

## Supporting Information

### Self-discriminating Fluorescent Hybrid Nanocrystals: Efficient and Accurate Tracking of Translocation via Oral Delivery

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Table S1. Physicochemical properties of QT-HNCs (mean  $\pm$  SD, n=3)

| Formulation     | Particle size<br>(nm) | PDI             | Zeta<br>potential<br>(mV) | TRE[p/s]/<br>[ $\mu$ W/cm <sup>2</sup> ] $\times$ 10 <sup>9</sup> | QT content<br>(mg/mL) | EE<br>(%)        | Loading<br>(%)    |
|-----------------|-----------------------|-----------------|---------------------------|---|-----------------------|------------------|-------------------|
| P2/QT-HNCs-280  | 274.23 $\pm$ 5.80     | 0.13 $\pm$ 0.03 | -20.3 $\pm$ 3.1           | 5.79 $\pm$ 0.09   | 18.46 $\pm$ 0.49      | 59.13 $\pm$ 0.97 | 0.13 $\pm$ 0.0029 |
| P2/QT-HNCs-550  | 541.20 $\pm$ 44.11    | 0.17 $\pm$ 0.01 | -21.4 $\pm$ 2.7           | 5.21 $\pm$ 0.25   | 18.00 $\pm$ 0.51      | 60.61 $\pm$ 0.23 | 0.13 $\pm$ 0.0033 |
| P2/QT-HNCs-1100 | 1163.00 $\pm$ 134.36  | 0.20 $\pm$ 0.03 | -23.2 $\pm$ 3.2           | 5.11 $\pm$ 0.15   | 17.90 $\pm$ 0.38      | 66.69 $\pm$ 1.30 | 0.15 $\pm$ 0.0009 |
| P4/QT-HNCs-280  | 283.41 $\pm$ 4.12     | 0.17 $\pm$ 0.02 | -24.1 $\pm$ 1.9           | 22.87 $\pm$ 0.24  | 18.06 $\pm$ 0.10      | 97.75 $\pm$ 1.04 | 0.22 $\pm$ 0.0020 |
| P4/QT-HNCs-550  | 558.14 $\pm$ 38.24    | 0.15 $\pm$ 0.04 | -22.8 $\pm$ 2.8           | 21.94 $\pm$ 0.08  | 17.68 $\pm$ 0.15      | 98.62 $\pm$ 0.63 | 0.22 $\pm$ 0.0033 |
| P4/QT-HNCs-1100 | 1093.52 $\pm$ 108.32  | 0.19 $\pm$ 0.03 | -23.7 $\pm$ 2.2           | 22.26 $\pm$ 0.21  | 18.45 $\pm$ 0.68      | 97.15 $\pm$ 0.88 | 0.2 $\pm$ 0.0086  |

Table S2. The composition of SGF, SIF, FaSSIF and FeSSIF.

| Component  | SGF     | SIF     | FaSSIF  | FeSSIF   |
|--|---------|---------|---------|----------|
| Sodium taurocholate                                | -       | -       | 6 mM    | 15 mM    |
| Lecithin   | -       | -       | 1.5 mM  | 3.75 mM  |
| Pepsin   | 3.2 g   | -       | -       | -        |
| Trypsin  | -       | 10 g    | -       | -        |
| NaH <sub>2</sub> PO <sub>4</sub> ·H <sub>2</sub> O | -       | 7.7 g   | 3.954 g | -        |
| NaOH   | -       | q.s.    | 0.348 g | 4.04 g   |
| NaCl   | 2.0 g   | -       | 6.186 g | 11.874 g |
| CH <sub>3</sub> COOH                               | -       | -       | -       | 8.65 g   |
| HCl  | 7.0 mL  | -       | -       | -        |
| H <sub>2</sub> O                                   | 1000 mL | 1000 mL | 1000 mL | 1000 mL  |

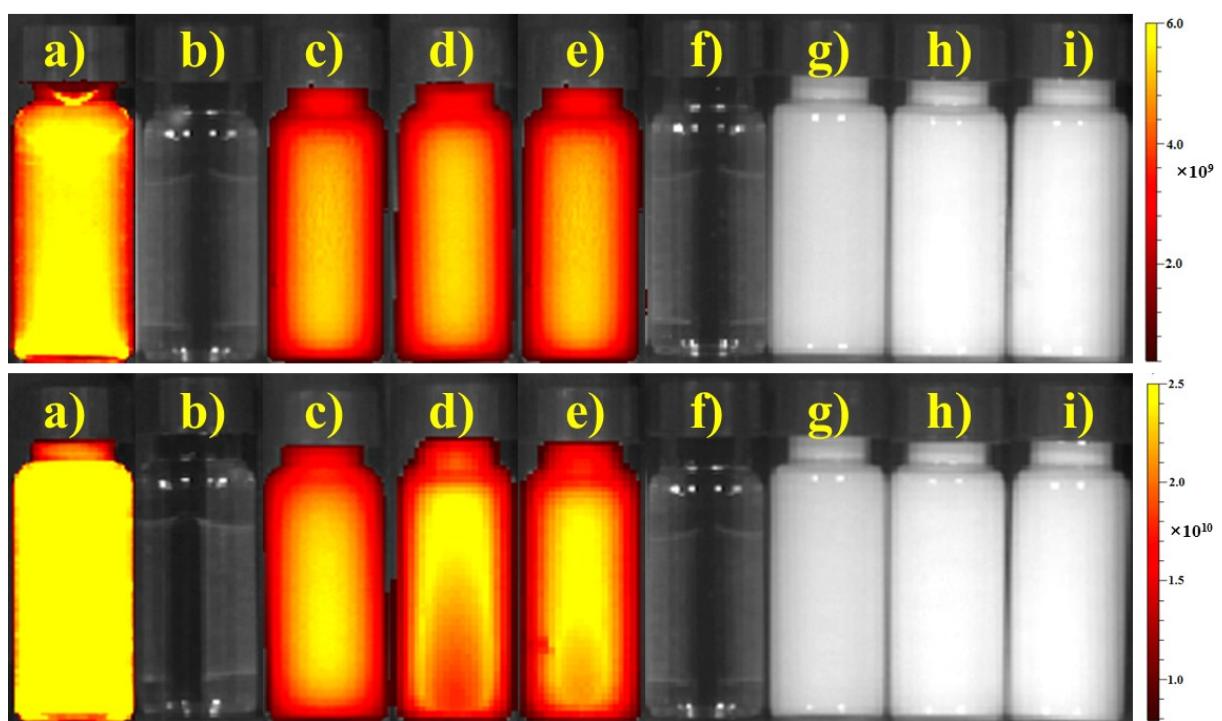


Figure S1. Fluorescence images of P2 samples (upper row) and P4 samples (lower row). The samples in each row from are listed left to right: a) probe solution in acetonitrile, b) probe solution in water, c) QT-HNCs-280, d) QT-HNCs-550, e) QT-HNCs-1100, f) probe solution in P188 (0.016%), g) QT-NCs-280, h) QT-NCs-550, i) QT-NCs-1100.