

## Supplementary Materials

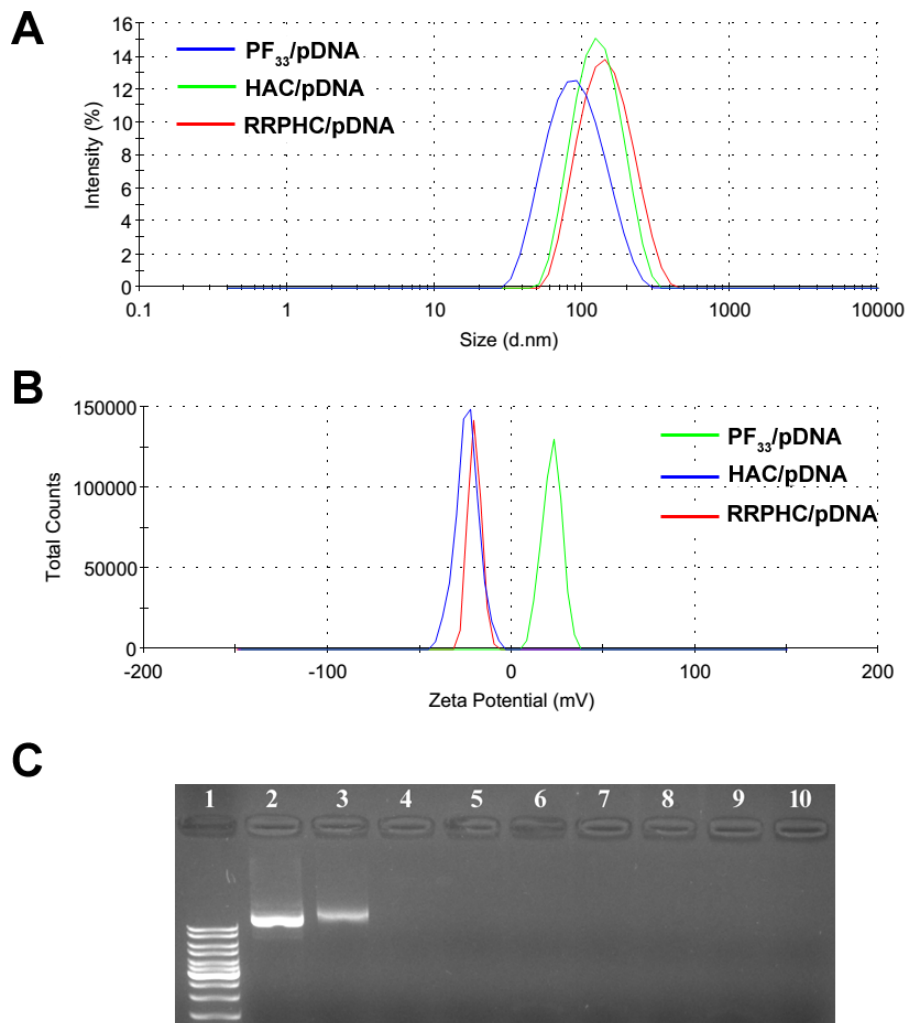
### **Multifunctional Nucleus-targeting Nanoparticles with Ultra-high Gene Transfection Efficiency for *In Vivo* Gene Therapy**

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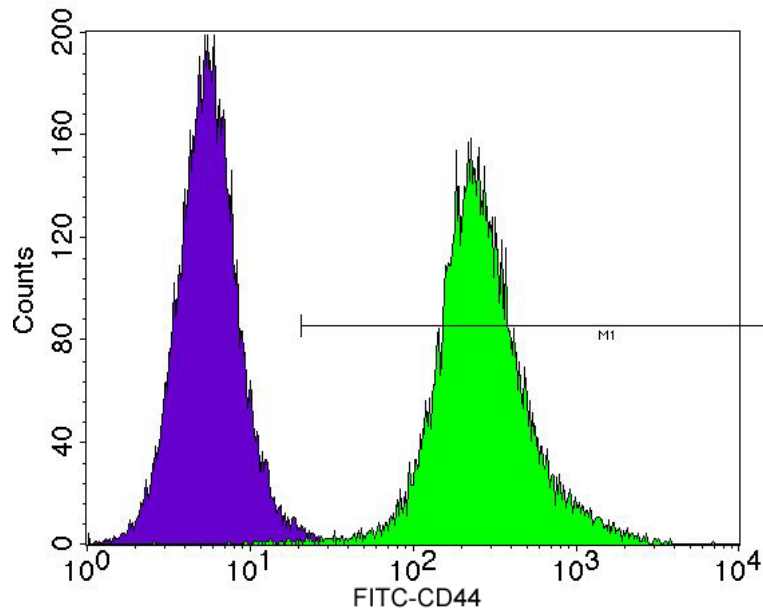
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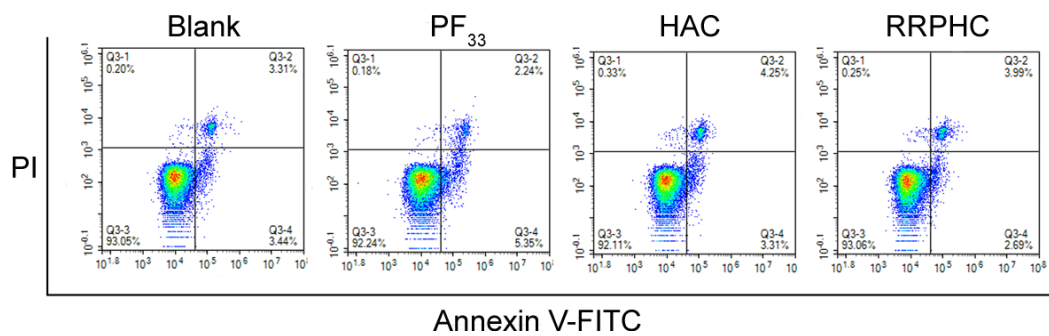
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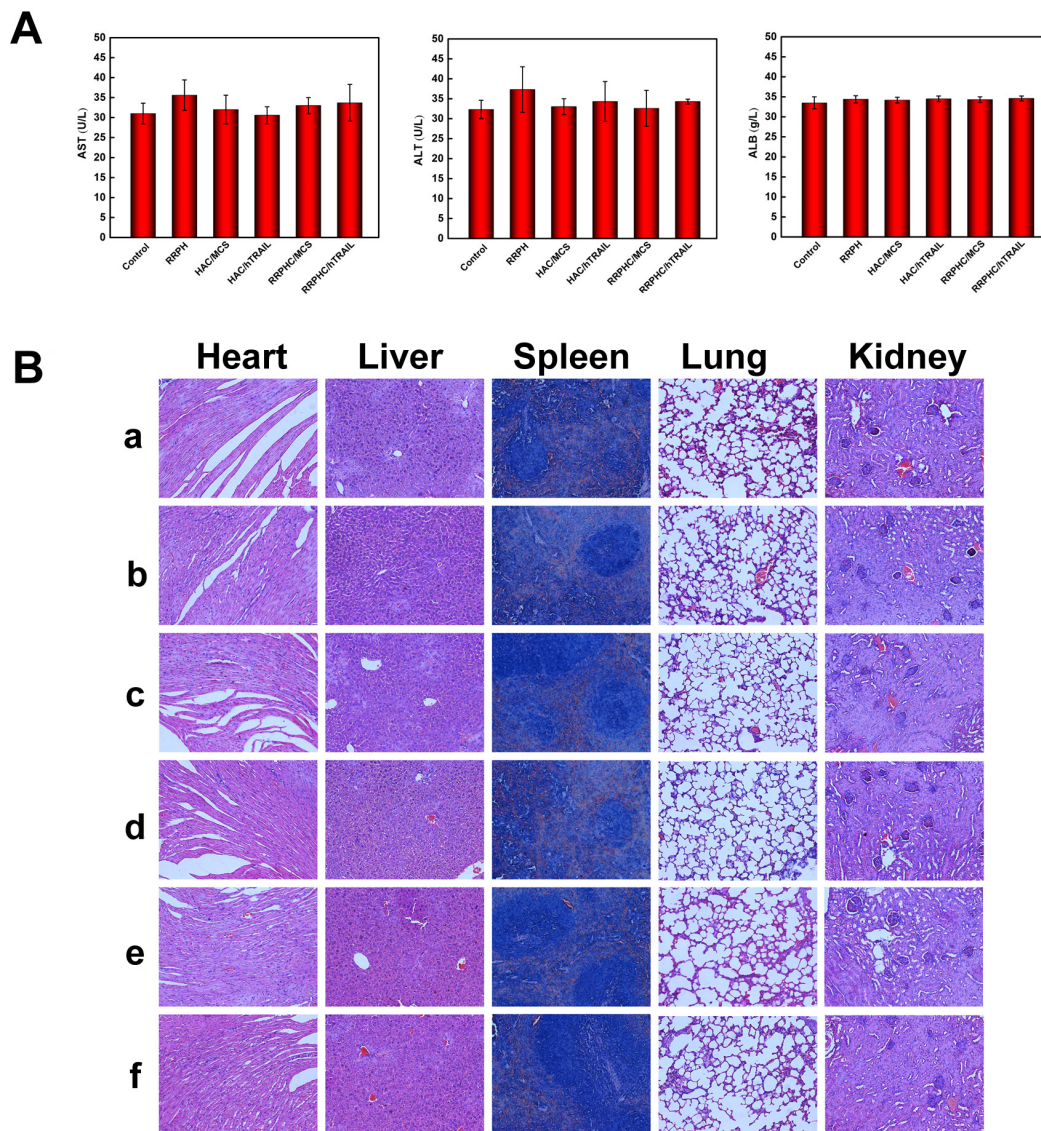
**Fig. S1.** (A, B) Particle size and zeta potential of the binary and ternary complexes measured by DLS. (C) Agarose gel electrophoresis of PF<sub>33</sub>/pDNA complexes, HAC/pDNA complexes and RRPHC/pDNA complexes. Lane 1, DNA ladder; lane 2, naked pDNA; lane 3-8, PF<sub>33</sub>/pDNA at mass ratios of 0.5:1, 1:1, 2:1, 4:1, 8:1, 10:1; lane 9, HAC/pDNA; lane 10, RRPHC/pDNA.



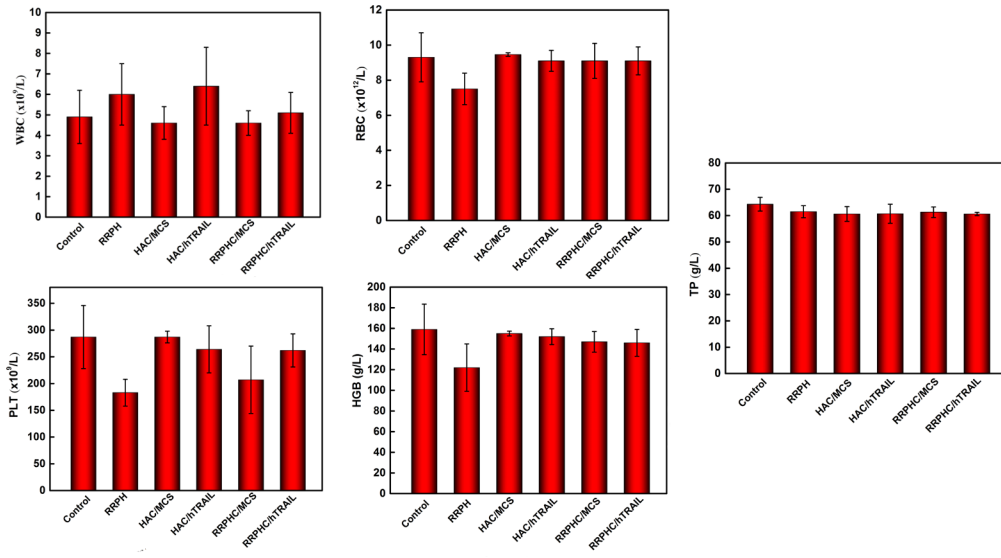
**Fig. S2.** Determination the expression level of CD44 receptor on the surface of HCT 116 cells by flow cytometry. Isotype control (purple).



**Fig. S3.** *In vitro* apoptosis-inducing effect in HCT 116 cells after treatment with PF<sub>33</sub>, RRPH, HA and RRPH materials at 24 h. \* $p < 0.05$ , \*\* $p < 0.01$  and \*\*\* $p < 0.001$ .



**Fig. S4.** Safety and toxicity evaluation. (A) Serum AST, ALT and ALB levels of mice after treatment with different formulations. (B) Histological examination of H&E staining of vital organ sections. PBS (a), RRPH (b), HAC/MCS (c), HAC/hTRAIL (d), RRPHC/MCS (e) and RRPHC/hTRAIL (f).



**Fig. S5.** Complete blood count (CBC) test after treatment with different formulations.