**Supporting Information for** 

## Non-invasive and Highly Selective Monitoring of Intracellular Glucose *via* Two-steps Recognition-based Nanokit

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## **EXPERIMENTAL SECTION**

**Instruments**. Transmission electron microscopy images obtained on JEM-100CXII microscope (JEOL, Ltd., Japan). UV-Vis absorption spectra were recorded on a Hitachi U-4100 UV/Vis spectrophotometer (Kyoto, Japan). Fluorescence images of cells were obtained using an Olympus FV1000-MPE multiphoton laser scanning confocal microscope (Japan). The pH was measured with a model 868 pH meter (Orion).

Entry	Sequence (5'-3')
Capture sequence Aptamer sequence	TGT CGT CCC GAG AGT TTT TTT-SH TAMRA-CTC TCG GGA CGA CAG CCG AGT TGA TTC AAC AGC CGA GTC GTC CC

## Table S1. Oligonucleotides Used in This Work\*

-	Sample	Added (µM)	Measured (µM)	Recovery (%)
_	1	0	0.27±0.3	
	2	50	51±1.4	101
	3	100	98±2.6	98

**Table S2.** Determination of Glucose in Cell Lysate

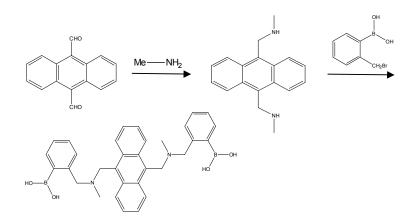


Figure S1. The synthetic route of the Shinkai's receptor.

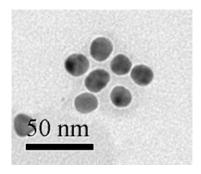
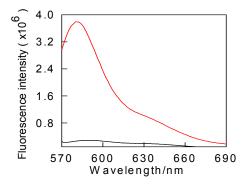


Figure S2. Representative TEM image of AuNPs.



**Figure S3**. Fluorescence spectrum of TAMRA-labeled aptamer before (red curve) and after (black curve) conjugation to the surface of AuNPs.

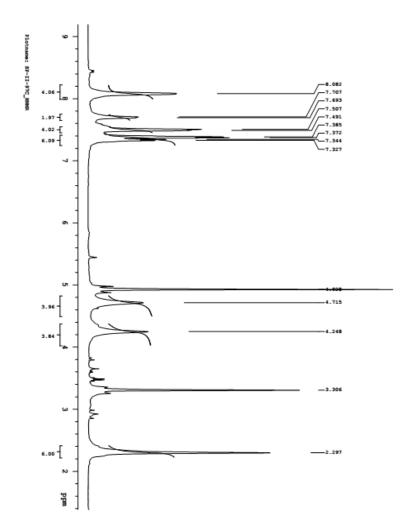
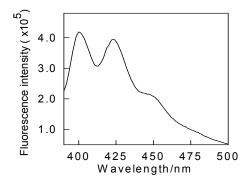
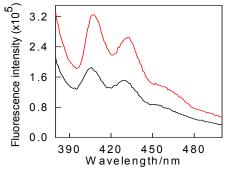


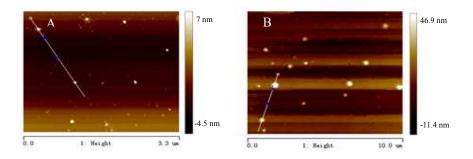
Figure S4. 1H NMR of the Shinkai's receptor.



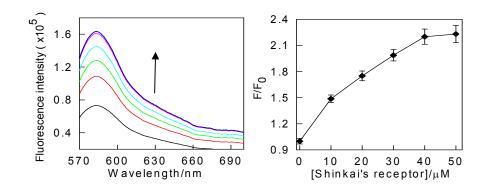
**Figure S5.** Fluorescence spectrum of the Shinkai's receptor. The excitation wavelength was 370 nm.



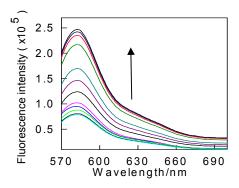
**Figure S6.** Fluorescence spectrum of the Shinkai's receptor-encapsulated liposomes before (black curve) and after (red curve) Triton X-100 treated. The excitation wavelength was 370 nm.



**Figure S7.** Representative AFM images of AuNPs (A) the synthetic Shinkai's receptor-encapsulated liposome (B).



**Figure S8.** The fluorescence recovery of AuNP@ODs upon 10 mM glucose addition with aid of different concentrations of the Shinkai's receptor.



**Figure S9.** Representative fluorescence spectrum of AuNP@ODs (10 nM) as functions of different concentrations of glucose with aid of the Shinkai's receptor (50  $\mu$ M). The arrows indicate the concentration from 0 to 2 mM (0, 0.1, 0.5, 1, 10, 100, 250, 500, 1000, 1500, 1750, 2000  $\mu$ M).

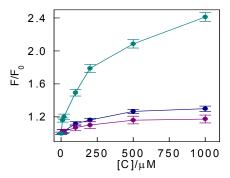


Figure S10. The fluorescence recovery of AuNP@ODs (10 nM) upon different concentration of glucose, galactose and fructose addition with aid of Shinkai's receptor (50  $\mu$ M).

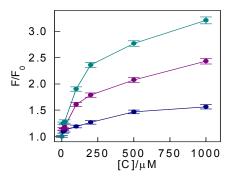
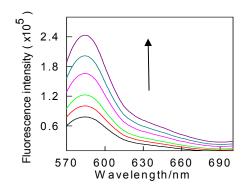
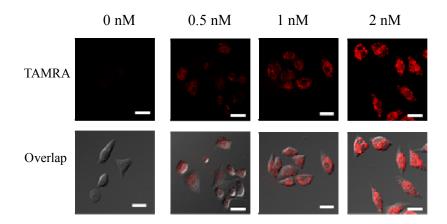


Figure S11. The fluorescence recovery of the Shinkai's receptor (50  $\mu$ M) upon different concentration of glucose, fructose and galactose.



**Figure S12.** Fluorescence spectra of nanokit upon addition of different concentrations of glucose after treated by Triton-100. The arrows indicated the concentration of glucose: 0, 10,100, 500, 1000, 2000  $\mu$ M.



**Figure S13**. Confocal microscopy images recorded at TAMRA channel of the nanokit-treated HeLa cells as a function of different concentrations of nanokit.

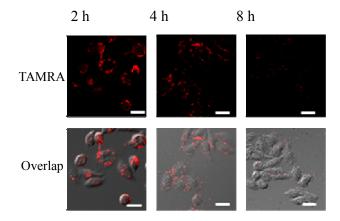


Figure S14. Intracellular glucose imaging of HeLa cells under 10% O<sub>2</sub> as functions of time. The concentration of the nanokit was 2 nM.