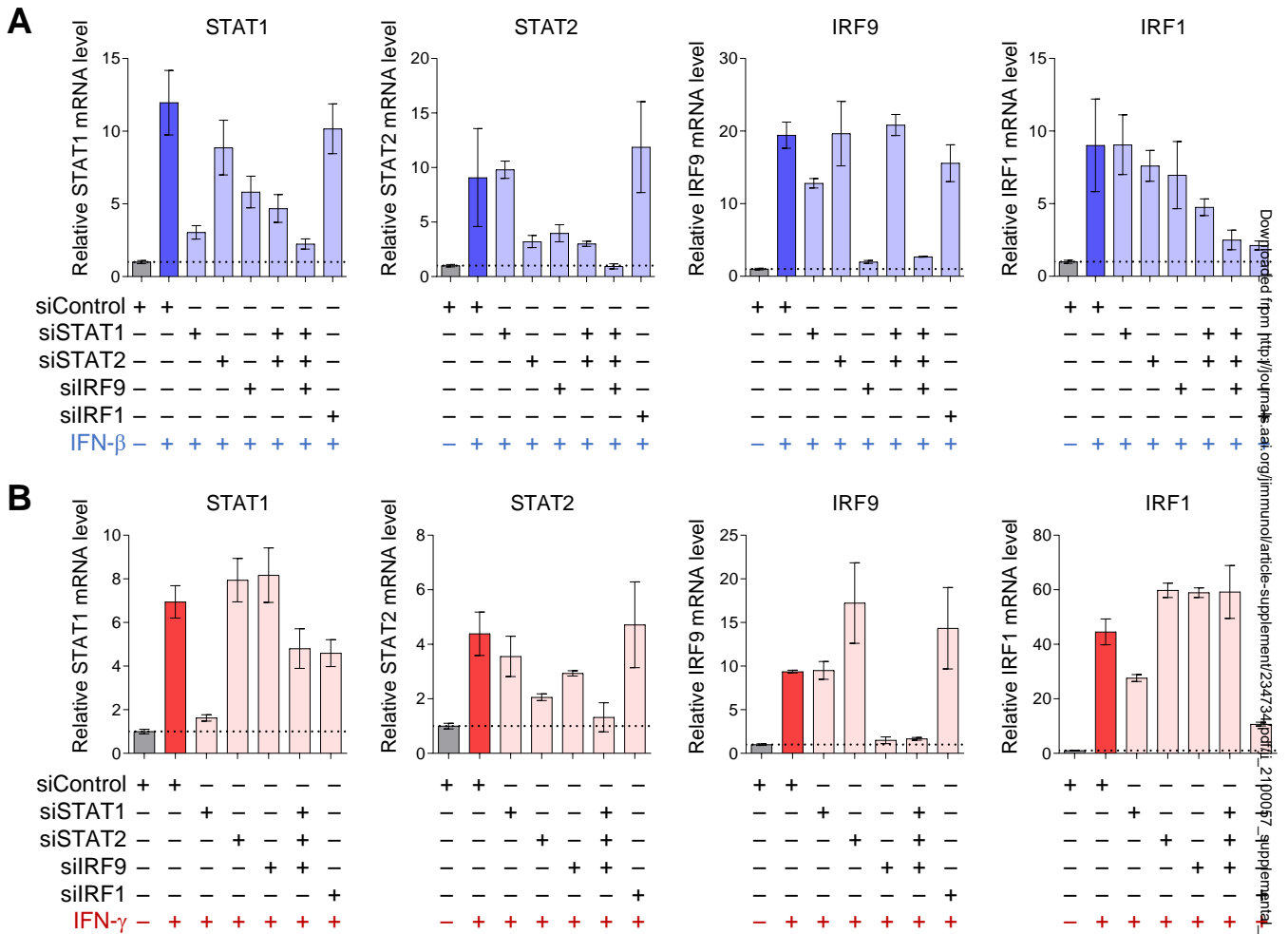
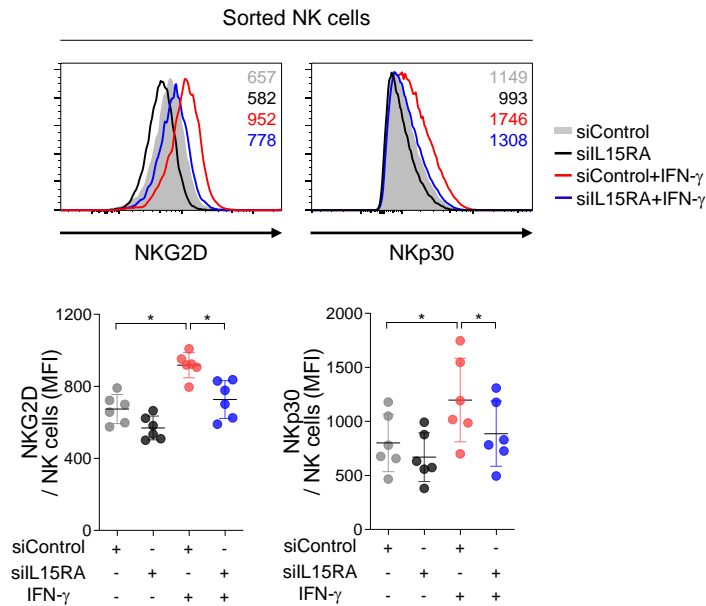


**Supplemental Figure 1 (related to Figure 1).** (A) Huh-7 cells were treated with various IFNs at a concentration of 20 ng/mL, except for IFN- $\lambda$ 3, which was given at 10 ng/mL to avoid cell death. After 24 hours, surface IL-15 levels were analyzed by flow cytometry. Shaded histograms represent isotype controls; red lines represent anti-IL-15 antibody. (B) Knockdown efficiency after siRNA transfection was examined by RT-qPCR (related to Figure 1G). IL15RA mRNA levels were normalized to an endogenous reference (GAPDH) and expressed as the relative level compared to negative controls, which were set to 1. Data are presented as the mean  $\pm$  SD. Statistical analysis was performed using one-way ANOVA followed by the Holm-Sidak multiple comparisons test. \*\*\*\*,  $p < 0.0001$ .

# Supplemental Figure 2

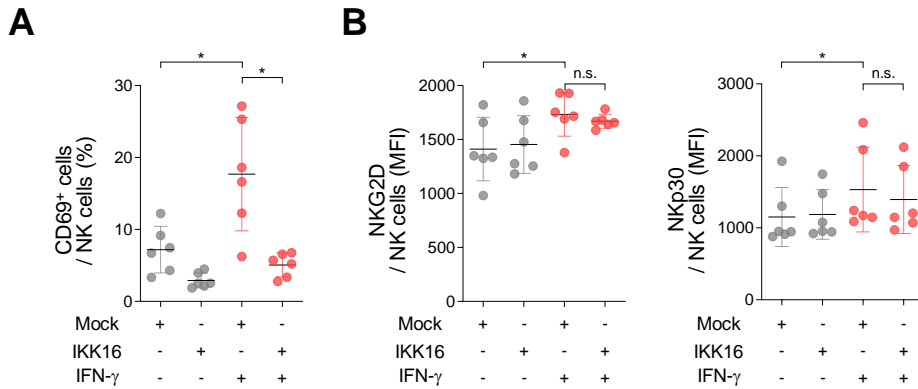


**Supplemental Figure 2. Knockdown efficiency after siRNA transfection (related to Figure 2C-D and 3C-D).** Huh-7 cells were transfected with the indicated siRNAs or combinations thereof. After 48 hours, Huh-7 cells were stimulated with IFN- $\beta$  (A) or IFN- $\gamma$  (B) at the final concentration of 20 ng/mL for 6 hours. Knockdown efficiencies for STAT1, STAT2, IRF9, and IRF1 were examined by RT-qPCR. mRNA levels were normalized to an endogenous reference (HPRT) and expressed relative to negative controls. Gray bars and horizontal dotted lines represent the reference. Data are presented as the mean  $\pm$  SD.



**Supplemental Figure 3. IFN- $\gamma$ -induced IL-15 on Huh-7 cells directly activates sorted NK cells (related to Fig. 5).** Huh-7 cells were transfected with control or IL-15R $\alpha$ -specific siRNA for 24 hours, stimulated with IFN- $\gamma$  (20 ng/mL) for an additional 24 hours, and co-cultured with magnetically sorted NK cells from healthy donors (n=6) for another 24 hours. The expression levels of NKG2D and NKp30 were analyzed in NK cells. Representative histograms (left) and summary data (right) are presented. Data are presented as the mean  $\pm$  SD. Statistical analysis was performed using the Wilcoxon signed-rank test. \*  $p < 0.05$ .

# Supplemental Figure 4



**Supplemental Figure 4 (related to Fig. 5).** (A and B) Huh-7 cells were pre-treated with IKK16 (1  $\mu$ M), stimulated with IFN- $\gamma$  (20 ng/mL) for an additional 24 hours, and co-cultured with PBMCs from healthy donors (n=6) for another 24 hours. The frequency of CD69<sup>+</sup> cells (A) and the expression level of NKG2D and NKp30 (B) were analyzed in NK cells. Data are presented as the mean  $\pm$  SD. Statistical analysis was performed using the Wilcoxon signed-rank test. \*,  $p < 0.05$ .