## **Supporting Information**

## The effect of RGD peptide on 2D and miniaturized 3D culture of HEPM cells, MSCs, and ADSCs with alginate hydrogel

Jenna Dumbleton, Pranay Agarwal, Haishui Huang, Nathaniel Hogrebe, Keith J. Gooch, and Xiaoming He\*

**Figure S1**. A schematic illustration of the microfluidic device used for producing the cell-laden coreshell microcapsules with the inset showing the design of the flow-focusing junction. After the core solution containing cells, shell solution containing sodium alginate, and mineral oil emulsified with aqueous solution of calcium chloride flow into the flow-focusing junction, the oil emulsion shears the core and shell solutions to generate core-shell structured droplets at the flow-focusing junction. The alginate solution in the droplets is cross-linked by  $Ca^{2+}$  in the oil emulsion into hydrogel at and after the flow-focusing junction. The extraction channel at the end of the microfluidic device helps to extract/transfer the microcapsules from the oil to aqueous solution of carboxymethyl cellulose for efficient collection of the cell-laden core-shell microcapsules. The oil channel was 400  $\mu$ m x 400  $\mu$ m, the shell channel was 300  $\mu$ m x 300  $\mu$ m, and the core channel was 200  $\mu$ m x 200  $\mu$ m.<sup>44</sup>

**Figure S2.** Schematic designs showing how mechanical properties including storage (G') and loss (G'') moduli were measured using rheometry. (A) Alginate solution was pipetted onto a circular PDMS mold, then cross-linked with CaCl<sub>2</sub>. The gel disk was then transferred to the parallel-plate platform of the rheometer. (B) The core solution, which was 2% carboxymethyl cellulose, was pipetted directly onto the cone-plate platform for measurements.

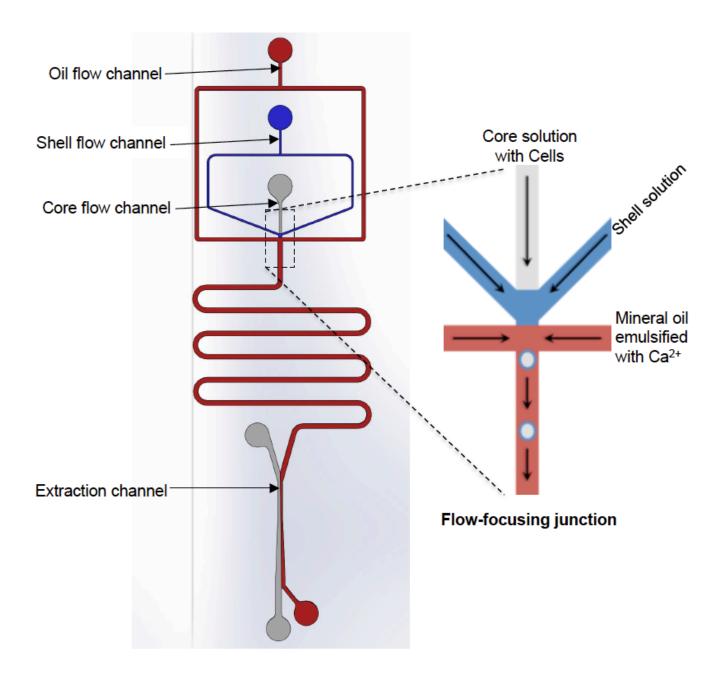
**Figure S3.** Human embryonic palatal mesenchyme (HEPM) cells, mesenchymal stem cells (MSCs), and adipose derived stem cells (ADSCs) cultured on flat hydrogels of 0.5% alginate-RGD, 2% alginate-RGD, and 2% unmodified alginate on days 0 and 4. The initial cell density was  $5 \times 10^4$  cells per well. Scale bar is 100 µm.

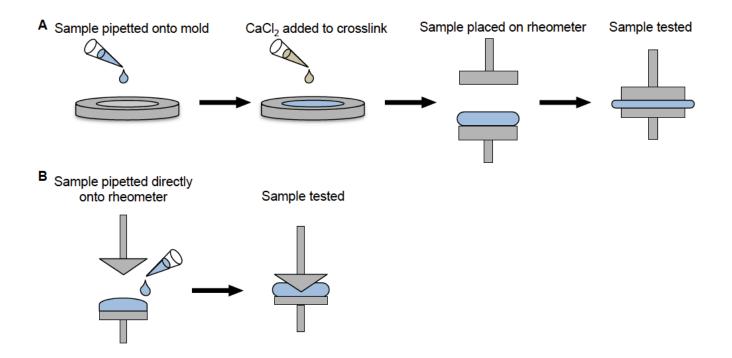
**Figure S4.** Typical images showing the morphology of HEPM cells cultured in the core of microcapsules with an alginate or alginate-RGD shell on days 0 and 4. Three different cores were studied: liquid core (0% alginate-RGD), hydrogel core of 0.5% alginate-RGD, and hydrogel core of 2% alginate-RGD. Scale bar is 100 μm.

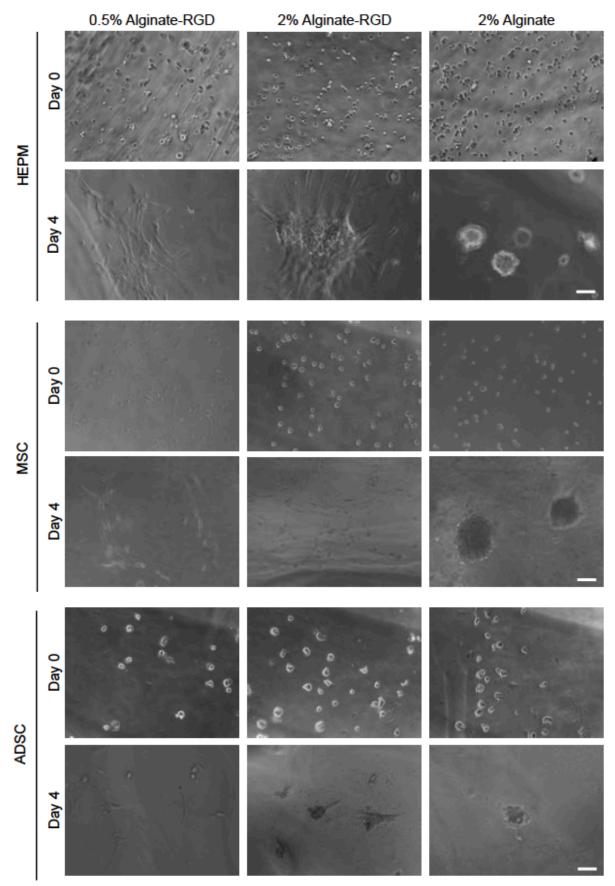
**Figure S5.** Typical images showing the morphology of MSCs cultured in the core of microcapsules with an alginate or alginate-RGD shell on days 0 and 4. Three different cores were studied: liquid core (0% alginate-RGD), hydrogel core of 0.5% alginate-RGD, and hydrogel core of 2% alginate-RGD. Scale bar is 100 µm.

**Figure S6.** Typical images showing the morphology of ADSCs cultured in the core of microcapsules with an alginate or alginate-RGD shell on days 0 and 4. Three different cores were studied: liquid core (0% alginate-RGD), hydrogel core of 0.5% alginate-RGD, and hydrogel core of 2% alginate-RGD. Scale bar is 100 µm.

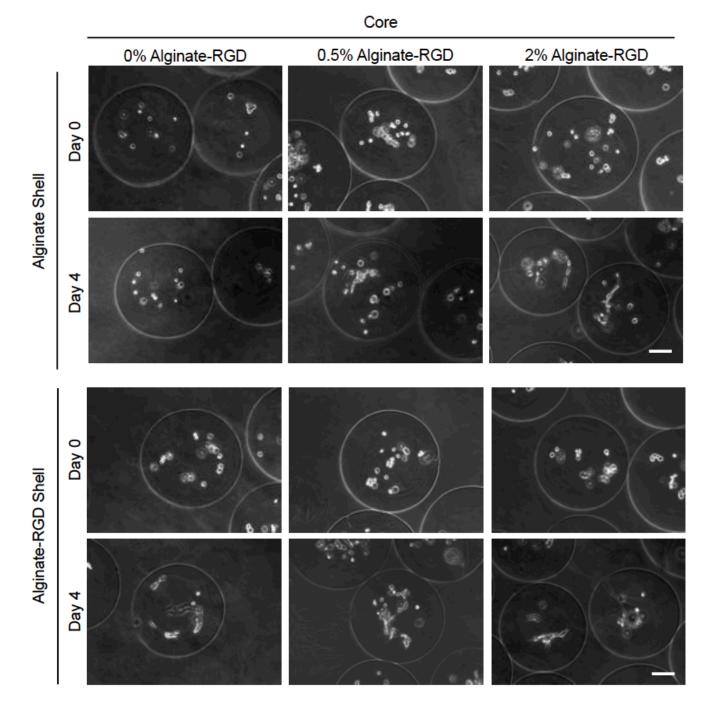
**Figure S7.** Human embryonic palatal mesenchyme (HEPM) cells, mesenchymal stem cells (MSCs), and adipose derived stem cells (ADSCs) cultured on flat hydrogels of 0.5% alginate-RGD, 2% alginate-RGD, and 2% unmodified alginate on days 6 with live/dead staining showing viability of all cells whether spread on the alginate-RGD hydrogels or aggregated on the alginate hydrogels. Scale bar is 100 μm.

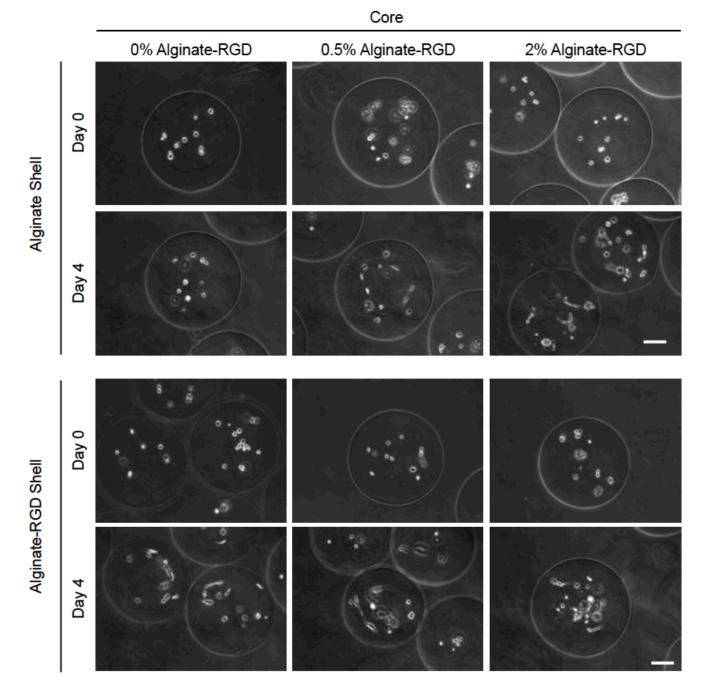




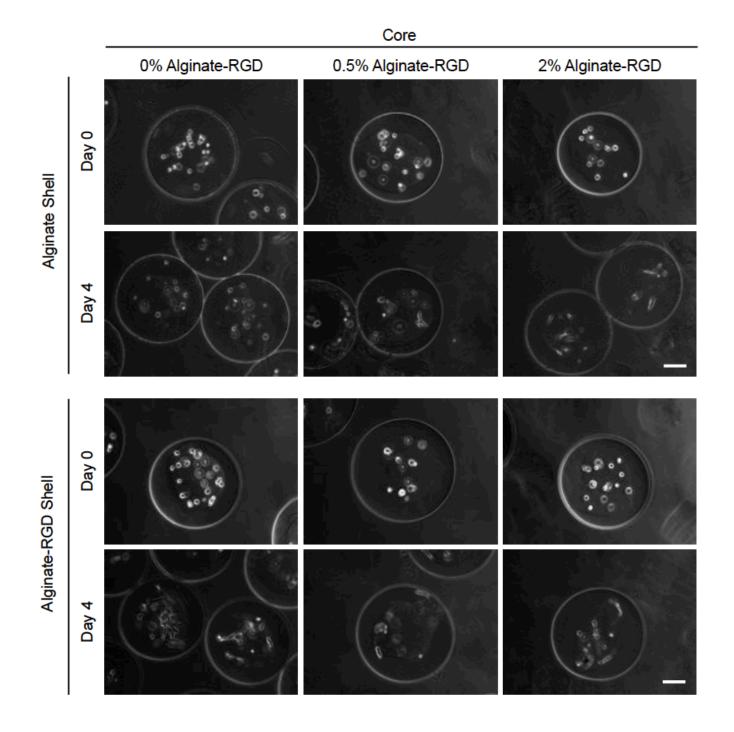


Dumbleton\_Figure S3





Dumbleton\_Figure S5



Dumbleton\_Figure S6

