SUPPORTING INFORMATION

Ni@4H-Chromene based Core-Shell Nanoparticles: Highly Sensitive and Selective Chemosensor for Radiosensitizer - Bromodeoxyuridine

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Table S1 Time resolved fluorescence data of receptor G1, ONPs, G2 and Ni@G1.

Table S2 Comparison table of present work with other analytical techniques.



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Fig. S18 FT-IR spectrum of ONP, Ni@G1, Ni@G1+BrdU.



Fig. S19 A) Cyclic voltammogram showing changes in cathodic and anodic peak when Ni(II) is added to ONPs. B) Cyclic voltammogram when $NaBH_4$ is added to the G2 complex formed.

S.No.	Compound	A ₁	$ au_{ m AV}$
1	Receptor G1	5339	3.99
2	ONPs	49075	0.23
3	G2	906.5	4.73
4	Ni@G1	3749.6	11.92

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S.No	Method used for	Real sample	Limitations	Reference
•	Detection	application		S
1	Flow cytometry	Rat blood serum	 Too slow Expensive Requires trained professional Complex instrumentatio n 	1
2	Immunocytochemic al detection	MCF-7 cells	The technique hassome limitations:• Stains are not	2

			 worldwide available Expensive instrument Result quantification is difficult
3	Flow cytometry	Chinese hamster ovary cells	 Too slow ³ Expensive Requires trained professional Complex instrumentatio n
4	Flow cytometry	 Keratinocytes Bone marrow cells 	 Too slow Expensive Requires trained professional Complex instrumentatio n
5	Flow cytometry	Cell lines including: • Chinese hamster embryo cells • Human skin fibroblasts • Friend erythroleukemi a cells	 Too slow Expensive Requires trained professional Complex instrumentatio n

		Human lymphocytes		
6	Immunocytochemic al detection	Serum samples of:CanariesQuailMice	The technique has some limitations: • Stains are not worldwide available • Expensive instrument • Result quantification is difficult	6
7	High performance liquid chromatography (HPLC)	Human serum	Thetechniquehassomeimitations:•Costly•Tedious•Timeconsuming•Complex	7
8	Fluorescencespectroscopy,UV-Visible-spectroscopy,andCyclic-voltammetry-	Human serum albumin		Present work

 Table S2 Comparison table of present work with other analytical techniques.

References

- 1 Z. Qiu, J. Shu and D. Tang, Anal. Chem., 2017, 89, 5152–5160.
- 2 F. Dolbeare, H. Gratzner, M. G. Pallavicini and J. W. Gray, *Proc. Natl. Acad. Sci. U. S. A.*, 1983, **80**, 5573–5577.

- J. Van Heusden, P. De Jong, F. Ramaekers, H. Bruwiere, M. Borgers and G. Smets, J. *Histochem. Cytochem.*, 1997, **45**, 315–319.
- 4 P. E. J. Van Erp, P. P. T. Brons, J. B. M. Boezeman, G. J. De Jongh and F. W. Bauer, *Cytometry*, 1988, **9**, 627–630.
- 5 J. A. Steinkamp, 1987, 0–5.
- 6 J. M. Barker, T. D. Charlier, G. F. Ball and J. Balthazart, *PLoS One*, 2013, **8**, 1–5.
- 7 T. J. Kinsella, J. B. Mitchell, A. Russo, M. Aiken, G. Morstyn, S. M. Hsu, J. Rowland and E. Glatstein, *J. Clin. Oncol.*, 1984, **2**, 1144–1150.