Supporting Information

Vitamin E based micelles with charge-reversible property for tumor growth inhibition and antimetastasis

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d

2.0

ppm

1.5

0.0

HN N N O

0.5

Ň

1.0

h f,g

2.5

3.0

3.5

R=





Scheme S2. ¹H spectrum of the charge-reversal micelles after incubation under pH 5.5 for 4 h with the detachment of cleavable carboxylic amide.



Scheme S3. ¹H spectrum of the cleavable carboxylic amide.



Scheme S4. (A and B) ¹H NMR spectra of the DPV micelles in CCI_3D (A) and D_2O (B). (C and D) ¹H NMR spectra of the SPV micelles in D_2O (C) and CCI_3D (D).



Figure S1. Characterization of SPVM. (A) The particle size distribution of SPVM. (B) TEM image of DPVM. (C) CMC of SPV. (D) Zeta potential of SPVM incubating at various pH (7.4, 6.8, 5.5) at 37 °C with the change of incubation time.



Figure S2. The stability of DPVM in FBS at pre-determined dilution factors.



Figure S3. TEM image and DLS data of DPVM incubated at pH 5.5 after 1 h. (A) The particle size distribution of DPVM after incubation under pH 5.5 after 1 h. (B) TEM image of DPVM after incubated at pH 5.5 after 1 h.



Figure S4. In vivo imaging studies of mice after injection via tail vein of DiR loaded SPVM and DPVM. Images of tumorbearing mice were recorded at 0, 4, 8, 12, 24, and 48 h.





Figure S5. Safety evaluation in S180 tumor model. (A) AST, ALT and LDH levels of four groups at the end of the treatment. (B) BUN and CRE levels. (C) H&E staining of organs including heart, liver, spleen, lung and kidney.

| рН | Average size (nm) | PDI |
|-----|---------------------|-------------------|
| 7.4 | 189.7±1.3nm | 0.075±0.002 |
| 6.8 | 681.0±14.4nm | 0.184 ± 0.067 |
| 5.5 | 1081.0 \pm 35.2nm | 0.378±0.078 |
| 4.5 | 1287.0 \pm 27.7nm | 0.228 ± 0.078 |

 Table S1.
 Particle size and PDI (polydispersity index) change of DPVM at pH 7.4, 6.8, 5.5 and 4.5 after 2 h.

| pН | Incubation | IC ₅₀ (µg/mL) | | |
|-----|------------|--------------------------|------|------|
| | time (h) | Taxol | SPVM | DPVM |
| 7.4 | 24 | 6.54 | 8.50 | 7.73 |
| | 48 | 2.09 | 2.76 | 2.86 |
| 6.8 | 24 | 6.58 | 8.70 | 5.54 |
| | 48 | 2.08 | 3.00 | 0.76 |

Table S2. IC_{50} values (μ g/mL) of Taxol, SPVM and DPVM after incubation with 4T1 cells for 24 and 48 h under pH 7.4 and 6.8, respectively.