## Supporting information for

## Probing nucleus-enriched proteins in single living cells via subcellular-resolved plasmonic immunosandwich assay

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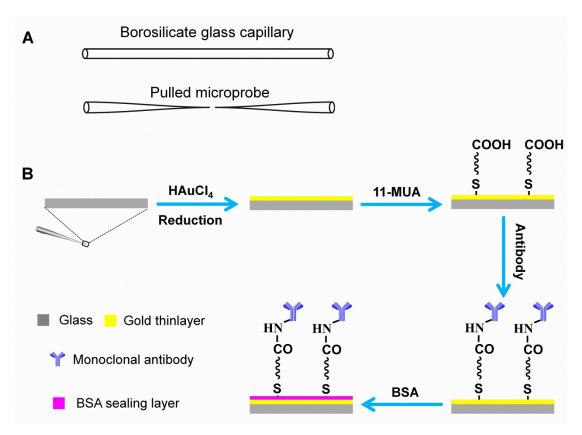
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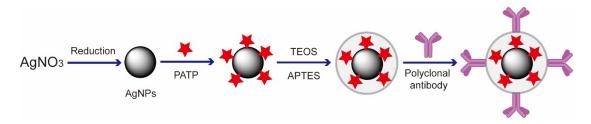
Supplementary Data

Fig. S1 to Fig. S5

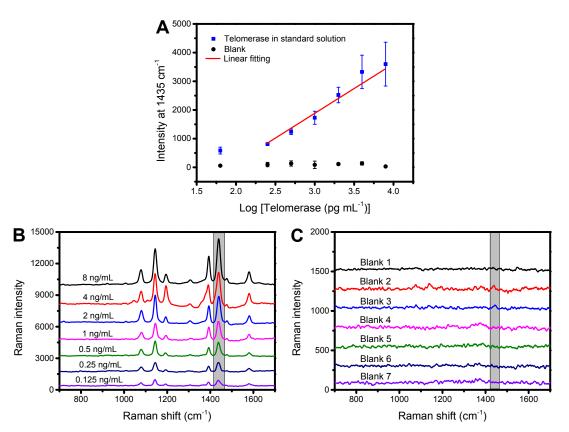
## **SUPPLEMENTARYDATA**



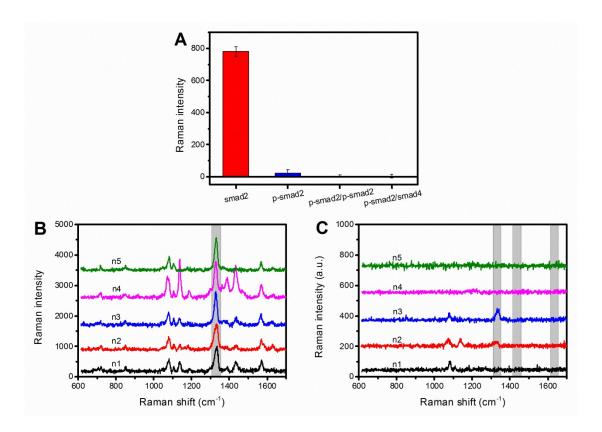
**Fig. S1** Schematic of the procedures for fabrication of extraction microprobes. (A) Preparation of pulled microprobe by a laser-based pipette puller. (B) Fabrication of monoclonal antibody-functionalized gold-based extraction microprobes.



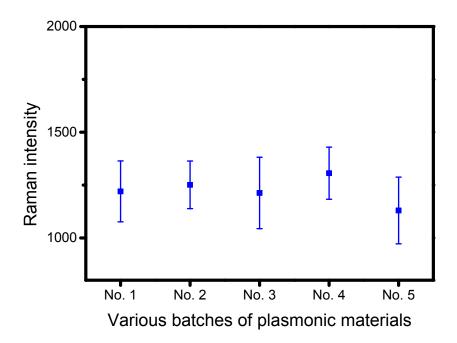
**Fig. S2** The synthesis route of polyclonal antibody-functionalized silver-based plasmonic nanotags.



**Fig. S3** Dependence of the Raman intensity at 1435 cm<sup>-1</sup> detected by srPISA with telomerase standard solution. Blank sample: 100 mM phosphate buffer, pH 7.4. Error bars represent standard deviations for three parallel measurements. (B, C) Representative Raman spectra for telomerase detection in standard solutions and blank samples, respectively.



**Fig. S4** Selectivity test of the srPISA approach in single living MCF-7 cells. (A) Raman intensity of target smad2 protein and its similar proteins, including monomeric phospho-smad2 (p-smad2), phospho-smad2 homomeric complexes (p-smad2/p-smad2) and phospho-smad2/smad4 heteromeric complexes (p-smad2/smad4). (B, C) Representative Raman spectra of selectivity test of the srPISA in single living MCF-7 cells for target smad2 protein and its similar proteins.



**Fig. S5** Raman intensity variation for various batches of plasmonic materials for detecting telomerase in standard solution. Sample concentration: 500 pg/mL. Error bars represent standard deviations for three parallel measurements.