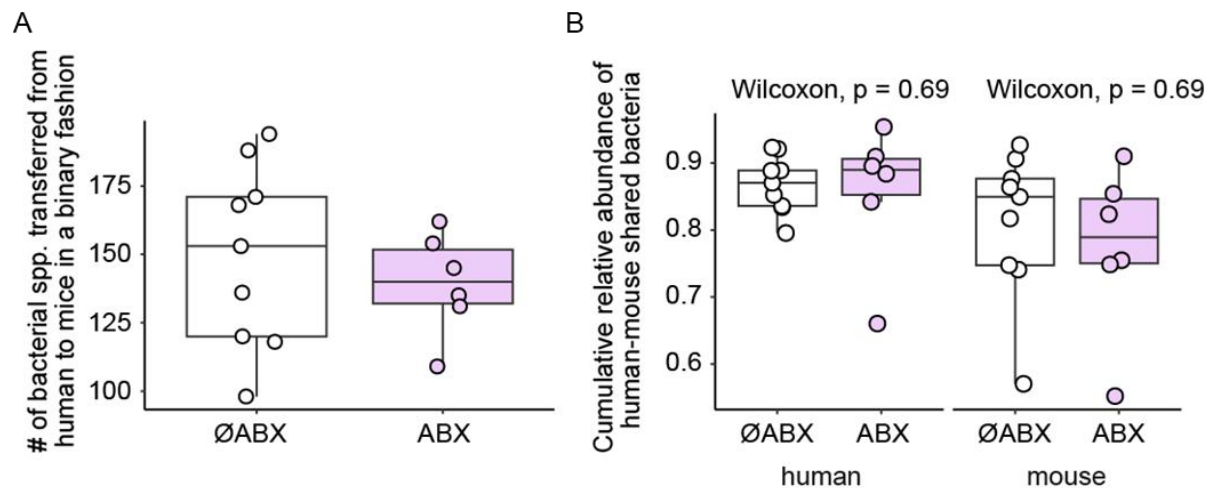
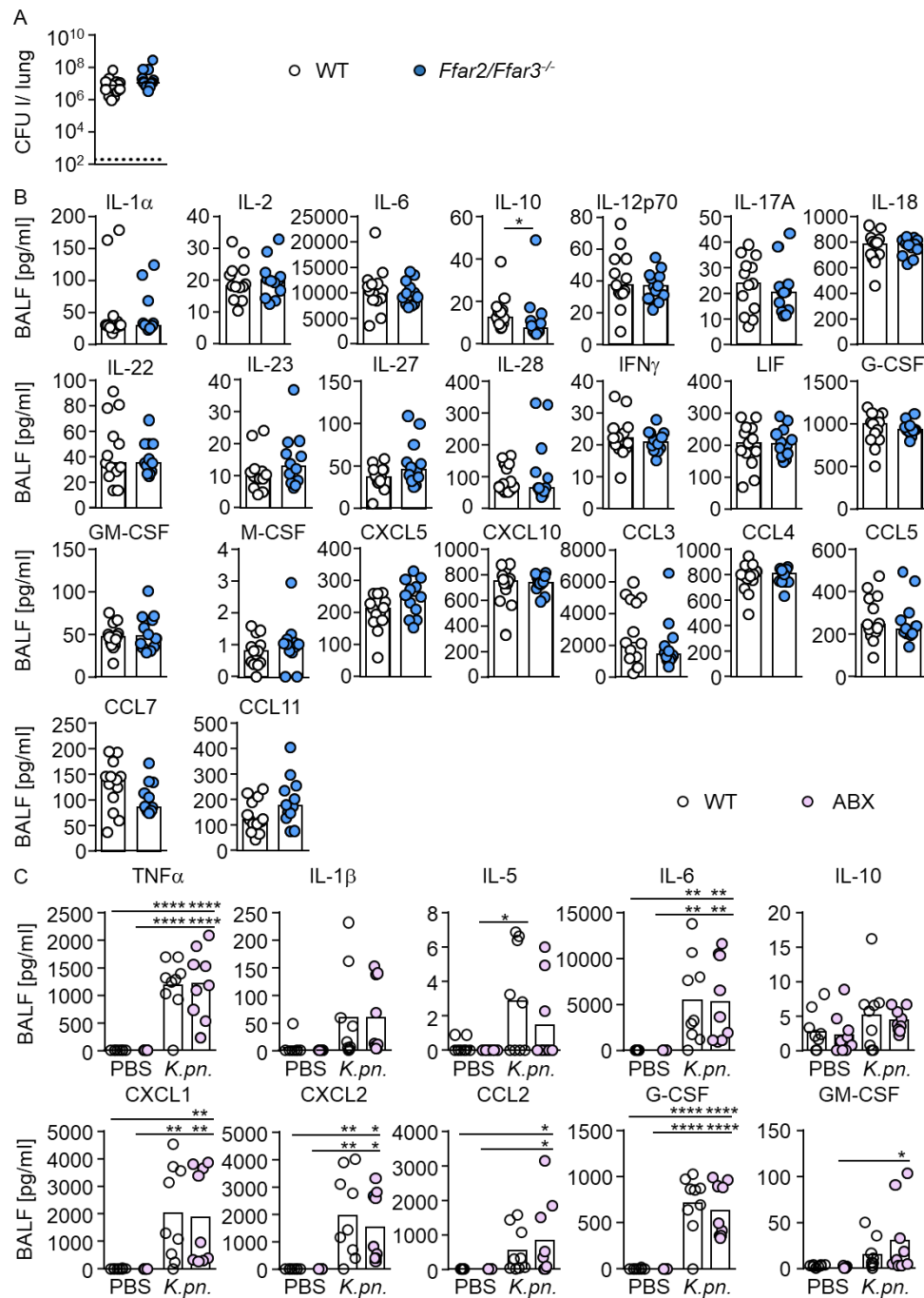


Suppl. Figure 1. Relative abundances of bacterial phyla in fecal samples of antibiotic-treated (ABX, n = 26) and untreated patients (ØABX, n=29).



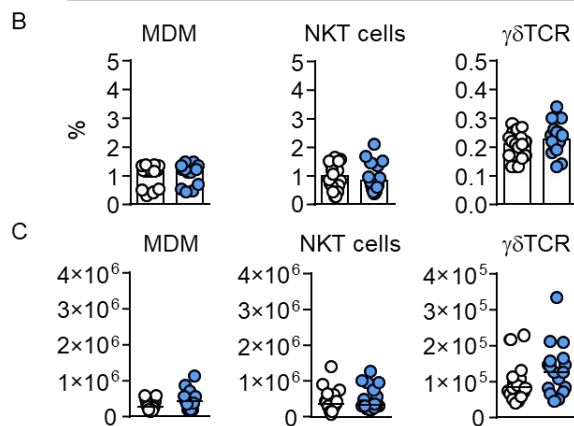
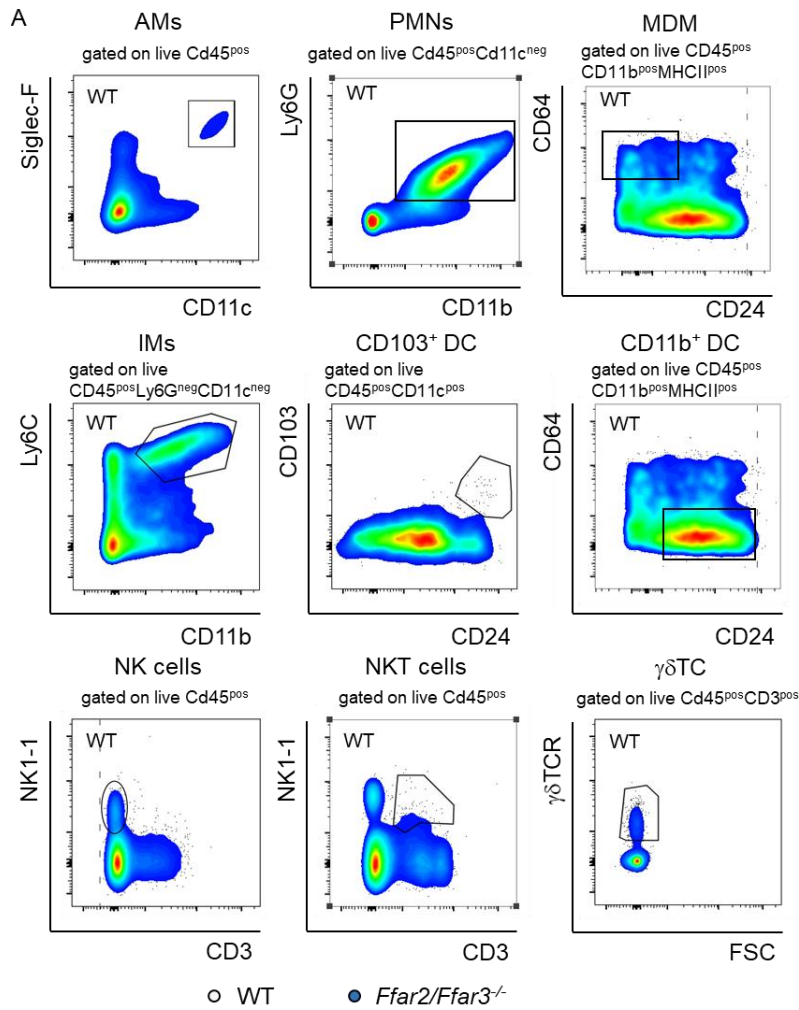
Suppl. Figure 2. Successful transfer of bacterial species from fecal samples of antibiotic-naïve patients (ØABX) or antibiotic-treated patients (ABX) into mice. **(A)** Number of bacterial species that were successfully transferred from patient fecal samples to mice. **(B)** Relative abundance of transferred bacterial species in the patient donor microbiota and in the mouse microbiota after transfer. Wilcoxon-Mann-Whitney U test (two-tailed) was used for relative abundance of transferred bacterial species. Each dot representing the data from one patient or mouse.



Suppl. Figure 3. The *K. pneumoniae*-induced inflammatory response seems not to be impaired in *Ffar2/Ffar3*^{-/-} or microbiota-depleted mice. **(A-B)** Conventionally housed WT and *Ffar2/Ffar3*^{-/-} mice were infected with *K. pneumoniae* for 12 h, bacterial loads (CFU) in lung tissues were counted ($n = 18$ for WT; $n = 19$ for *Ffar2/Ffar3*^{-/-}) **(A)** and cytokine levels in bronchoalveolar lavage fluid (BALF) were measured ($n = 12$ for WT; $n = 13$ for *Ffar2/Ffar3*^{-/-}) **(B)**. **(C)** Conventionally colonized (CONV) and antibiotic-treated WT animals were infected

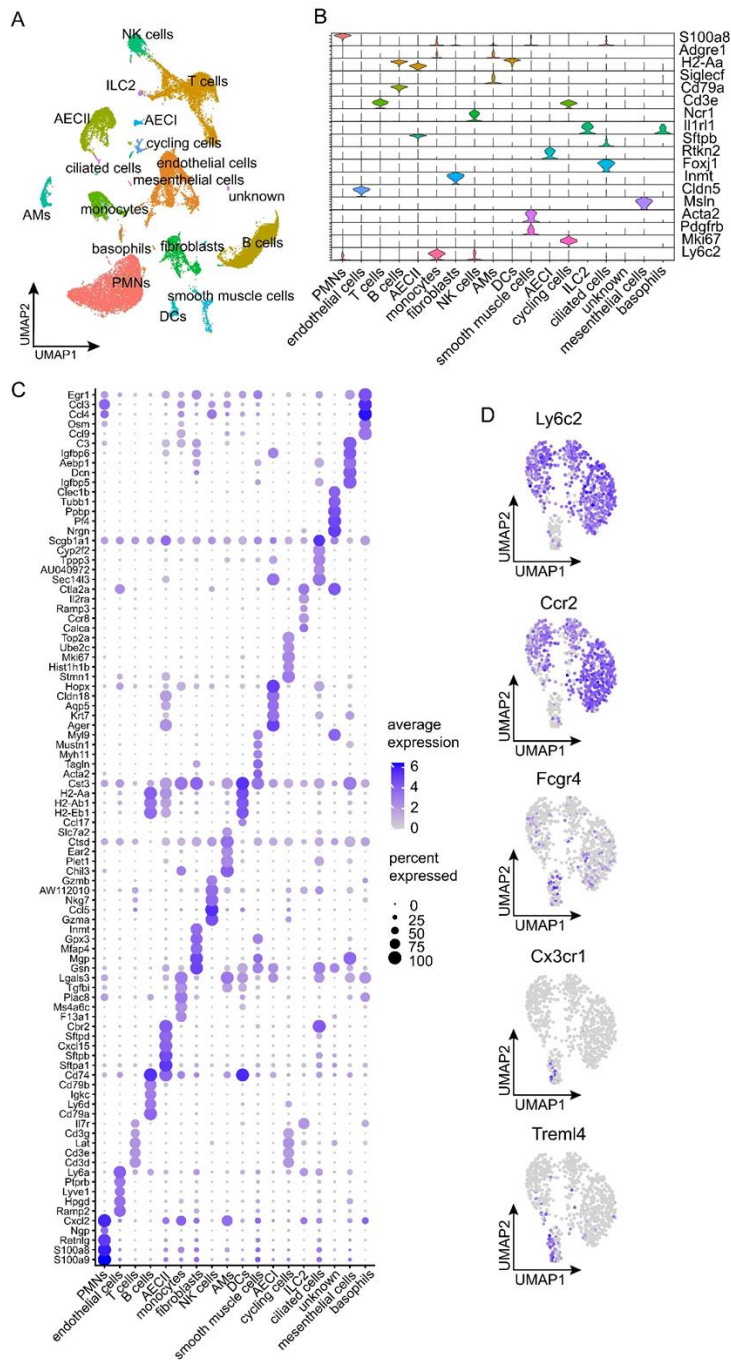
with *K. pneumoniae* or treated with PBS for 12 hours, and cytokine levels in BALF were assessed ($n = 9$ for CONV PBS, $n = 9$ for ABX PBS, $n = 9$ for CONV *K. pneumoniae* $n = 9$ for ABX *K. pneumoniae*). Mann-Whitney U test (two-tailed) was determined for bacterial loads (**A**) and inflammatory mediators' (**B**) from WT and *Ffar2/Ffar3*^{-/-} mice. Kruskal-Wallis test followed by Dunn's multiple comparison was applied to cytokine dataset from CONV and ABX mice (**C**). Values are shown as median (**A-C**), each dot represents the data from a single mouse.

* $P < 0.05$, ** $P < 0.01$, *** $P < 0.005$, ***** $P < 0.001$.

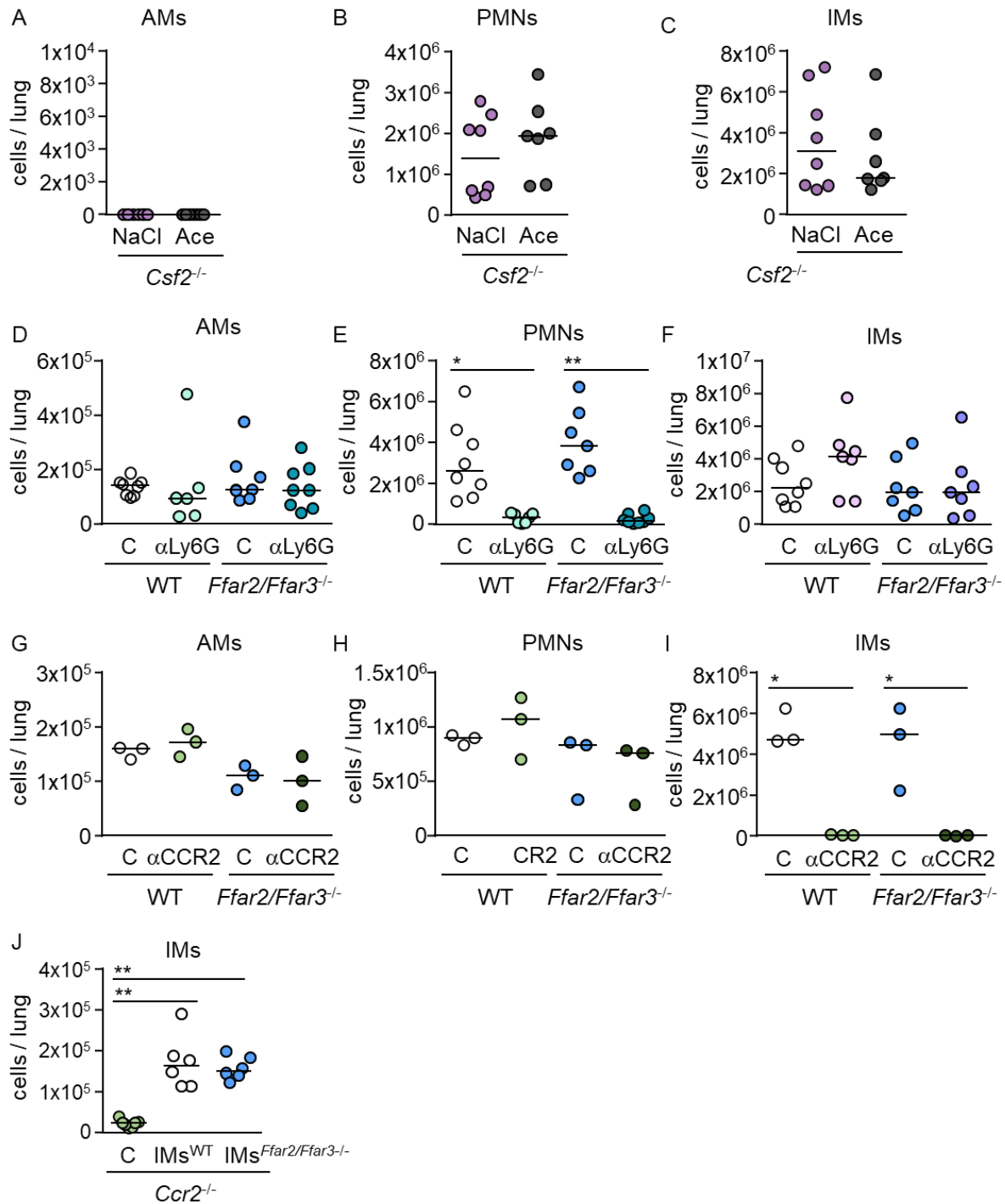


Suppl. Figure 4. Composition of lung immune cell populations in WT- and *Ffar2/Ffar3*^{-/-} animals do not differ upon *K. pneumoniae* infection. (A-C) WT (*n* = 19) and *Ffar2/Ffar3*^{-/-} (*n* = 18) mice were infected with *K. pneumoniae* for 12 hours. (A) FACS gating strategy of lung cells after exclusion of cell doublets and gating on live cells (CD45⁺) for AMs, PMNs, eosinophils, IMs, CD11b⁺ DCs, CD103b⁺ DCs, monocyte derived macrophages (MDMs), NK

cells, natural killer T (NKT) cells and $\gamma\delta$ T cells ($\gamma\delta$ TC). **(B-C)** Percentage and numbers of MDMs, NKT and $\gamma\delta$ T cells were counted by FACS after staining of lung tissues. Mann-Whitney *U* test (two-tailed) was applied for lung cell populations **(B-C)**. Values are shown as median, each dot represents the data from one mouse. **P* < 0.05, ***P* < 0.01, ****P* < 0.005.



Suppl. Figure 5. scRNAseq of lung cells of WT and *Ffar2/Ffar3*^{-/-} mice infected with *K. pneumoniae* for 12 h ($n = 4$ for WT PBS, $n = 4$ for WT infected, $n = 4$ for *Ffar2/Ffar3*^{-/-} PBS, $n = 4$ for *Ffar2/Ffar3*^{-/-} infected). **(A)** Two-dimensional embedding computed by UMAP on 30,215 computationally identified cells, **(B)** violin plots depicting representative marker genes for each cell type, and **(C)** dot plots of most differentially expressed genes for each cell population. **(D)** Normalized expression of marker genes of IMs and patrolling monocytes projected onto the UMAP embedding in Fig. 4G.



Suppl. Figure 6. Leukocyte numbers in lung tissues of *K. pneumoniae*-infected *Csf2*^{-/-}, WT and *Ffar2/Ffar3*^{-/-} mice. (**A-C**) Numbers of AMs, PMNs, and IMs in *Csf2*^{-/-} mice (NaCl; $n = 8$; $n = 7$ for acetate). (**D-F**) Numbers of AMs, PMNs, and IMs in WT and *Ffar2/Ffar3*^{-/-} mice treated with control antibodies (C) or α Ly6G ($n = 8$ for WT C, $n = 7$ for *Ffar2/Ffar3*^{-/-} C, $n = 7$ for WT α Ly6G; $n = 8$ for *Ffar2/Ffar3*^{-/-} α Ly6G). (**G-I**) Numbers of AMs, PMNs, and IMs in WT and *Ffar2/Ffar3*^{-/-} mice treated with control antibodies (C) or α CCR2 ($n = 3$ for WT C, $n = 3$ for

Ffar2/Ffar3^{-/-} C, n = 3 for WT α CCR2 ; n = 3 for *Ffar2/Ffar3^{-/-}* α CCR2). **(J)** Number of IMs in *K. pneumoniae*-infected *Ccr2^{-/-}* mice treated intravenously with PBS (C) or transplanted with IMs of WT or *Ffar2/Ffar3^{-/-}* animals (n = 7 for PBS , n = 6 for WT α CCR2; n = 6 for *Ffar2/Ffar3^{-/-}*). Values are shown as median, each dot represents the data from a single mouse. **P* < 0.05, ***P* < 0.01, ****P* < 0.005.