

Supporting Information

A water-soluble nanoconjugate for enhanced cellular
delivery
of receptor-targeted MR contrast agents

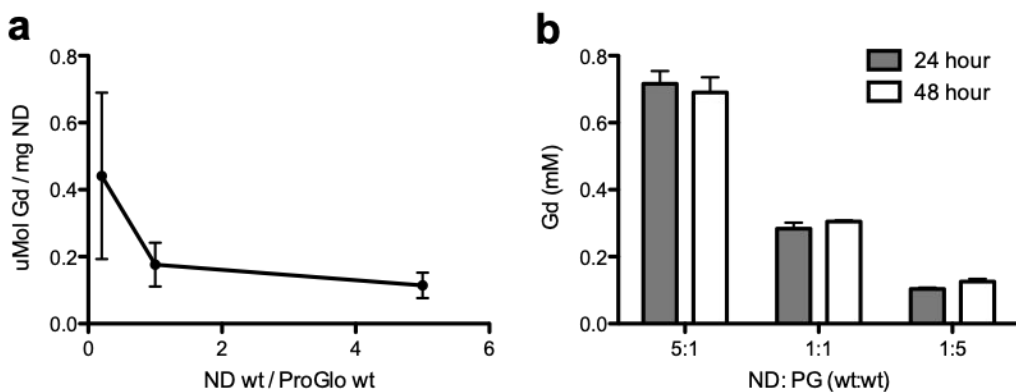
*Laura K. Moore,^{1†} Michael A. Caldwell,^{2†} Taryn R. Townsend,² Keith
W. MacRenaris,² Georgette Moyle-Heyrman,⁴ Nikhil Rammohan,² Erika
K. Schonher,² Joanna E. Burdette,⁴ Dean Ho,^{5‡} Thomas J. Meade^{1,2‡}*

1 Department of Biomedical Engineering, Feinberg School of
Medicine, Northwestern University, Chicago, IL 60611

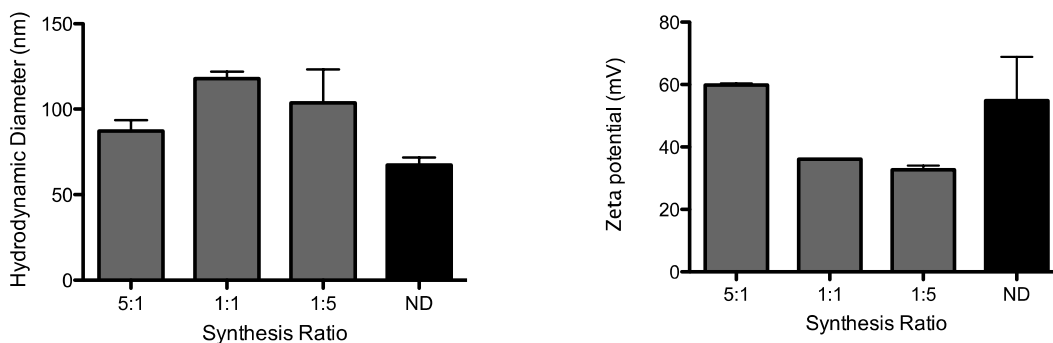
2 Departments of Chemistry, Molecular Biosciences, Neurobiology,
and Radiology, Northwestern University, Evanston, IL, 60208

4 Department of Medicinal Chemistry and Pharmacognosy,
University of Illinois at Chicago, Chicago, IL, 60607

5 The N.1 Institute for Health (N.1) National University of
Singapore, Singapore 117556
Department of Biomedical Engineering: NUS Engineering
National University of Singapore, Singapore, 117583



S1. A 24 hr coincubation at a 5:1 weight to weight ratio of ND to ProGlo produced the highest total loading (**b**) and loading per mg ND (**a**) of ProGlo into ND-ProGlo. The concentration of ProGlo utilized for loading was approximately 1.8 mM and the incubation concentration of ND was reduced from 8.3 mg/mL to 0.3 mg/mL across the series.

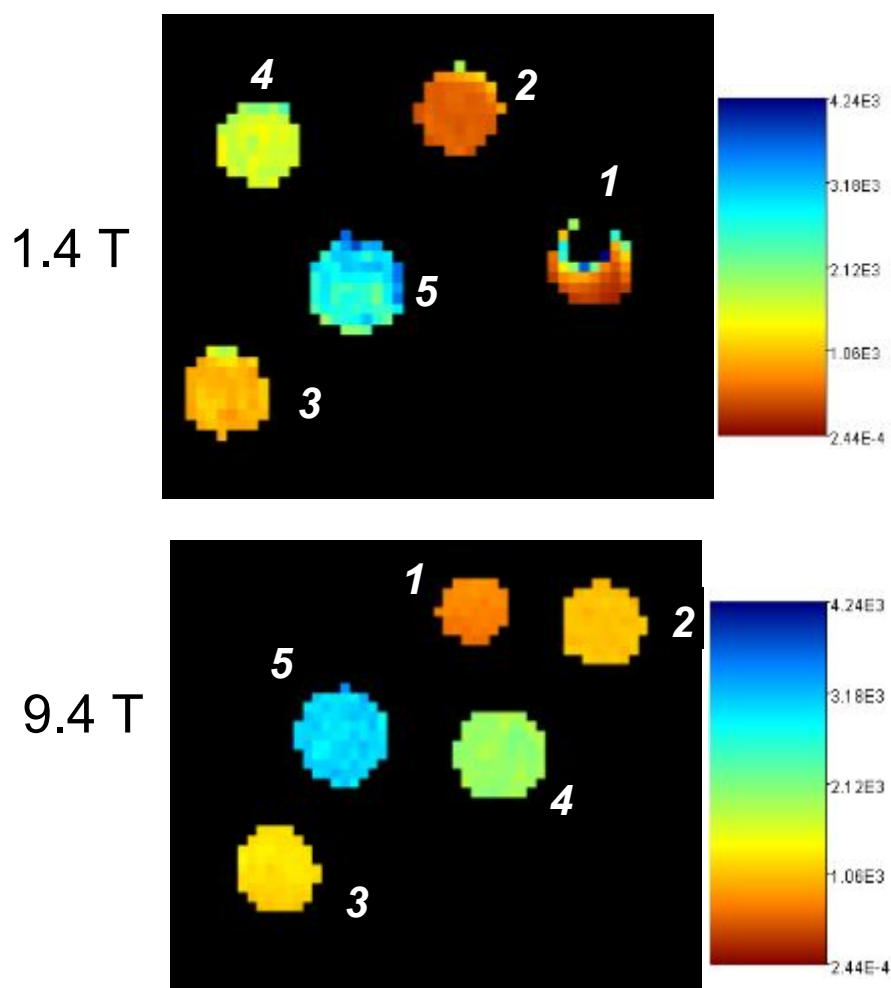


S2. a. The hydrodynamic diameter of ND-ProGlo aggregates synthesized by varying the wt:wt ratio of ND to ProGlo (n=3). **b.** Zeta-potential of ND-ProGlo synthesized at varying ratios (n=3). Hydrodynamic diameters and zeta-potentials were measured for native NDs and ND-ProGlo modified with varying ratios of ND to ProGlo. Addition of ProGlo produced an increase in aggregate size that varied little with increasing ProGlo loading. Zeta potentials showed a decrease with increasing stoichiometric ratios (and loading) of ProGlo.

Agent	r_1^a	r_2^a	r_2/r_1	LogP
ProGlo	5.4 ± 0.7	6.1 ± 0.8	1.15	1.40 ± 0.08
ND- ProGlo	22.7 ± 1.5	37.9 ± 2.3	1.67	-2.40 ± 0.26

^a r_1 and r_2 measured in $\text{mM}^{-1}\text{s}^{-1}$ at 1.41 T, 37 °C

S3. Supplementary Table 1. Relaxivities of ProGlo and ND-ProGlo were measured at 1.41 T in a Bruker bench-top relaxometer in 1% DMSO in water and pure water, respectively. ND-ProGlo showed a significant increase in relaxivity over molecular ProGlo, as expected from the increase in τ_r associated with immobilization on a particle. The logP values for of ProGlo and ND-ProGlo reflect the change from hydrophobic to hydrophilic character upon loading of ProGlo into ND clusters.



S4. MR phantom images of ND-ProGlo solutions: 1.4 T (top) and 9.4 T (bottom). The tubes containing the samples are: **(1)** ProGlo-ND [Gd(III) = 0.14 mM, 5 mg ND], **(2)** ProGlo-ND [Gd(III) = 0.07 mM, 2.5 mg ND] **(3)** ProGlo [Gd(III) = 0.10 mM, in 1% DMSO], **(4)** ND-COOH (5 mg ND), and **(5)** water. The images are consistent with the observed relaxivities, where ProGlo-ND solutions show significantly faster T_1 times than ProGlo in phantom images. NDs alone have only a small effect on observed T_1 s.