# **Electronic Supplementary Information**

#### Multifunctional magnetic nanoparticles for enhanced intracellular drug transport

### C. Tudisco,<sup>a</sup> M.T. Cambria,<sup>b</sup> F. Sinatra,<sup>b</sup> F. Bertani,<sup>c</sup> A. Alba,<sup>b</sup> A.E. Giuffrida,<sup>a</sup> S. Saccone,<sup>d</sup> E. Fantechi,<sup>e</sup> C. Innocenti,<sup>e</sup> C. Sangregorio,<sup>f,e</sup> E. Dalcanale,<sup>c</sup> G.G. Condorelli<sup>a,\*</sup>

<sup>a</sup>Dipartimento di Scienze Chimiche, Università di Catania and INSTM UdR di Catania, V.le A. Doria 6, 95125 Catania, Italy. E-mail: guido.condorelli@unict.it

<sup>b</sup>Dipartimento di Scienze Biomediche e Biotecnologiche, Università di Catania, Via S.Sofia 64, 95100 Catania, Italy

<sup>c</sup>Dipartimento di Chimica, Università di Parma and INSTM UdR di Parma, Parco Area delle Scienze 17/A, 43124 Parma, Italy

<sup>d</sup>Dipartimento di Scienze Biologiche, Geologiche e Ambientali, Università di Catania, Via A. Longo, 19, 95125 Catania

<sup>e</sup>Dipartimento di Chimica "U. Schiff", Università di Firenze and INSTM UdR Firenze, via della Lastruccia 3-13, Sesto Fiorentino, 50019 Firenze, Italy

<sup>f</sup>CNR-ICCOM, via Madonna del piano 10, 50019 Sesto Fiorentino, Firenze, Italy

#### **Table of contents**

Methods	<b>S</b> 1
XRD pattern of bare MNPs	S2
TEM image of bare MNPs	S3
Thermogravimetric measurement	S4
DLS measurement	S5
Cell Viability	S6
Synthesis of 3-PA@MNPs	S7
FT-IR spectra of 3-PA@MNPs.	S7
XPS spectra of 3-PA@MNPs	<b>S</b> 8

#### Methods

X-Ray powder diffraction (XRD) measurements were performed with a  $\theta$ - $\theta$  5005 Bruker-AXS diffractometer (Zeiss, Oberkochen, Germany) using Cu *Ka* radiation operating at 40 kV and 30 mA. The average diameter and size distribution were determined by Transmission Electron Microscopy (TEM), using a CM12 PHILIPS microscope operating at 100 kV. A dilute suspension of MNPs in water was drop dried onto 200 mesh carbon-coated copper grids. The statistics of MNPs diameter and size distribution was obtained considering about 400 particles, analyzing the recorded images with the Image Prop plus © software.

Thermogravimetric analysis (TGA) was performed on a Mettler Toledo TGA/SDTA 851<sup>e</sup> under air atmosphere. A 5-10 mg portion of nanoparticles was heated to 90 °C at 10 °C/min and kept at 90 °C for 30 min to remove all adsorbed solvent. The sample was then heated to 600 °C at 20 °C/min to determine the amount of organic coating on the nanoparticle surface.

Dynamic Light scattering (DLS) and zeta-potential measurements of MNPs were performed with a Nano Zetasizer (Malvern Instruments, Malvern, UK).





Figure S1. XRD pattern of bare MNPs.

## TEM image of bare MNPs



Figure S2. Selected low magnification TEM image of bare MNPs. The scale bar is 100 nm.

## Thermogravimetric measurement



Figure S3. Thermogram of 1-PA@MNPs.

### **DLS** measurement



Figure S4. Size distribution in water of 1-PA@MNPs.

Cell Viability



Figure S5. Cell viability of hMSCs evaluated by MTT after 72h of incubation with 1-PA@MNPs and 2-PA@MNPs at different concentrations.

#### Synthesis and characterization of 3-PA@MNPs

**3**-PA@MNPs were obtained with the same procedure as described for **1**-PA@MNPs but without using folic acid in the last step. Briefly, Tiiii functionalized PA@MNPs were dispersed in DMSO (20 ml) and 5(6)-Carboxy-X-rhodamine N-succinimidyl ester (Rhod-NHS, 1.5 mg) and PEG-NHS (50 mg) were added. The solution was mixed overnight at 25°C and the obtained particles were separated magnetically and washed as described for **1**-PA@MNPs.

**3-**PA@MNPs were characterized by FTIR and XPS spectroscopy. FT-IR and XPS spectra are similar to those of **1-**PA@MNPs since similar functional groups are present.



Figure S6. FT-IR spectra of 3-PA@MNPs.



Figure S7. XPS spectra of 3-PA@MNPs a) N1s b) C1s.

b)

a)