

***In vivo* fate of free and encapsulated iron oxide nanoparticles after injection of labelled stem cells**

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Supplementary Information

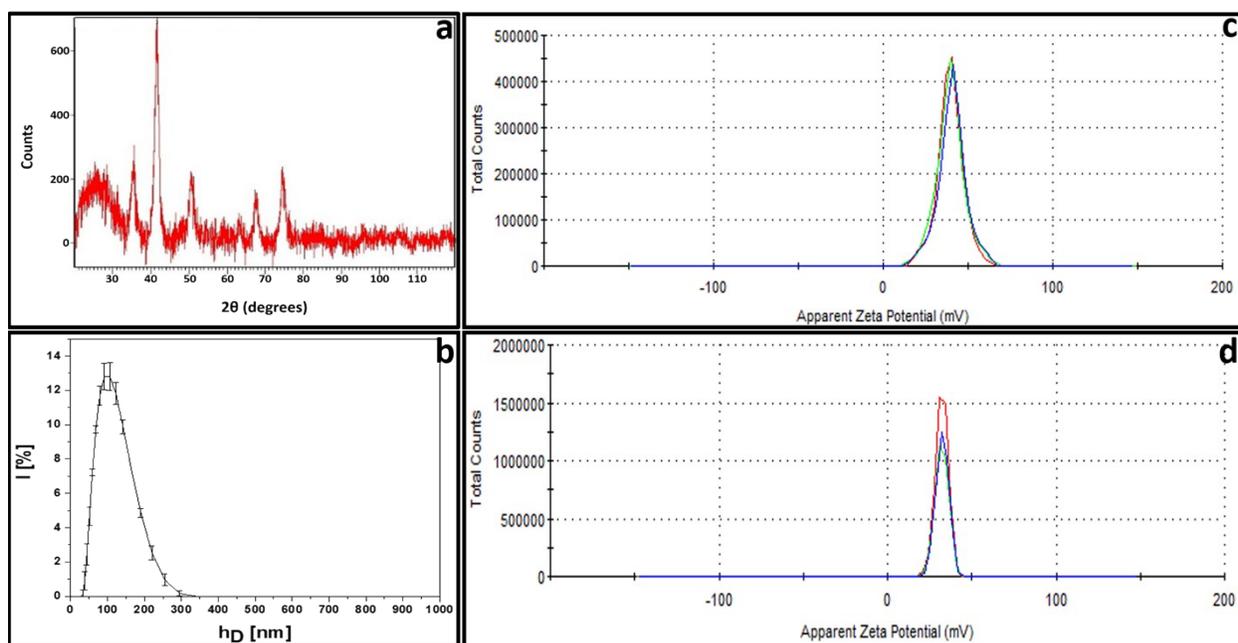


Figure SI.1: Characterization of free and encapsulated SPIONs by means of powder X-ray diffraction (pXRD), dynamic light scattering and zeta potential (ZP). pXRD of free SPIONs (a), hydrodynamic diameter (% intensity) of free SPIONs in water (b), and zeta potential measurements of free (39.5 mV) and encapsulated SPIONs (32.1 mV) in water is presented in (c) and (d), respectively. Three repeat measurements are shown for (c) and (d). The ZP measurements for large sedimenting objects (encapsulated SPIONs; d) is not reliable as sedimentation during the measurement period affects the accuracy of the data.

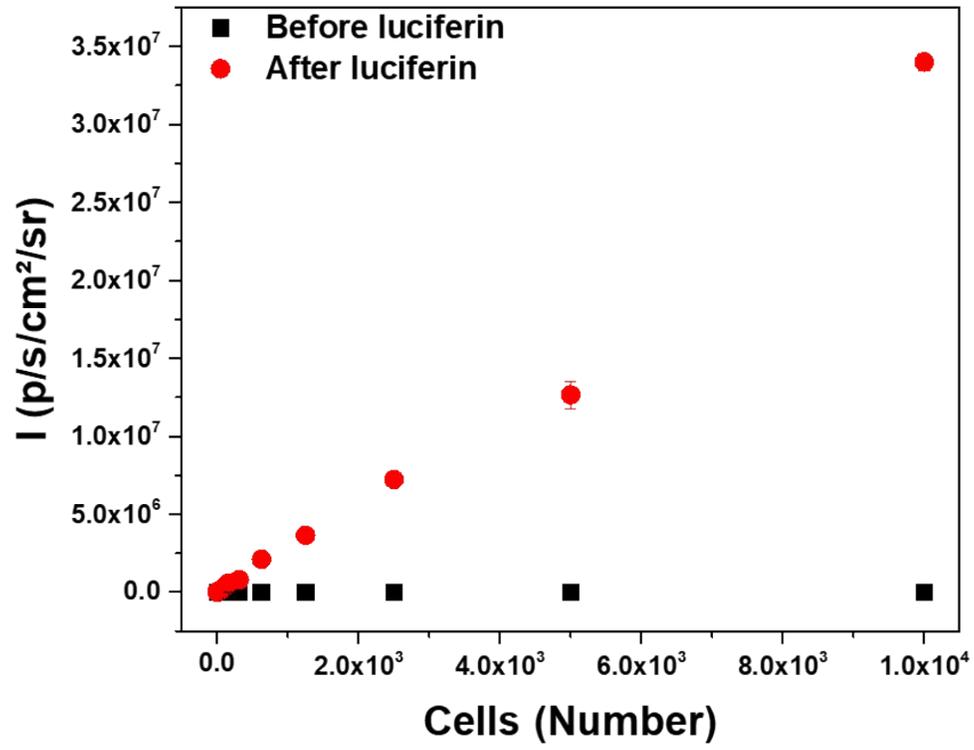


Figure SI.2: In vitro bioluminescence of mMSCs before and after the addition of D-luciferin.

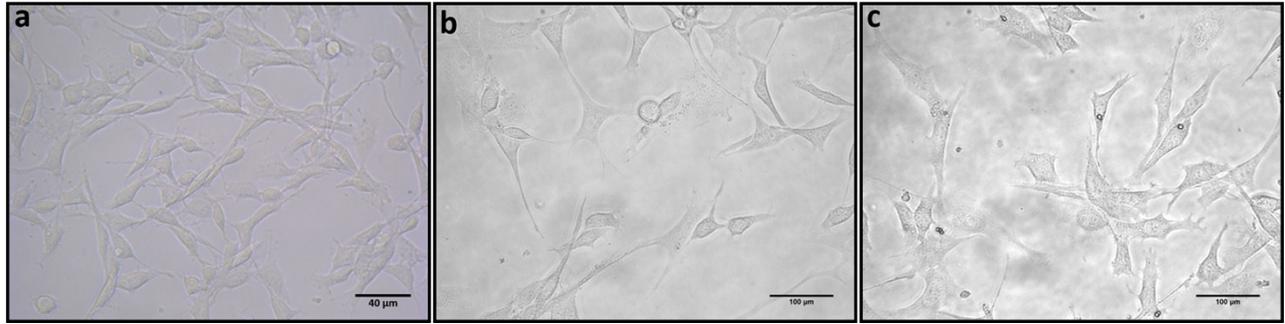


Figure SI.3: Morphology of mMSCs after labelling with particles. Unlabelled cells (a), cells labelled with free (b) and encapsulated SPIONs (c). After trypsinization the particle labelled cells were kept in ice cold PBS for 5 - 6 hrs and re-seeded. Cells were observed 3 days post seeding using 40 and 20 x objectives for unlabelled and particle-labelled cells, respectively.

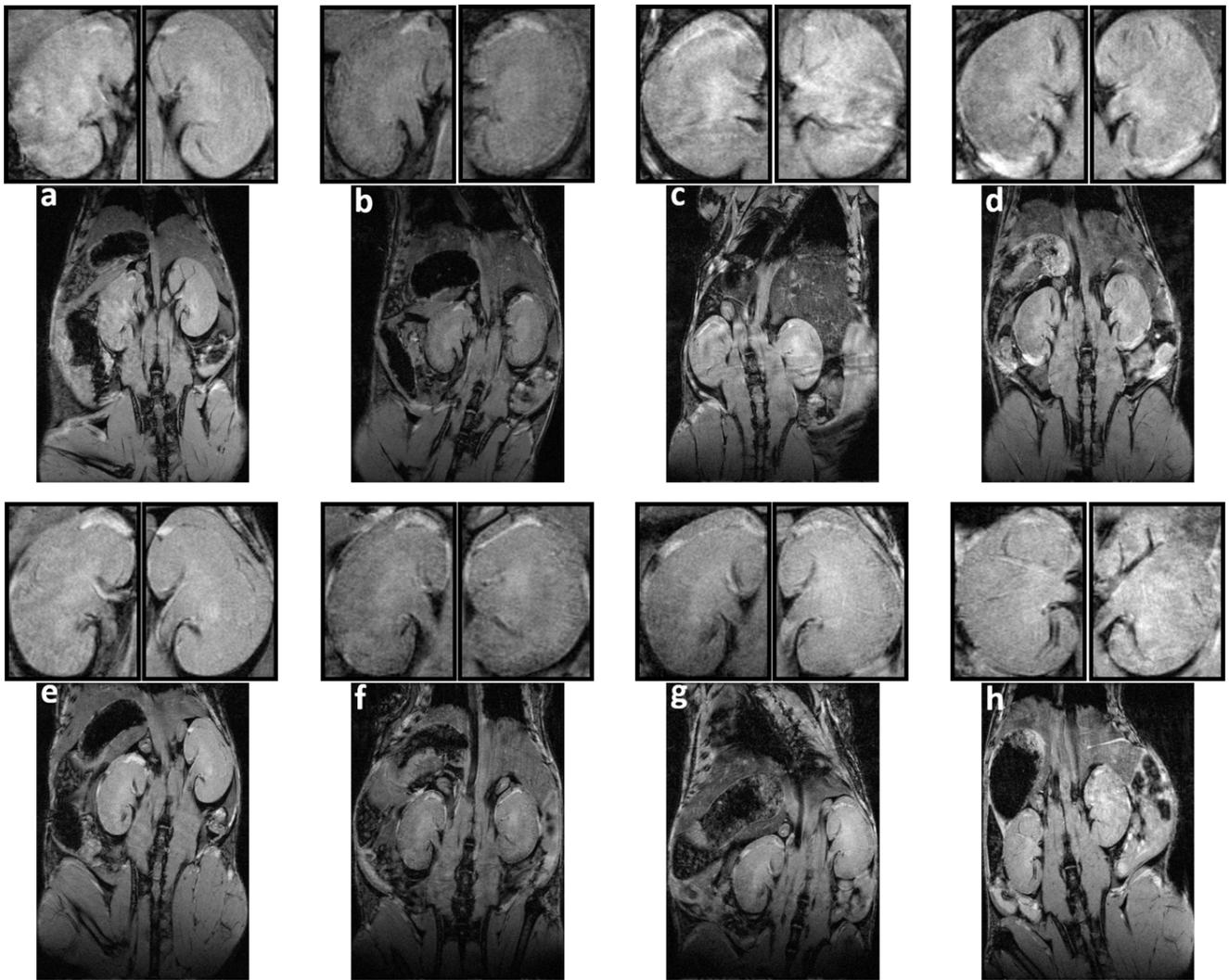


Figure SI.4: MR images of mice organs at selective days are presented. For clarity one mouse representing each treatment group (free SPIONs; a-d, and encapsulated SPIONs; e-h) are presented with zoomed in kidneys above each mouse (to see the contrast of SPIONs in kidney cortices) are shown. Mice abdominal MR scans before injection (baseline; a, e), day 0 (injection day; b, f), day 1 (post injection; c, g) and day 14 (post injection; d, h) are presented. Detailed abdominal scans of all mice at each imaging/scanning day can be found from <https://doi.org/10.5281/zenodo.1203991>.

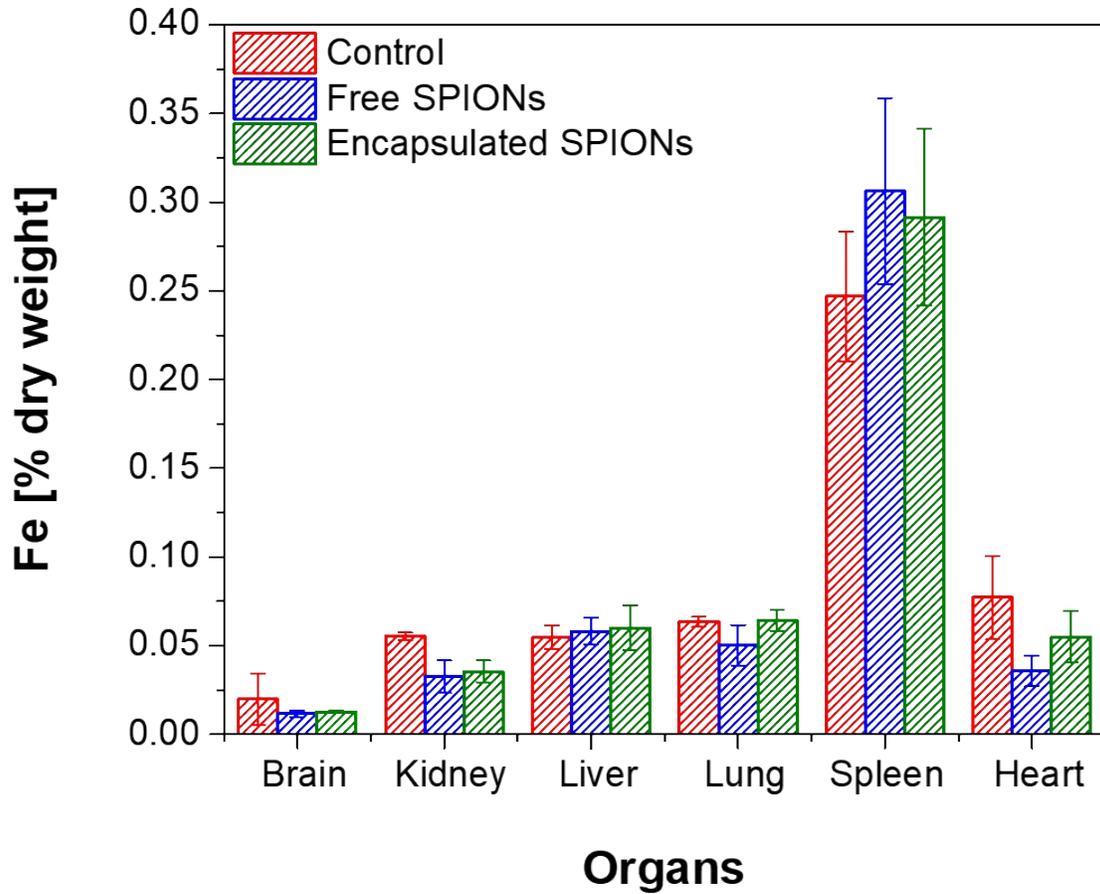


Figure SI.5: ICP-OES based Fe (iron) quantification of dried organs, collected at the end of experiments (day 14 post injection). Control group of mice did not receive any injection.

Sequence	FLASH T2*	B0 Map	MGE T2* Map
Purpose	T2* imaging	T2* Map Shimming	T2* relaxation
Echo time (ms)	4.9	1.59	4.5
Repetition time (ms)	253	10	800
Flip angle	20	15	50
Matrix size (pixels)	326 x 326	64 x 64	256 x 256
Image resolution	92 $\mu\text{m}/\text{pixel}$	625 $\mu\text{m}/\text{pixel}$	117 $\mu\text{m}/\text{pixel}$
Field of view (mm)	30 \times 30	40 \times 40	30 \times 30
Number of averages	8	3	2
Slices	10	1	1
Slice thickness (μm)	500	40000	1000
Echo images	N.A	2	8
Echo spacing (ms)	N.A	3.569	5.5

Acquisition time	3 min, 13 sec, 915 ms	2 min, 2 sec, 880 ms	8 min, 32 sec
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Table SI.1. MR acquisition parameters for SPIONs based phantoms. All images and mapping were axially performed.

Sequence	T2 Map MSME	B0 Map	MGE T2* Map	RARE T2	FLASH (axial and coronal)
Purpose	T2 relaxation	Shimming	T2* relaxation	T2 imaging	T2* imaging
Echo time (ms)	9	1.59	4.5	36	7
Repetition time (ms)	2000	10	800	2200	330
Flip angle	N.A	15	50	N.A	30
Matrix size (pixels)	256 x 256	64 x 64	256 x 256	512 x 512	512 x 512
Image resolution	117 $\mu\text{m}/\text{pixel}$	625 $\mu\text{m}/\text{pixel}$	117 $\mu\text{m}/\text{pixel}$	59 $\mu\text{m}/\text{pixel}$	59 $\mu\text{m}/\text{pixel}$
Field of view (mm)	30 x 30	40 x 40	30 x 30	30 x 30	30 x 30
Number of averages	1	3	2	6	6
Slices	5	1	1	19	19
Slice thickness (μm)	1000	40000	1000	500	500
Echo images	25	2	8	1	N.A
Echo spacing (ms)	9	3.569	5.5	12	N.A

Acquisition time	8 min, 32 sec	2 min, 2 sec, 880 ms	5 min, 7 sec, 200 ms	14 min, 4 sec, 800 ms	16 min, 53 sec, 760 ms
Rare factor	N.A	N.A	N.A	8	N.A
Excitation angle	90	N.A	N.A	90	N.A
Refocusing angle	180	N.A	N.A	180	N.A

Table SI.2. MR acquisition parameters for labelled mMSCs based phantoms.

Sequence	FLASH T2*	B0 Map	MGE T2* Map
Purpose	T2* imaging	Shimming	T2* relaxation
Echo time (ms)	5.5	1.59	4.5
Repetition time (ms)	305	10	1050
Flip angle	20	15	50
Matrix size (pixels)	386 x 386	64 x 64	256 x 256
Image resolution	91 $\mu\text{m}/\text{pixel}$	703 $\mu\text{m}/\text{pixel}$	137 $\mu\text{m}/\text{pixel}$
Field of view (mm)	35 x 35	45 x 45	35 x 35
Number of averages	3	3	2
Slices	23	1	23
Slice thickness (μm)	500	45000	500
Echo images	N.A	2	8
Echo spacing (ms)	N.A	2.855	4.5
Acquisition time	5 min, 53 sec, 190 ms	2 min, 2 sec, 880 ms	6 min, 43 sec, 200 ms

Reconstruction MAP	N.A	20	N.A
Signal to noise			

Table SI.3. MR acquisition parameters for *in vivo* MRI.