

**Figure S1**: MHC-II<sup>-/-</sup> mice have normal B cell and  $CD8^+$  T cell populations, but lack  $CD4^+$  T cells. Flow cytometry of the deep cervical lymph nodes of wild type (**A**) and MHCII<sup>-/-</sup> (**B**) mice showing CD4<sup>+</sup> and CD8<sup>+</sup> lymphocytes in the TCR $\beta^+$  population and CD19<sup>+</sup>B220<sup>+</sup> B cells in the CD45<sup>+</sup>TCR $\beta^-$  population. Numbers indicate percent CD4<sup>+</sup> (upper left) and CD8<sup>+</sup> (lower right), as a percentage of TCR $\beta^+$  cells and CD19<sup>+</sup>B220<sup>+</sup> B cells as a percentage of the CD45<sup>+</sup>TCR $\beta^-$  population.



**Figure S2**: No difference in contralateral, uninjured retinas of all mouse strains and experimental manipulations examined in this manuscript. (A)  $MHCII^{-/-}$  mice injected with 3 x 10<sup>6</sup> wild type CD4<sup>+</sup> T cells on the day of injury exhibit no difference in neuronal counts in the uninjured retinas compared to those injected with saline. Retinal ganglion cell counts (% of  $MHCII^{-/-}$ ) of  $MHCII^{-/-}$  mice and  $MHCII^{-/-}$  mice injected with wild type

 $CD4^+$  T cells as assessed by Fluoro-Gold staining (n = 3, wild type thymus; n = 3) MHCII<sup>-/-</sup> thymus; Student's t-test). (B) MHCII<sup>-/-</sup> mice implanted with wild type thymi exhibit no difference in neuronal counts in the uninjured retinas compared to those implanted with MHCII<sup>-/-</sup> thymi. Retinal ganglion cell counts (% of MHCII<sup>-/-</sup> + MHCII<sup>-/-</sup> thymus) of MHCII<sup>-/-</sup> mice implanted with either a MHCII<sup>-/-</sup> or wild type thymi (six weeks after implantation) as assessed by Fluoro-Gold staining (n = 5, wild type thymus; n = 8 MHCII<sup>-/-</sup> thymus; Student's t-test). (C) OTII/Rag1<sup>-/-</sup> mice exhibit no difference in retinal ganglion cell number in the uninjured retinas compared to Rag1<sup>-/-</sup> mice. Retinal ganglion cell counts (% of Rag1<sup>-/-</sup>) of Rag1<sup>-/-</sup> and OTII/Rag1<sup>-/-</sup> mice assessed by Fluoro-Gold staining (n = 3, Rag1<sup>-/-</sup>; n = 3, OTII/Rag1<sup>-/-</sup>; Student's t-test). (D) IL-4<sup>-/-</sup> mice exhibit no difference in retinal ganglion cell number in the contralateral retinas compared to wild type mice. Bar graphs represent retinal ganglion cell counts (% of wild type) of IL-4<sup>-/-</sup> or wild type mice, assessed by Fluoro-Gold staining (n = 5, wild type; n = 5, IL-4<sup>-</sup> <sup>/-</sup>; Student's t-test). (E) Mice transplanted with IL-4<sup>/-</sup> bone marrow exhibit no difference in retinal ganglion cell number in the contralateral retinas compared to those transplanted with wild type bone marrow. Retinal ganglion cell counts (% of wild type  $\Rightarrow$  wild type) of wild type  $\Rightarrow$  wild type or IL-4<sup>-/-</sup>  $\Rightarrow$  wild type bone marrow chimeras. Bone marrow was allowed to engraft for 6 weeks before optic nerve injury, and retinal ganglion cell counts were assessed by Fluoro-Gold staining (n = 3, wild type bone marrow; n = 3, IL- $4^{-/-}$  bone marrow; Student's t-test). (F) Rag $1^{-/-}$  mice receiving IL- $4^{-/-}$  CD $4^+$  T cells demonstrate no difference in retinal ganglion cell number in the contralateral retinas compared to Rag1<sup>-/-</sup> mice receiving wild type T cells. Retinal ganglion cell counts (% of Rag1<sup>-/-</sup> mice injected with wild type T cells) of Rag1<sup>-/-</sup> mice injected with either wild type or IL-4<sup>-/-</sup> CD4<sup>+</sup> T cells 3 weeks before optic nerve injury. Retinal ganglion cell counts were assessed by Fluoro-Gold staining (n = 7, wild type; n = 7, IL-4<sup>-/-</sup> T cell injected; Student's t-test). (G) MyD88<sup>-/-</sup> mice exhibit no difference in retinal ganglion cell number in the contralateral retinas compared to wild type mice. Retinal ganglion cell counts (% of wild type) of wild type and MyD88<sup>-/-</sup> mice were assessed by Fluoro-Gold staining (n = 3, wild type; n = 3, MyD88<sup>-/-</sup>; Student's t-test). (H) Mice receiving MyD88<sup>-</sup> <sup>/-</sup> bone marrow exhibit no difference in retinal ganglion cell number in the contralateral retinas compared to those receiving wild type bone marrow. Retinal ganglion cell counts (% of wild type  $\Rightarrow$  wild type) of wild type  $\Rightarrow$  wild type or MyD88<sup>-/-</sup>  $\Rightarrow$  wild type bone marrow chimeras. Bone marrow was allowed to engraft for 6 weeks before optic nerve injury, and retinal ganglion cell survival was assessed by Fluoro-Gold staining (n = 3, n)wild type bone marrow recipients;  $n = 3 \text{ MyD88}^{-/-}$  bone marrow recipients; Student's ttest).



**Figure S3:** Classical TLR ligands do not increase Gata3 mRNA expression in  $CD4^+$  T cells. Gata3 expression (mean ± s.e.m.) in sorted wild type  $CD4^+$  T cells co-cultured with Pam3CSK4 (A), zymosan (B), LPS (C), flagellin (D), FSL-1 (E), ssRNA (F), CpG (G), or HMGB1 (H) at the indicated concentrations for 3 days (n = 3 per group, representative of two experiments).



**Figure S4**: *IL-4* induces axon elongation in cultured neurons (**A**, **B**) The application of a single dose of 100 pg/ml recombinant IL-4 significantly increased the length of axons (mean  $\pm$  s.e.m.) of isolated primary neurons. Representative microphotographs (**A**) and quantification of axonal length (**B**) of isolated neurons treated with PBS or a single dose of 100 pg/ml recombinant IL-4. Scale bar: 20 µm. (n[well] = 4 per group. \*\*\*, p < 0.001; One-way ANOVA with Bonferroni's post-test).



**Figure S5:** *IL-4R expression on spinal cord axons.* Axons in longitudinal sections of the spinal cord show tubular, membrane-related IL-4R expression (green) pattern, adjacent to the stained axonal neurofilament (SMI, red). The presumed membrane is indicated by dotted lines. Two representative axons (left and right panels) are provided. Scale bar: 1µm.



Perilesional (tangential section)

Caudal (5mm)

*Figure S6: T* cells migrate caudally from the lesion site after injection. CFDA<sub>SE</sub> labeled T cells were injected into the site of injury immediately after spinal cord injury. Spinal cords were isolated and visualized for CFDA<sub>SE</sub> labeled T cells 6 days after the lesion. (Scale bars: 15 µm). T cells were found at least 5 mm away from the site of injection, where axonal regrowth was observed.