

## Supplementary Information

# Doxorubicin-Polysorbate 80 Conjugates: Targeting Effective and Sustained Delivery to the Brain

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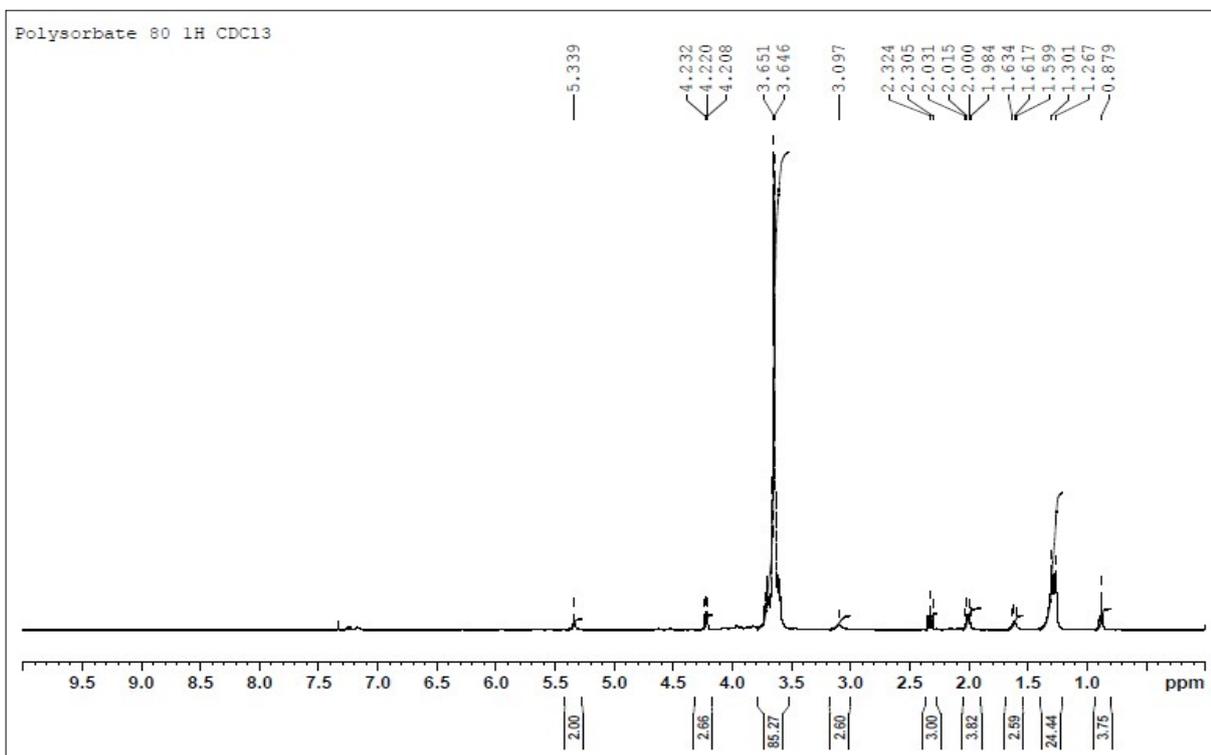
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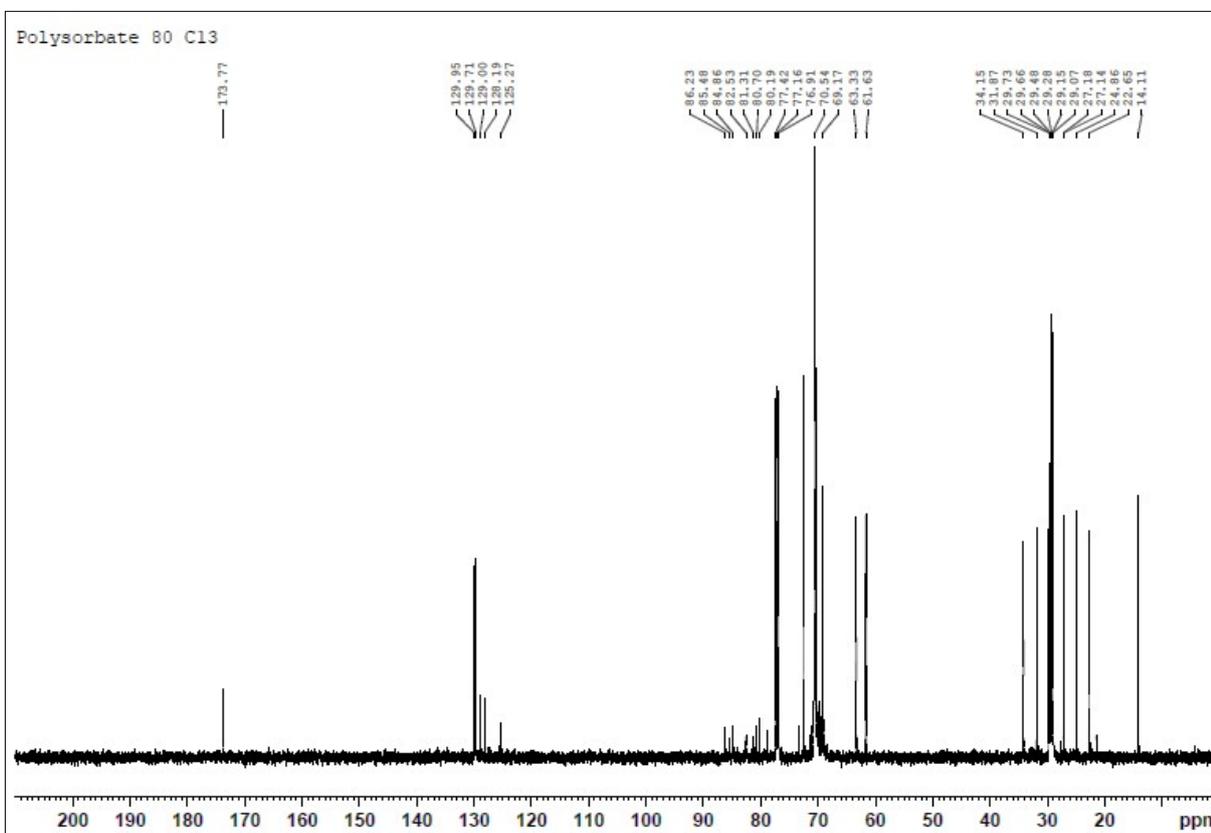
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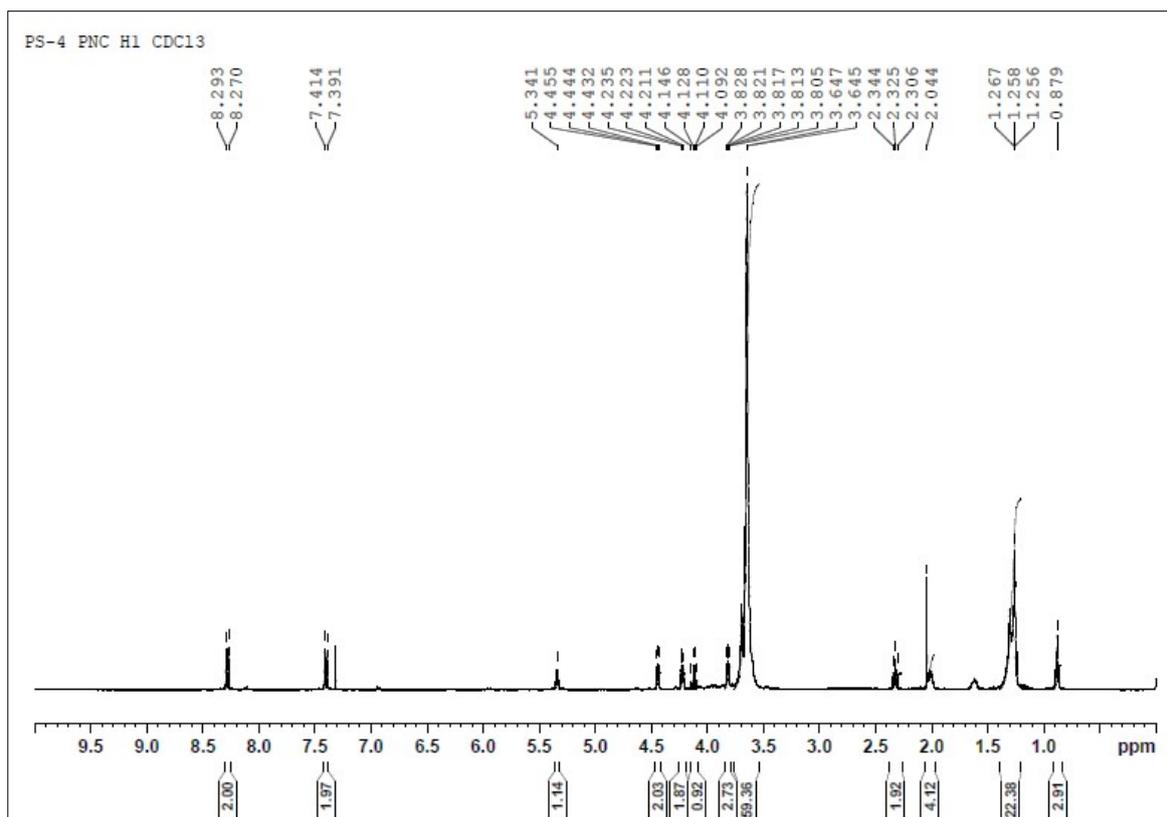
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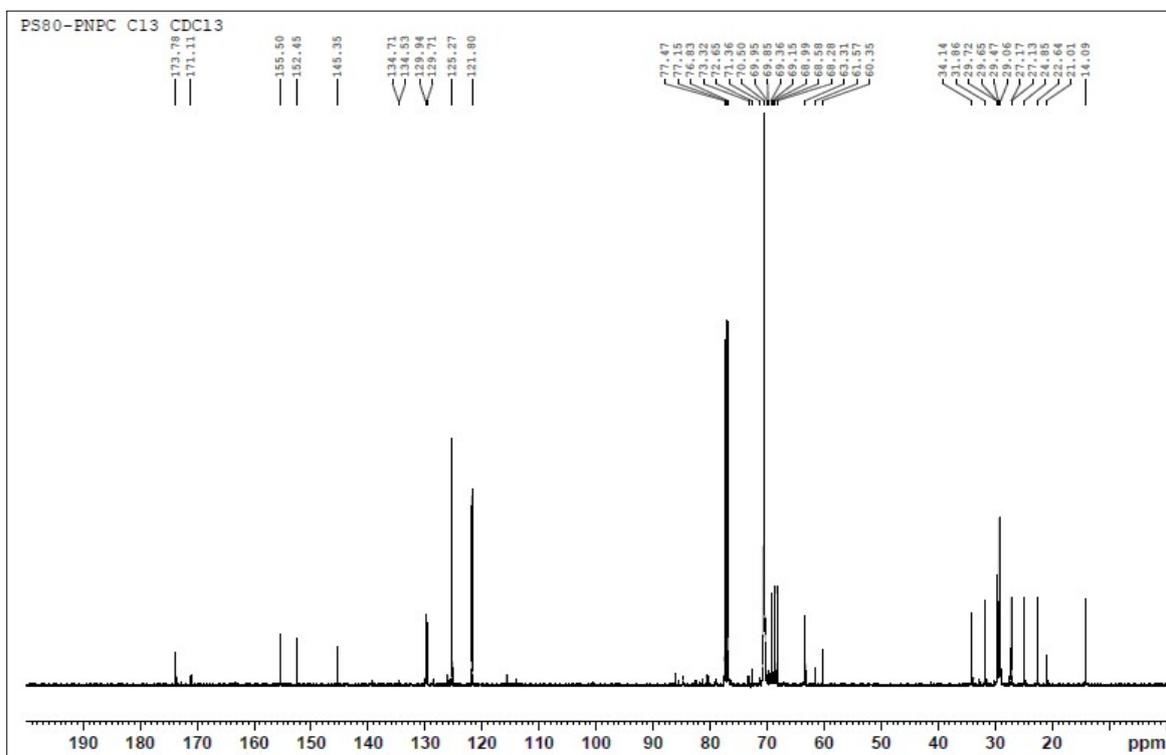
**Fig. S1.**  $^1\text{H}$  NMR spectrum of Polysorbate 80



**Fig. S2.**  $^{13}\text{C}$  NMR spectrum of Polysorbate 80



**Fig. S3.**  $^1\text{H}$  NMR spectrum of Polysorbate 80-(4-nitrophenyl carbonate)



**Fig. S4.**  $^{13}\text{C}$  NMR spectrum of Polysorbate 80-(4-nitrophenyl carbonate)

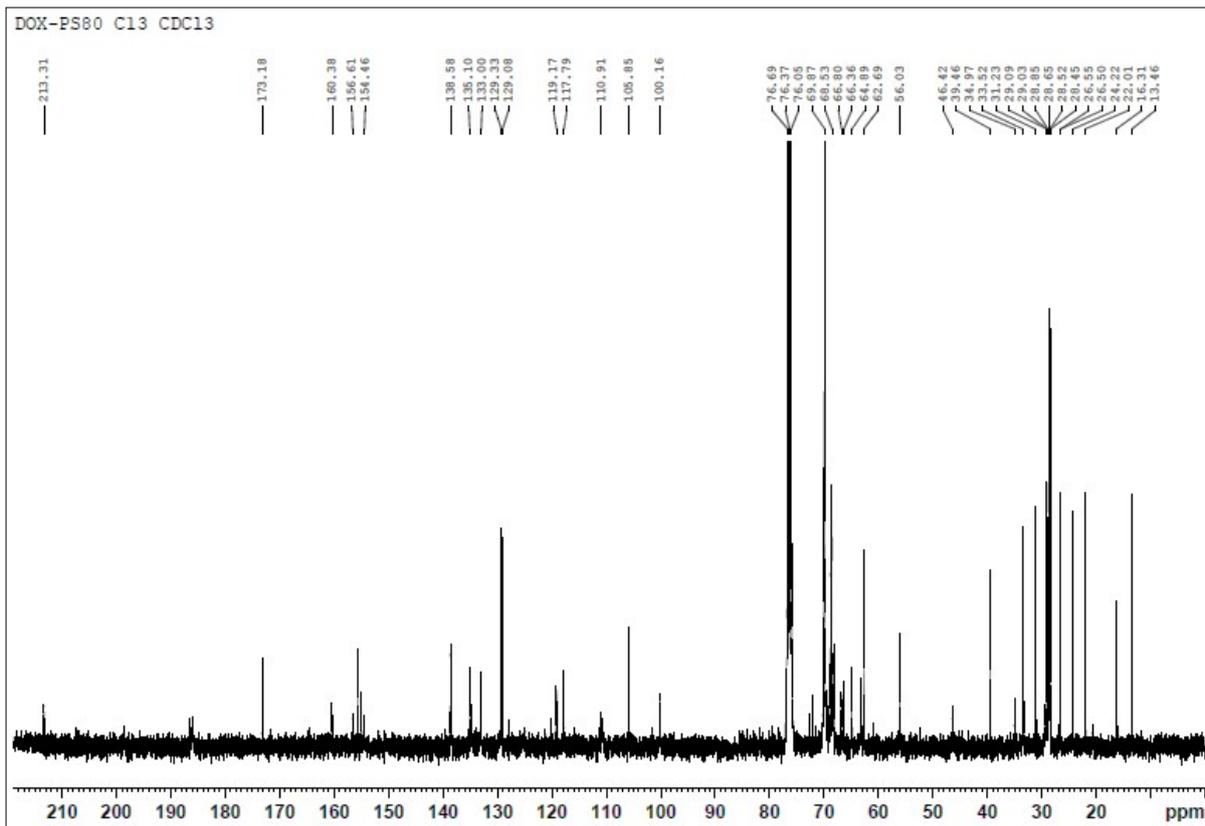
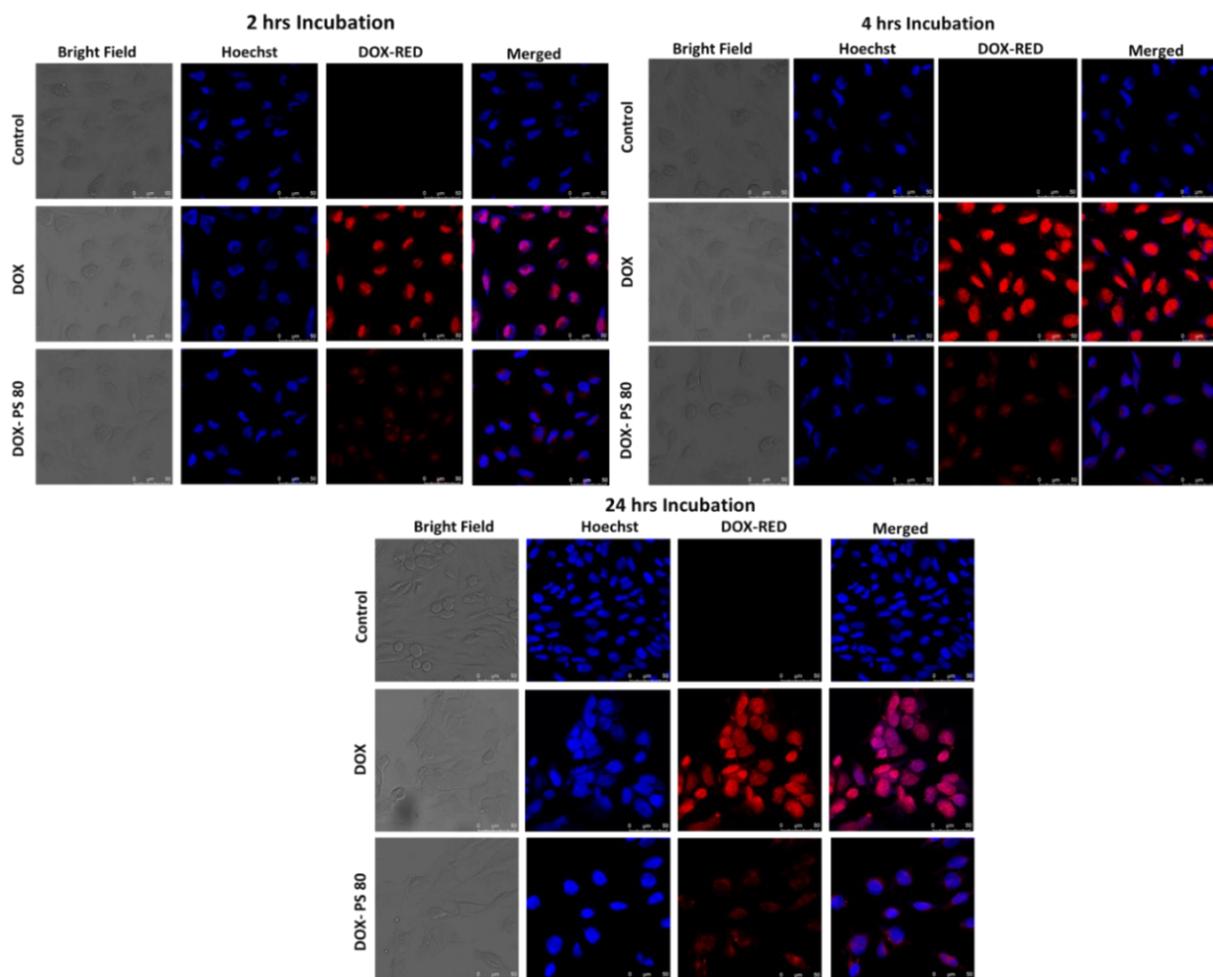
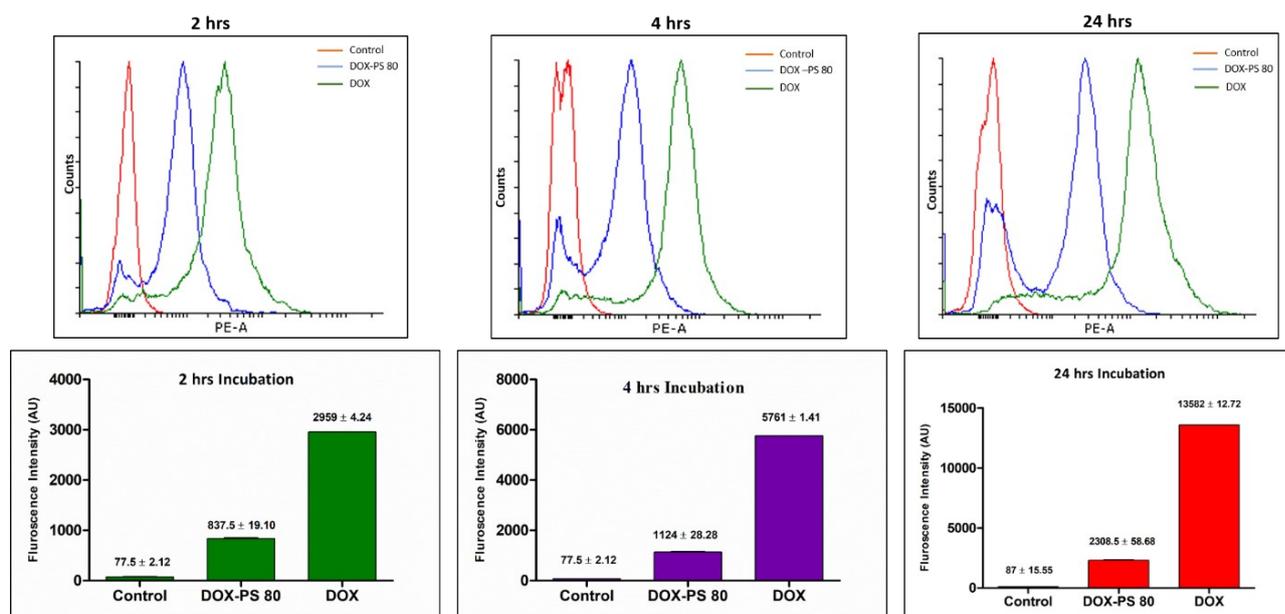


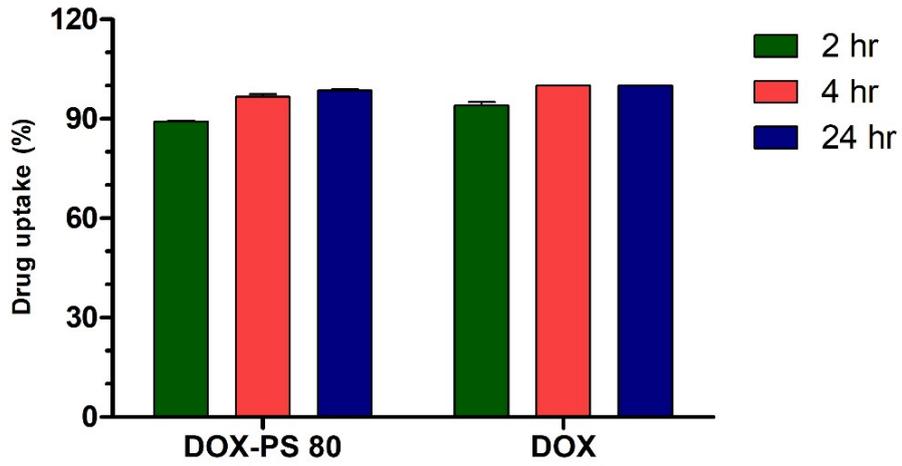
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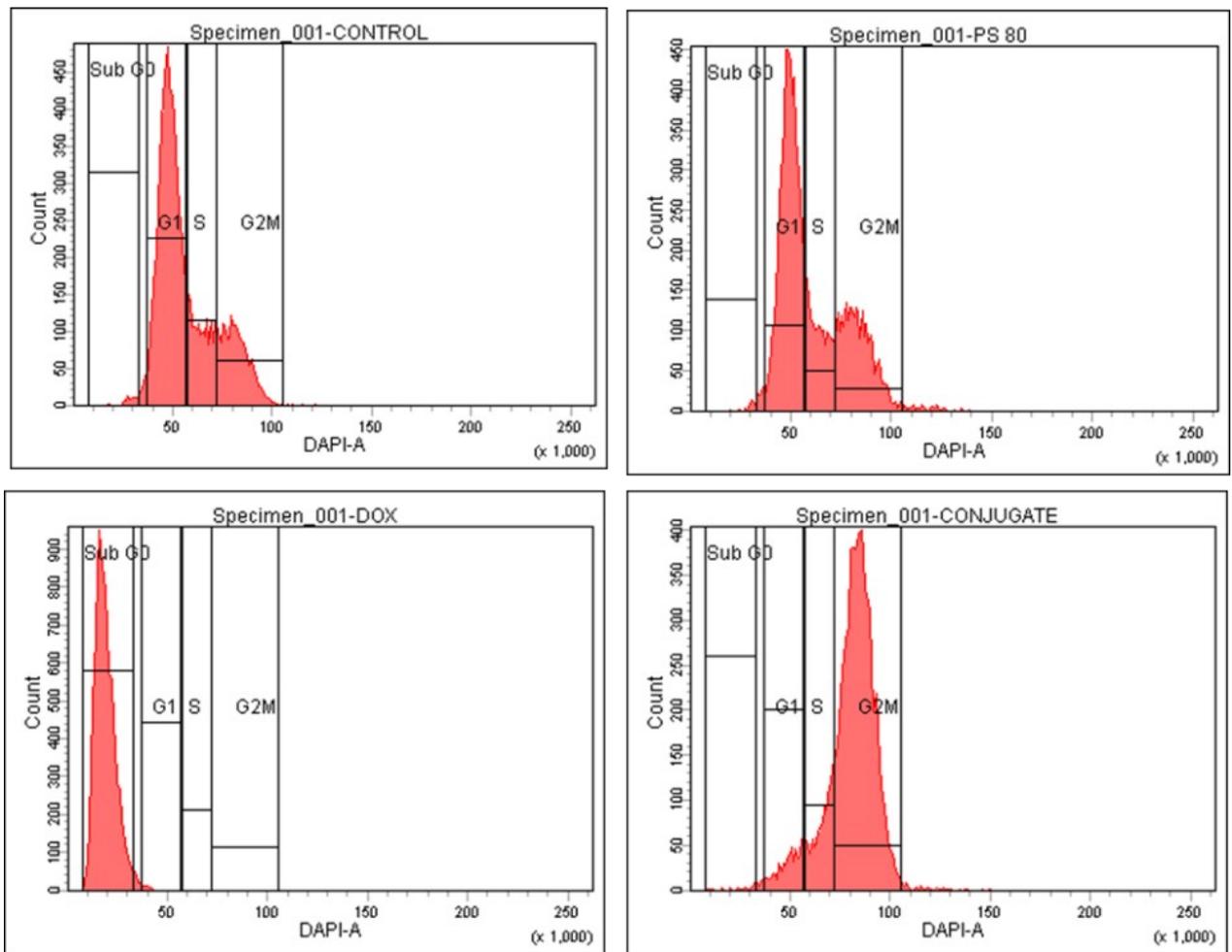
**Fig. S6.** Cellular uptake studies at 2, 4 and 24 hr incubation by confocal imaging



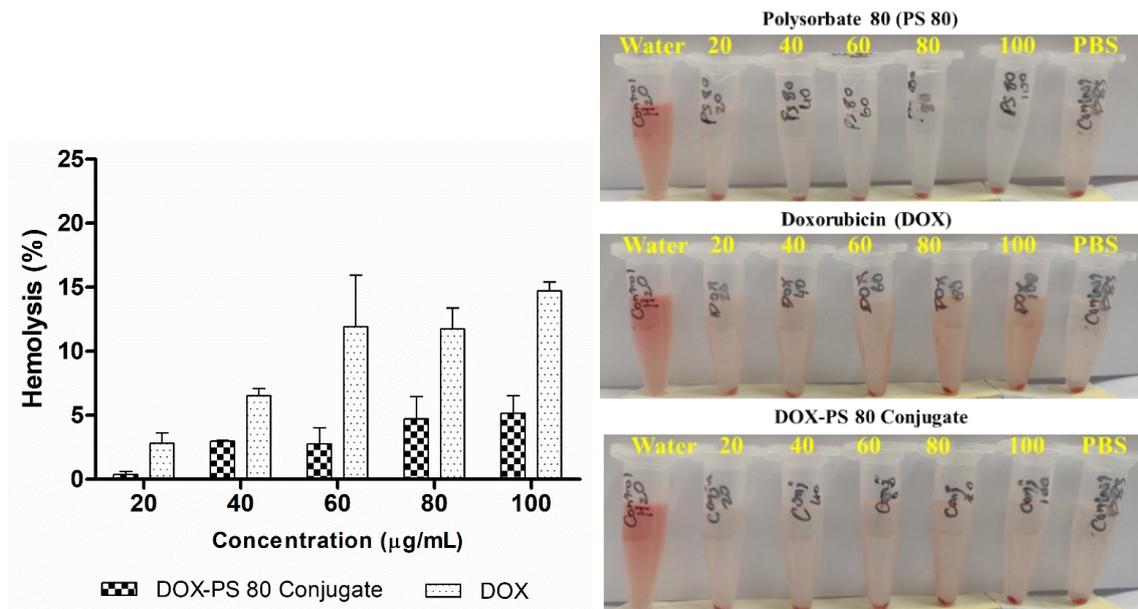
**Fig. S7.** The intra-cellular uptake and mean fluorescence intensity for DOX-PS 80 and DOX using flow cytometry.



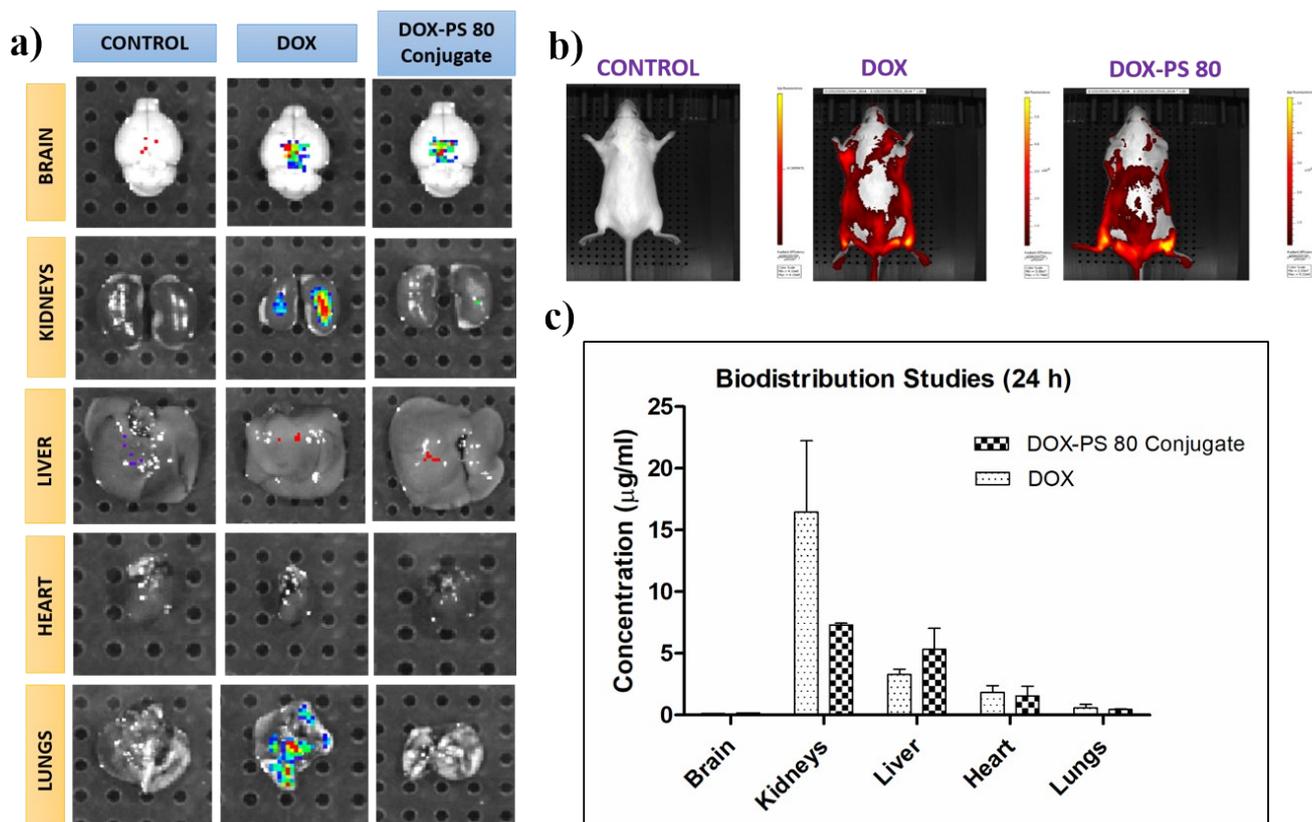
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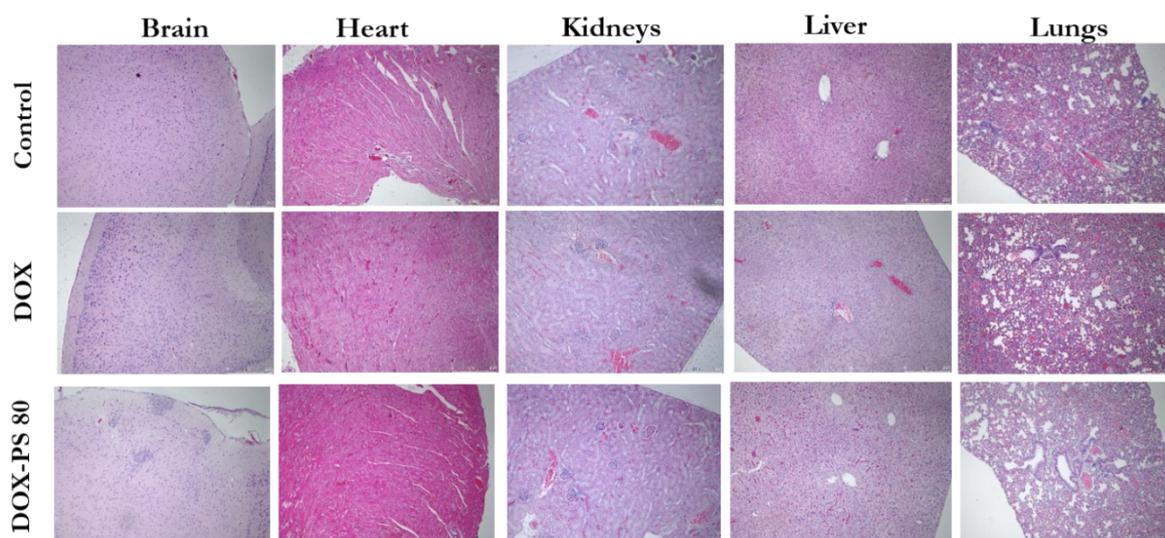
**Fig. S9.** Cell Cycle Analysis by Flow cytometry



**Fig. S10.** Hemolytic assay for DOX-PS 80 and DOX at different concentration



**Fig. S11.** IVIS fluorescence image of different organs after 24 h drug treatment (a), whole live animal image at 24 h (b), Quantitative estimation of DOX and DOX-PS 80 in organs (c)



**Fig. S12.** Histopathological Observations of brain, heart, kidneys, liver, and lungs tissues after 24 h drug treatment in Swiss albino mice.

**Table S1.** Concentration of DOX and DOX-PS 80 in Organs at 24 h

Organs	Doxorubicin (µg/ml)	DOX-PS 80 Conjugate (µg/ml)
<b>Brain</b>	0.11 ± 0.022	0.16 ± 0.03
<b>Kidneys</b>	16.48 ± 8.14	7.30 ± 0.21
<b>Liver</b>	3.29 ± 0.62	5.32 ± 2.42
<b>Heart</b>	1.85 ± 0.77	1.54 ± 1.11
<b>Lungs</b>	0.58 ± 0.40	0.44 ± 0.09