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Double-sided effect of tumor microenvironment on platelets targeting nanoparticles

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## ACCEPTED MANUSCRIPT

- 1 Double-sided effect of tumor microenvironment on platelets targeting nanoparticles
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15 Hypopermeable; Photothermal therapy

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17 Abstract: The cancer cells and stromal cells in tumor microenvironment secrete cytokines and 18 recruit "homing" cells (macrophage, lymphocytes, platelets, etc.). Platelets can interact with tumor 19 microenvironment and specifically aggregate at tumor sites. Surprising, we observed different 20 "homing" effects of activated platelets in breast cancer model and pancreatic cancer model which 21 is highly related with the blood supply of tumors. Besides, platelets targeting magnetic 22 nanoparticles (MNPs) can home to breast cancer but be repelled from pancreatic cancer. We 23 observed the targeting effect of MNPs is highly related with the expressions of collagen  $\Box$  (marker 24 of extracellular matrix) and CD34 (marker of tumor neovascularization). The homing 25 nanoparticles such as platelets targeting MNPs could realize the tumor targeting ability, 26 photo-thermal effect and tumor immunotherapeutic ability in the accessible tumor (e.g. breast 27 cancer) but not the hypovascular tumor (e.g. pancreatic cancer).

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