## Obviously Enhanced Fluorescent Signal of Core-Shell Nanostructures through Simultaneous Regulation of Spectral Overlap and Shell Thickness for Imaging and Photothermal Therapy of Ovarian Cancer Cells

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In this research, by simultaneously regulating the two major factors affecting the plasmonic enhanced fluorescence (PEF), spectral overlap and the distance between the fluororophores and the noble metal nanoparticles, a significantly enhanced fluorescent signal is achieved. Core-shell nanostructures composed of aspect ratio (AR) adjustable gold nanorods (GNRs) and various thickness of  $SiO_2$  are prepared and the decorated fluorophores are realized optimized PEF. A typical stimuli-responsive conjugated polymer, polydiacetylene (PDA), and a near-infrared (NIR) dye Cy5.5 are selected as fluorophores and their fluorescent signal are enhanced 7.26 and 4.41 times, respectively. Based on the optimized optical properties, a multifunctional antibody modified Mab-Cy5.5-GNRs@SiO<sub>2</sub> is successfully demonstrated the targeting, imaging, and photothermal therapy (PTT) effects on SKOV-3 ovarian cancer cells.

## 1. Introduction

Noble metal nanoparticles and nano-rough films possess localized surface plasmon resonances (LSPRs) that imbue these materials with unique optical properties.<sup>[1-3]</sup></sup> Especially, owing to

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tunable absorption bands based on their aspect ratios (ARs), excellent photothermal conversion efficiency, biocompatibility and low toxicity, gold nanorods (GNRs) are considered promising photothermal agents in photothermal therapy (PTT).<sup>[4-6]</sup> Additionally, elongated nanostructures penetrate tumor cells more rapidly due to excellent transmembrane transport and diffusion rates than spherical nanoparticles (NPs).<sup>[7-9]</sup> Based on these factors, various nanostructures containing GNRs have recently been vigorously developed in biomedicine.<sup>[10-14]</sup> Precise and effective PTT need to be supplemented by a clear imaging guidance.<sup>[15]</sup> The GNR itself has imaging capabilities, namely photoacoustic

imaging.<sup>[16,17]</sup> However, due to the principle of imaging, photoacoustic imaging has two main fatal disadvantages. First, the heat energy caused by constant light irradiation can lead to the deformation of GNR and a sharp decrease in the imaging and PTT effects.<sup>[18]</sup> Second, photoacoustic imaging is only suitable for a dense tissue environment.<sup>[19]</sup> For the above reasons, the combination of traditional luminescent materials to the protected GNR is still recommended.<sup>[12,20,21]</sup> In these cases, another effect based on LSPR, plasmon-enhanced fluorescence (PEF) has been extensively studied by scholars. The enhanced fluorescent signal could be contributed to the bio-imaging of the GNR-containing nanostructures by being decorated with various fluorophores or quantum dots.<sup>[22–24]</sup>

From the perspective of the imaging function, due to PEF could increase the quantum yield and improve the photo-stability of fluorophores, hence it ensures imaging effects in complex biological environments.<sup>[25–27]</sup> In general, spectral overlap and distance between fluorophores and plasmon resonant novel metal NPs are considered the main factors of the PEF.<sup>[28–31]</sup> However, until now, few studies have simultaneously regulated both factors and achieved an optimized PEF effect.<sup>[32–36]</sup>

In this research, we developed GNR-containing nanostructures decorated with fluorophores, a stimuli-responsive polydiacetylene (PDA), and a near-infrared (NIR) Cy5.5 dye, respectively. Optimized fluorescent signals were achieved through simultaneous regulation of spectral overlap and distance between