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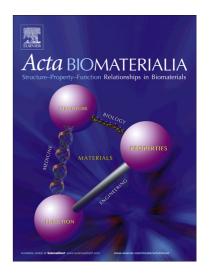
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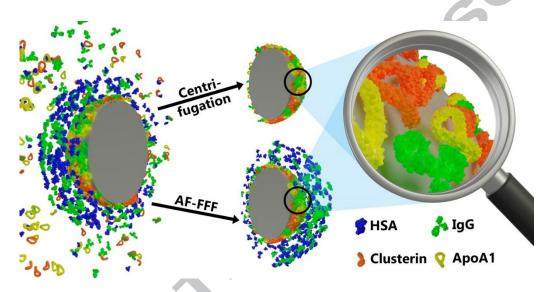
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ACCEPTED MANUSCRIPT

Preservation of the soft protein corona in distinct flow allows identification of weakly bound proteins

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Abstract

Nanocarriers that are used for targeted drug delivery come in contact with biological liquid and proteins will adsorb to the nanocarriers' surface to form the so called 'protein corona'. The protein corona defines the biological identity and determines the biological response towards the nanocarriers in the body. To make nanomedicine safe and reliable it is required to get a better insight into this protein corona and, therefore, the adsorbed proteins have to be characterized. Currently, centrifugation is the common method to isolate the protein corona for further investigations. However, with this method it is only possible to investigate the strongly bound proteins, also referred to as 'hard protein corona'. Therefore, we want to introduce a new separation technique to separate nanoparticles including the soft protein corona containing also loosely bound proteins for further characterizations. The used separation technique is the asymmetric flow filed-flow fractionation (AF4). We were able to separate the nanoparticles with proteins forming the soft protein corona and were able to show that only the hard protein corona directly influences the cell uptake behavior.

Keywords

soft and hard protein corona; nanocarriers; asymmetric flow field-flow fractionation; biological identity

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