

**Figure S1. Antigen-specific T cell responses in mice with deletion of TNF in T cells and myeloid cells. A, B.** Frequency of IFN $\gamma$ - and IL-17A – producing CD4 T cells in WT, T-TNF KO, M-TNF KO and TNF KO splenocytes at 14<sup>th</sup> day after second immunization in the absence of type II collagen. Proliferation of CD4 T cells in WT, T-TNF KO, M-TNF KO and TNF KO splenocytes at 14<sup>th</sup> day after second immunization activated with type II collagen (40 mcg/ml) **(C)** or anti-CD3/anti-CD28 **(D)**. CFSE dilution was analysed 24 hours after stimulation. **E.** Cell proliferation in WT and TNF KO splenocytes measured by <sup>3</sup>H-thymidin incorporation at day 14 after 2<sup>nd</sup> immunization. Cells were activated with 10 mcg/ml of chicken type II collagen for 72 hours, <sup>3</sup>H-Thymidine was added for last 16 hours of incubation. Data are representative of two independent experiments.

**Figure S2. The role of myeloid cell-derived TNF during collagen induced arthritis. A.** Soluble TNF levels produced by bone marrow derived macrophages (BMDM) generated from WT (tmTNFKI/TNF<sup>flox</sup>×Mlys-Cre<sup>wt/wt</sup>) and tm-M-TNF KI (tmTNFKI/TNF<sup>flox</sup>×Mlys-Cre<sup>Cre/wt</sup>) mice and stimulated with LPS (100 ng/ml) for 4 hours. **B.** Mean fluorescence intensity of TNF produced by WT and tm-M-TNF KI BMDM stimulated with LPS (100 ng/ml) for 4 hours in the presence of Brefeldin A (5 mcg/ml) and stained intracellularly for TNF. **C.** Frequency of TNF producing cells in WT, T-TNF KO, M-TNF KO and TNF KO mice at 14<sup>th</sup> day after second immunization. **D.** Expression of Ly6C and Gr1 on TNF-producing CD11b<sup>+</sup> cells in WT, T-TNF KO, M-TNF KO and TNF KO mice at 14<sup>th</sup> day after second immunization. Splenocytes from various mouse strains were activated with LPS (100 ng/ml) for 4 hours in the presence of Brefeldin A (5 mcg/ml) and stained intracellularly for TNF. **E.** YFP expression in blood monocytes (CD11b<sup>+</sup>CD115<sup>+</sup> cells) in ROSA26-STOP-YFP×Mlys-Cre mice. **F.** TNF mRNA expression levels in synovial tissue of

WT, M-TNF KO, T-TNF KO and TNF KO mice at 21<sup>st</sup> day after immunization. Data are representative of two independent experiments. \* $P < 0.05$ , \*\* $P < 0.01$ , \*\*\* $P < 0.001$ , as calculated by Students' *t*-test.

**Figure S3. Analysis of infiltrating cells in the inflamed paws during CIA.**

Representative dot plots of embryonically derived joint macrophages (F4/80<sup>+</sup>CD11b<sup>-</sup>) (A), inflammatory monocytes (CD11b<sup>+</sup>F4/80<sup>+</sup>Gr1<sup>low</sup>) and neutrophils (CD11b<sup>+</sup>F4/80<sup>-</sup>Gr1<sup>high</sup>) (B), CD4 T cells (C) in arthritic WT, T-TNF KO and M-TNF KO mice and CII-immunized TNF KO mice at 14<sup>th</sup> day after second immunization. Data are representative of two independent experiments.

**Figure S4. Analysis of T cell compartment in mice with cell-type specific TNF deletion during arthritis.**

**A.** Frequency of T regulatory T cells in spleens of WT, T-TNF KO, M-TNF KO and TNF KO mice at day 14 after second immunization. **B.** Mean fluorescent intensity of FOXP3 in Treg cells in spleens of WT, T-TNF KO, M-TNF KO and TNF KO mice at day 14 after second immunization. **C.** Incidence of arthritis in T-TNF KO mice treated with antibodies directed against TNFR2 (Clone: TR75-54.7; 100 mcg/mouse, i.p., twice per week) or IFN $\gamma$  (Clone: XMG1.2; 100 mcg/mouse, i.p., twice per week). Frequency of inflammatory monocytes (CD11b<sup>+</sup>Ly6C<sup>+</sup>Gr1<sup>low</sup>) (D) and IL-12p40 producing DCs (CD11c<sup>+</sup>MHCII<sup>+</sup>) (E) in mice with conditional ablation of TNF during arthritis (day 6 after immunization). **E.** Data are representative of two independent experiments. \* $P < 0.05$ , \*\* $P < 0.01$ , \*\*\* $P < 0.001$ , as calculated by Students' *t*-test.