SUPPLEMENTARY FIGURE LEGENDS

Figure S1: **I-BET151 inhibits growth of PDAC cells in three-dimensional collagen.** A, B. AsPC1, CD18 and Panc1 cells were grown in three-dimensional collagen gels and fresh serum-containing medium supplemented with DMSO or I-BET151 was added every other day for 7 days. The effect on colony size was examined by phase contrast microscopy (A), and size of the individual colonies measured (B). The results are representative of three independent experiments. *, p < 0.05.

Figure S2: **I-BET151 decreases growth of CD18-Snail cells in three-dimensional collagen.** CD18-Vector and CD18-Snail cells were grown in three-dimensional collagen gels and fresh serum-containing medium supplemented with DMSO or I-BET151 was added every other day for 7 days. The effect of I-BET151 on colony size in three-dimensional collagen was examined by phase contrast microscopy and size of the individual colonies measured. The results are representative of at least three independent experiments. *, p < 0.05.

Figure S3: **JQ1 and I-BET151 decrease growth of AsPC1-Snail cells.** A. Lysates from AsPC1-Vector and AsPC1-Snail cells were analyzed for Snail by Western blotting, and the effect on E-cadherin and vimentin were analyzed by Western blotting. B,C. AsPC1-Vector and AsPC1-Snail cells were grown in three-dimensional collagen gels and fresh serum-containing medium supplemented with DMSO, JQ1 or I-BET151 was added every other day for 7 days. The effect of JQ1 (B) and I-BET151 (C) on colony size in three-dimensional collagen was examined by phase contrast microscopy and size of the individual colonies measured. The results are representative of three independent experiments. *, p < 0.05.

Figure S4: **I-BET151 decreases growth of chemoresistant CD18 cells in three-dimensional collagen.** CD18-CR cells were grown in three-dimensional collagen gels and fresh serum-containing medium supplemented with DMSO or I-BET151 was added every other day for 7 days. The effect of JQ1 on colony size in three-dimensional collagen was examined by phase contrast microscopy, and size of the
individual colonies measured. The results are representative of at least three independent experiments. *, p < 0.05.

Figure S5: A. I-BET151 decreases expression of c-MYC in AsPC1 and CD18 cells, but not in Panc1 cells. PDAC cells were grown in three-dimensional type I collagen gels in the presence of DMSO (vehicle control), or I-BET151 (1 μM) for 48 hours and c-MYC mRNA expression analyzed by qRT-PCR using GAPDH as normalization control. **B. Effect of JQ1 treatment for 8 hours and 24 hours on c-MYC mRNA expression in PDAC cells.** PDAC cells were grown in three-dimensional type I collagen gels in the presence of DMSO (vehicle control), or JQ1 (0.5 μM) for 8 hours or 24 hours and c-MYC mRNA expression analyzed by qRT-PCR using GAPDH as normalization control. *, p < 0.05. **C, D. JQ1 represses c-MYC in Snail-expressing PDAC cells and chemo-resistant PDAC cells.** PDAC-vector, PDAC-Snail and CD18-CR cells were grown in three-dimensional type I collagen gels in the presence of DMSO (vehicle control), or JQ1 (0.5 μM) for 48 hours and c-MYC mRNA expression analyzed by qRT-PCR using GAPDH as normalization control. The results are representative of at least two independent experiments. *, p < 0.05.

Figure S6: A. I-BET151 decreases expression of FOSL1 in PDAC cells. PDAC cells were grown in three-dimensional type I collagen gels in the presence of DMSO (vehicle control), or I-BET151 (1 μM) for 48 hours and FOSL1 mRNA expression analyzed by qRT-PCR using GAPDH as normalization control. **B. siRNA BRD4 decreases FOSL1 mRNA in PDAC cells.** Effect of siBRD4 on FOSL1 mRNA expression analyzed by qRT-PCR using GAPDH as normalization control. **C. Effect of JQ1 treatment for 8 hours and 24 hours on FOSL1 mRNA expression in PDAC cells.** PDAC cells were grown in three-dimensional type I collagen gels in the presence of DMSO (vehicle control), or JQ1 (0.5 μM) for 8 hours or 24 hours and FOSL1 mRNA expression analyzed by qRT-PCR using GAPDH as normalization control. *, p < 0.05.
Figure S7: Effect of JQ1 on BRD4 and RNA polymerase II occupancy on the c-MYC I and FOSL1 loci in Panc1 cells. A, B. Panc1 cells were treated with JQ1 (0.5 μM) for 24 hours and chromatin immunoprecipitation was performed across MYC and FOSL1 loci with control IgG antibody, anti-BRD4 antibody or anti-RNA polymerase II antibody. Purified DNA was then analyzed by PCR using primers specific for c-MYC or FOSL1 locus, and the PCR products visualized on 2% agarose gels. The results are representative of two independent experiments.
Fig. S1

A.

<table>
<thead>
<tr>
<th></th>
<th>0</th>
<th>0.25</th>
<th>0.5</th>
<th>1</th>
</tr>
</thead>
<tbody>
<tr>
<td>AsPC1</td>
<td><img src="image1.png" alt="Image" /></td>
<td><img src="image2.png" alt="Image" /></td>
<td><img src="image3.png" alt="Image" /></td>
<td><img src="image4.png" alt="Image" /></td>
</tr>
<tr>
<td>CD18</td>
<td><img src="image5.png" alt="Image" /></td>
<td><img src="image6.png" alt="Image" /></td>
<td><img src="image7.png" alt="Image" /></td>
<td><img src="image8.png" alt="Image" /></td>
</tr>
<tr>
<td>Panc1</td>
<td><img src="image9.png" alt="Image" /></td>
<td><img src="image10.png" alt="Image" /></td>
<td><img src="image11.png" alt="Image" /></td>
<td><img src="image12.png" alt="Image" /></td>
</tr>
</tbody>
</table>

B.

- **Average Colony Size (µM)**
  - AsPC1
  - CD18
  - Panc1

![Graphs](image13.png)
Fig. S4

I-BET151 [μM]

CD18-CR

Collagen

Average Colony

I-BET151 [μM]
Fig. S5

A. c-MYC

B. c-MYC

C. c-MYC

D. c-MYC
Fig. S6

A. FOSL1

B. FOSL1

C. FOSL1