## Supplementary Data



SUPPLEMENTARY FIG. S1. IFN- $\lambda$ 4 signals through JAK tyrosine kinases and the shared receptors of type-III interferons (IFNLR1 and IL10R2) in stable HepG2-ISRE-Luc cells. (A) The effects of JAK inhibitor on signaling of IFN-λ4 (transient transfection) or treatment with recombinant IFN- $\lambda$ 3 (10 ng/mL) or IFN- $\alpha$  (2 ng/mL), all in 8 biological replicates. Results are presented as fold change to corresponding controls—cells transfected with control-Halo or treated with 0.1% DMSO (mock). (B) Effects of transient siRNA knockdown of IFNLR1 transcript on IFN-λ4 signaling, in siRNA untreated samples (cells) and cells treated with negative control si-Scr (scrambled siRNA) or si-IFNLR1, all in 6 biological replicates. After 24 h, cells were treated with IFN- $\lambda$ 3 (10 ng/mL), IFN- $\alpha$  (2 ng/mL) or mock treated. Results are presented as fold change to cells treated with scrambled siRNA (si-Scr). (C) Effects of treatment with a blocking  $\alpha$ -IL10R2 antibody on signaling of IFN- $\lambda$ 4 (transient transfection) or treatment by IFN- $\lambda$ 3 (10 ng/mL) or IFN- $\alpha$  (2 ng/mL). For all experiments, the cells were assayed 48 h post-transfection for expression of the ISRE-Luc reporter. \*\*\*\*P < 0.0001 based on *t*-tests. (D) Efficiency of siRNA knockdown of IFNLR1 in samples presented on (B). siRNA untreated cells and cells treated with negative control (si-Scr) and si-IFNLR1, all in 6 biological replicates. Analysis was performed with qRT-PCR for IFNLR1 mRNA and normalized by ACTB expression in the same samples. Expression is presented as IFNLR1 expression, in % to the level of untreated samples, taken as 100%. There was an average of 76% in the group of si-Scr samples and 42% in the group of si-IFNLR1 samples; group means are marked by red dashed lines. Error bars represent SEM. P values are based on *t*-tests. DMSO, dimethyl sulfoxide: IFN- $\lambda$ 4, interferon lambda 4; ISRE, interferon-stimulated response element; qRT-PCR, quantitative reverse-transcriptase-polymerase chain reaction; SEM, standard error of the mean.