

**Supplementary Table 1: Comparison of characteristics for patients with matching score <50% versus ≥50% and first versus second line or greater.**

<b>Characteristics</b>	<b>Matching Score &lt;50% (N = 7)</b>	<b>Matching Score ≥50% (N = 11)</b>	<b>P value</b>
<b>Age &gt; 67 years* (N (%))</b>	3 (42.8%)	6 (54.5%)	1.0
<b>Number of men (N (%))</b>	3 (42.9%)	4 (36.4%)	1.0
<b>Matched therapy as first line therapy (N (%))</b>	1 (14.2%)	4 (36.4%)	0.60
	<b>1<sup>st</sup> line of therapy</b>	<b>Later lines</b>	
<b>Matching score ≥ 50%</b>	4/5 (80%)	7/13 (54%)	0.3

\*Age 67 was the median age of the overall cohort.

**Supplementary Table 2:** Molecular alterations and agents administered for 13 patients who did not achieve clinical benefit

<b>Molecular alterations</b>	<b>Drugs Administered</b>	<b>Matching Score (%) ≥50% versus &lt;50%</b>
<i>CDKN2A</i> alteration <i>KRASG12V</i> <i>TP53</i> mutation	Palbociclib and trametinib	≥50%
<i>CDKN2A</i> alteration GNASR201H <i>KRAS G12D</i> <i>TP53</i> mutation	Palbociclib and trametinib	≥50%
<i>CDKN2A</i> alteration <i>ERBB2</i> amplification <i>KRAS G12D</i>	Palbociclib and trametinib	≥50%
<i>CDKN2A</i> alteration GNASR201C <i>KRAS Q61H</i> <i>SMAD4</i> alteration	Palbociclib and trametinib	≥50%
Alterations in <i>CDKNA</i> , <i>CCND1</i> , <i>CCNE1</i> , <i>KRAS</i>	Palbociclib, trametinib, and bevacizumab	≥50%
Alterations in <i>CDKN2A</i> , <i>KRAS</i> , <i>TP53</i>	Palbociclib, trametinib, and bevacizumab	≥50%
<i>MDM2</i> amplification Alterations in <i>AKT2</i> , <i>AXL</i> , <i>KRAS</i> , <i>TP53</i> ,	Trametinib and everolimus	<50%
<i>CDKNA</i> loss Alterations in <i>BRCA2</i> , <i>IRS2</i> , <i>KRAS</i> . <i>SMAD4</i> , <i>TP53</i>	Olaparib and trametinib	<50%
Alterations in <i>BRCA1</i> , <i>KRAS</i> , <i>MEN1</i> , <i>MSH6</i>	Oxaliplatin* and capecitabine	<50%
<i>CDKNA</i> loss Alterations in <i>ARID1A</i> , <i>BRCA1</i> , <i>CDKN1B</i> , <i>ERBB2</i> , <i>KRAS</i> , <i>TP53</i>	Olaparib	<50%
Alteration in <i>ATM</i> , <i>CCND2</i> , <i>CDH1</i> , <i>FGFR2</i> , <i>FGF23</i> , <i>GNAS</i> , <i>MYC</i> , <i>NOTCH1</i> , <i>SMAD4</i> , <i>TP53</i>	Erlotinib, cetuximab, palbociclib, and trametinib	<50%
Alterations in <i>CDK4</i> , <i>CDKN2A</i> , <i>ERBB2</i> , <i>ERBB3</i> , <i>GLI1</i> , <i>KRAS</i> , <i>TP53</i>	Trastuzumab and pertuzumab	<50%
<i>BRAF V600E</i> Alterations in <i>CDH1</i> , <i>TP53</i> ,	Trametinib and erlotinib	<50%

\*Platinum agents target DNA repair and hence were considered as matched targeted agents for *BRCA*-mutant tumors

**Supplementary Figure 1:** Consort Diagram: A total of 123 patients with pancreatic cancer had NGS results in the database.

