## SERS Microscopy: Tissue Diagnostics with Rationally Designed Nanoparticle Probes

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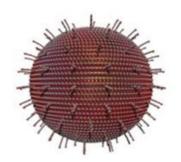
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Surface-enhanced Raman scattering microscopy (µSERS) [1] employs target-specific colloidal SERS probes in combination with Raman microspectroscopy. For example, SERS-labeled antibodies allow the selective and sensitive localization of the corresponding antigen in tissue specimens [2].

The physical and chemical properties of the colloidal SERS probes are crucial for the success of SERS microscopic experiments [3]. Stability and robustness, sensitivity as well as steric accessibility for bioconjugation are few very important aspects. Figure 1 left shows a hydrophilic SERS probe stabilized by many short and few longer hydrophilic spacer units [4]. A second approach, based on the glass encapsulation for stabilizing SERS labels, is schematically depicted in Figure 1 right. This type of SERS probes was optimized for red laser excitation in order to improve image contrast by minimizing unwanted autofluorescence of biological specimens [5]. Two different routes to the glass encapsulation of a complete monolayer of Raman reporters will be described [5-6]. Small clusters (dimers, trimers) of gold nanocrystals even exhibit single-particle sensitivity and enable rapid mapping experiments with only 30 msec acquisition time per pixel [7]. Future directions of this innovative Raman/SERS microspectrosopic technique for tissue-based tumor diagnostics will be discussed.

## References

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**FIGURE 1.** (Left) SERS labels [3] stabilized by many short and few longer hydrophilic spacer molecules for controlled bioconjugation [4]. (Right) SERS nanoprobes optimized for red laser excitation for minimizing tissue autofluorescence. Gold/silver nanoshells are covered by a complete self-assembled monolayer (SAM) of Raman labels. The SERS particle is protected and stabilized by a glass shell [5-6].