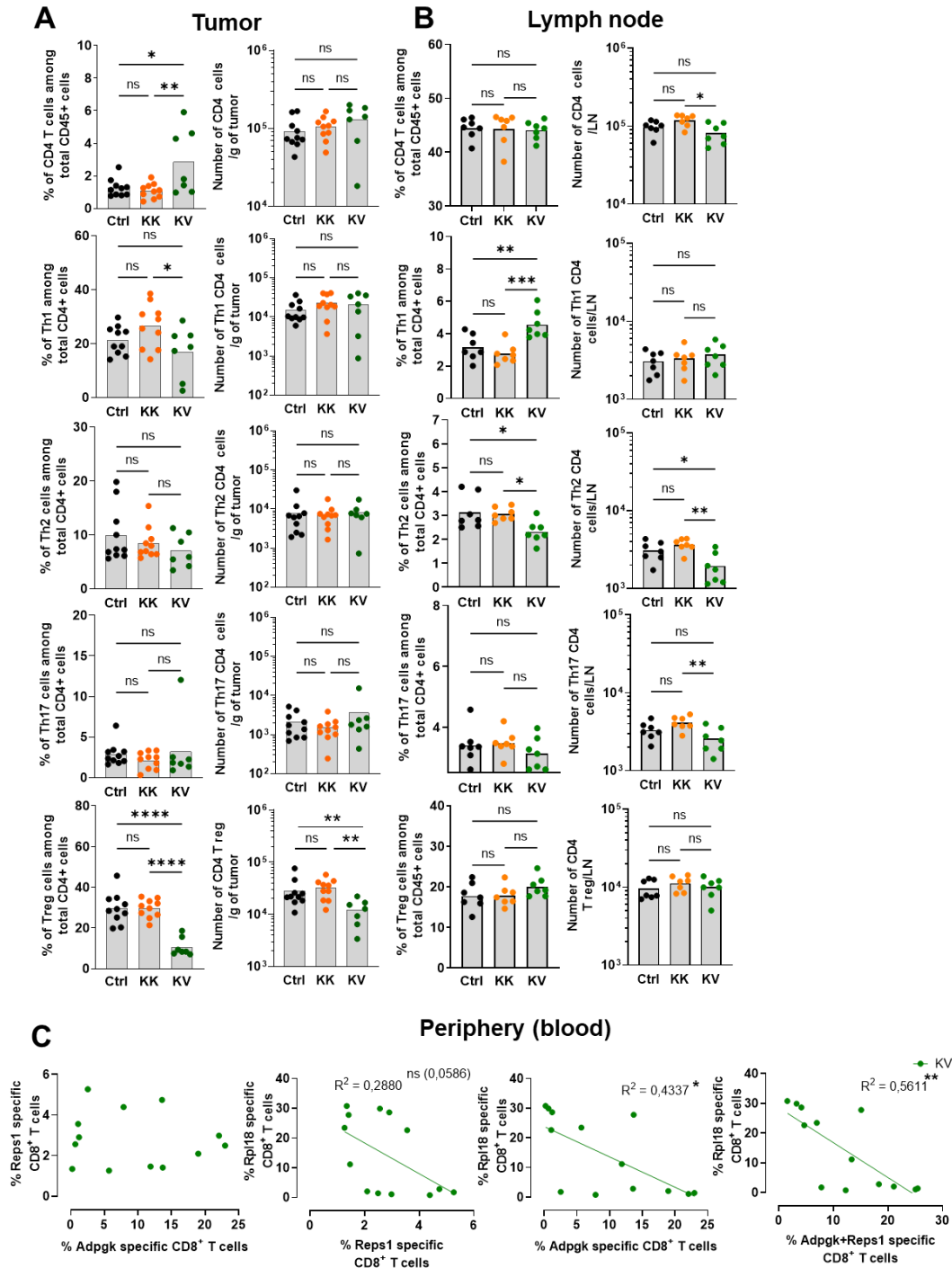


**Figure S1.** Tumor growth curves, intratumoral NK and NKT cells, intratumoral and intranodal dendritic cell characterization and immune cell ratios in TC-1 tumor-bearing C57BL/6 mice. **(A)** Individual TC1 tumor-growth curves in mm<sup>3</sup> post-tumor implantation for untreated (control), homologous KISIMA-Mad25-treated (KK) and heterologous KISIMA-Mad25- and VSV-GP-HPV-treated (KV) groups up to two days before tumor harvest for FACS analysis. **(B)** Frequencies and absolute numbers of tumor-infiltrating NK (left) and NKT (right) cells were analyzed on day 21 post-TC-1 tumor implantation. **(C)** From top to bottom: Frequencies (left) and absolute numbers (right) of tumor-infiltrating cDC2s, pDCs and moDCs on day 18 post-MC-38 tumor implantation. **(D)** From top to bottom: Frequencies (left) and absolute numbers (right) of intranodal cDC2s, pDCs and moDCs on day 18 post-MC-38 tumor implantation. **(E)** Ratio of intranodal Tconv to Treg (left side) or intranodal CD8<sup>+</sup> T cells to Treg derived from TC-1 tumor-bearing mice on day 21 post-tumor implantation. Data in **(A)** to **(E)** are shown as means (grey bar) and are derived from at least two independent experiments. \*  $p < 0.05$ , \*\*  $p < 0.01$ , \*\*\*  $p < 0.001$ , \*\*\*\*  $p < 0.0001$  One-Way ANOVA followed by Tukey's multiple comparison test.

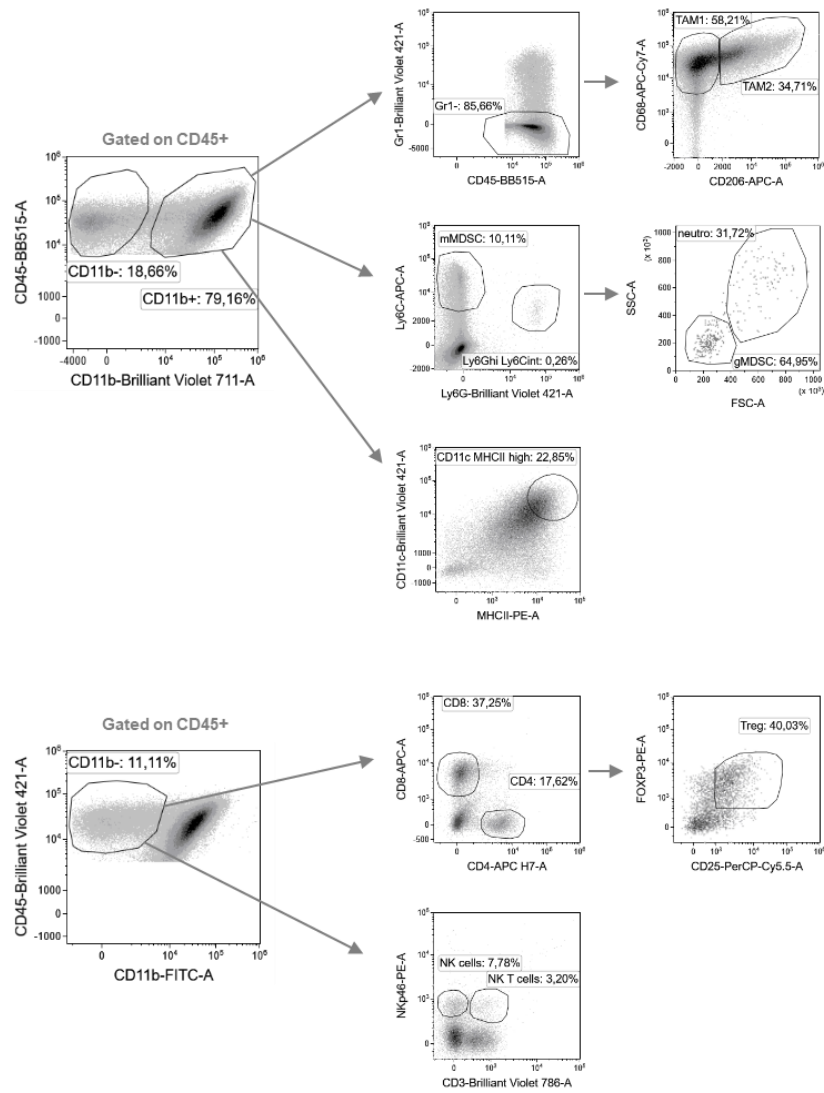
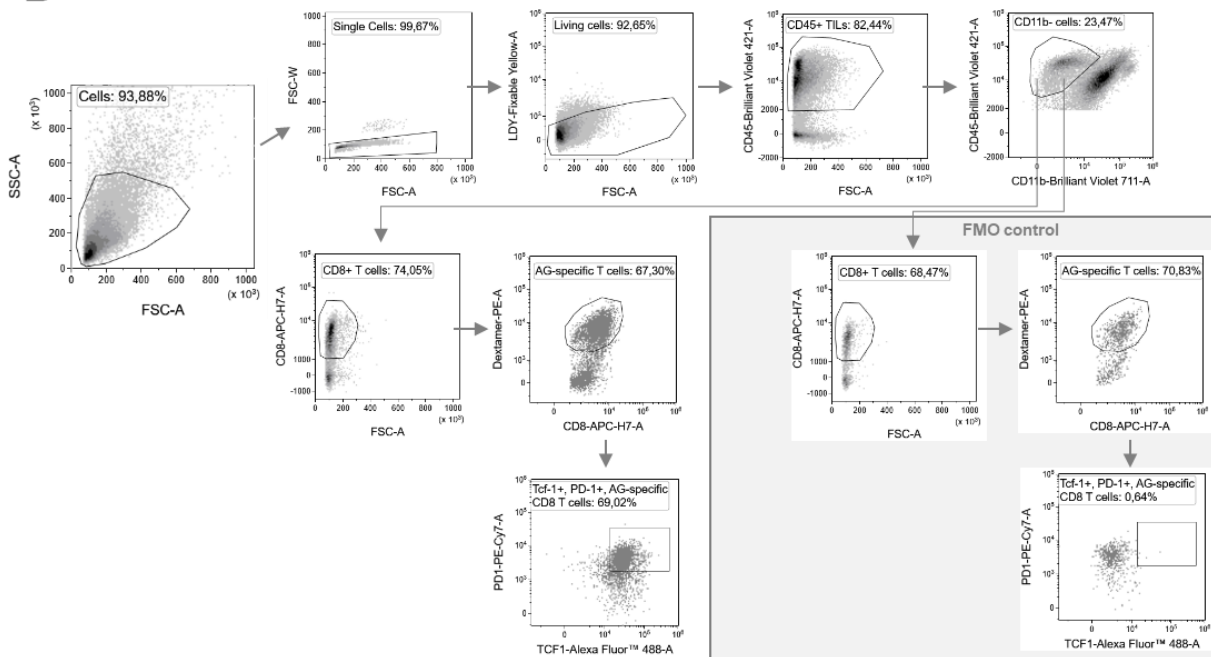


**G****H**

**Figure S2.** CD8+ T-cell characteristics in TC-1 tumor-bearing C57BL/6 mice. Tumor-growth curves, intratumoral NK and NKT cells, intratumoral and intranodal dendritic cell in MC-38 tumor-bearing C57BL/6 mice. (A) Left: Tumor-infiltrating CD8+ T cells were stimulated for 6 hours with E7 peptide and stained for IFN- $\gamma$ , TNF- $\alpha$  and CD107 $\alpha$ . Middle: The exhaustion status was examined on tumor-infiltrating E7-tetramer-positive CD8+ T cells in regards to expression of PD-1, Tim-3 and the effector marker KLRG1. Right: Tumor-derived E7-specific CD8+ T cells were stained for the co-expression of PD-1 and Tcf-1. Tumors were harvested on day 21 post-TC-1 tumor implantation. (B) Individual MC-38 tumor-growth curves in mm<sup>3</sup> post-tumor implantation for heterologous KISIMA-Mad46- and VSV-GP-Mad46-treated (KV) groups two days before tumor harvest (arrow) for FACS analysis. (C) Frequencies (top) and absolute numbers (bottom) of tumor-infiltrating NK (left) and NKT (right) cells were analyzed on day 18 post-MC-38 tumor implantation. (D) Frequencies (left) or absolute numbers (right) of either migratory CD103+ cDC1s (upper graphs) or tissue-resident CD8+ cDC1s (lower graphs) derived from inguinal lymph nodes on day 18 post-MC-38 tumor implantation. (E) From left to right: Frequencies and absolute numbers of tumor-infiltrating cDC2s, pDCs and moDCs on day 18 post-MC-38 tumor implantation. (F) From left to right: Frequencies and absolute numbers of intranodal migratory CD103+ cDC1s, cDC2s and moDCs on day 18 post-MC-38 tumor implantation. (G) From left to right: Frequencies and absolute numbers of tumor-infiltrating CD4 T cells, Th1, Th2, Th17 and Tregs on day 18 post-MC-38 tumor implantation. (H) From left to right: Frequencies and absolute numbers of intranodal CD4 T cells, Th1, Th2, Th17 and Tregs on day 18 post-MC-38 tumor implantation. Data in (A) to (H) are shown as means (grey bar) and are derived from at least two independent experiments. \*  $p < 0.05$ , \*\*  $p < 0.01$ , \*\*\*  $p < 0.001$ , \*\*\*\*  $p < 0.0001$  One-Way ANOVA followed by Tukey's multiple comparison test.

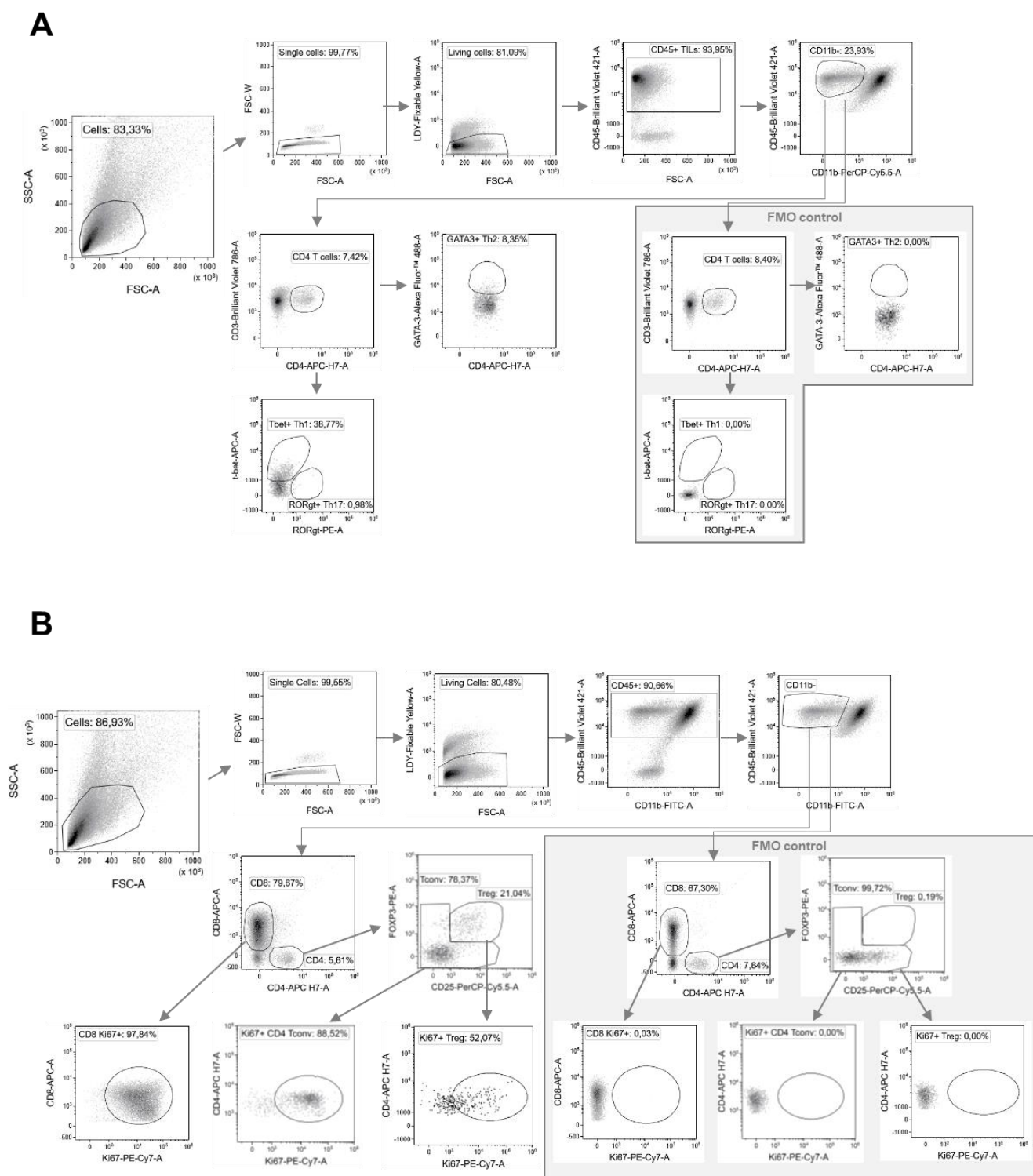


**Figure S3.** CD4<sup>+</sup> T cell characterization of tumor and tumor-dLN and blood Ag correlation graphs in MC-38 tumor-bearing C57BL/6 mice. (A) From top to bottom: Frequencies (left) or numbers (right) of intratumoral CD4<sup>+</sup> T cells, CD4<sup>+</sup> Th1 cells, CD4<sup>+</sup> Th2 cells, CD4<sup>+</sup> Th17 cells or CD4<sup>+</sup> Treg cells on day 18 post-MC-38 tumor implantation. (B) From top to bottom: Frequencies (left) or numbers (right) of intranodal CD4<sup>+</sup> T cells, CD4<sup>+</sup> Th1 cells, CD4<sup>+</sup> Th2 cells, CD4<sup>+</sup> Th17 cells or CD4<sup>+</sup> Treg cells on day 18 post-MC-38 tumor implantation. (C) From left to right: Correlation graphs (KV-treated mice) of blood-derived Rps1-specific CD8<sup>+</sup> T cell % to Adpgk-specific CD8<sup>+</sup> T cell %, blood-derived Rpl18-specific CD8<sup>+</sup> T cell % to Rps1-specific CD8<sup>+</sup> T cell %, blood-derived Rpl18-specific CD8<sup>+</sup> T cell % to Adpgk-specific CD8<sup>+</sup> T cell % and blood-derived Rpl18-specific CD8<sup>+</sup> T cell % to the sum of Rps1- and Adpgk-specific CD8<sup>+</sup> T cell % on day 17 post-MC-38 tumor implantation. Data in (A) and (B) are shown as means (grey bar), (A) is derived from two independent experiments and (B) from one experiment. \*  $p < 0.05$ , \*\*  $p < 0.01$ , \*\*\*  $p < 0.001$ , \*\*\*\*  $p < 0.0001$  One-Way ANOVA followed by Tukey's multiple comparison. Data in C) are derived from two independent experiments. \*  $p < 0.05$ , \*\*  $p < 0.01$  are calculated with simple linear regression.

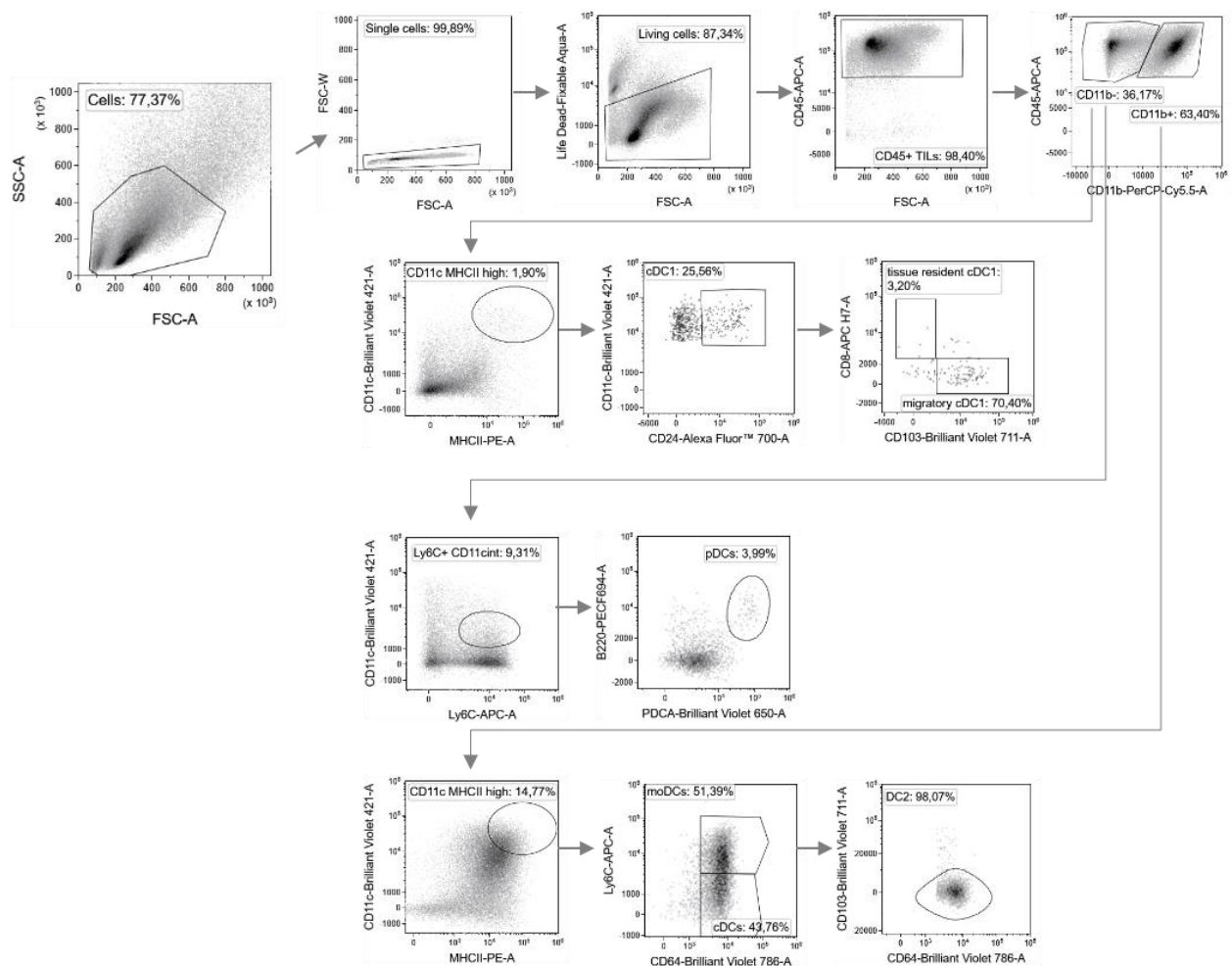
**A****B**

**Figure S4. TIL characterization and Tcf1<sup>+</sup>/PD-1<sup>+</sup> CD8<sup>+</sup> T-cell stemness gating strategies. (A) FACS gating strategy for the in-depth TIL characterization. (B) FACS gating strategy for the detection of Tcf1<sup>+</sup>/PD-1<sup>+</sup> antigen-specific CD8<sup>+</sup> T cells.**



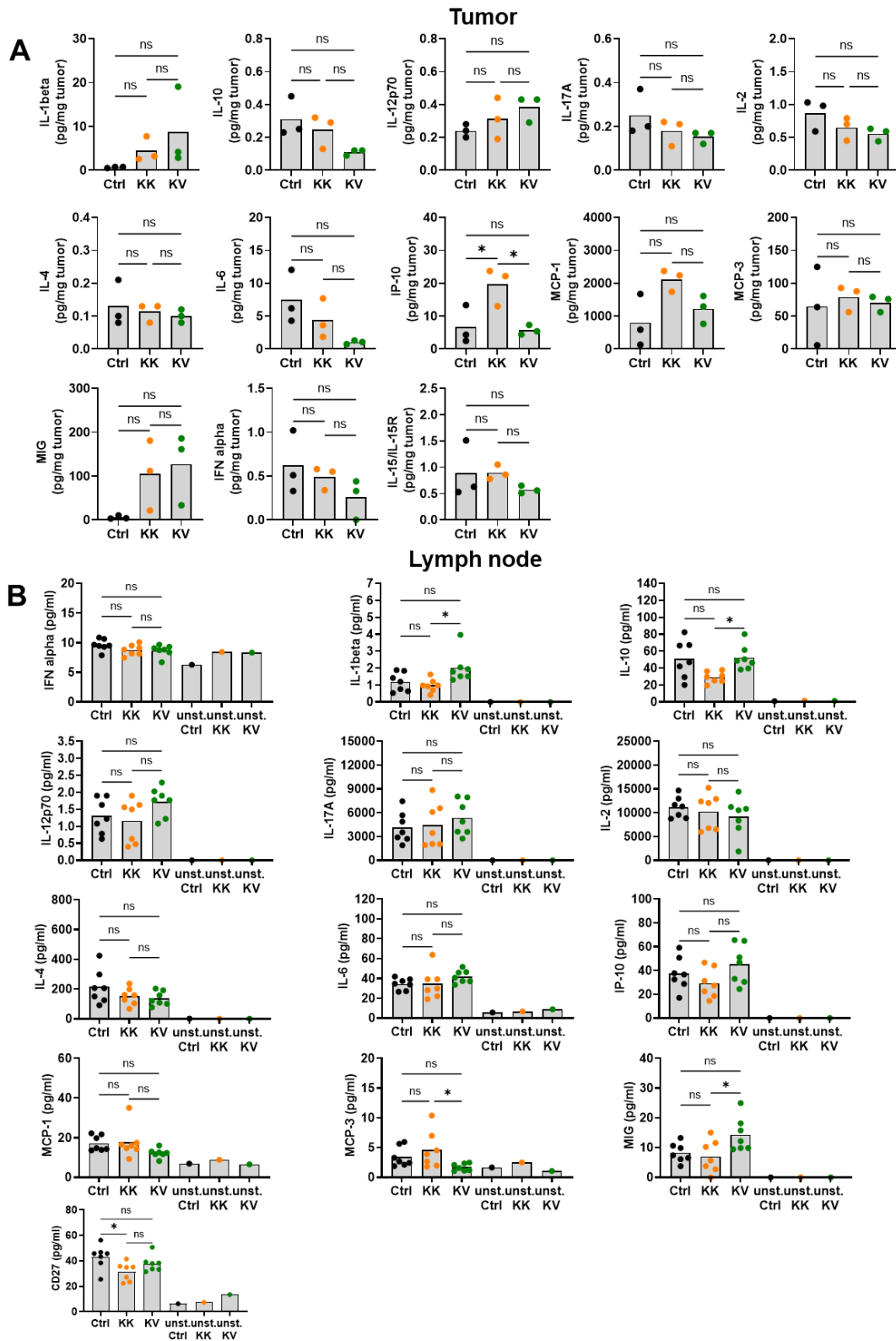


**Figure S5.** CD4<sup>+</sup> T-helper and CD4<sup>+</sup> regulatory T-cell gating strategies. (A) FACS gating strategy for the characterization of Th1, Th2 and Th17 CD4<sup>+</sup> cells. (B) FACS gating strategy for the detection of CD4<sup>+</sup> regulatory T cells and the investigation of proliferation via the proliferation marker Ki67.



**Figure S6. DC characterization gating strategies.** FACS gating strategies for the characterization of tissue-resident CD8+ cDC1s, migratory CD103+ cDC1s, moDCs, cDC2s and pDCs.





**Figure S7. Cytokine concentrations from tumor and tumor-dLNs of TC-1 tumor-bearing C57BL/6 mice.** (A) Cytokine (IL-1beta, IL-10, IL-12p70, IL-17A, IL-2, IL-4, IL-6, IP-10, MCP-1, MCP-3, MIG, IFN- $\alpha$ , IL-15/IL-15R) concentrations in pg/mL derived from tumor lysates that were harvested on day 21 post-tumor implantation and measured by Luminex. (B) Cytokine (IFN- $\alpha$ , IL-1beta, IL-10, IL-12p70, IL-17A, IL-2, IL-4, IL-6, IP-10, MCP-1, MCP-3, MIG, CD27) concentrations in pg/mL derived from inguinal lymph node cell suspension supernatants 24 hours after re-stimulation with  $\alpha$ -CD3/ $\alpha$ -CD28 beads or respective controls, which were harvested on day 21 post-tumor implantation and measured by Luminex. Data in (A) and (B) are shown as means (grey bar) and are derived from one experiment. \*  $p < 0.05$  One-Way ANOVA followed by Tukey's multiple comparison test.

**Table S1.** Mean values and statistical analysis of TIL characterization from Figure 4C on day 18 post-MC-38 tumor implantation. Statistical analysis was performed using One-Way ANOVA followed by Tukey's multiple comparison test.

Tumor	Mean % ( $\pm$ SD)			<i>p</i> -value		
	Ctrl	KK	KV	Ctrl vs KK	Ctrl vs KV	KK vs KV
CD8+ T cells	3.6 (2.6)	4.3 (1.9)	42.6 (23.6)	0.9893	<0.0001	<0.0001
CD4+ Tregs	0.4 (0.3)	0.4 (0.13)	0.36 (0.37)	0.991	0.9274	0.965
CD4+ Tconv	1.04 (0.17)	0.93 (0.18)	2.75 (1.63)	0.9503	<b>0.001</b>	<b>0.0005</b>
mMDSCs	9.5 (4.8)	7.3 (4.1)	5.22 (3.15)	0.4661	0.1123	0.5793
gMDSCs	0.38 (0.37)	0.15 (0.12)	0.19 (0.24)	0.1646	0.3499	0.9526
Neutrophils	0.22 (0.18)	0.08 (0.04)	0.05 (0.04)	<b>0.0309</b>	<b>0.0188</b>	0.8837
TAM1	33.3 (13.1)	42.3 (11.7)	20.7 (12.7)	0.2636	0.1248	<b>0.0052</b>
TAM2	41.5 (13.9)	35.7 (13.1)	24.2 (12.4)	0.5973	<b>0.0371</b>	0.2083
NKT	0 (0)	0 (0)	0.27 (0.47)	0.9999	0.0672	0.0672
NK	0.74 (0.33)	0.61 (0.22)	1.46 (0.67)	0.7568	<b>0.0046</b>	<b>0.0009</b>
B	0.46 (0.25)	0.36 (0.21)	0.68 (0.91)	0.9035	0.6597	0.4268
DCs	12.8 (6.3)	13.2 (6.5)	6.98 (4.39)	0.9857	0.1372	0.1042

**Table S2.** Antibodies and Co-Stimulators.

Antibody specificity	Clone	Manufacturer
CD45	30-F11	BD
CD11b	M1/70	BD
PD1	29F.1A12	Biologend
CD8	53-6.7	BD
KLRG1	2F1	BD
Tim3	B8.2C12	Biologend
CD103	M290	BD
CD49a	Ha31/8	BD
Ly6C	AL-21	BD
Ly6G	1A8	BD
PD-L1	10F.9G2	Biologend
CD103	M290	BD
MHCII	2D4	BD
CD11c	HL3	BD
CD40	HM40-3	eBioscience
B220	RA3-6B2	BD
CD24	M1/69	BD
CD64	X54-5/7.1	BD
Pdca	927	BD
CD4	RM4-5	BD
CD4	GK1-5	BD
CD25	PC61.5	TONBO Biosciences
FoxP3	FJK-16s	eBioscience
Ki67	solA15	eBioscience
Gr1	RB6-8C5	BD
LAP (TGF-beta)	TW 7-16B4	BioLegend
CD3	500A2	BD
CD335 (NKp46)	29A1.4	BD
CD206	C068C2	BioLegend
CD68	Fa-11	BioLegend
T-bet	eBio4B10 (4B10)	eBioscience
GATA-3	TWAJ	eBioscience
RORgt	AFKJS-9	eBioscience
CD69	H1.2F3	BD
TNFa	MP6-XT22	BD
IFNg	XMG1.2	BD
IL-2	JES6-5H4	BD
Isotype rat IgG1k	R3-34	BD
Isotype rat IgG1k	G234-2356	BD
Isotype rat IgG2bk	A95-1	BD
Anti-mouse CD28 (for co-stimulation)	Mouse EL-4 (T-cell lymphoma Cells)	BD
Anti-mouse CD3e (for co-stimulation)	H-2Kb-specific cytotoxic T lymphocyte clone BM10-37	BD

**Table S3.** Multimers.

Multimers	Cat. N°	Manufacturer
E7 MHC dextramer (H-2Db RAHYNIVTF - PE)	JA2195-PE	Immudex
Adpgk MHC dextramer (H-2Db ASMTNMELM - PE)	JA3803-PE	Immudex
Reps1 MHC tetramer (H2-Db AQLANDVVL - PE)	TB-5114-1	MBL
Rpl18 MHC tetramer (H2-Kb KILTFDRL - PE)	MKb-017	Tetramer Shop