

## **PCL-PEG graft copolymers with tunable amphiphilicity as efficient drug delivery systems**

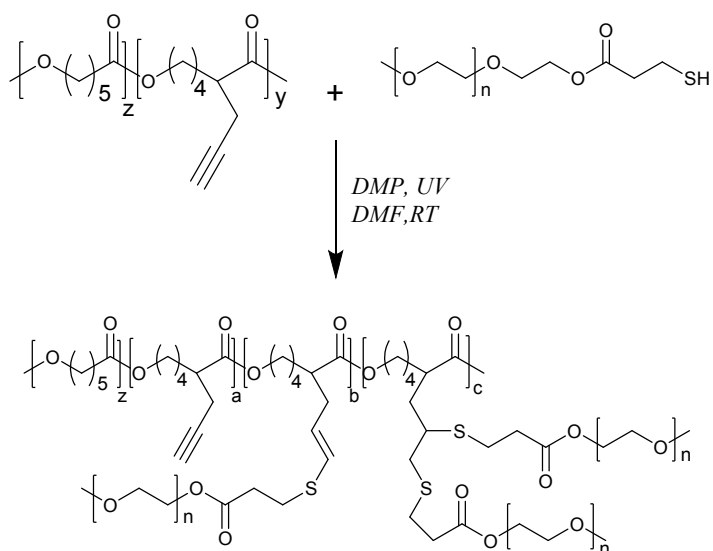
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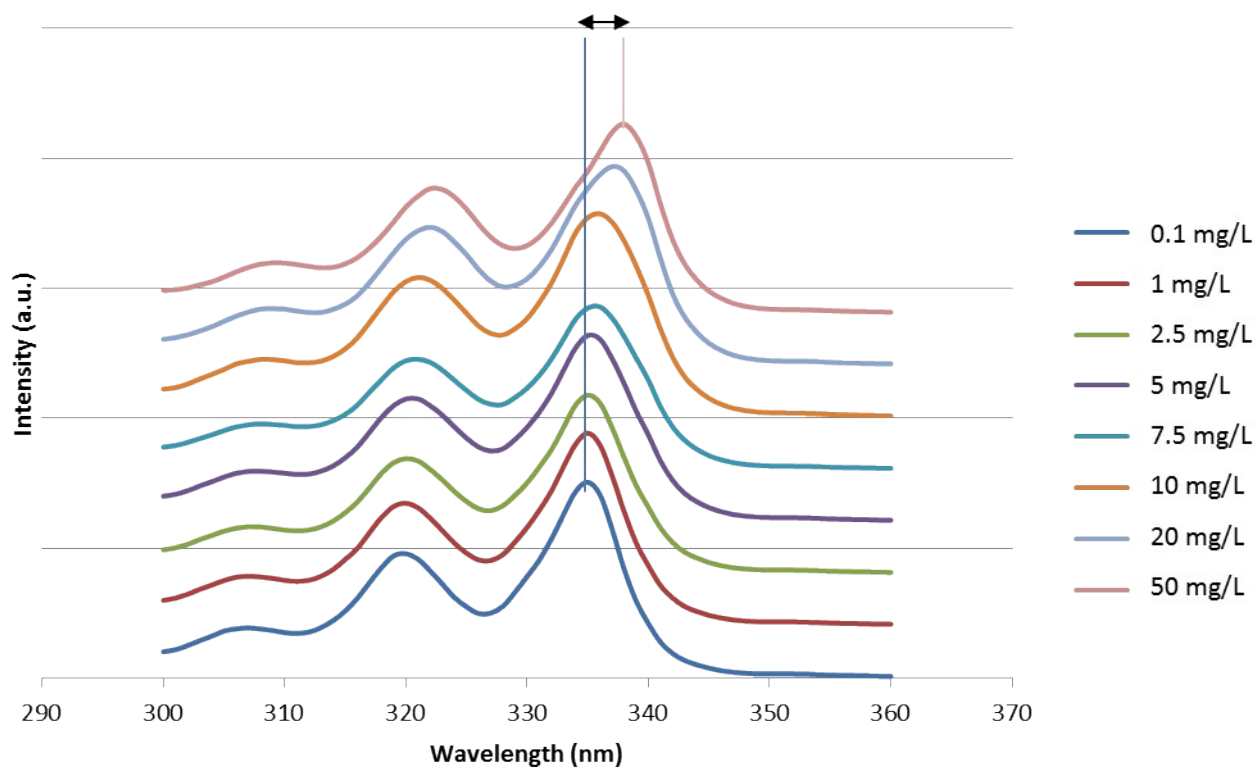
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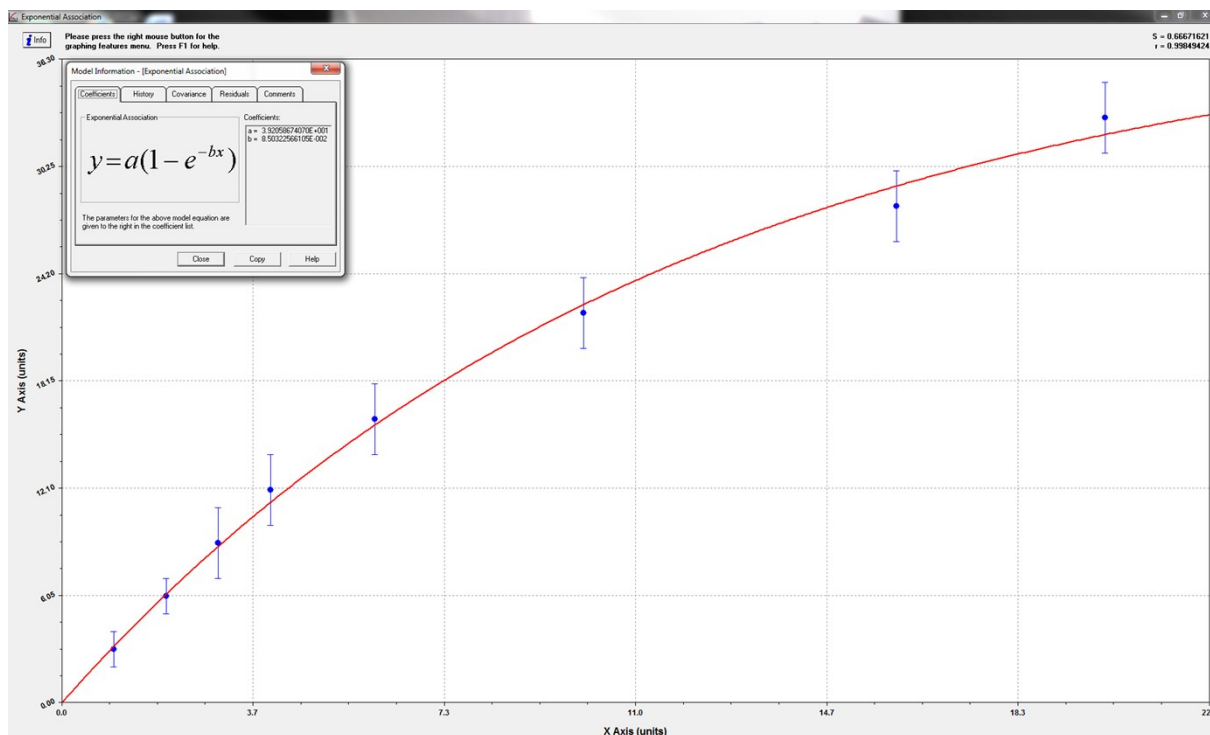
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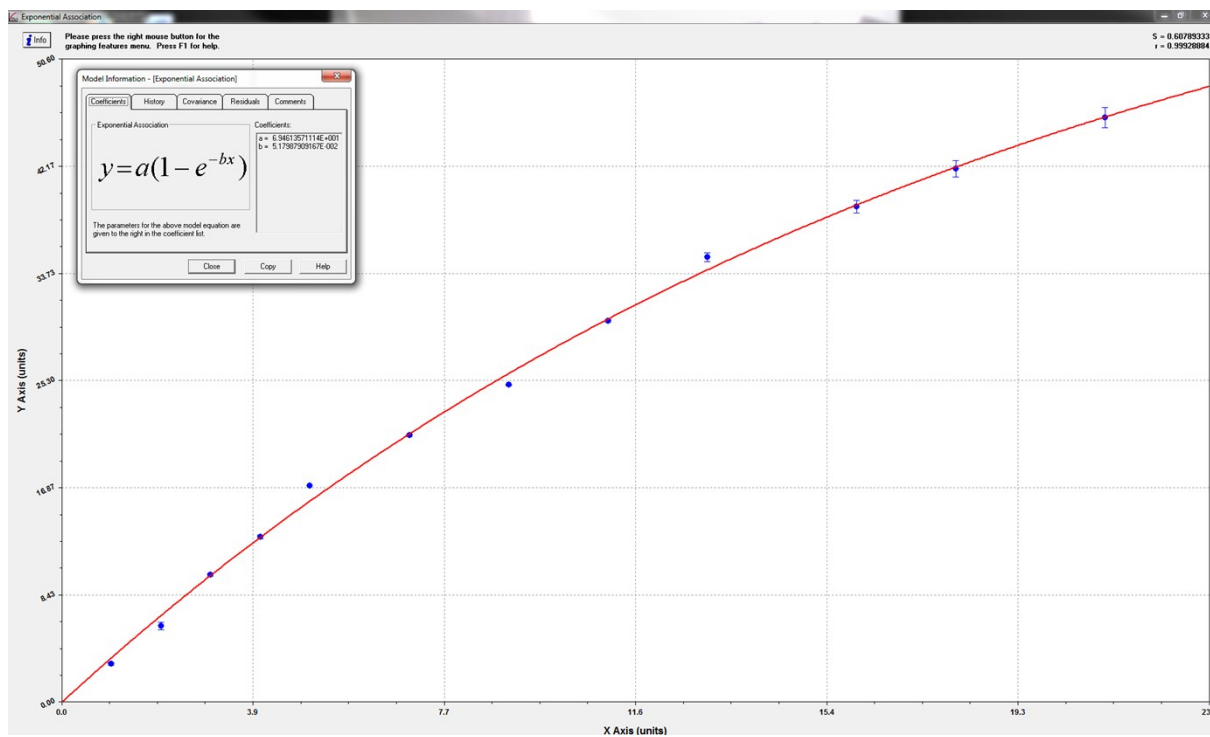
**Scheme S1.** Schematic representation of the preparation of the PCL-*g*-PEG copolymers by thiol-yne approach.



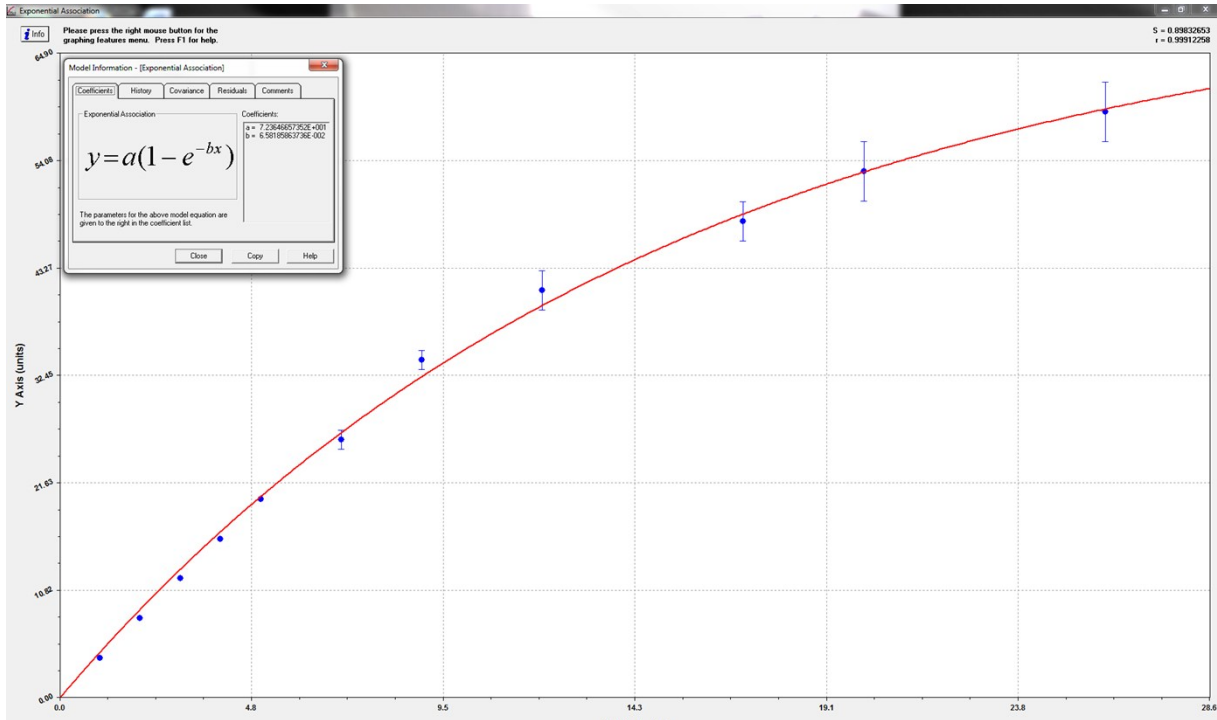
**Figure S1.** Excitation spectra of pyrene ( $\lambda_{em} = 371$  nm) as a function of the concentration of PCL-g-PEG2k<sub>3,2</sub>.



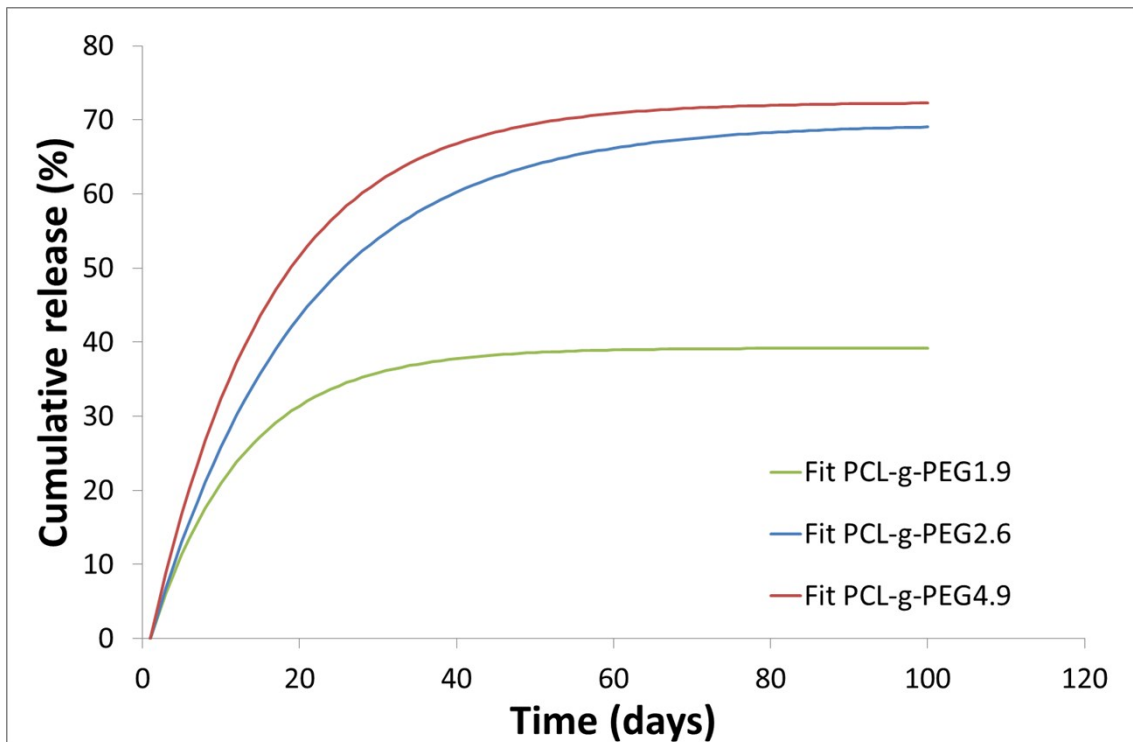
**Figure S2a.** Fit curve for the curcumin release from PCL-g-PEG0.7k<sub>1.9</sub> (software Curve Expert1.4)



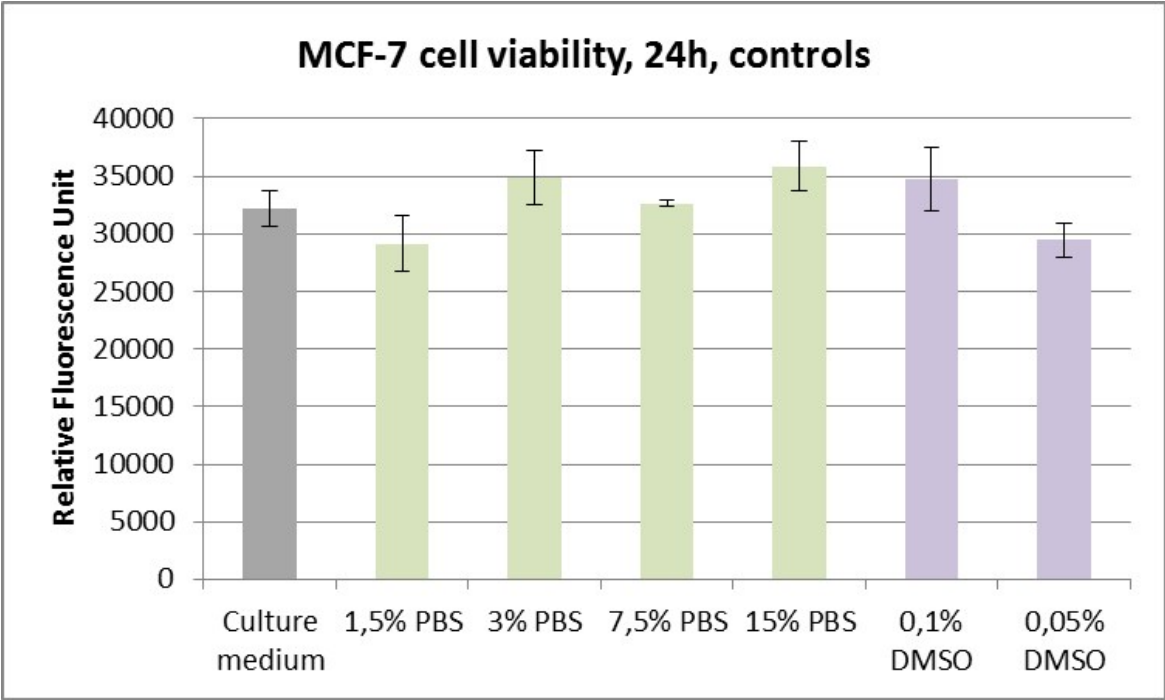
**Figure S2b.** Fit curve for the curcumin release from PCL-g-PEG2k<sub>2.6</sub> (software Curve Expert1.4)



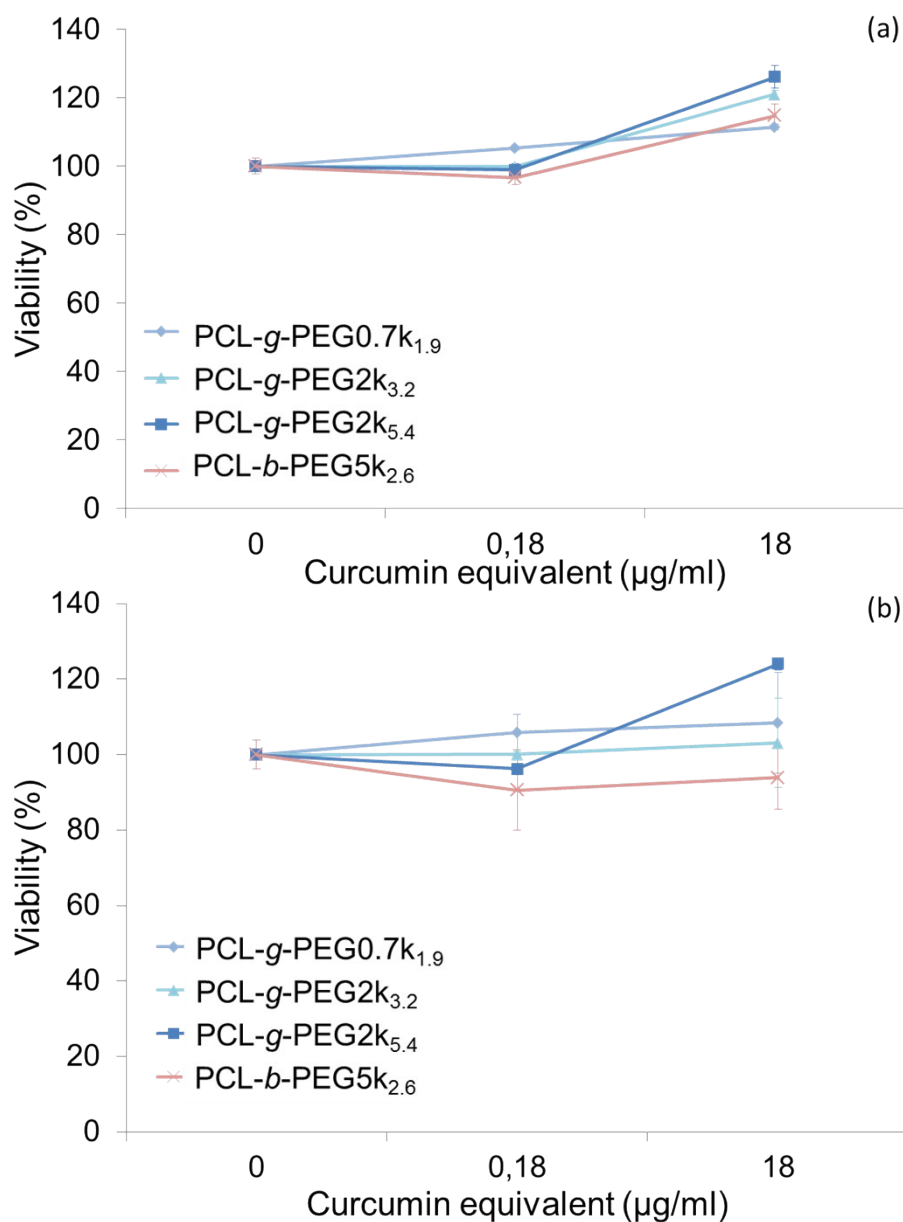
**Figure S2c.** Fit curve for the curcumin release from PCL-g-PEG2k<sub>4.9</sub> (software Curve Expert1.4)



**Figure S2d.** Extrapolation of fit curves over 100 days for the release curcumin from PCL-g-PEG



**Figure S3.** Viability of MCF-7 cells in different media. The grey bar represent the control in culture medium, green bars represent the culture medium supplemented with PBS (1.5% to 15%) and purple bars represent the culture medium supplemented with DMSO (0.05% and 0.1%)



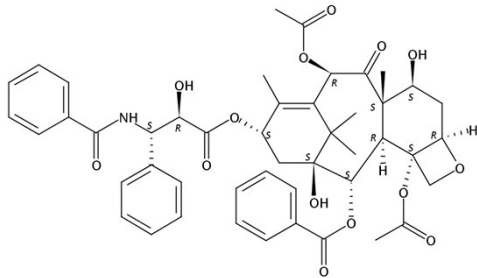
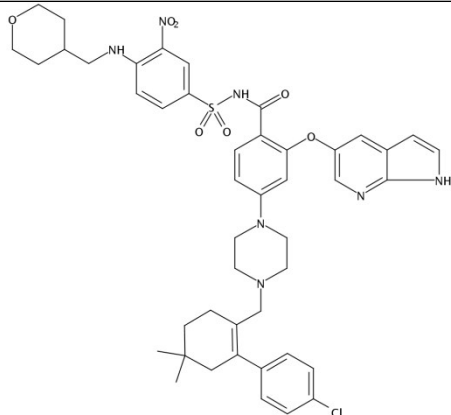
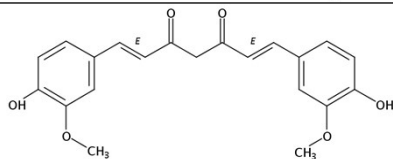
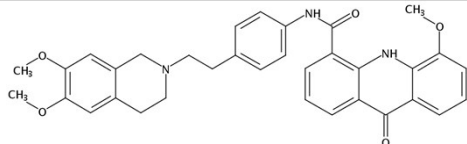
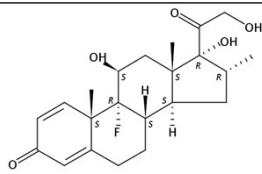
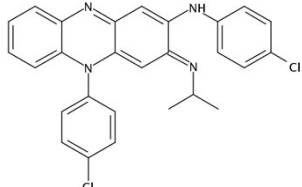
**Figure S4.** Viability of MCF-7 cells in the presence of non-loaded nanoaggregates. The concentration of blank nanoaggregates used in this assay was equal to the concentration of loaded nanoaggregates used to reach curcumin concentrations of 0.18 and 18 µg/L. (Data are expressed as means ± SD and correspond to measurements in triplicate)

**Table S1.** SEC analysis of PCL-*g*-PEG during degradation at 37°C in PBS. (Mn values expressed as means ± SD and correspond to measurements in triplicate)

Copolymer / Time	PCL- <i>g</i> -PEG0.7k <sub>1,2</sub>		PCL- <i>g</i> -PEG2k <sub>5,4</sub>	
	Mn (g/mol)	Đ	Mn (g/mol)	Đ
<b>T<sub>0</sub></b>	19400 ± 1400	2,1 ± 0,2	27000 ± 3700	1,6 ± 0,2
<b>1 W</b>	27000 ± 1150	2,6 ± 0,2	20000 ± 7200	1,5 ± 0,1
	1700 ± 100	1,2 ± 0,2		
<b>1 M</b>	27500 ± 1800	2,8 ± 0,1	17400 ± 5500	1,2 ± 0,1
	1200	1,2	2900 ± 100	1,0 ± 0,1
<b>3 M</b>	16400 ± 500	1,9 ± 0,1	37100 ± 2300	1,4 ± 0,1
	960 ± 60	1,1	2600 ± 100	1,0 ± 0,1
<b>6 M</b>	12800 ± 400	1,6 ± 0,1	26000 ± 4900	1,4
	900 ± 50	1,1	2500 ± 100	1,1



**Table S2.** Chemical structure and solubility parameters of tested APIs

Drug	Chemical structure	Solubility	logP*
Paclitaxel		1 mg/L Zhang JA. <i>et al. Eur J Pharm Biopharm</i> <b>2005</b> , <i>59</i> , 177.	3.95
ABT-199 (Venetoclax)		< 0.4 mg/L 3. Choo E.F. <i>et al. Drug Metab Dispos</i> <b>2014</b> , <i>42</i> , 207.	8.05
Curcumin		0.6 mg/L B.T. Kurien <i>et al. Assay Drug Dev Technol</i> , <b>2007</b> , <i>5</i> , 567.	3.07
Elacridar		120 µg/L Sane R. <i>et al. J Pharm Sci</i> <b>2013</b> , <i>102</i> , 1343.	4.43
Dexamethasone		0.1 g/L Dilova V. <i>et al. Boll Chim Farm</i> <b>2004</b> , <i>143</i> , 20.	2.03
Clofazimine		0.3 g/L Peters K. <i>et al. J. Antimicrob. Chemother.</i> <b>2000</b> , <i>45</i> , 77.	7.46

\*From Scifinder, calculated using Advanced Chemistry Development (ACD/Labs) Software V11.02