

S1 Table. Main model parameters and associated values. All other model parameters are as in [1].

(*) Value is rescaled by the square of the simulation system characteristic length (1 cm) and divided by the system characteristic time (1 sec) multiplied by the oxygen diffusivity [2] ($1 \times 10^{-5} \text{ cm}^2 \text{ s}^{-1}$).

Parameter	Value	Reference
Tumor proliferation rate	1 day^{-1}	[3]
Tumor necrosis threshold	0.5700	[3]
Tumor hypoxic threshold	0.5750	[3]
Oxygen diffusivity	1 (*)	[4]
Oxygen transfer rate from vasculature	5 (*)	[4]
Oxygen uptake rate by proliferating tumor cells	1.5 (*)	[4]
Oxygen uptake rate by hypoxic tumor cells	1.3 (*)	[4]
Oxygen uptake rate by tumor microenvironment	0.12 (*)	[4]
Oxygen decay rate	0.35 (*)	[4]
NP diameter d	100; 600; 1,000 nm	[5]
NP drug load	(variable)	Proportional to d
NP drug release	(variable)	Proportional to \sqrt{d}
Drug diffusivity	0.6 (*)	Initial value; varied over [0.01,1.00]
Drug decay rate	6hr half-life	Similar to Paclitaxel

References:

- Wu, M., Frieboes, H.B., McDougall, S.R., Chaplain, M.A., Cristini, V., Lowengrub, J. The effect of interstitial pressure on tumor growth: Coupling with the blood and lymphatic vascular systems. *J Theor Biol* 320, 131-151 (2013).
- Nugent, L.J., Jain, R.K. Extravascular diffusion in normal and neoplastic tissues. *Cancer Res* 44, 238-244 (1984).
- van de Ven, A.L., Wu, M., Lowengrub, J., McDougall, S.R., Chaplain, M.A., Cristini, V., et al. Integrated intravital microscopy and mathematical modeling to optimize nanotherapeutics delivery to tumors. *AIP Adv* 2, 11208 (2012).
- Macklin, P., McDougall, S., Anderson, A.R., Chaplain, M.A., Cristini, V., Lowengrub, J. Multiscale modelling and nonlinear simulation of vascular tumour growth. *J Math Biol* 58, 765-798 (2009).
- Frieboes, H.B., Wu, M., Lowengrub, J., Decuzzi, P., Cristini, V. A computational model for predicting nanoparticle accumulation in tumor vasculature. *PLoS One* 8, e56876 (2013).