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Supporting Information

## Heterogeneous multi-compartmental DNA hydrogel particles through microfluidic assembly for lymphocyte inspired precision medicine

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## Table S1. Sequences of the DNA and RNA oligonucleotides. Acryd: Acrylic phosphoramidite added to oligonucleotides as a 5'-modification ROXN: ROXN fluorophore (attached to a guanine residue) BHQ-2: Black Hole Quencher®-2 (attached to a cytosine residue) CholTEG: CholTEG (attached to a thymine residue) TAMRA: TAMRA fluorophore (attached to a guanine residue)

Name of	Saguanaa	
oligonucleotide	Sequence	
Sequence 1	5'-5Acryd/TTT TTG ATC TCA TCA ACA TCA	
b-a	GTC TGA TAA GCT A-3'	
Sequence 2	5'-/56-ROXN/rGrGrA rGrUrA rCrCrC rUrGrA	
b'-c	rUrGrA rGrArU rCTT-3'	
Sequence 3	5'-TCA GAC TGA TGT GGG TAC TCC/3BHQ_2/-	
a'-c'	3	
Sequence 3'	5'-TCA GAC TGA TGT GGG TAC TCC-3'	
a'-c'		
Sequence 4		
d'-b'	5 - TAG CTT ATC AGA CTG ATG TTG ATG-5	
Sequence 5	5'-rGrArU rCrUrC rArUrC rArGrG rGrUrA rCrUrC	
b-c'	rCTT/3CholTEG/-3'	
Sequence 5'	5'-/TAMRA/rGrArU rCrUrC rArUrC rArGrG	
b-c'	rGrUrA rCrUrC rCTT/-3'	
Sequence 6	5'-/5Acryd/TTT TTG GAG TAC CCA CAT CAG	
c-d	TCT GAT AAG C-3'	
Synthetic		
miRNA-21	5'-rUrArG rCrUrU rArUrC rArGrA rCrUrG rArUrG	
target	rUrUrG rA-3'	
a'-b'		

**Table S2:** The radius and volume of core-shell particles fabricated are summarized here. The core and shell. The core-shell volume ratio was altered from 1:1 to 1:4 by controlling the flow rates.

R:r	V <sub>core</sub> :V <sub>shell</sub>	<b>C</b> <sub>D1</sub> : <b>C</b> <sub>D2</sub>
<b>√</b> 2:1	1:1	1:1
<b>√</b> 3:1	1:2	1:0.5
<b>√</b> 4:1	1:3	1:0.33
√5:1	1:4	1:0.25



**Figure S1.** Microfluidic droplet generator to form monodispersed water-in-oil with pregel monomers as templates to fabricate the hydrogel particles as the cores in the coreshell hydrogel particles. The dimension of the channel is in the following (W: 25  $\mu$ m, H: 30  $\mu$ m).



**Figure S2.** Microfluidic device to produce the pre-gel droplets for core PEG hydrogel particle encapsulation forming core-shell PEG hydrogel particles via polymerization. The DNAs can be compartmentalized into the core and shell accordingly for logical control of siRNA releasing. The channel diameter is indicated here (W:  $30 \mu m$ , H:  $30 \mu m$ ).