

**Supplementary Information**

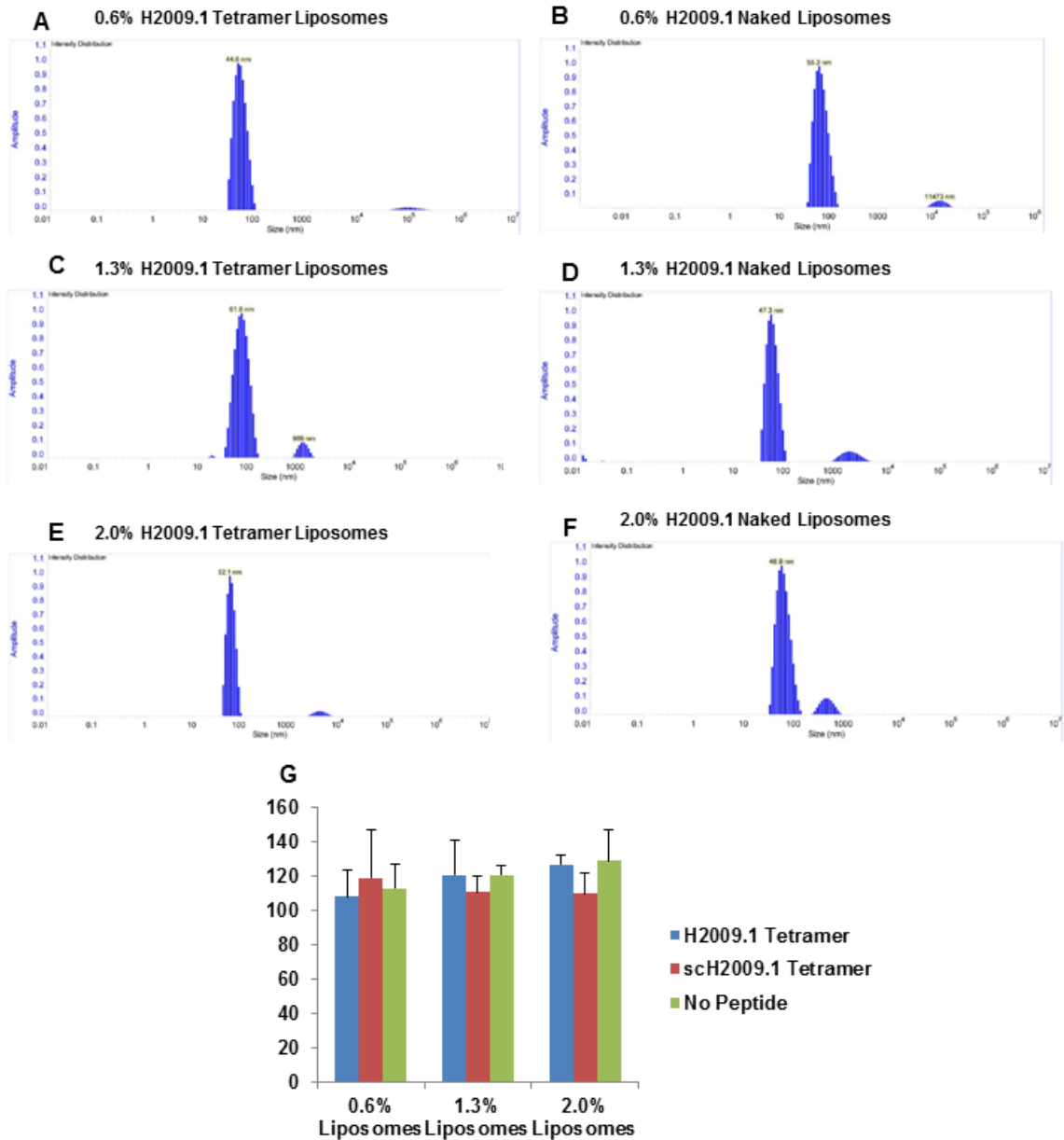
**From Phage Display to Nanoparticle Delivery: Functionalizing  
Nanoparticles with Multivalent Peptides Improves Targeting to a  
Cancer Biomarker**

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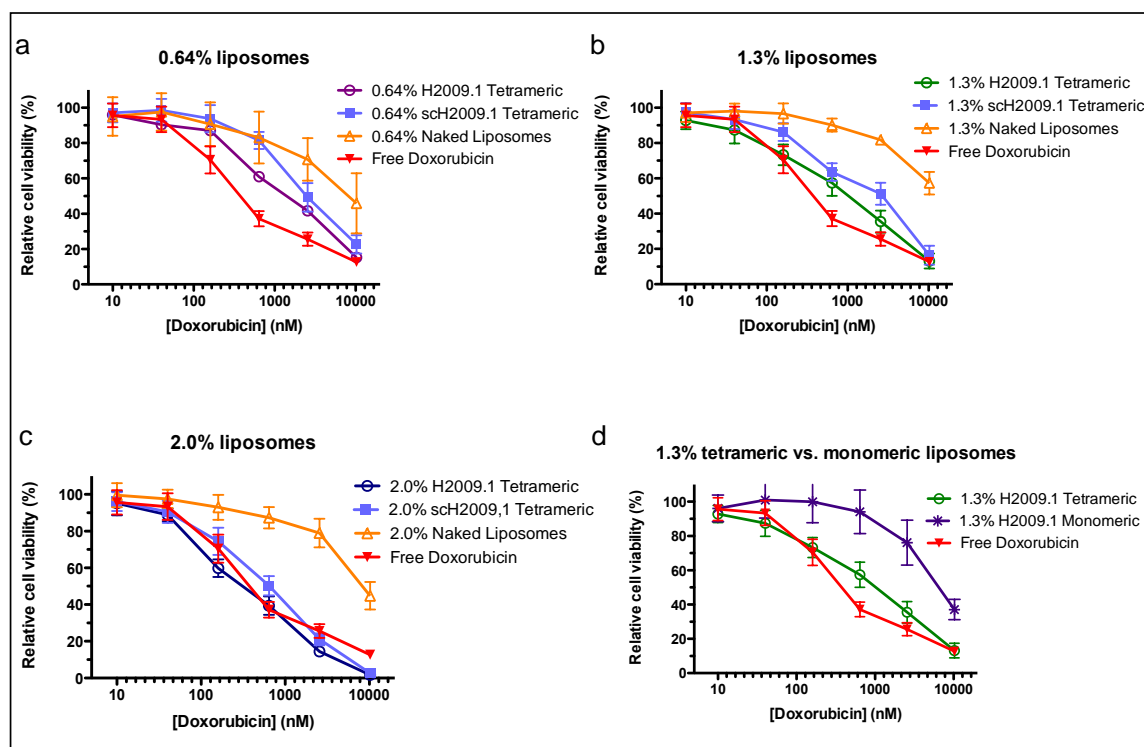
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**Figure 1S.** *The liposomes are approximately 100 nm in diameter.* Representative intensity graphs displaying the size distribution of the liposomes as determined by dynamic light scattering. A: 0.6% H2009.1 tetrameric liposomes B: 0.6% naked liposomes C: 1.3% H2009.1 tetrameric liposomes D: 1.3% naked liposomes E: 2.0% H2009.1 tetrameric liposomes F: 2.0% naked liposomes G: Average diameter of the liposomes. Error bars represent standard deviation of at least 3 independent liposome preparations.

Table S1. Liposome Formulations			
Formulation (molar percentages)			
Lipid	0.64%	1.3%	2.0%
HSPC <sup>a</sup>	65%	65%	65%
Cholesterol	32%	32%	32%
DSPE-PEG <sub>2000</sub> <sup>b</sup>	2.5%	1.9%	1.2%
DSPE-PEG <sub>2000</sub> -Maleimide <sup>c</sup>	0.64%	1.3%	2.0%

<sup>a</sup> Hydrogenated soy phosphatidylcholine. <sup>b</sup> 1,2-distearoyl-*sn*-glycero-3-phosphoethanolamine-N-[carbonyl-methoxypolyethylene glycol-2000]. <sup>c</sup> 1,2-distearoyl-*sn*-glycero-3-phosphoethanolamine-N-[maleimide(polyethylene glycol)-2000].



**Figure S2. H2009.1 tetrameric liposomes are selectively cytotoxic against  $\alpha_v\beta_6$ -expressing cells.** H2009 cells were incubated with liposomes for 1 hour before washing to remove non-internalized drug and adding fresh media. At 120 hours, cell viability was determined and normalized to non-treated cells using the ATP-based CellTiter-Glo® Luminescent Cell Viability Assay. (a-c) Viability for cells treated with tetrameric peptide liposomes. (d) Viability for cells treated with 1.3% H2009.1 tetrameric versus 1.3% H2009.1 monomeric liposomes.

**Table S2. IC<sub>50</sub> Values of Different Liposome Formulations on  $\alpha_v\beta_6$ -Negative H1299 Cells<sup>a</sup>**

<b>Drug Formulation</b>	<b>IC50 on H1299 cells (nM)</b>
1.3% H2009.1 Tetrameric	3700 ± 130
1.3% scH2009.1 Tetrameric	3500 ± 400
1.3% Naked	-
Free Doxorubicin	1400 ± 58

<sup>a</sup> Cells were incubated with liposomes for 1 hour followed by a 120 hour recovery in media.