Kramer et al.:

Modeling Chemotherapy-Induced Peripheral Neuropathy Using a Nerve-on-a-Chip Microphysiological System

Supplementary Data

Tab. S1: Calculated values of nerve conduction velocity (NCV) and amplitude (AMP) presented as mean ±SEM

	Vehicle							
NCV (m/s)	0.303 ±0.017							
AMP (µV)	104.9 ±13.5							
	Paclitaxel							
	1 nM	10 nM	100 nM	200 nM	300 nM			
NCV (m/s)	0.288 ±0.016	0.320 ±0.018	0.221 ±0.013	0.237 ±0.014	0 ±0			
AMP (µV)	103.8 ±12.7	87.2 ±10.9	69.1 ±6.5	54.6 ±3.80	0 ±0			
	Oxaliplatin							
	0.01 μM	0.1 μΜ	1 μM	10 μM	100 μΜ			
NCV (m/s)	0.323 ±0.026	0.404 ±0.022	0.304 ±0.041	0.042 ±0.013	0 ±0			
AMP (µV)	84.0 ±7.9	100.4 ±15.8	94.2 ±19.8	25.8 ±8.4	0 ±0			
	Bortezomib							
	1 nM	2 nM	5 nM	10 nM	100 nM			
NCV (m/s)	0.357 ±0.018	0.412 ±0.022	0.367 ±0.011	0.022 ±0.013	0 ±0			
AMP (µV)	108.5 ±12.5	96.7 ±10.3	115.0 ±13.1	6.7 ±3.7	0 ±0			
		Vincristine						
	0.1 nM	1 nM	5 nM	10 nM	100 nM			
NCV (m/s)	0.293 ±0.017	0.307 ±0.017	0.286 ±0.016	0.223 ±0.027	0 ±0			
AMP (µV)	103.5 ±9.5	113.8 ±10.5	94.2 ±8.5	64.4 ±7.6	0 ±0			
		Acetaminophen						
	0.5 μΜ	5 μM	50 μM	500 μM	5000 μM			
NCV (m/s)	0.358 ±0.032	0.282 ±0.013	0.258 ±0.014	0.336 ±0.026	0.328 ±0.028			
AMP (µV)	107.5 ±11.6	97.3 ±19.1	94.2 ±9.2	86.7 ±14.2	87.5 ±9.2			
		Amoxicillin						
	0.5 μΜ	5 μM	50 μM	500 μM	5000 μM			
NCV (m/s)	0.285 ±0.039	0.286 ±0.008	0.261 ±0.029	0.273 ±0.018	0.228 ±0.041			
AMP (µV)	93.3 ±10.8	124.2 ±20.0	100.0 ±17.1	78.3 ±9.3	42.5 ±5.2			

Tab. S2: Summary of all experimental metrics measured at the NCV IC50 concentration

	Ptx (245 nM)	Ox (2.9 µM)	Bz (9.0 nM)	Vn (32 nM)
NCV	0.153 m/s	0.179 m/s	0.176 m/s	0.149 m/s
AMP	34 μV	90 μV	57 μV	39 µV
CCK-8	_	_	_	_
LDH	†	_	_	_
MBP staining	$\downarrow\downarrow\downarrow$	_	_	$\downarrow\downarrow$
Myelinated fiber density	↓	\	\downarrow	$\downarrow\downarrow\downarrow$
Nonmyelinated fiber density	$\downarrow\downarrow$	\	\downarrow	$\downarrow\downarrow\downarrow$
G-Ratio	_	_	_	$\downarrow\downarrow\downarrow$
% Irregular myelin	†	↑	_	↑
% Eccentric nuclei	↑ ↑↑	$\uparrow \uparrow \uparrow$	↑ ↑	↑
% Degenerated fibers	↑	_	1	$\uparrow \uparrow \uparrow$
Nonmyelinated fiber diameter	↑ ↑↑	↑	$\uparrow \uparrow$	↑
Myelinated fiber diameter	↑ ↑	_	↑	_
TMRE dye	_	_	1	_

[&]quot;—" indicates similar values. Three arrows indicate dramatic, readily apparent deviations from vehicle controls. Two arrows point to a moderately appreciable deviation, while one arrow shows a trend away from control values.

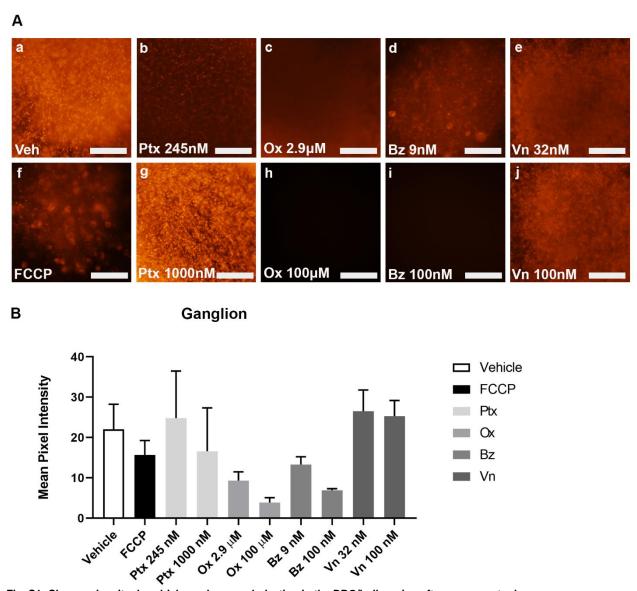


Fig. S1: Changes in mitochondrial membrane polarization in the DRG/bulb region after exposure to drugs

A) Live cells stained with TMRE mitochondrial membrane potential dye demonstrate changes in mitochondrial membrane polarization after exposure to vehicle control (a), Ptx (b, g), Ox (c, h), Bz (d, i), Vn (e, j), or positive control FCCP (f). Representative images taken from the DRG/bulb region of the construct. Scale bar: 100 μm. B) Mean pixel intensity calculated from N = 4 TMRE images. Data are mean + SEM.