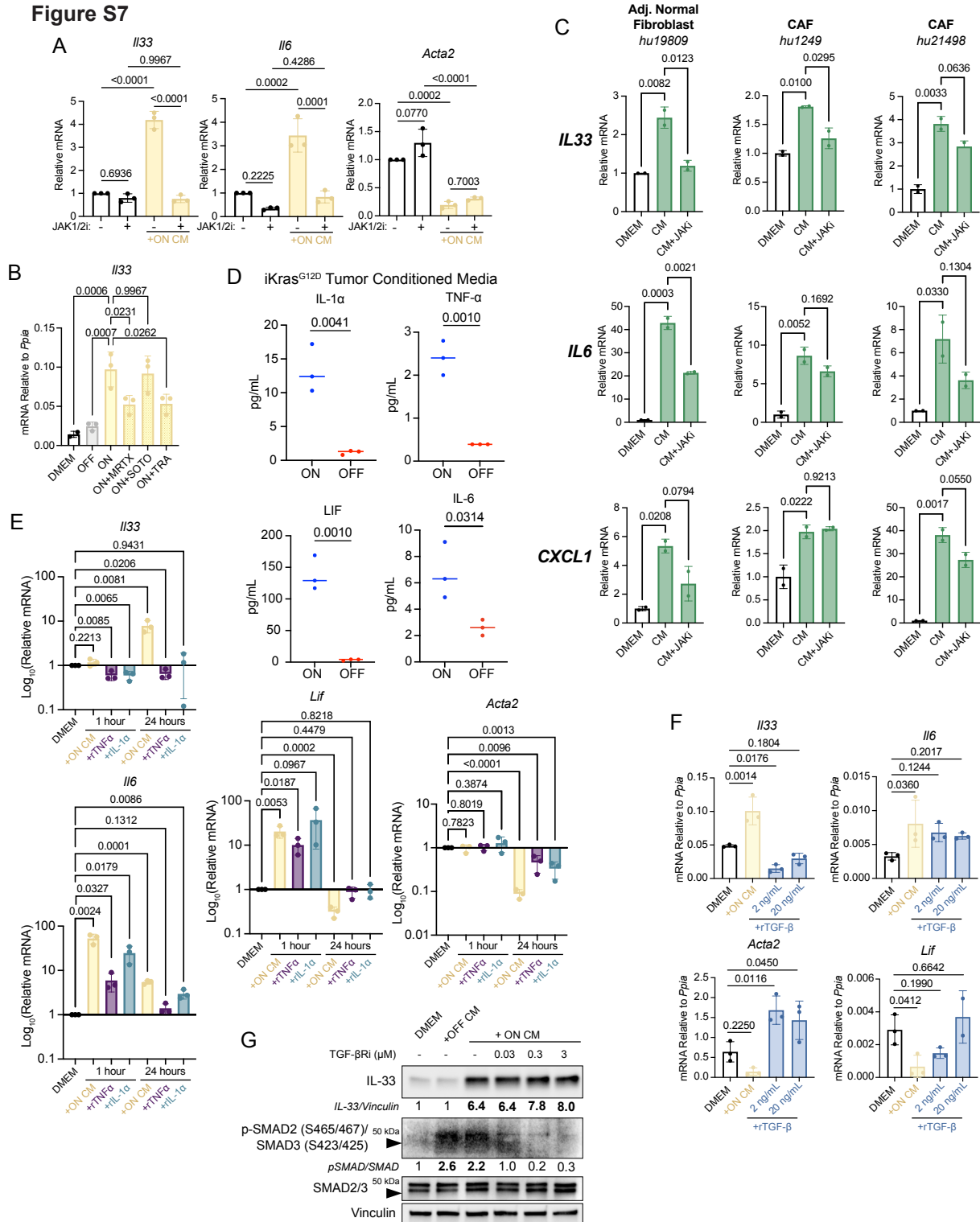**Figure S7**

Figure S7: Tumor cell-initiated autocrine signaling drives IL-33 upregulation in pancreatic fibroblasts. (A) qRT-PCR of CD1WT after 24h of treatment with DMEM, JAK1/2i, iKras^{G12D} "ON" CM, or concurrent iKras^{G12D} "ON" CM and JAK1/2i (0.3 µM/mL). Values are relative to DMEM group. Groups were compared with ordinary one-way ANOVA. (B) qRT-PCR of CD1WT after 24h of treatment with DMEM, iKras^{G12D} "OFF" CM, iKras^{G12D} "ON" CM, or concurrent iKras^{G12D} "ON" CM and the KRAS^{G12D}i MRTX1133 (0.5 µM/mL), the KRAS^{G12C}i Sotorasib (0.5 µM/mL), or the MEK1/2i Trametinib (50 nM/mL). Values are relative to DMEM group. Ordinary one-way ANOVA was performed to compare each condition to iKras^{G12D} "ON" CM group. (C) qRT-PCR of primary human pancreatic fibroblasts after 24h of treatment with DMEM, PDA organoid CM, or concurrent PDA organoid CM and JAK1/2i (0.5 µM/mL). Values are relative to DMEM group. Groups were compared with ordinary one-way ANOVA. (D) Concentration of cytokines in iKras^{G12D} "ON" CM and iKras^{G12D} "OFF" CM as measured by ELISA. Two-tailed Student's *t* test was performed to compare groups. IL-1β was probed for but not detected in our samples. (E) qRT-PCR of CD1WT after treatment with DMEM, iKras^{G12D} "ON" CM, rTNF-α (20 ng/mL), or rIL-1α (20 ng/mL) for 1h or 24h. Values are relative to DMEM group and Log$_{10}$ transformed to better visualize the large changes in gene expression level. Ordinary one-way ANOVA was performed to compare each condition to DMEM group. (F) qRT-PCR of CD1WT after treatment with DMEM, iKras^{G12D} "ON" CM, rTGF-β (2 ng/mL or 20 ng/mL) for 24h. Values are relative to DMEM group. Ordinary one-way ANOVA was performed to compare each condition to DMEM group. (G) Western blot of CD1WT whole cell lysates treated with DMEM, iKras^{G12D} "OFF" CM, iKras^{G12D} "ON" CM, or concurrent iKras^{G12D} "ON" CM and TGF-βRi. In all experiments with iKras^{G12D} CM, doxycycline is used as a vehicle control. In all experiments with JAK1/2i (Ruxolitinib) or TGF-βRi (Galunisertib), DMSO was used as a vehicle control. qRT-PCR values are normalized to *Ppia* (Cyclophilin A) in murine experiments and *GAPDH* in human experiments. All replicates represent complete, independent experiments.