Electronic Supplementary Information (ESI)

Controlling Nanoparticle-Induced Endothelial Leakiness with The Protein Corona

Aparna Nandakumar,^a Huayuan Tang,^{b,c} Nicholas Andrikopoulos,^{a,d} John F. Quinn,^a Feng

Ding, * ^c Pu Chun Ke^{* a,d} and Yuhuan Li^{* a,e}

 ^a Drug Delivery, Disposition and Dynamics, Monash Institute of Pharmaceutical Sciences, Monash University, 381 Royal Parade, Parkville, VIC 3052, Australia

^b Department of Engineering Mechanics, Hohai University, Nanjing 211100, China

^c Department of Physics and Astronomy, Clemson University, Clemson, SC 29634, United States

^d Nanomedicine Centre, The Great Bay Area National Institute for Nanotechnology Innovation, 136 Kaiyuan Avenue, Guangzhou, 510700, China

^e Liver Cancer Institute, Zhongshan Hospital, Key Laboratory of Carcinogenesis and Cancer Invasion, Ministry of Education, Fudan University, Shanghai, 200032, China

Corresponding Authors

Feng Ding, fding@clemson.edu; Pu Chun Ke, pu-chun.ke@monash.edu; Yuhuan Li,

li.yuhuan1@zs-hospital.sh.cn



Figure S1. Hydrodynamic size of citrate-capped AuNPs.



Figure S2. Primary size distributions of the AuNPs and their corresponding protein coronas. A total of 50 individual particles from TEM images were analyzed using ImageJ and the histogram was mapped in Minitab statistical package.



Figure S3. (A) Absorbance measurement at 562 nm for different AuNP/protein coronas. The mass (B) and molar amount (C) of the proteins bound to AuNPs (1 mg) were determined from the BCA protein assay interpolating the BSA standards.



Figure S4. SDS-PAGE confirmed the presence of coronal proteins for the fixed mass ratio of 1:5 AuNP/protein used for the cell culture experiments.



Figure S5. Mixtures of the AuNPs and the four types of plasma proteins at 0 h and 24 h incubation under constant shaking at 37 °C.



Figure S6. Transwell assay revealed no leakiness in the HUVEC monolayers due to the introduction of the plasma proteins alone.



Figure S7. Steady-state graph of different concentrations of Lyz interacting with VE-cadherin coated on NTA sensors examined with the BLI assay.



Figure S8. Schematic illustration of the binding of multiple proteins with AuNPs.

Table S1. Amino acid sequence,	PDB	structure	and	net	charge	of the	plasma	proteins
used in binding analysis.								

Protein	Molecular weight (kDa)	Sequence from UniProt	PDB for docking	Net charge
Alb	66.5	P02768	1AO6.pdb	-11
Fib	340	Alpha chain: P02671 Beta chain: P02675 Gamma chain: P02679	3GHG.pdb	-10
IgG	150	Immunoglobulin heavy constant gamma 2: P01859 Immunoglobulin heavy variable: P01814 Immunoglobulin light chain: P0DOX8	1HZH.pdb	0
Lyz	14.3	P61626	1REX.pdb	+9

 Table S2. Physiochemical characteristics of the AuNP-protein coronas.

Sample	Peak absorbance (nm)		Primary size (nm)	ζ – potential (mV)		
	0 h	24 h	24 h	W/o wash	With wash	
AuNP	519	524	15.0±0.7	-28.3±0.2	-27.3±1.6	
AuNP-Alb	527	551	17.5±1.3	-0.6±0.3	-10.2±0.5	
AuNP-Fib	541	542	16.5±1.2	-6.6±0.4	-8.6±0.9	
AuNP-IgG	527	535	17.3±1.2	-4.0±0.5	-6. 7±0.7	
AuNP-Lyz	558	570	17.2±1.0	3.0±0.4	-6.7±0.7	