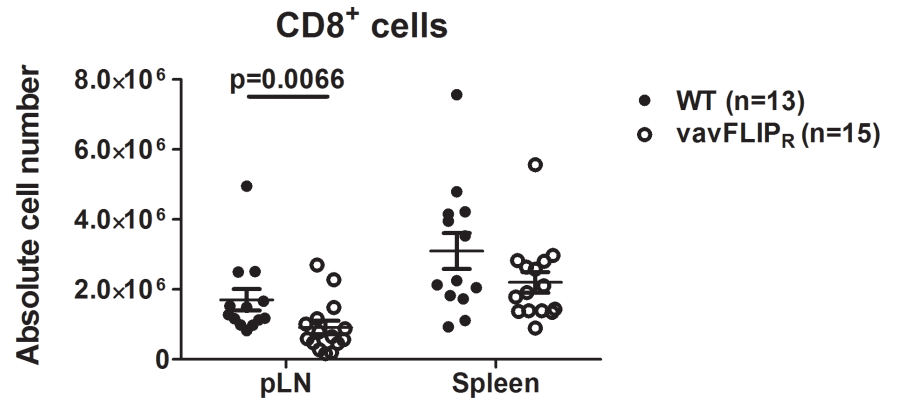
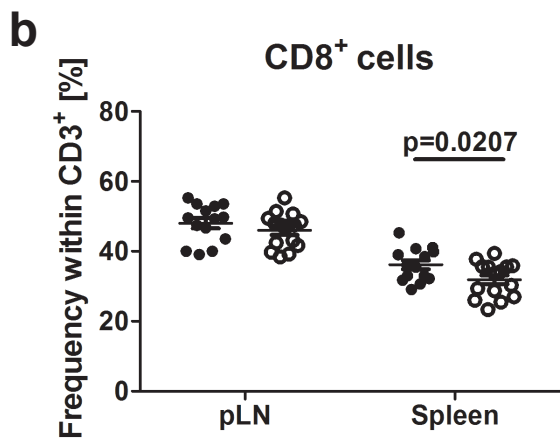
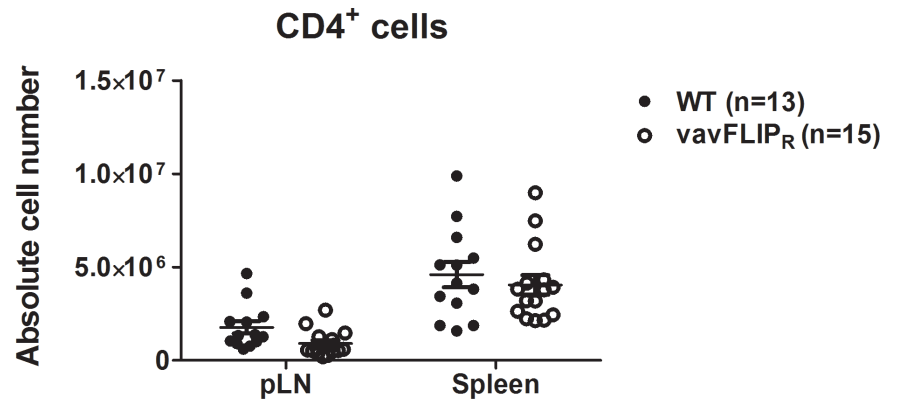
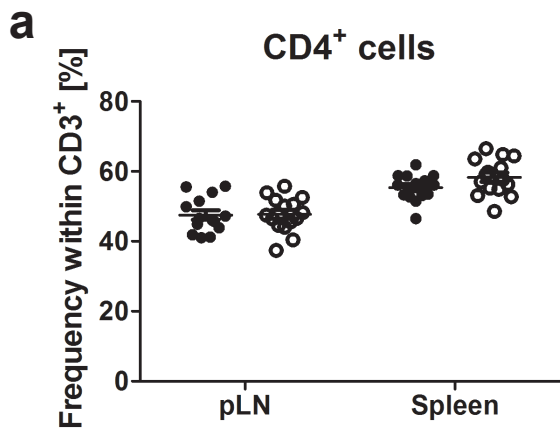


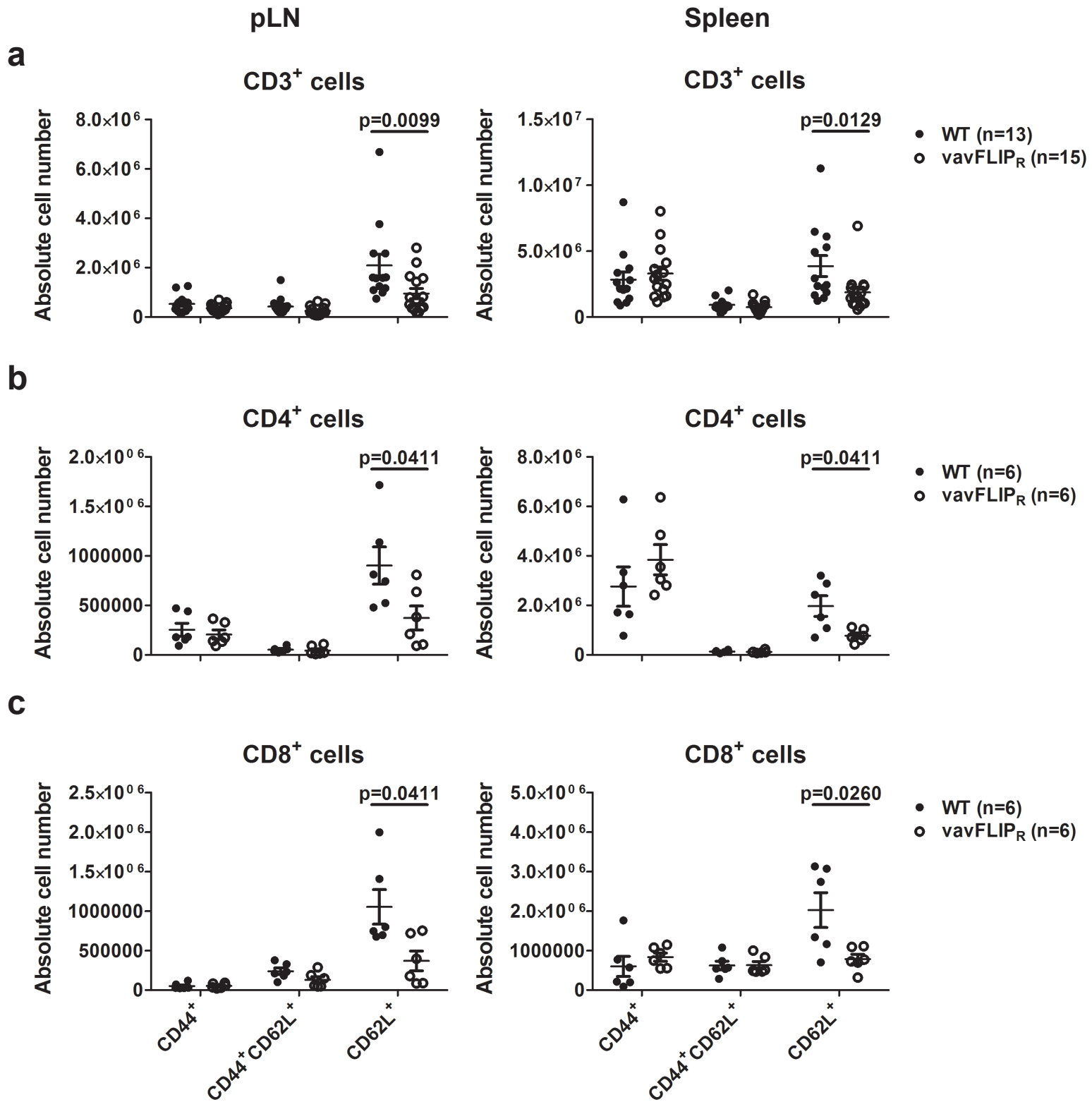
Supplementary table 1. Frequencies of CD11c⁺, F4/80⁺, Gr1⁺ (WT n=4, vavFLIP_R n=6) and CD49b⁺ (WT: n=2, vavFLIP_R: n=4) cells within the CD3⁻CD19⁻ compartment in spleen and bone marrow (BM) from one-year-old mice.

Mouse strain	Organ	Frequency within CD3 ⁻ CD19 ⁻ subset (\pm SD) [%]			
		CD11c ⁺	CD49b ⁺	F4/80 ⁺	Gr1 ⁺
WT	spleen	3.2 (2.7)	12.9 (2.8)	8.5 (7.4)	2.7 (1.7)
vavFLIP_R	spleen	3.8 (3.4)	12.0 (3.9)	8.5 (7.3)	5.0 (3.1)
WT	BM	0.8 (0.2)	4.8 (1.6)	16.1 (7.5)	51.1 (8.5)
vavFLIP_R	BM	0.9 (0.5)	6.9 (2.5)	19.8 (14.8)	55.7 (21.2)

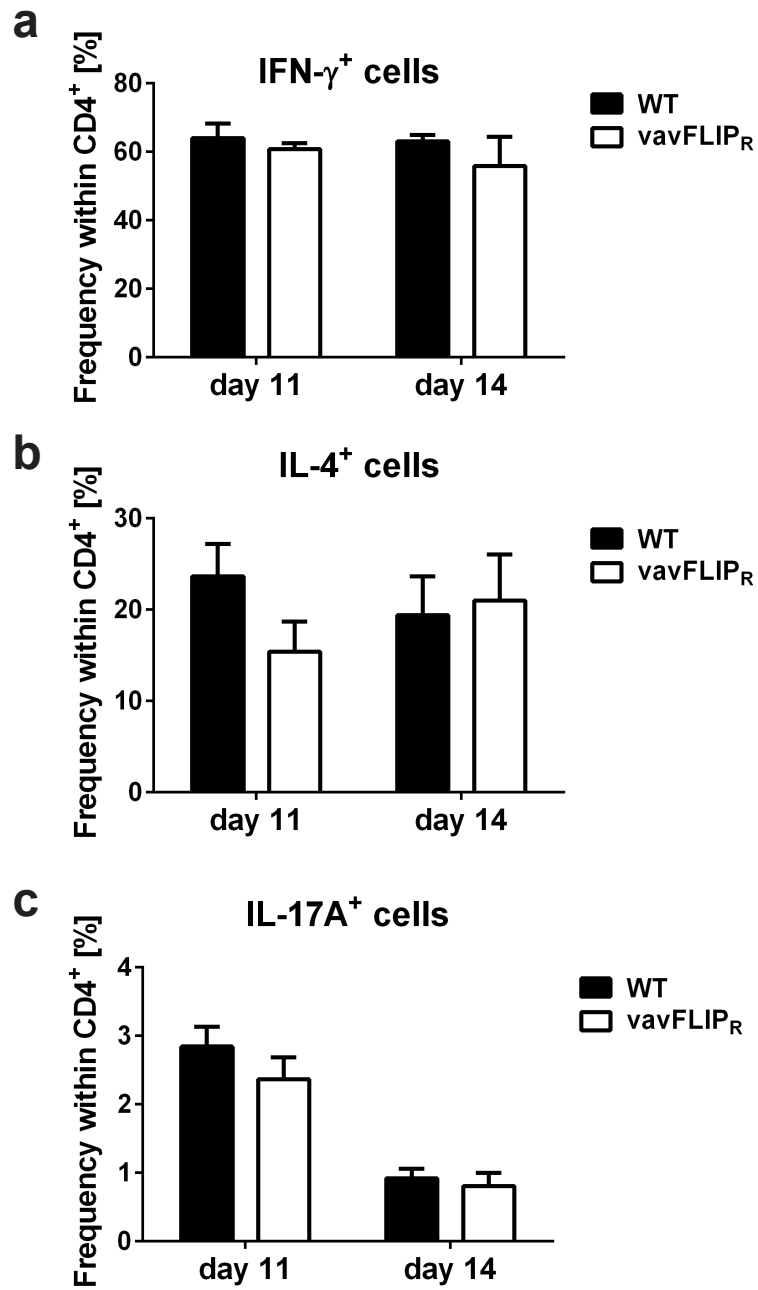
(SD = standard deviation)



Supplementary figure 1. Analysis of lymphocyte populations in 12-14 months old WT and vavFLIP_R mice. Frequency and absolute cell number of CD4⁺ cells (**a**) and CD8⁺ cells (**b**) in freshly isolated peripheral lymph nodes (pLN) and spleen from WT (n=13) and vavFLIP_R (n=15) littermates. Individual mice are represented as separate symbols. Horizontal lines show the mean pooled from five independent experiments; error bars display the SEM. Statistical analyses were performed with two-tailed nonparametric Mann-Whitney *U* tests.



Supplementary figure 2. T cell activation status in pLN and spleen from mice at 12-14 months of age. Absolute cell numbers of CD44⁺, CD44⁺CD62L⁺ and CD62L⁺ cells within CD3⁺ cells (**a**), CD4⁺ cells (**b**) and CD8⁺ cells (**c**) from peripheral lymph nodes (left panel) and spleen (right panel). Symbols represent individual WT (a: n=13, b-c: n=6) and vavFLIP_R (a: n=15, b-c: n=6) mice. Horizontal lines display (**a**) the mean pooled from five independent experiments ± SEM, (**b-c**) the mean of one experiment representative for two independent experiments ± SEM. Statistical analyses were performed with two-tailed nonparametric Mann-Whitney *U* tests.



Supplementary figure 3. Anti-CD3/anti-CD28 restimulation of peripheral lymph node cells from MOG-immunised WT and vavFLIP_R mice. WT and vavFLIP_R animals were injected with the MOG₃₅₋₅₅-peptide in complete Freund's adjuvant. Peripheral lymph node cells isolated on day 11 (WT n=4, vavFLIP_R n=4) and day 14 (WT n=5, vavFLIP_R n=3) after immunisation were restimulated with anti-CD3 and anti-CD28 (both 2 µg/ml) for 24 h, followed by flow cytometry analysis of IFN-γ, IL-17A and IL-4 producing T cells.