

Supplementary information for

Thermosensitive polynorbornene poly(ethylene oxide) nanoparticles  
loaded with oligoDNAs : an innovative approach for acting on cancer-  
associated pain

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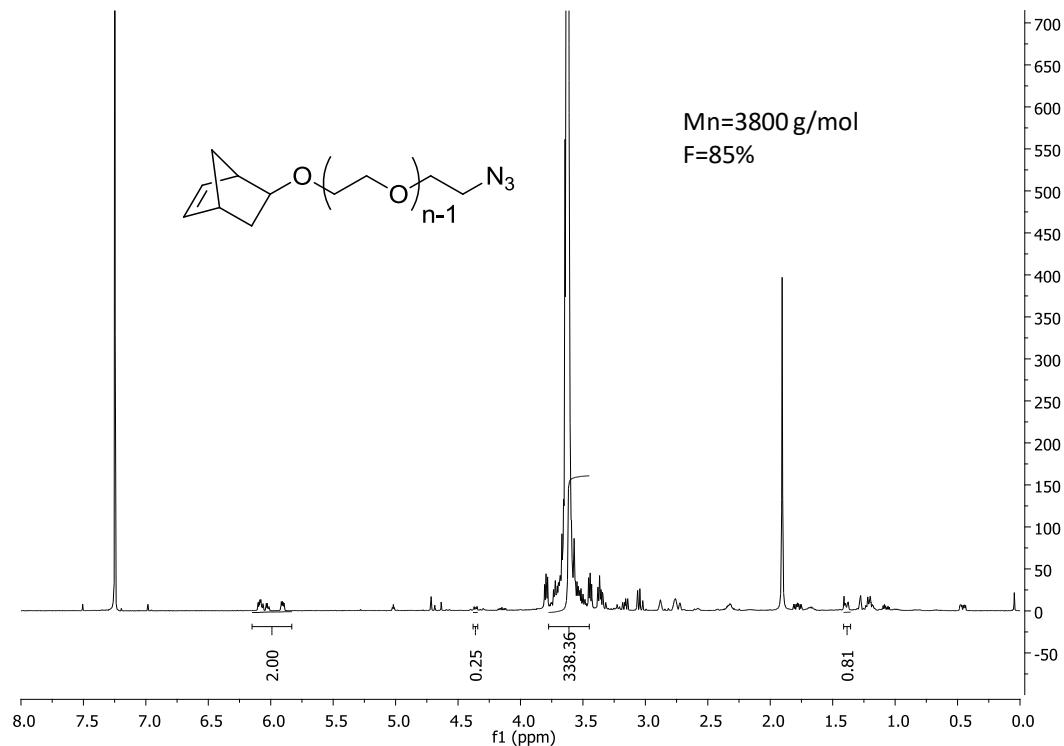
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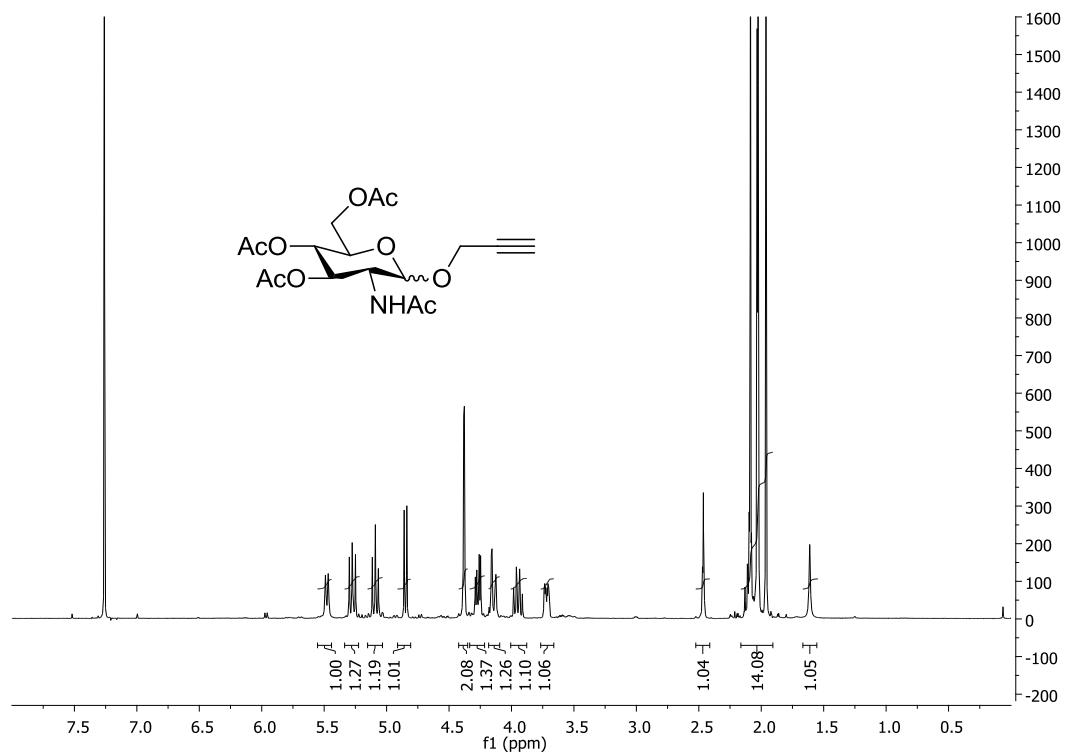
- **S1:  $^1\text{H}$  NMR spectra**
- **S2: Calculation of the amount of active molecules per NP (Glucosamine; Rhodamine)**
- **S3: UV calibration curve Absorbance at 260 nm vs oligoDNA concentration**
- **S4: Calculation of the amount of entrapped oligoDNA molecules per NP**
- **S5: ACSF preparation and final NPs concentration in ACSF medium**

**S1:  $^1\text{H}$  NMR spectra:**

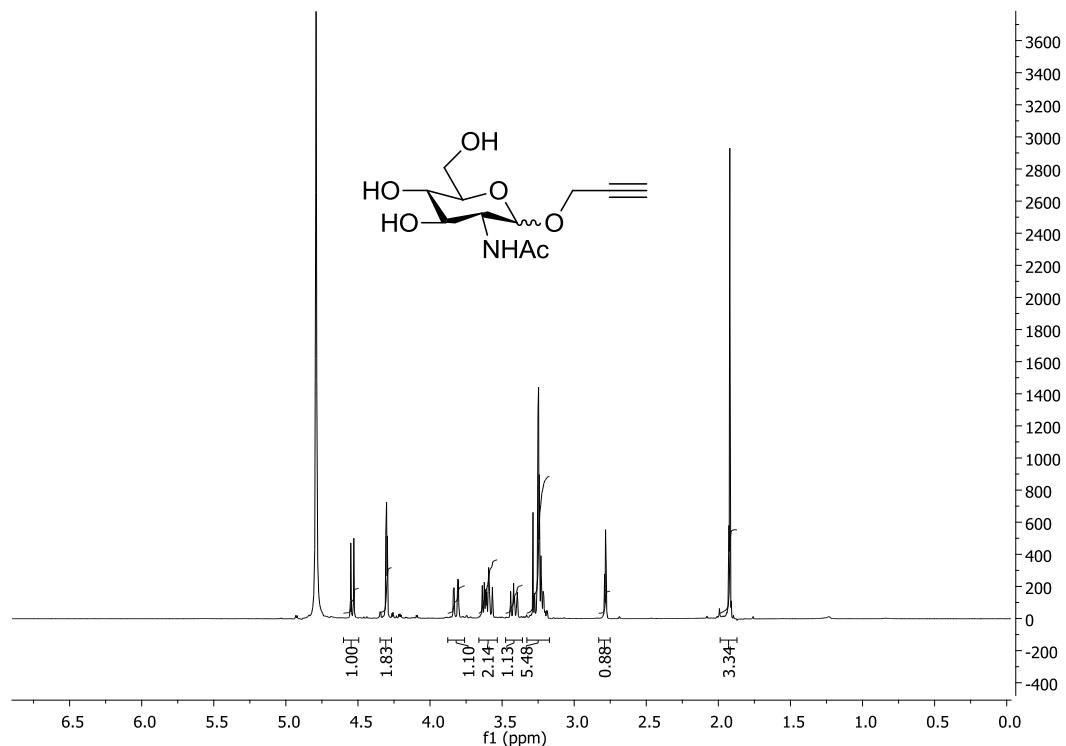
- $\alpha$ -norbornenyl- $\omega$ -azide-poly(ethylene oxide) (3)



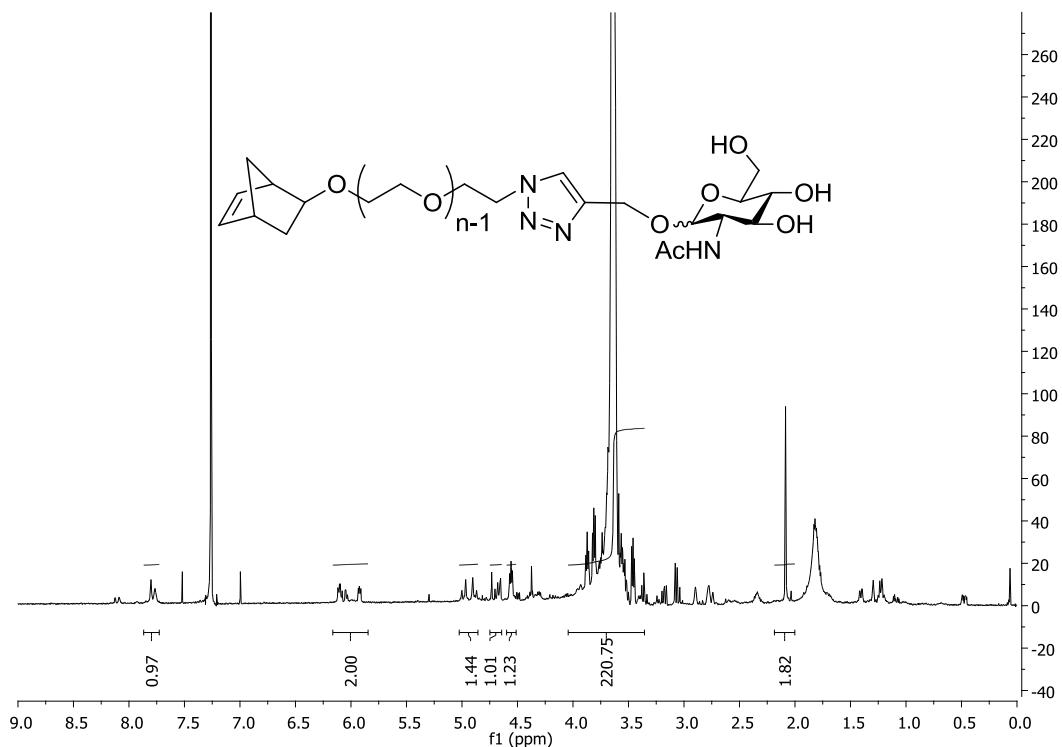
- Propargyl 2-acetamido-3,4,6-tri-O-acetyl-2-deoxy- $\alpha$ -D-glucopyranoside (5)



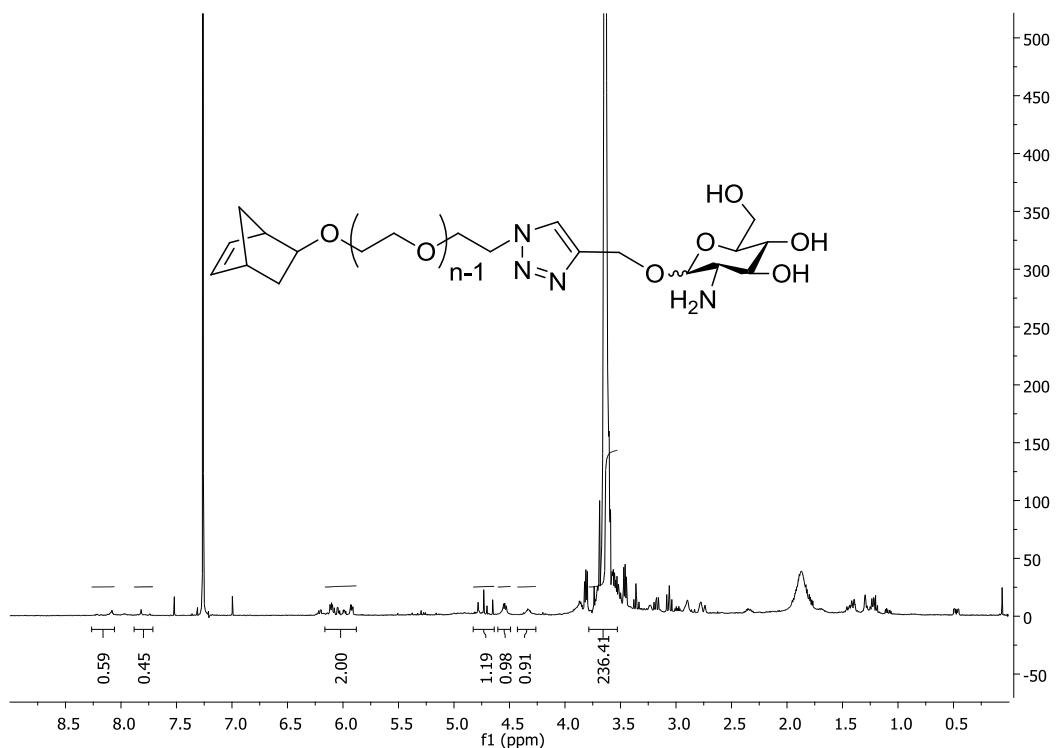
- Propargyl 2-acetamido-2-deoxy- $\beta$ -D-glucopyranoside (6)



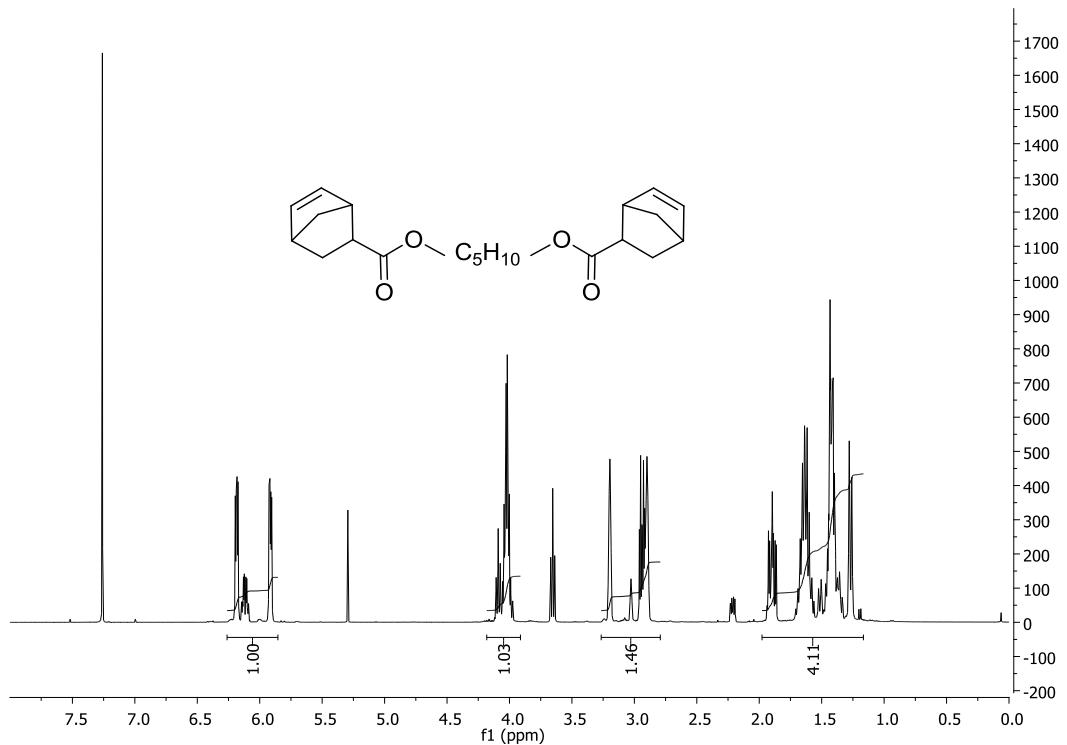
- Acetyl protected  $\alpha$ -norbornenyl- $\omega$ -glucosamine-poly(ethylene oxide) (7)



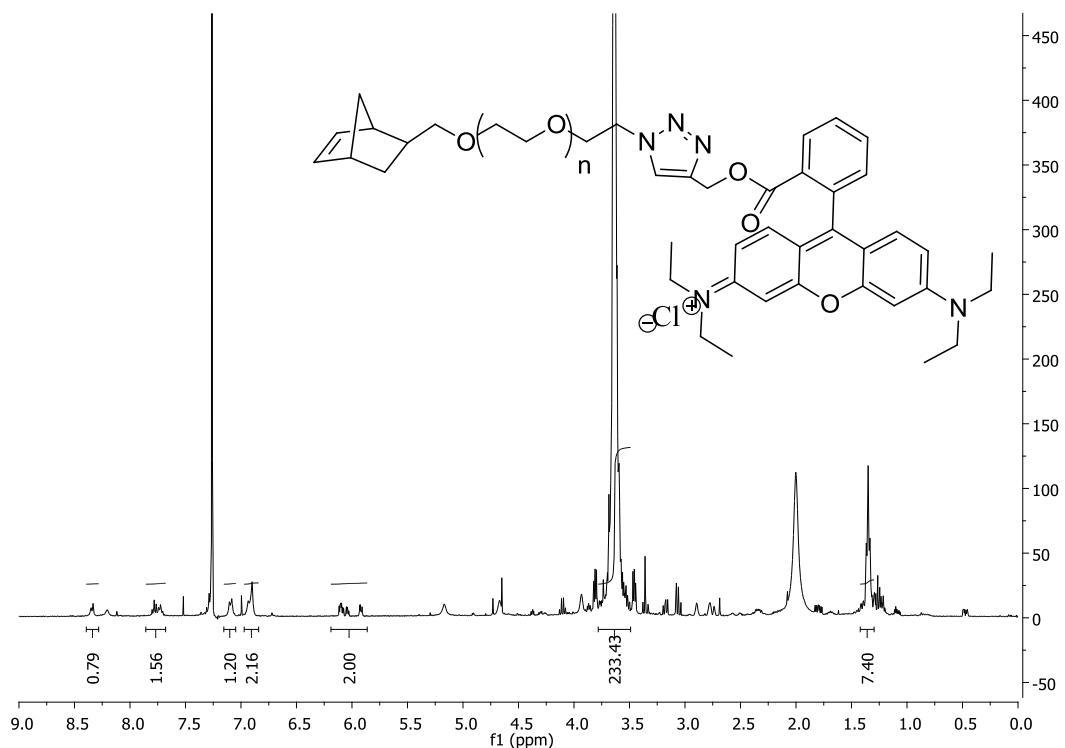
-  $\alpha$ -norbornenyl- $\omega$ -glucosamine-poly(ethylene oxide) (8)



- Pentane-1,5-diyl dinorbornene



-  $\alpha$ -norbornenyl- $\omega$ -Rhodamine-poly(ethylene oxide) (9)



## S2: Calculation of the amount of active molecules per NP (Glucosamine; Rhodamine)

To determine the amount of active molecules per NP, the active molecule concentration in the latex is first calculated with the following equation:

$$C_i = \frac{\pi \times F_i \times n_{Macro;i} \times M_i}{m_{Nb} + \pi \times m_{Macro;i}} \quad (\text{in gram of active molecule } i \text{ per gram of latex})$$

( $i$  = Glucosamine or Rhodamine)

With:

- $C_i$ : concentration of active molecule  $i$  in the latex (in gram of active molecule  $i$  per gram of latex)
- $\pi$ : conversion of macromonomer  $i$
- $F_i$ :  $\omega$  functionalization of macromonomer  $i$
- $n_{Macro;i}$ : initial amount of macromonomer  $i$
- $M_i$ : molecular weight of active molecule  $i$
- $m_{Nb}$ : initial weight of norbornene
- $m_{Macro;i}$ : initial weight of macromonomer  $i$

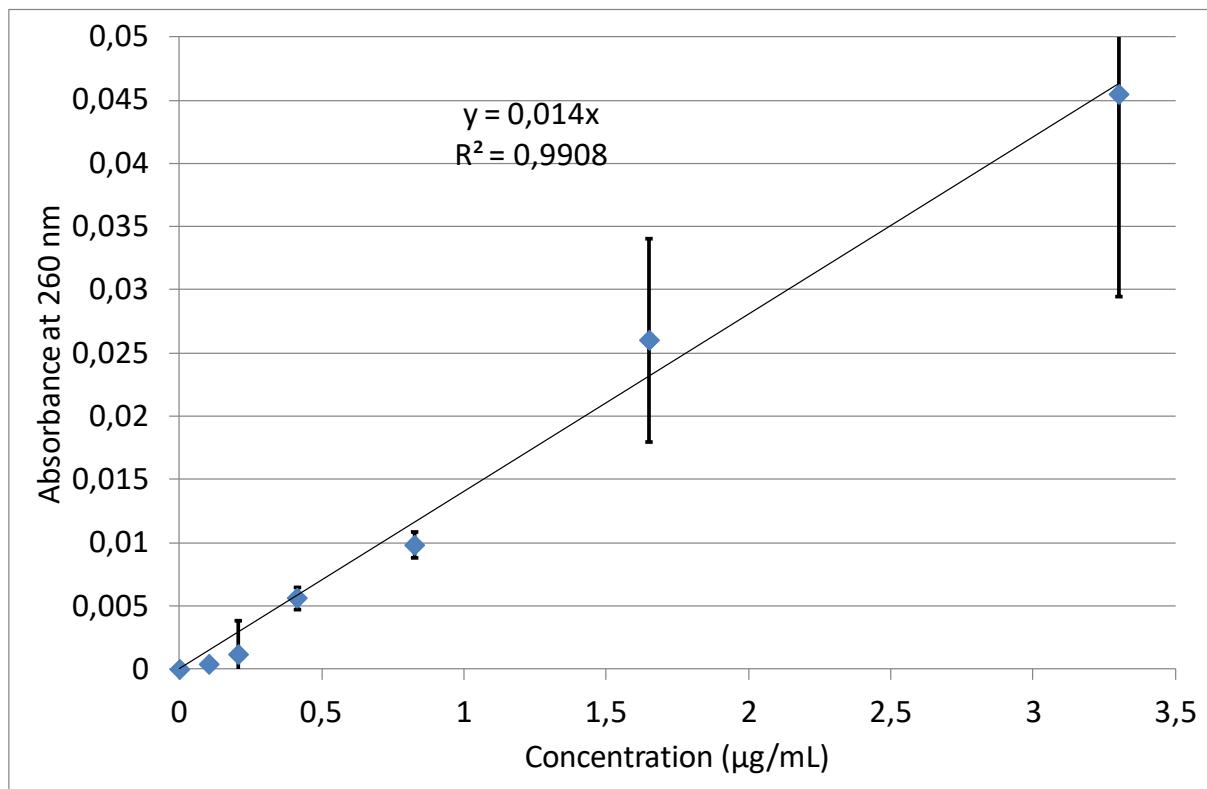
Then, the amount of molecule  $i$  per NP can be approximated with the following equation:

$$N_i/NP = \frac{C_i \times \rho_{NP} \times V_{NP} \times N_A}{M_i} \quad (\text{molecules per NP})$$

With :

- $N_i/NP$ : amount of molecule  $i$  per NP
- $\rho_{NP}$ : latex density approximated to equal to 1 g/mL
- $V_{NP}$ : volume of a NP ( $V_{NP} = \pi D_{NP}^3 / 6$ ) ( $D_{NP}$ : diameter of a NP measured by DLS)
- $N_A$ : Avogadro number
- $M_i$ : molecular weight of active molecule  $i$

**S3: UV calibration curve Absorbance at 260 nm vs oligoDNA concentration**



#### S4: Calculation of the amount of entrapped oligoDNA molecules per NP

The amount of entrapped oligoDNA molecules per NP is calculated with the following equation:

$$N_{\text{oligoDNA}}/\text{NP} = \frac{\%_{\text{ent}} \times V_{\text{NP}} \times \rho_{\text{NP}} \times [\text{oligoDNA}] \times V_{\text{oligoDNA}} \times N_A}{M_{\text{oligoDNA}} \times [m_{\text{Nb}} + \pi_{\text{CL}} \times m_{\text{CL}} + \pi_{\text{Macro}} \times m_{\text{Macro}}]}$$

With:

- $\%_{\text{ent}}$ : percentage of entrapped oligoDNA molecules during the reaction
- $N_{\text{oligoDNA}}/\text{NP}$ : the amount of entrapped oligoDNA molecules per NP
- $\rho_{\text{NP}}$ : latex density approximated to equal to 1 g/mL
- $V_{\text{NP}}$ : volume of a NP ( $V_{\text{NP}}=\pi D_{\text{NP}}^3/6$ ) ( $D_{\text{NP}}$ : diameter of a NP measured by DLS)
- $[\text{oligoDNA}]$ : concentration of of oligoDNA solution (1445 µg/mL)
- $V_{\text{oligoDNA}}$ : volume of the oligoDNA solution injected during the reaction
- $N_A$ : Avogadro number
- $M_{\text{oligoDNA}}$ : molecular weight of the oligoDNAs
- $m_{\text{Nb}}$ : initial weight of norbornene
- $\pi_{\text{CL}}$ : conversion of the crosslinker
- $m_{\text{CL}}$ : initial weight of crosslinker
- $\pi_{\text{Macro}}$ : conversion of the macromonomer
- $m_{\text{Macro}}$ : initial weight of macromonomer

## S5: ACSF preparation

products	Concentration (mM)
NaCl	130.5
KCl	2.4
MgSO <sub>4</sub> ;7H <sub>2</sub> O	1.3
KH <sub>2</sub> PO <sub>4</sub>	1.2
HEPES	1.25
CaCl <sub>2</sub>	2.4
Glucose	10
NaHCO <sub>3</sub>	19.5

The final NPs concentration in the ACSF medium ( $[NPs]_{ACSF}^f$ ; NPs/mL) can be calculated with the following equation:

$$[NPs]_{ACSF}^f = \frac{(m_{Nb} + \pi_{CL}m_{CL} + \pi_{macro}m_{macro})}{V_{NP}\rho_{NP}V_{ACSF}}$$

With:

- $m_{Nb}$ : initial weight of norbornene
- $\pi_{CL}$ : conversion of the crosslinker
- $m_{CL}$ : initial weight of crosslinker
- $\pi_{Macro}$ : conversion of the macromonomer
- $m_{Macro}$ : initial weight of macromonomer
- $\rho_{NP}$ : latex density approximated to equal to 1 g/mL
- $V_{NP}$ : volume of a NP ( $V_{NP}=\pi D_{NP}^3/6$ ) ( $D_{NP}$ : diameter of a NP measured by DLS)
- $V_{ACSF}$ : volume of ACSF solution

And the values of these concentrations for each system are summarized in the following table:

System	NP-OH	NP-OH-CL 5w-%	NP-GA
$[NPs]_{ACSF}^f$ (NPs/mL)	$5.5 \cdot 10^{11}$	$2.4 \cdot 10^{11}$	$4.1 \cdot 10^{11}$