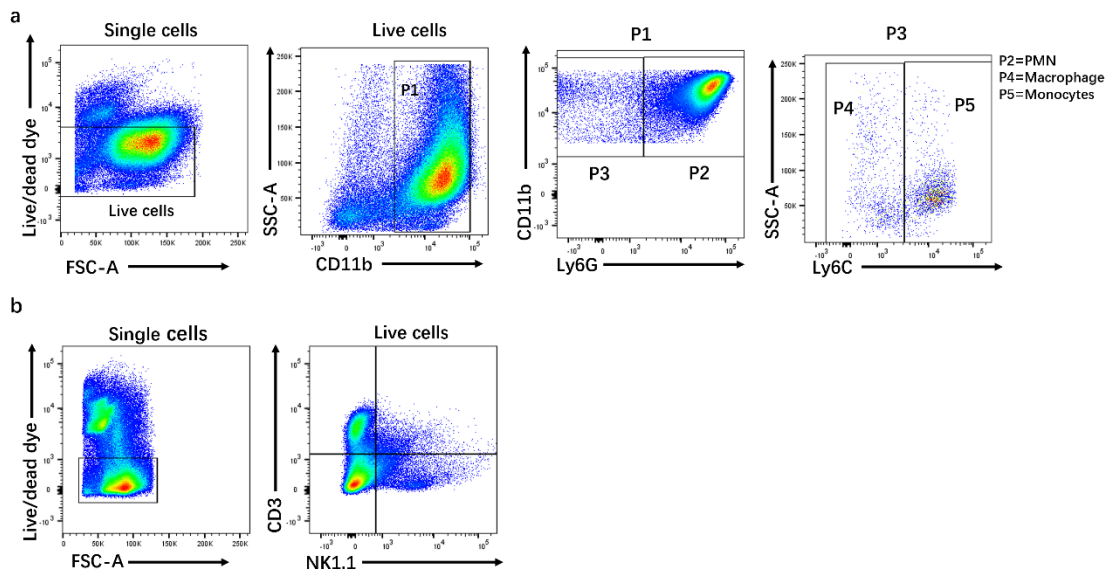
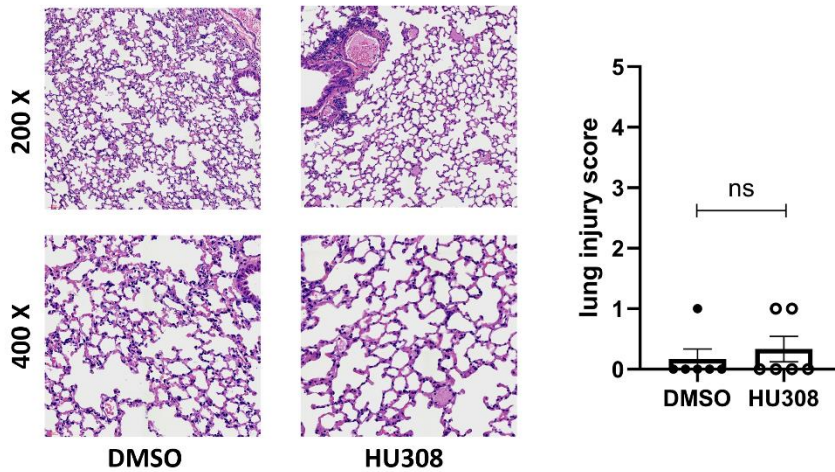


Supplementary figure 1. Quantitative real-time PCR and Western-blot were used to confirmed the CNR2 conditional knockout mice in specific cells. (a) CNR2 mRNA expression were compared between CNR2^{fl/fl} and Lyz2 Cre CNR2^{fl/fl} mice in BMDM cells and CD4 Cre CNR2^{fl/fl} mice in CD4⁺ cells. (b) MAPK signaling pathway activated by CB2 receptor were determined in CNR2^{fl/fl} and Lyz2 Cre CNR2^{fl/fl} mice in BMDM cells. BMDM, bone marrow derived macrophage. MAPK, mitogen activated protein kinase.



Supplementary figure 2. Gating strategy for analysis of immune cells in lung, spleen and peritoneal lavage fluids from septic mice. (a) PMN, macrophage, monocytes were gated based on CD11b, Ly6G, Ly6C. (b) T cell and NK cell were gated according to CD3, NK1.1.



Supplementary figure 3. HU308 did not result in significant harm to mice. CNR2^{fl/fl} mice were treated with 4.8 mg/kg HU308 or DMSO, intraperitoneally administered. 24 h after treatment, lung histopathology comparison was performed by H&E staining. Data were representative of at least two independent experiments with n = 6 per group. P valued by Mann-Whitney *U* test. ** p < 0.01.