Supplementary Information

Cotargeting Plk1 and androgen receptor enhances the therapeutic sensitivity of paclitaxel-resistant prostate cancer

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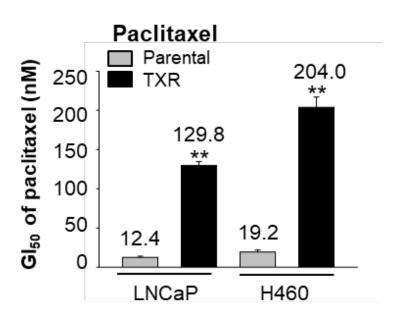
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Running Title

Cotargeting Plk1 and AR in taxol-resistant prostate cancer

Supplementary Figure 1.

Cells were treated with paclitaxel in a concentration-dependent manner in LNCaP, paclitaxel-resistant LNCaP (LNCaP^{TXR}), NCI-H460, and paclitaxel-resistant NCI-H460 (H460^{TXR}) cells for 48 h. The numbers of viable cells were then measured by a cell viability assay. The bar graph presents the mean values of half maximal growth inhibitory concentration (GI₅₀, nM for paclitaxel).



Supplementary Table 1. A combination of BI 2536 and bicalutamide in paclitaxel- resistant LNCaP^{TXR} **cells.** LNCaP and LNCaP^{TXR} cells were treated with BI 2536 and bicalutamide in a concentration-dependent manner for 48 h. The percentages of viable cells were measured by a cell viability assay.

			BI 2536 (nM)					
			0	1	5	10	25	50
	0	LNCaP	100.0 ± 0.1	89.0 ± 0.1	70.4 ± 1.3	48.9 ± 0.7	37.3 ± 0.4	29.5 ± 0.1
		LNCaP ^{TXR}	100.0 ± 1.1	96.0 ± 1.6	79.7 ± 0.5	70.6 ± 1.1	48.6 ± 1.2	33.1 ± 0.8
Bicalutamide	2.5	LNCaP	82.3 ± 0.9	67.3 ± 0.9	56.9 ± 0.8	43.7 ± 0.7	34.6 ± 1.1	25.0 ± 1.3
(μΜ)		LNCaP ^{TXR}	73.2 ± 1.6	67.4 ± 0.9	58.8 ± 1.6	50.0 ± 1.4	38.1 ± 0.9	27.0 ± 1.1
	5.5	LNCaP	72.3 ± 1.3	54.8 ± 1.3	51.5 ± 1.3	36.9 ± 1.1	22.7 ± 1.8	17.3 ± 0.8
		LNCaP ^{TXR}	67.6 ± 1.2	60.1 ± 1.0	53.2 ± 1.4	38.7 ± 1.0	28.6 ± 0.6	18.4 ± 0.4

Supplementary Table 2. A combination of volasertib and bicalutamide in paclitaxel- resistant LNCaP^{TXR} **cells.** LNCaP and LNCaP^{TXR} cells were treated with volasertib and bicalutamide in a concentration-dependent manner for 48 h. The percentages of viable cells were measured by a cell viability assay.

			Volasertib (nM)					
			0	1	5	10	25	50
	0	LNCaP	100.0 ± 0.1	89.0 ± 0.1	68.3 ± 1.1	50.9 ± 0.6	43.3 ± 0.3	29.5 ± 0.1
	-	LNCaP ^{TXR}	100.0 ± 1.1	92.3 ± 1.6	75.7 ± 0.5	69.8 ± 0.9	60.2 ± 1.0	49.1 ± 0.5
Bicalutamide	2.5	LNCaP	81.1 ± 0.4	77.9 ± 0.7	57.1 ± 0.2	46.8 ± 0.9	39.8 ± 0.4	24.4 ± 0.4
(μΜ)		LNCaP ^{TXR}	72.1 ± 0.3	66.0 ± 1.4	60.4 ± 1.7	51.7 ± 1.1	44.4 ± 0.7	30.6 ± 1.1
	5.5	LNCaP	71.8 ± 1.4	58.2 ± 1.6	53.4 ± 0.9	39.9 ± 2.0	28.1 ± 0.4	17.5 ± 0.8
		LNCaP ^{TXR}	66.5 ± 1.4	63.1 ± 0.9	55.7 ± 1.5	40.4 ± 1.4	33.5 ± 1.1	19.7 ± 0.8

Supplementary Table 3. A combination of genistein and bicalutamide in paclitaxel- resistant LNCaP^{TXR} **cells.** LNCaP and LNCaP^{TXR} cells were treated with genistein and bicalutamide in a concentration-dependent manner for 48 h. The percentages of viable cells were measured by a cell viability assay.

			Genistein (μM)					
			0	1	5	10	25	50
	0	LNCaP	100.0 ± 1.4	87.9 ± 1.3	79.9 ± 1.2	70.4 ± 1.1	60.8 ± 1.0	38.4 ± 1.4
		LNCaP ^{TXR}	100.0 ± 0.8	77.7 ± 2.3	64.0 ± 1.3	50.5 ± 0.5	46.1 ± 1.8	37.3 ± 1.1
Bicalutamide	2.5	LNCaP	82.0 ± 1.1	66.2 ± 1.2	61.0 ± 1.5	54.2 ± 1.3	47.8 ± 1.3	37.3 ± 2.2
(μΜ)		LNCaP ^{TXR}	74.0 ± 1.2	69.8 ± 1.6	59.3 ± 1.6	48.2 ± 0.9	35.3 ± 1.5	25.7 ± 0.6
	5.5	LNCaP	71.0 ± 1.6	65.4 ± 1.5	55.6 ± 1.2	48.4 ± 1.2	40.5 ± 0.8	36.1 ± 0.6
		LNCaP ^{TXR}	68.5 ± 1.2	55.2 ± 2.3	48.6 ± 0.9	37.9 ± 0.6	30.3 ± 1.2	19.8 ± 0.3

Supplementary Table 4. Oligonucleotide sequence of PCR primers used in this study

Gene	Forward primer (5'-3')	Reverse primer (5'-3')		
MDR1	5'-ATATCAGCAGCCCACATCAT -3'	5'- GAAGCACTGGGATGTCCGGT -3'		
MRP1	5'-CAGTGACCTCTGGTCCTTAAAC-3'	5'-ATCCACCTTGGAACTCTCTTTC-3'		
PLK1	5'- AAGAGATCCCGGAGGTCCTA -3'	5'- TCATTCAGGAAAAGGTTGCC -3'		
AR	5'-CGCTGAAGGGAAACAGAAGTA-3'	5'-CCTCTCCTTCCTCCTGTAGTT-3'		
PSA	5'-CTTGTAGCCTCTCGTGGCAG-3'	5'-GACCTTCATAGCATCCGTGAG-3'		

Supplementary Figure 2

Figure 1D

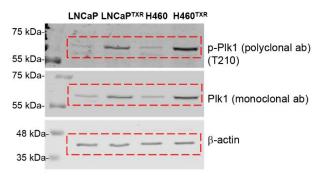


Figure 4F

