### **SUPPLEMENTARY FIGURES**

# Senescence promotes in vivo reprogramming through $p16^{INK4a}$ and IL-6

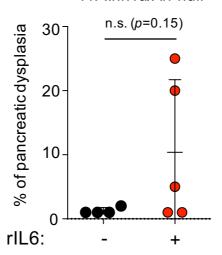
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Supplementary Figure S1

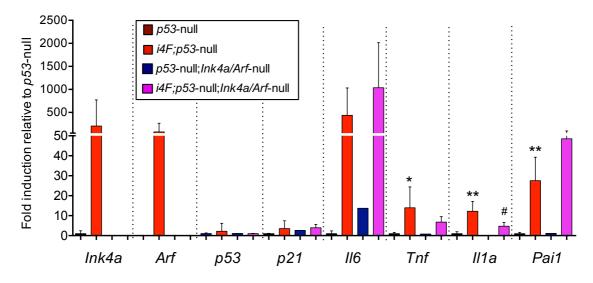
Supplementary Figure S2

#### i4F:Ink4a/Arf-null



## Supplementary Figure S1. Treatment with rIL6 increases in vivo reprogramming in i4F;Ink4a/Arf-null pancreas

Percentage of dysplasia in the pancreas of i4F;Ink4a/Arf-null male mice of 8-10 weeks of age, non-treated (-) and treated (+) with rIL6. Mice were treated with 0.2 mg/ml doxycycline for 7 days and analyzed at the end of the treatment. Treatment with recombinant IL6 (Abyntek Biopharma, S.L, #AI081) was concomitant to doxycycline treatment and was performed by intraperitoneal injection with 5  $\mu$ g of rIL6 three times during a period of 7 days. Values correspond to % of pancreatic area affected (evaluated blindly), the average  $\pm$  s.d. are also indicated. Statistical significance was assessed using the unpaired two-tailed Student's t-test with Welch's correction (n.s.: non-significant).



Supplementary Figure S2. Expression levels of senescence genes and SASP cytokines in *i4F;p53*-null;*Ink4a/Arf*-null pancreas

Levels of mRNA of the indicated genes in the pancreas of p53-null mice (n=5); p53-null; Ink4a/Arf-null mice (n=1); i4F; p53-null mice (n=6) and i4F; p53-null; Ink4a/Arf-null mice (n=2). All the mice tested are males of 10-14 weeks of age. Statistical significance compared to p53-null controls was assessed using the unpaired two-tailed Student's t-test with Welch's correction: p<0.05, \*; p<0.01, \*\*. Comparisons between i4F; p53-null and i4F; p53-null; Ink4a/Arf-null are indicated in the same manner but using the symbol "#".