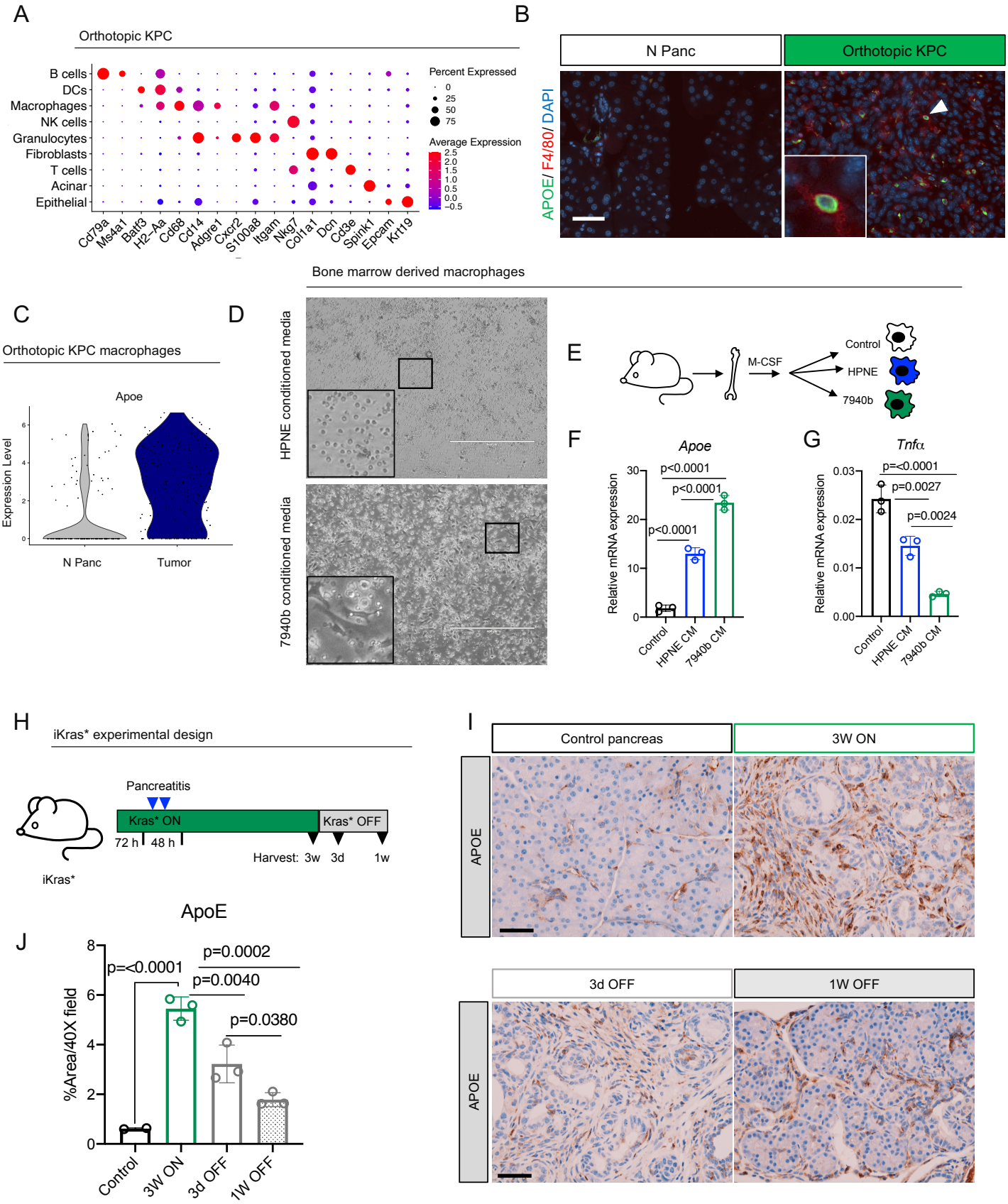
A
Orthotopic KPC
B cells
DCs
Macrophages
NK cells
Granulocytes
Fibroblasts
T cells
Acinar
Epithelial
Cd79a
Ms4a1
Batf3
H2-Aa
Cd68
Cd14
Adgre1
Cxcr2
S100a8
Itgam
Nkg7
Col1a1
Dcn
Cd3e
Spink1
Epcam
Krt19
Percent Expressed
0
25
50
75
Average Expression
2.5
2.0
1.5
1.0
0.5
0.0
−0.5
B
N Panc
Orthotopic KPC
APOE/ F4/80/ DAPI
C
Orthotopic KPC macrophages
Apoe
Expression Level
N Panc
Tumor
D
Bone marrow derived macrophages
HPNE conditioned media
7940b conditioned media
E
M-CSF
Control
HPNE
7940b
F
Apoe
Relative mRNA expression
p<0.0001
p<0.0001
p<0.0001
Control
HPNE CM
7940b CM
G
Tnfα
Relative mRNA expression
p=<0.0001
p=0.0027
p=0.0024
Control
HPNE CM
7940b CM
H
iKras* experimental design
Pancreatitis
Kras* ON
Kras* OFF
72 h
48 h
Harvest: 3w 3d 1w
iKras*
I
Control pancreas
3W ON
APOE
3d OFF
1W OFF
APOE
J
ApoE
%Area/40X field
p=<0.0001
p=0.0002
p=0.0040
p=0.0380
Control
3W ON
3d OFF
1W OFF

Supplementary Figure 2. APOE is highly expressed by tumor-associated macrophages. **(A)** Dot plot of lineage markers used to define identified cell populations in orthotopic KPC tumors (n=2). Color of the dot represents average expression. Size of the dot represents expression frequency. **(B)** Co-immunofluorescence staining on normal mouse pancreas (N Panc) and orthotopic KPC tumor for APOE (green), F4/80 (red) and DAPI (blue). White arrowhead denotes APOE and F4/80 colocalization. White box represents higher magnification. Scale bars, 100 μm. **(C)** Violin plot of normalized ApoE expression in mouse normal pancreas (grey) and tumor (navy) samples. **(D)** Bright field images of bone marrow cells cultured with either HPNE (top) or 7940b (bottom) conditioned media for 5 days. Scale bar, 400 μm. Inset represents higher magnification of boxed area. **(E)** Experimental design schematic. **(F)** qRT-PCR analysis of *Apoe* and **(G)** *Tnfa* mRNA levels relative to *Cyclophilin A* housekeeping in untreated macrophages (control), and macrophages treated with either HPNE or 7940b conditioned media for 24 hours. Statistical significance was determined using one-way ANOVA with Tukey's test for multiple correction. **(H)** iKras* experimental design. **(I)** Immunohistochemical staining for APOE in mouse normal pancreas, pancreatic tissue after 3 weeks of oncogenic *Kras* expression (3W ON), and extinguishment of oncogenic *Kras* expression for 3 days (3d OFF) and 1 week (1W OFF). Scale bars, 50 μm. **(J)** Quantitation of positive APOE staining as percent area in a 40x field. 5 images per mouse were averaged. Control (n=2), 3W ON (n=3), 3d OFF (n=3), and 1W OFF (n=3). W=week. d=day. Statistical significance was determined using one-way ANOVA with Tukey's test for multiple correction.