

Fig S1- phenotypic characterization of splenic and BM pDCs in *SLAMF9*^{-/-} mice and illustration of deleted sequence in *SLAMF9*^{-/-} mice.

(A) Sorted pDCs (CD19-CD11C^{inter}B220+PDCA⁺), B cells (CD19⁺ B220⁺) and cDCs (CD19-B220-CD11B+CD11C^{high}) were analyzed for the mRNA levels of TCF-4 by qRT-PCR. Graphs show relative expression of target gene/reference gene (L32); n=3 mice. **(B)** Schematic illustration of 211 bp deletion on chromosome one within the genomic region of the *SLAMF9* gene. The blue bars indicate exon1 and exon2. Green vertical line within exon1 indicates the location of the initiator methionine. The deleted sequence is indicated between the inner red diagonal line. **(C-D)** Analysis of BM pDCs (CD19-CD11C^{inter}B220+PDCA⁺) in WT and *SLAMF9*^{-/-} mice shown in a representative dot plot and a graphical view. **(E)** Graph shows the percent of pDCs in the spleen. Results are a summary of 6 independent experiments. n= 22 mice. **(F-G)** Analysis of lymph node WT and *SLAMF9*^{-/-} pDCs. Graph shows the percent of LN pDCs from total live cells (F) and absolute numbers of pDCs in the iLNs (G). Results are a summary of at least three independent experiments. n= 12-22 mice. **(H-J)** Analysis of LN B cells (CD19⁺ B220⁺) (H) and LN cDCs (CD19-B220-CD11B+CD11C^{high}) (I) and macrophages (CD19-B220-CD11C-CD11B+Ly6G- F480+) (J) in WT or *SLAMF9*^{-/-} mice.

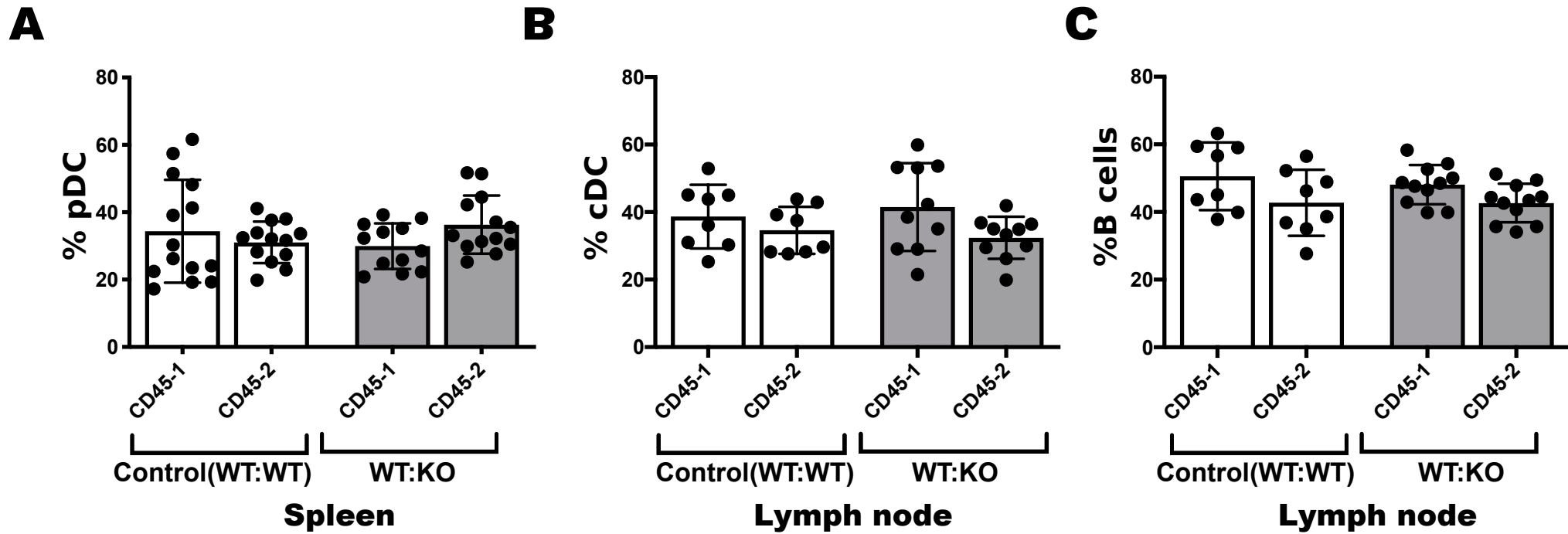


Fig S2 -B cell and CDC frequencies in chimeric mice

Lethally irradiated CD45.1 mice were transplanted with a total of 2×10^6 BM cells from CD45.1 (WT) and CD45.2 (WT) mice, or from CD45.1 (WT) and CD45.2 (KO: SLAMF9^{-/-}) mice at a 1:1 ratio. Mice were harvested 8 weeks post-transplant and splenic pDCs and cDC and B cell populations in the lymph node were analyzed. **(A)** Graph shows the percentage of pDCs in the spleen derived from CD45.1 and CD45.2 lineages in chimeric mice. **(B-C)** Graph shows the percentage of cDCs (B), or B cells (C) derived from CD45.1 and CD45.2 lineages in chimeric mice. Results are representative of experiments from three independent chimera. n=11-13 mice.

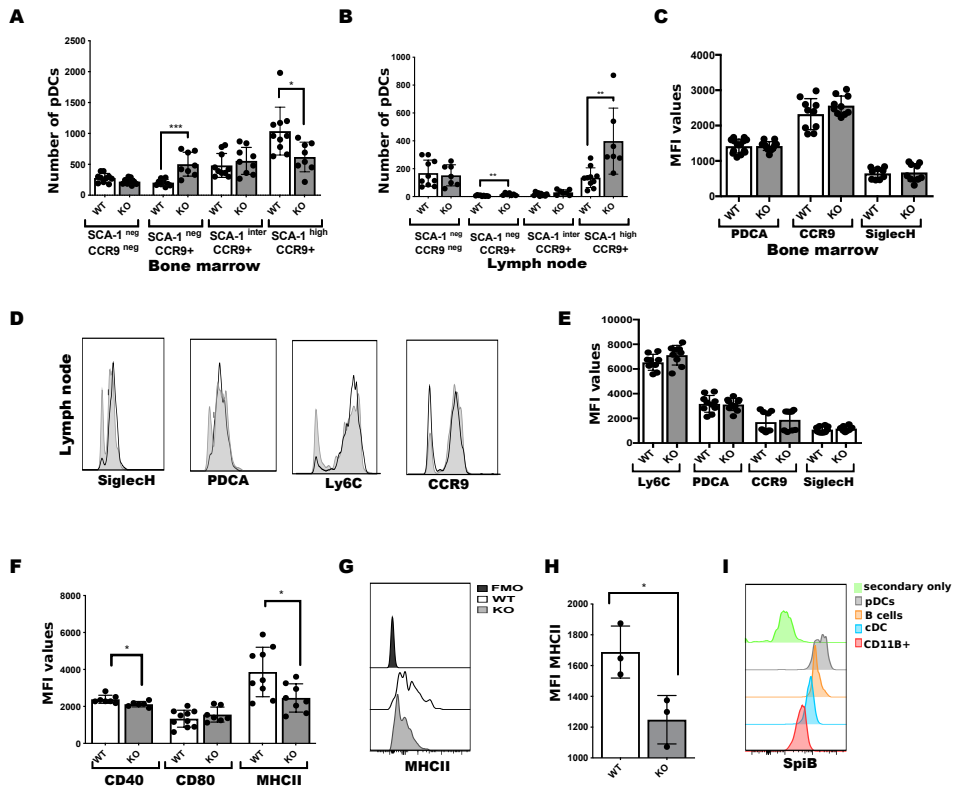


Fig S3- SLAMF9^{-/-} LN pDCs have similar activation markers

(A-B) pDCs were harvested from the BM and LN of WT and SLAMF9^{-/-} mice (KO). Graph shows the absolute numbers of pDCs subsets per 300,000 cells in the BM (A) and in the iLNs (B). **(C)** Graph shows the MFI values of pDC markers in the BM. **(D-F)** pDCs were harvested from the LN of WT and SLAMF9^{-/-} mice (KO). Representative histograms of staining and MFI values for pDC markers in LN of WT and SLAMF9^{-/-} pDCs (D-E) and MFI values of co-stimulatory markers in LN pDCs (CD19⁻ CD11C^{inter} B220⁺ PDCA⁺) (F). Results are a summary of three independent experiments. n=8-11 mice. **(G-H)** Analysis of MHC class II expression in pDCs from the LN (CD19⁻ CD11C^{inter} B220⁺ CCR9⁺ SCA-1⁺ CD11B⁻). Representative histogram of staining for MHC class II in WT and SLAMF9^{-/-} pDCs compared with fluorescence minus one (FMO) control (G) and MFI values of MHC class II of WT and SLAMF9^{-/-} pDCs (H). **(I)** Analysis of SPIB expression in immune cell population in the LN. Representative histogram of staining in B cells, cDC cells, pDCs and CD11B cells compared with negative control (only secondary).

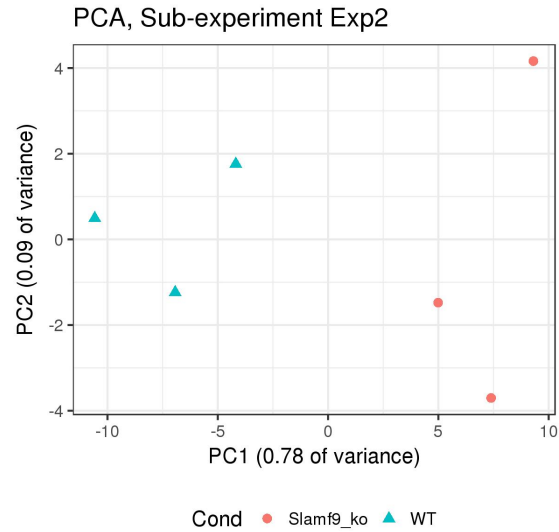
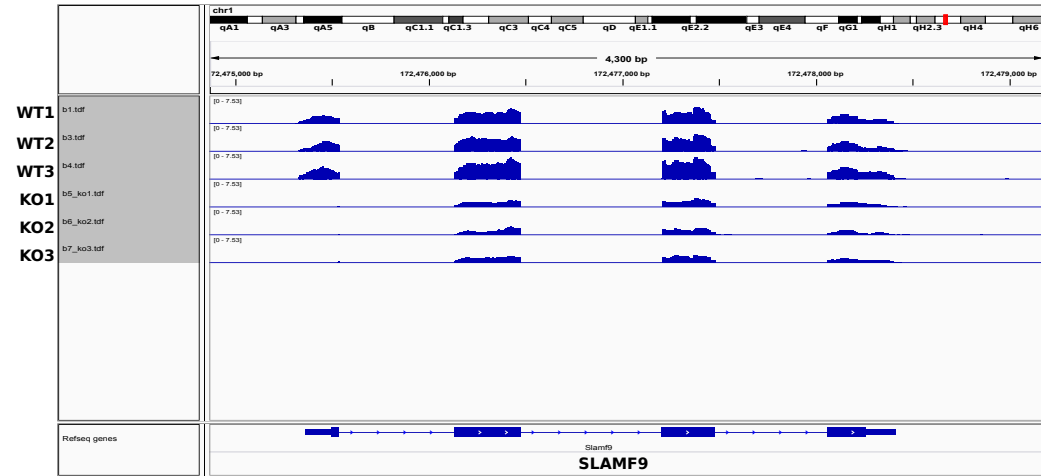
A**B**

Fig S4-RNA-seq PCA analysis and IGV plots

BM pDCs (CD19-CD11B-CD11C^{inter} B220+PDCA+) were sorted from WT and SLAMF9^{-/-} mice (KO). RNA was extracted and subjected to sequencing. Differentially expressed genes were identified by p value < 0.05 and fold change greater or equal to 1.5. **(A)** Principal component analysis (PCA) demonstrating distinct clusters for pDC samples obtained from WT and SLAMF9^{-/-} mice. **(B)** IGV image demonstrating deleted exon1 region in knockout samples after normalization of the coverage to the library size.

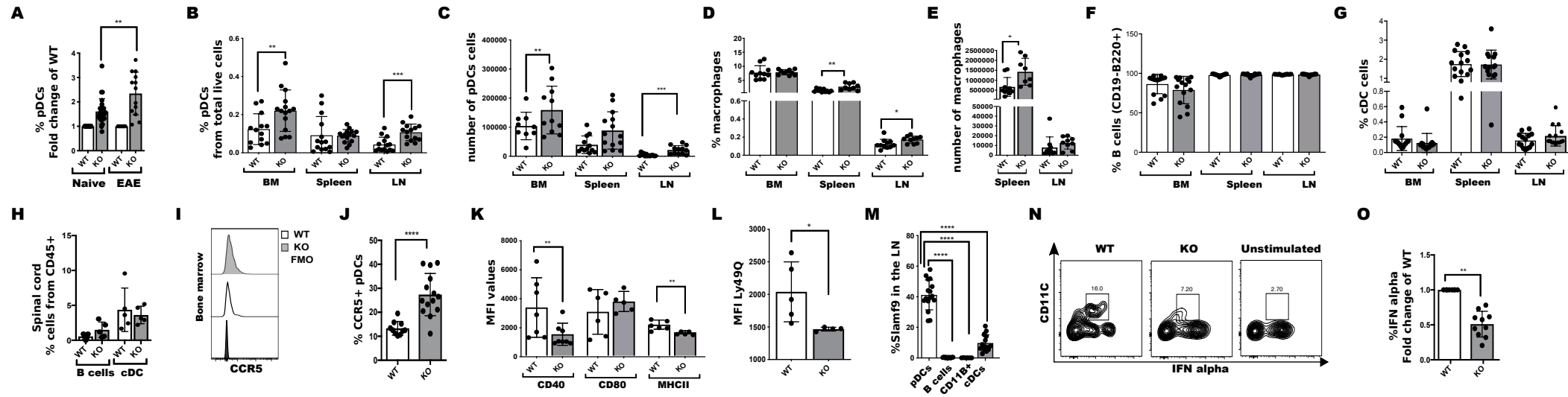


Fig S5-Analysis of cells following 14 days of EAE induction

EAE was induced in C57BL/6 WT and SLAMF9 ^{-/-} mice by subcutaneous injection of MOG33-55 peptide in complete Freund's adjuvant and further intraperitoneal injection of Pertussis toxin on days 0 and 2. **(A)** Graph shows the increase in SLAMF9 ^{-/-} pDCs (KO) compared to WT cells in healthy animals, and during EAE. **(B-C)** Analysis of pDCs in SLAMF9 ^{-/-} compared with WT mice in EAE. Graph shows the percent of pDCs (CD19- CD11C^{inter} B220+PDCA+ CD11B-) from total live cells (B) and absolute numbers in the iLNs (C). **(D-E)** Analysis of macrophages in SLAMF9 ^{-/-} compared with WT mice. Graph shows the absolute numbers of macrophages in the spleen and lymph nodes (D) and the percent of macrophages (CD19-B220-CD11C-CD11B+Ly6G-F480) from total live cells (E) **(F-G)** Graph shows the percentages B cells (CD45+B220+CD11B-) and the percent of cDC (CD19-B220-CD11B+CD11C^{high}) (G). **(H)** Graph shows the percentages of cDC (CD45+CD11C^{high} CD11B+) and B cells (CD45+B220+CD11B-) in the spinal cord, 16 days after EAE induction. Results are representative of two independent experiments; each dot represents two mice.

(I-J) Analysis of CCR5 levels in BM pDCs following 14 days of EAE induction. Representative histograms of CCR5 on WT and SLAMF9(-/-) pDCs compared with fluorescence minus one (FMO) control(I) Graph shows the percent of CCR5+ pDCs in the BM (J). Results are representative of two independent experiments; n=10-14 mice; p****<0.0001. **(K-L)** Analysis of MFI values in WT and SLAMF9^{-/-} pDCs in the LN. Graph shows MFI values of co-stimulatory markers in LN pDCs (K) and MFI values for Ly49Q (L). **(M)** SLAMF9 protein levels were analyzed in following 14 days post EAE induction in the LN. Graph shows protein expression in pDCs (CD19-CD11C^{inter} B220+PDCA+ CD11B-), B cells (CD19+ B220+), cDCs (CD19-B220-CD11B+CD11C^{high}) and in macrophages (CD11B+CD11C^{neg} CD19-SSC^{low}). **(N-O)** pDCs were freshly isolated at day 16 following EAE induction and stimulated for 18h with 1 μM of ODN1585. (N) Representative plot of IFNα levels on WT and SLAMF9(-/-) pDCs compared with unstimulated control. (O) Graph shows the fold of change in IFNα percent from WT following TLR9 stimulation. Results are representative of two independent experiments n=8-10 ;p****<0.0001.