

## Salt-assisted biofunctionalization protocol for the ligand exchange of CTAC-capped gold nanocubes

Nicole Slesiona, Sophie Thamm, Lisa Stolle, Andrea Csáki, Wolfgang Fritzsche

Leibniz Institute of Photonic Technology, Albert-Einstein-Str. 9, 07745 Jena, Germany

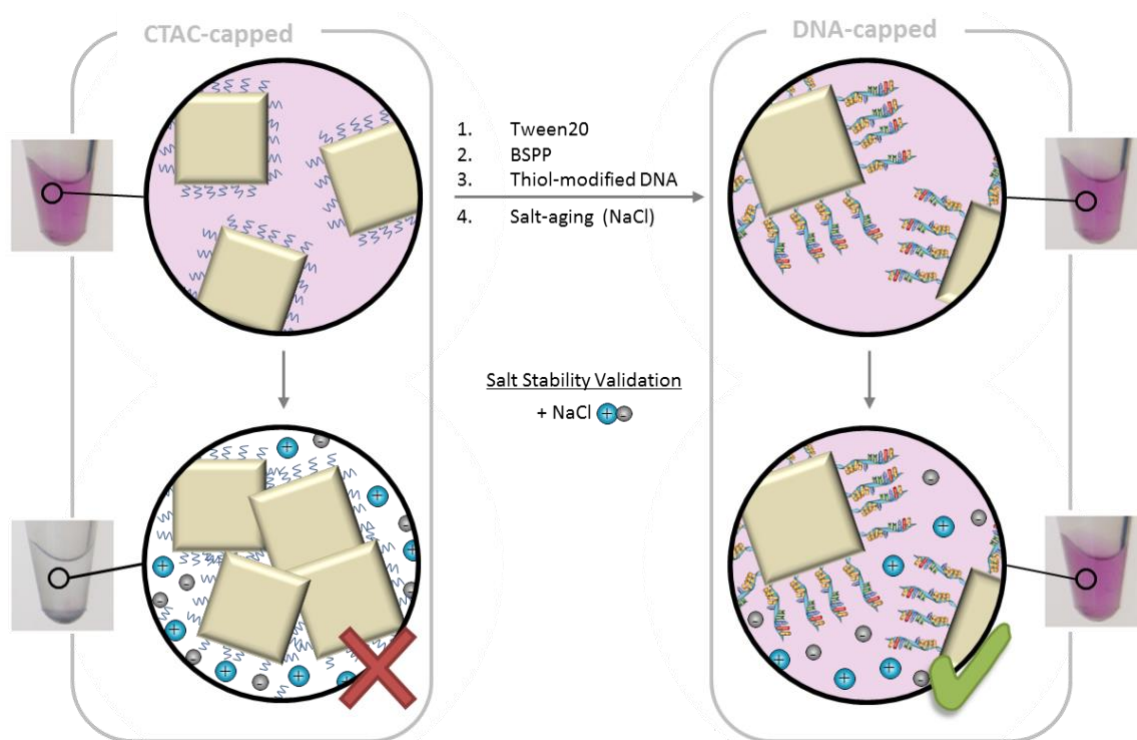
Due to their strong interaction with light in the visible/IR spectrum and the strong field enhancements that can be found at edges and corners of shape-anisotropic gold nanoparticles<sup>1</sup>, these nanostructures have become an integral part in various biomedical detection devices and nanotherapeutics<sup>2</sup>. The high biocompatibility of gold nanoparticles especially facilitates their incorporation into biosensor designs that exhibit enhanced analytical performance.<sup>2</sup>

Shape-anisotropic gold nanoparticles are synthesized with the help of facet blocking agents, such as CTAB/CTAC (cetyltrimethylammonium bromide/chloride) that selectively adsorb on particular crystallographic planes.<sup>3, 4, 5</sup> Oftentimes, biocompatible groups such as DNA cannot sufficiently penetrate through a dense layer of capping agents, preventing their transfer into existing protocols of biologically relevant applications. In order to merge shape-anisotropic gold nanoparticles into analytic sensing devices, it is necessary to establish a reliable modification protocol to reduce their cytotoxicity, stabilize them against aggregation, and to functionalize them with other biomolecules to increase their biocompatibility for analytical applications.

Here, a two-step protocol is presented that combines the approaches of

1. ligand exchange with BSPP (bis(p-sulfonatophenyl) phenylphosphine) as the intermediate capping-agent, and
2. electrostatic interaction with a salt-aging step.

This protocol allows for a reliable and efficient conjugation of thiol-modified DNA strands to anisotropic particles with great potential for biosensing applications and nanotherapeutics. The combination with DNA Nanotechnology allows for the self-assembly of the particles directed by a DNA origami scaffold.



- (1) Agrawal, A.; Kriegel, I.; Milliron, D. J., Shape-Dependent Field Enhancement and Plasmon Resonance of Oxide Nanocrystals. *The Journal of Physical Chemistry C* **2015**, *119*, 6227-6238.
- (2) Li, Y.; Schluesener, H. J.; Xu, S., Gold nanoparticle-based biosensors. *Gold Bulletin* **2010**, *43*, 29-41.
- (3) Scarabelli, L.; Sánchez-Iglesias, A.; Pérez-Juste, J.; Liz-Marzán, L. M., A “Tips and Tricks” Practical Guide to the Synthesis of Gold Nanorods. *The Journal of Physical Chemistry Letters* **2015**, *6*, 4270-4279.
- (4) Wu, H.-L.; Kuo, C.-H.; Huang, M. H., Seed-Mediated Synthesis of Gold Nanocrystals with Systematic Shape Evolution from Cubic to Trisoctahedral and Rhombic Dodecahedral Structures. *Langmuir* **2010**, *26*, 12307-12313.
- (5) Thiele, M.; Knauer, A.; Malsch, D.; Csáki, A.; Henkel, T.; Köhler, J. M.; Fritzsche, W., Combination of microfluidic high-throughput production and parameter screening for efficient shaping of gold nanocubes using Dean-flow mixing. *Lab on a Chip* **2017**, *17*, 1487-1495.