Near-Infrared SERS Nanoprobes with Plasmonic Au/Ag Hollow-Shell Assemblies for In Vivo Multiplex Detection

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Nanostructures of noble metal such as gold and silver exhibit an optical phenomenon known as surface-enhanced Raman scattering (SERS), which enhances Raman scattering of molecules adsorbed on the surface of them. When SERS is used as a coding method, it has advantages for bioassays over other optical tools: A narrow spectral band of SERS peaks (less than 0.5 nm), which is able to make a large number of coded, flexible selectivity of photoexcitation line covering UV to NIR region, and no-photobleaching feature. [1–5]. Owing to these advantages, SERS active bio-probe can be utilized for multiplexed detection of numerous bio-targets. Nevertheless, SERS nanoprobes still need to be improved in sensitivity and tissue penetration ability for *in vivo* SERS application. In this regard, novel metal nanoshells have been extensively studied to prepare the SERS active nanoparticles because of their high surface area and unique optical properties. In particular, hollow Au/Ag nanostructures have tunable extinction band from visible to near infrared (NIR) region (*ca.* 400-1200 nm), exhibiting both surface plasmon resonance and scattering [6-8].

In this study, we introduce the preparation of plasmonic Au/Ag hollow-shell assemblies (Au/AgHSA) as high sensitive SERS active nanoprobes in NIR region. The Au/Ag nanoshells were formed on the surface of silica sphere from silver nanoparticles embedded silica sphere via galvanic replacement reaction. Raman reporter molecules were then absorbed on the Au/Ag surface, and subsequently encapsulated with silica shell to prevent dissociation of label compounds and aggregation of probes. The Au/Ag-HSA exhibited highly sensitive signals due to their localized surface electromagnetic field and the assembled multiple Au/Ag nanoshells on silica sphere. Further it shows extinction (absorption/scattering) in NIR region, the transparent window of biological tissue. These results show that Au/AgHSA can be applied to *in vivo* optical imaging as a SERS nanoprobe.

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