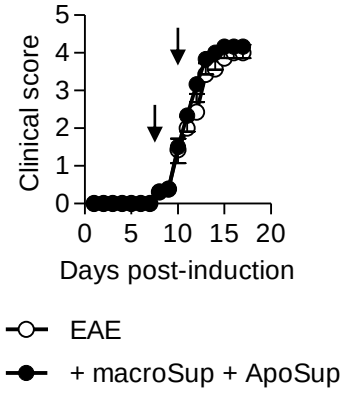

FIGURE S1. Apoptotic cell supernatant and macrophage supernatant injections do not improve EAE. Clinical score of EAE mice treated or not (EAE) twice (black arrows) with supernatant of apoptotic cells plus the supernatant of macrophages (1 mL) (+ macroSup + ApoSup). Data representative of 2 independent experiments as mean +/- SEM, with 5 mice per group. Statistical significance was assessed using ANOVA two-way and Bonferroni posttest

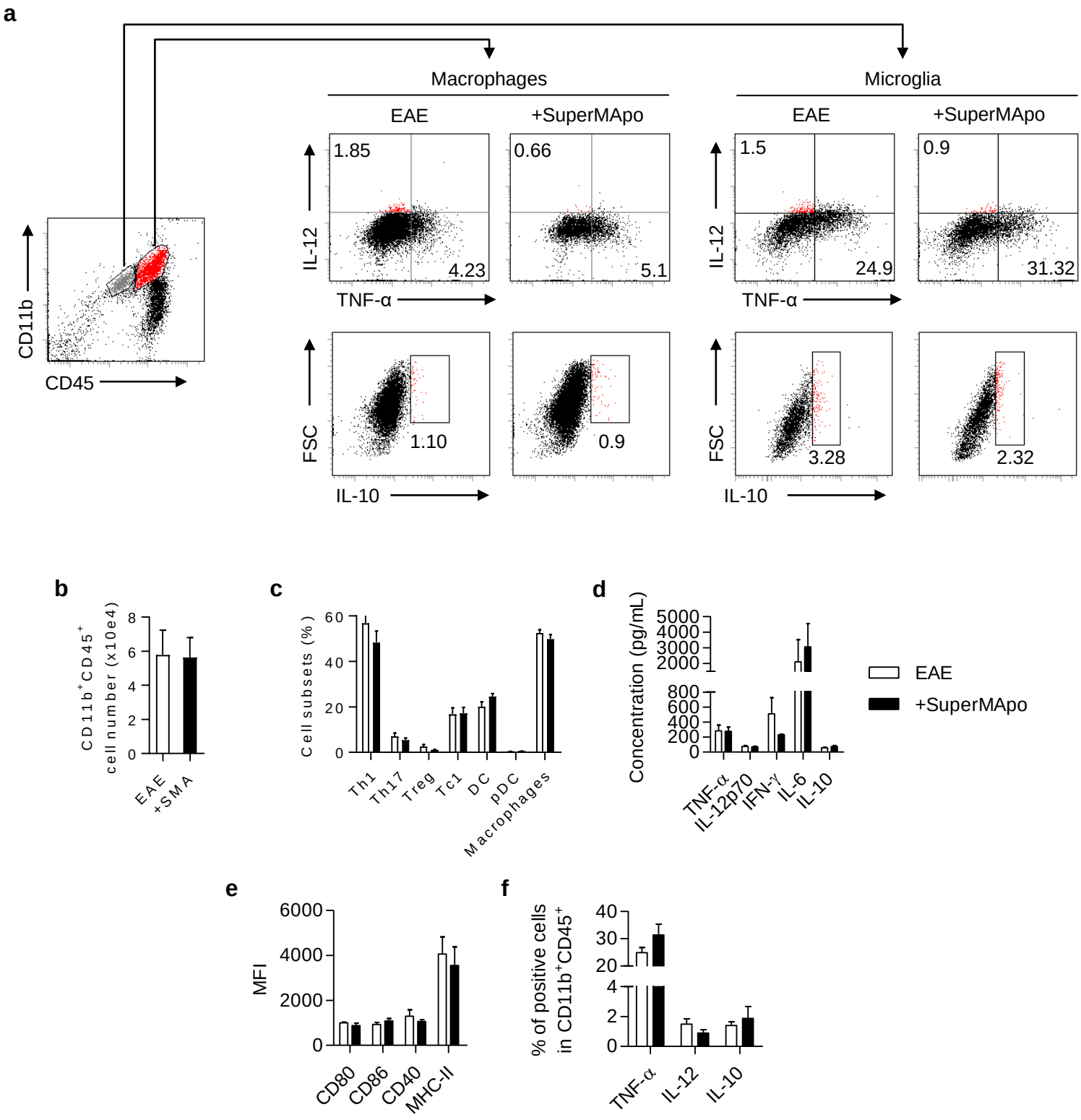

FIGURE S2. SuperMApo treatment modulates CNS infiltrating macrophage but not of microglia activation profiles. (**a**) Gating strategy for CNS macrophages and microglia. (**b**) Absolute number of CD11b^{med}CD45^{+} microglia in spinal cords of EAE mice 72 h after SuperMApo (+SAM) or vehicle (EAE) injection. Data representative of 3 independent experiments shown as mean +/- SEM with 5 mice per group. (**c**) Percentages of CNS cell subsets in mice from **b**. (**d**) Spin cord fluid concentrations of TNF-α, IL-12p70, IFN-γ, IL-6 and IL-10 in mice from **b**. (**e**) Mean Fluorescence Intensity (MFI) of CD80, CD86, CD40 and MHC-II expressed by microglia in mice from **b**. (**f**) Percentages of TNF-α, IL-12 and IL-10 positive cells microglia in the spinal cords of mice from **b**.

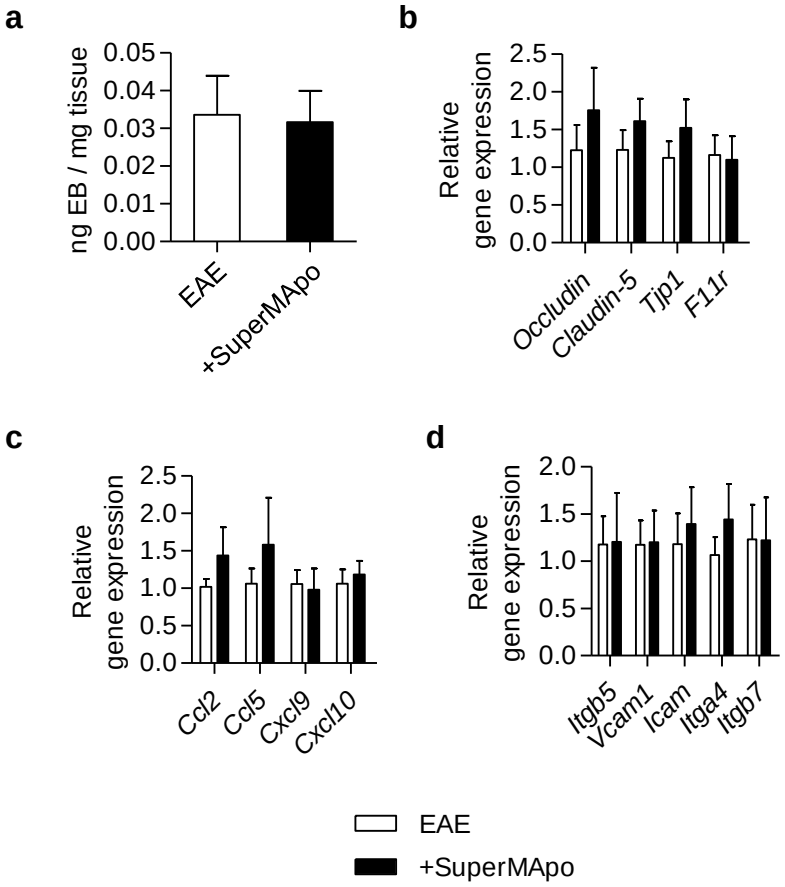

FIGURE S3. SuperMApo treatment does not modulate directly the blood brain barrier. (**a**) Evan's Blue (EB) dye quantification in EAE mice receiving SuperMApo or not (EAE) (n= 5 mice). Data are given as ng of EB/mg of tissue, 5 mice per group. Data representative of 3 independent experiments shown as mean +/- SEM. Spinal cord of EAE mice receiving SuperMApo or not (EAE) were analysed for the mRNA expression of *Occludin*, *Claudin5*, *Tjp1*and *F11r* (**b**), *Ccl2*, *Ccl5*, *Cxcl9* and *Cxcl10* (**c**) and *Itgb5*, *Itgb7*, *Vla-4*, *Icam-1* and *Vcam* (**d**). Data are representative of 3 independent experiments, 5 mice per group, and shown as mean +/- SEM.

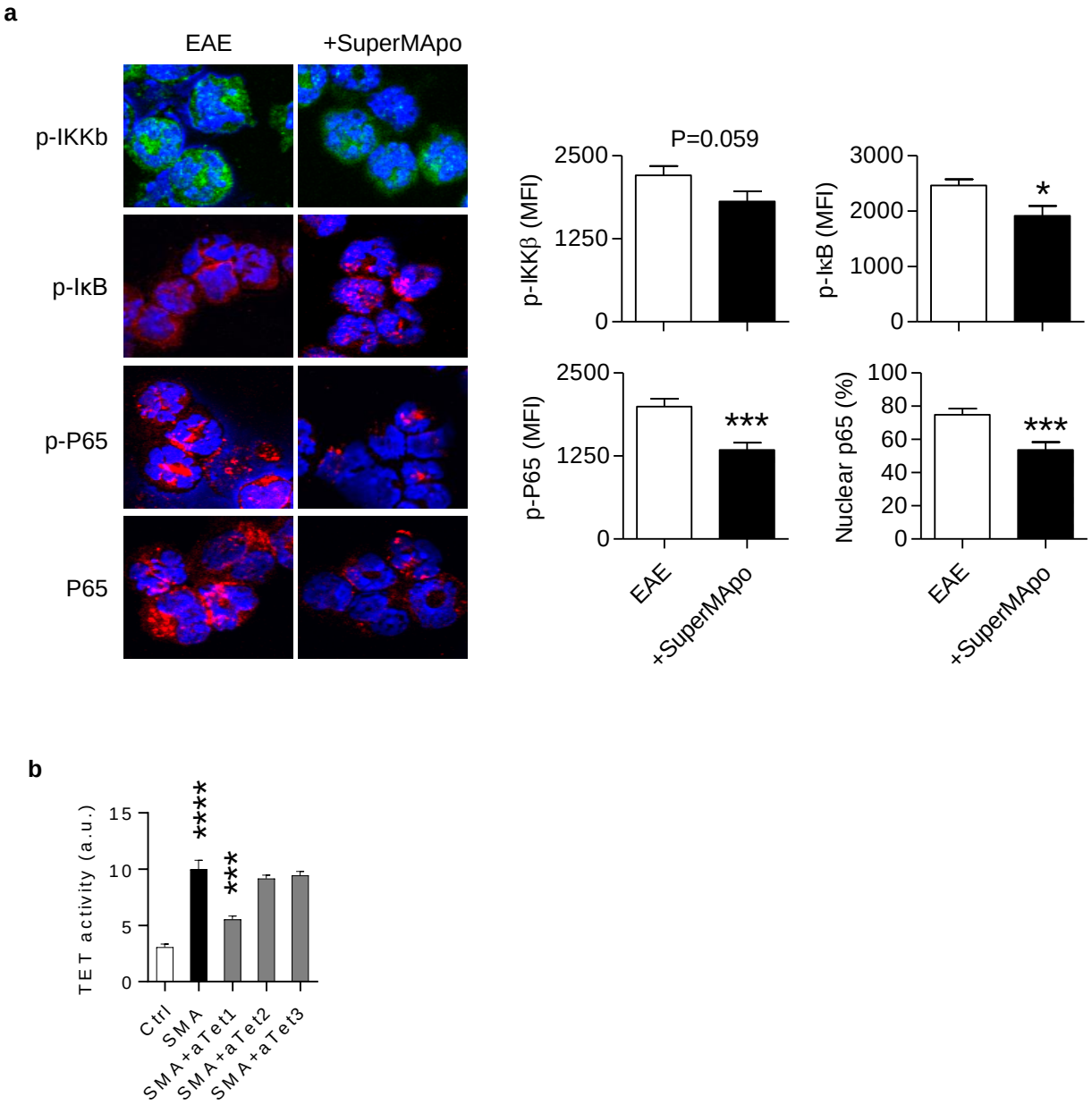

FIGURE S4 SuperMApo treatment decreases macrophage inflammatory state by blocking NF-κB activation. (**a**) Immunofluorescence staining of p-IKKb, p-IκB, p-P65 and P65 subunits on CD11b^{+} macrophages sorted from the spleen of SuperMApo- or vehicle-treated EAE mice. Data, representative of three independent experiments, are shown as representative pictures (left panels) or cumulative bar graphs (right panels) as mean +/- SEM. *=p<0.05, ***=p<0.001, unpaired two-tailed Student's t test. (**b**) TET activity in macrophages from EAE mice receiving vehicle (Ctrl) or SuperMApo (SMA) in the presence or not of blocking antibodies against TET1 (aTet1), TET2 (aTet2) or TET3 (aTet3). Data are shown as mean +/- SEM. ***=p<0.001, ****=p<0.0001, one-way ANOVA plus Tukey's multiple comparisons test.