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NMR relaxivity on coated and non-coated size-sorted maghemite nanoparticles

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Statement of article significance

We investigate the NMR relaxation of iron oxide nanoparticles (MNPs). We sort by size MNPs to obtain numerous samples with diameter ranging from 4.5 to 12.5 nm with low polydispersity. We confirm that r_1 and r_2 NMR relaxivities increase with nanoparticle diameter. We also analyze the role of polydispersity for nanoparticles with the same mean size. Complementarily, we quantitatively investigate the role of coating on nanoparticles NMR relaxivity between bare and poly(sodium acrylate-co-maleate) coated nanoparticles (PAAMA). At last, we highlight that activation energy E_a decreases with nanoparticle diameter when determined from T_1 , but increases for T_2 determination.

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Supplementary information

SI-1 : scheme of size sorting process

Size sorting process was achieved to obtain the more monodispersed MNPs batches (Fig. SI-1a). From the original unsorted ferrofluid, 8 secondary batches are produced. Batches 2SC, 4SC, 5SC, 6SC, 7SC, and 8SC are sorted again to reach different sizes with similar polydispersity. Sample 4SC9SC was sorted again to get 10.7 nm large MNPs, compared to 11.2 for 4SC9SC sample.

Figure SI-1a : size sorting process of the first batch used in this study.

To reach very small nanoparticles, another batch was used and sorted thoroughly to have two sample denoted YP6SC and YP 7SC with diameter of 5.6 nm and 4.5 nm, respectively (Fig. SI-1b).

Figure SI-1b : size sorting process of the first batch used in this study.

SI-2 : properties of MNPs synthesized in this study

Table 1 : VSM, TEM, diameters with their correlated polydispersity (lognormal distribution) and DLS diameter of samples used in this study

Uncoated MNPs	VSM diameter (nm)	σ	TEM diameter (nm)	σ	DLS diameter (nm)
4SCC			12.72	0.29	28.8
4SCSC			11.88	0.28	26
4SC9SC	9.8	0.24	11.17	0.24	26.5
5SC13SC	7.8	0.16	9.4	0.17	6.2
7SC3SC	8.5	0.2	9.99	0.2	16.7
8SCC	8.6	0.21	9.23	0.21	15.5

8SC4SC	7.36	0.22	8.54	0.22	11.6
8SC5SC	7.4	0.19	7.73	0.19	12
4SCS9C4SC			10.68	0.17	15.3
7SC12SC			7.61	0.18	9.9
6SC26SC			7.21	0.18	9.4
2SC31SC			12.14	0.35	
YP1416SC			5.6	0.25	7.8
YP1417SC			4.5	0.16	9.2
Coated MNPs	VSM diameter (nm)	σ	TEM diameter (nm)	σ	DLS diameter (nm)
4SCC-PAAMA			12.72	0.29	21.5
4SCSC-PAAMA			11.88	0.28	34.2
4SC9SC-PAAMA	9.8	0.24	11.17	0.24	34.7
5SC13SC-PAAMA	7.8	0.16	9.4	0.17	21.9
7SC3SC-PAAMA	8.5	0.2	9.99	0.2	22.6
8SCC-PAAMA	8.6	0.21	9.23	0.21	22
8SC4SC-PAAMA	7.36	0.22	8.54	0.22	20
8SC5SC-PAAMA	7.4	0.19	7.73	0.19	55.2
4SC9SC4SC-PAAMA			10.68	0.17	25.5
2SC31SC-PAAMA			12.14	0.35	
YP1416SC-PAAMA			5.6	0.25	15.5
YP1417SC-PAAMA			4.5	0.16	17.6

NMR relaxivity on coated and non-coated size-sorted maghemite nanoparticles

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NMR relaxivity on coated and non-coated size-sorted maghemite nanoparticles

Relaxation dispersion profile, i.e. frequency dependence of the proton longitudinal relaxation times, were recorded for numerous samples of iron oxide nanoparticle dispersion with narrow size dispersity and diameters varying from 4 to 12.5 nm. We demonstrated that r_1 and r_2 NMR relaxivities increase with nanoparticle diameter, as expected by the models. We also analyze the role of polydispersity for nanoparticles with the same mean size on the dispersion curves. Then, we compared intensively the role of coating on nanoparticles NMR relaxivity between bare and poly(sodium acrylate-co-maleate) coated nanoparticles. At last, we investigated the influence of nanoparticle size on the activation energy E_a . Interestingly, while E_a decreases with nanoparticle diameter when determined from T_1 , it increases for T_2 determination. The influence is more important for small particles (<9 nm) than for big particles (>9 nm). More, the PAAMA coating changes the energy E_a obtained from T_2 : E_a becomes independent of the nanoparticle diameters.

Keywords: maghemite nanoparticles; polymer coating; NMR relaxivity

Introduction

The magnetic nanoparticles (MNPs) are widely studied and used for their high efficiency as contrast agent in MRI combined with their therapeutic properties¹⁻³. There are two major contributions to the relaxation of the magnetic moment of MNPs. First, their magnetic moment is proportional to the volume of the particles. The relaxation rates are thus largely ruled by the size of the particles, as well as by their anisotropy⁴. Second, the role of the coating on iron oxide nanoparticles is often neglected but can

drastically modify the MNPs relaxation process^{5,6}. Indeed, the coating can contribute to slow down the rotation of MNPs and modify the diffusion of solvent molecules compared to the medium. The relaxivity properties r_1 and r_2 , i.e. the relaxation rate of the surrounding nuclei R_1 and R_2 divided by the concentration of iron, are the two parameters that are determined experimentally. They are usually obtained at a unique frequency, but more informations can be obtained by their determination at various frequencies⁷. Experimental results highlight that r_1 and r_2 varies with nanoparticle diameter⁸. The particle with the optimal diameter is thus an important parameter to be taken into account for bio-applications for magnetic resonance imaging (MRI)⁷. However, only few to no systematic experimental works were conducted to measure the relaxivity of iron oxide nanoparticles on a large range of particle size with controlled (and low) polydispersity. On the more, one important factor can impact the relaxivities of water in presence of MNPs. Indeed, it has also been evidenced that the aggregation state of the particle plays a crucial role in their relaxivity properties⁹⁻¹¹. However, once incorporated in living animals, the MNPs may agglomerate or simply not be in a bulky state (adsorption on membrane cell...) ¹². Thus, the main recent developments are devoted to the design of efficient coatings that avoid or at least reduce magnetic dipolar interactions between MNPs^{3,13}. Once again, the coating can modify significantly the relaxation process by slowing the rotation of the particles and perturbing the diffusion of solvent molecules in the vicinity of the MNPs.

Complementarily, the response depends on the nature of MNPs. The example of maghemite and cobalt ferrite MNPs is illustrative. The magnetic anisotropy energy constant K that reflects the way the magnetic moment of the MNPs fluctuates along the easy magnetic axes, is very different in these two cases¹⁴. For maghemite, its value is quite low (about 6 kJ/mol) and is ruled by surface effects^{15,16}. For cobalt ferrite, its

value is about 10 times higher than for maghemite and originate from the core of MNPs, surface effects being negligible. Hence, adsorption of molecules on maghemite MNPs might lead to important consequences on their relaxivity properties¹³. On the contrary, the cobalt ferrite MNPs relaxivity properties should be much less affected by coating.

In this article, we have been interested in the maghemite MNPs. The study of relaxivity properties is often hindered by polydispersity of MNPs batch that are produced and used in classical studies. Herein we focus on a drastic size sorting process to obtain large MNPs batches with controlled size and low polydispersity (less than 0.2, according to a lognormal size distribution). We systematically compared the relaxivities of bare MNPs and poly(acrylate-co-maleate) coated nanoparticles, and explored the activation energy of uncoated and coated particles.

Experimental and method

Maghemite nanoparticles synthesis and coating

Iron oxide nanoparticles with bulk mass density $\rho = 5.10^3 \text{ kg.m}^{-3}$ were synthesized according to the Massart's pathway by aqueous alkaline co-precipitation of iron (II) and iron (III) salts and oxidation of the magnetite (Fe_3O_4) into maghemite ($\gamma\text{-Fe}_2\text{O}_3$) to prevent further oxidation¹⁷. At pH = 1.8, the bare particles are positively charged, with nitrate counter-ions. They have a diameter size distribution centered on 8 nm according to a lognormal distribution. The resulting inter-particle interactions are repulsive and impart an excellent colloidal stability to the dispersion. However, the nanoparticle polydispersity is high (typically 0.45 following the lognormal distribution). As the magnetic properties are proportional to the volume of the nanoparticles, the polydispersity induce an average of the magnetic properties, thus the relaxivities. To

highlight more precisely the influence of size nanoparticles on relaxivity, a size sorting has been achieved.

Size sorting is used to select sample with controlled size and lower size polydispersity. A well-established method was followed to obtain different sample with narrow size distribution¹⁸. Electrostatic repulsion between nanoparticles can be screened by the addition of salt. The repulsion will depend on the volume of the nanoparticles, and a phase separation is obtained where the concentrated phase contains the largest particles and the dilute one the smallest. Multiple steps of size sorting were achieved to obtain samples with the narrowest size distribution ($\sigma < 0.2$) and diameters going from 4.5 nm to 12 nm (see figure SI-1 for the size sorting pathway for the samples used in this study).

Experimentally, a given volume of nitric acid is added to the ferrofluid, that is placed on a magnet to separate the condensed phase (C) from the supernatant (S). The dense phase is washed with acetone and diethyl ether and redispersed in water. Another amount of nitric acid is added to the supernatant phase (S) and a second condensed phase (SC) is recovered and washed to obtain a second sample. The protocol is reproduced (samples 2SC, 3SC...) until the final supernatant is free from iron oxide nanoparticles. If the polydispersity of each sample is too large, each sample is sorted again by size following the same process. For instance, 5SC sorting processes give samples which are named 5SC1SC, 5SC2SC...

Ferrofluids were characterized with Vibrating Sample Magnetization (VSM), Transmission Electron Microscopy (TEM), atomic absorption spectrometry, and UV-Visible spectrometry (see table 1 in SI-2 for complete informations about the samples used in this study).

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3 Bare nanoparticles were coated with poly(acrylic acid-co-maleic acid) polymer chains
4 (PAAMA) using electrostatic interactions between carboxylate functions and the
5 positive charges on the nanoparticles, similarly to a previous pathway used to coat
6 MNPs with poly(sodium acrylate) ¹⁹. PAAMA was purchase at Sigma Aldrich and was
7 used without any purification. It consists of chains with average molecular weight of
8 3000 g/mol with a molar ratio 1:1 between each monomers. A solution at 0.5 %wt. of
9 bare nanoparticles is added to a 0.5%wt. of PAAMA (1:2 volume ratio) at acidic pH
10 value (1.8). A flocculation is induced by the interaction between the PAAMA and the
11 nanoparticles. The transparent supernatant is removed and the pH of the condensed
12 phase is increased up to 10 to redispersed the nanoparticles. This coating is particularly
13 efficient to stabilize the nanoparticles in brine or complex media ²⁰.
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30 *Maghemite nanoparticles characterization*

31 TEM is used to analyze physical size distribution of each nanoparticle sample. From
32 different images, diameters of nanoparticles are measured. A statistic is achieved to
33 obtain the size distribution of the nanoparticles. A lognormal distribution is used to fit
34 the experimental data and obtain a mean diameter and a size polydispersity. This size is
35 different from hydrodynamic diameter obtained by dynamic light scattering and the one
36 obtained by VSM. This diameter resulting from the TEM image analysis is used to
37 compare the relaxivity properties.
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49 The concentration of ferrofluid are determined using both atomic absorption and UV-
50 visible spectroscopies ²¹. Both methods lead to comparable results.
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53 Magnetic properties were obtained from a homemade VSM at room temperature,
54 operating between -1 Tesla and +1 Tesla. From magnetization curves, magnetization at
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saturation and magnetic susceptibility were obtained. These values are used to modelize the relaxometry dispersion curves.

The measurements of the water ^1H relaxation times (T_1 and T_2) were carried out on four different NMR apparatus. At 2 T (100 MHz for ^1H resonance frequency), the experiments were carried out using DSX100 Bruker spectrometer equipped with a diffusion probe (diff30 Bruker). The T_1 was measured using an inversion-recovery sequence and T_2 using a Hahn echo, with 0.2s recycle delay. At 1.2 T (60MHz) and 0.4 T (20MHz), the experiments were carried out using a 60 and a 20 Bruker Minispec, respectively. T_1 was measured using an inversion-recovery sequence with 16 recovery delays ranging from 40 μs to 10 T_1 approximately. T_2 using a CPMG sequence with 50 to 500 echoes separated by 80 μs , a recycle delay of 0.2 s.

The low frequency domain from 10 kHz to 15 MHz is explored on a Stelar Spin Master relaxometer. In this case, only T_1 has been measured using a PP sequence from 10 kHz to 8 MHz and a NP sequence from 8 MHz to 15 MHz²².

The relaxivities $r_1 = (T_1 [\text{Fe}])^{-1}$ and $r_2 = (T_2 [\text{Fe}])^{-1}$ were determined using five MNPs dispersions with the iron concentration $[\text{Fe}]$ ranging from 0.5 to 20mM, approximately. In this range, the relaxation rates $R_1 = 1/T_1$ and $R_2 = 1/T_2$ vary linearly with $[\text{Fe}]$, and the r_1 and r_2 values are obtained with a linear regression (figure S1). On the Stelar Spin Master, the measurements were performed for only one concentration because of the duration of the experiments. At 0.47 T, the relaxivities r_1 and r_2 were measured as a function of five different temperatures ($T=10, 17, 25, 33$ and 40°C).

Results and discussion

Effect of polydispersity

The size polydispersity of the nanoparticles dispersion has an obvious effect on relaxivity as the latter is influenced by nanoparticles size and shape. To illustrate this influence, the r_1 and r_2 at 20 MHz has been recorded for a large panel of maghemite samples and the r_1 profile has been recorded for two samples of the same medium size but with different polydispersity.

The polydispersity σ of our samples was determined from the analysis of TEM images using ImageJ software and fitted using a log-normal function fitted over several thousand of MNPs.

$$f(x; \mu, \sigma) = \frac{1}{x\sigma\sqrt{2\pi}} \exp\left(-\frac{(\ln x - \mu)^2}{2\sigma^2}\right)$$

The effect of polydispersity is first presented at one Larmor frequency (20 MHz) in order to show its effect depending on the MNPs size. Figure 3 shows the value of r_1 and r_2 as a function of the MNPs size for several set of samples with different polydispersity: $\sigma < 0.2$, $\sigma \approx 0.24$, $\sigma \approx 0.3$, $\sigma \approx 0.35$, and $\sigma > 0.4$.

With no surprise, the greater size polydispersity, the greater r_1 and r_2 value dispersion. The effect is much more pronounced for r_1 than for r_2 , for which the all points are relatively grouped. It can also be noticed that the increase of the polydispersity leads to lower relaxivity values for a given medium size. At first sight, it seems surprising because the increase of σ in the log-normal function for a given medium size implies that the proportion of bigger MNPs is higher in the sample and an increase of the MNPs size leads to higher r_1 and r_2 relaxivities as shown in the same figure 3.

In order to clarify this phenomenon, we have recorded the relaxation dispersion curve for two samples of the same medium size but with very different polydispersity: $d_0 = 10.8$ nm with $\sigma = 0.17$ and $\sigma = 0.53$ i.e. non sorted sample (figure 4). For the

maghemite sample with low polydispersity, the curve presents a well-defined profile with a plateau at low frequency, a depression just before the peak around 7 MHz and the decrease at high frequency. For the maghemite sample with large polydispersity, the characteristic figure of maghemite dispersion profile is shaded. The depression before the peak has disappeared and the peak is somehow crushed, i.e. broader and less intense. Moreover, its position is shifted toward low frequency. Hence, it can also be noticed that at high frequency, i.e. around 20 MHz, it leads to a lower r_1 value as compared to the sample with low polydispersity.

Effect of size

We now present only the results for the highly sorted samples, i.e. for which the polydispersity is lower than 0.2. Their NMR relaxivities r_1 et r_2 are plotted as a function of the TEM diameter in figure 5. An important increase is observed for both the longitudinal and the transversal components. The slope are about $6.4 \text{ s}^{-1} \text{ mM}^{-1}$ per nm for r_1 and $17.8 \text{ s}^{-1} \text{ mM}^{-1}$ per nm for r_2 . This difference is due to the dependence of the difference in the Curie term for r_1 and for r_2 .

Our values are well in line with the huge compilation study of Vuong et al.⁷ where the variation of r_2 with the size of MNPs has been studied over a very wide range of size (using also aggregates of MNPs). The authors show the linear dependence of r_2/M_v with the particle diameter where M_v is the saturation magnetization and this dependence pertains up to $0.4 \mu\text{m}$.

The effect of size on r_1 is not usually studied because the targeted application for MNPs is T_2 contrast agent. r_1 continuously increases with the diameter and tends to be linear in this range of size. This variation differs from the one observed for MNPs with high magnetic anisotropy constant like cobalt ferrite MNPs. In the latter case, r_1 increases with MNPs size but the slope decreases with the diameter.

Tabletop relaxometer allows us to record rapidly the r_1 and r_2 and ergo to be able to study a lot of samples relaxivities. Considering the data discrepancies in the literature, it is crucial to get statistics to investigate parameters like size, coating etc. This is an obvious advantage of this kind of relaxometers. However, the effect of the various parameters is better revealed by NMR relaxation profile.

On figure 6 are presented the r_1 profiles for non-coated MNPs with ($\sigma \leq 0.2$) and with diameter ranging from 4 to 11 nm. By increasing the size of spherical superparamagnetic nanoparticles several features occurs in the r_1 profile: (a) the maximum of the bump is shifted toward lower frequency, (b) the height of this maximum is increased, (c) the value of the plateau at low frequency is increased and (d) the hollow just before the bump is decreased. As already explained by Roch et al.²³ in their model, these variations are the result of a balance between the different dynamical characteristic times of the system (Neel time, diffusion time, rotational time).

Effect of coating

The PAAMA coating is achieved through electrostatic anchoring of the carboxylate functions at the nanoparticle surfaces. The molecular weight of the PAAMA chains (3,000 g/mol) corresponds to qualitatively 16 equivalent monomers of acrylate-maleate. This corresponds to a total length of the polymer chain of qualitatively 10-12 nm when completely elongated. Part of the monomers is adsorbed on the surface of the nanoparticles, and the chains are not completely straight in solution. This reduced drastically the thickness of the PAAMA corona around nanoparticles. Dynamic light scattering experiment comparing bare and PAAMA coated MNPs demonstrated an

increase of diameter of 4-5 nm for almost all MNPs batches, that is in good agreement with what was observed for poly(sodium acrylate) coated nanoparticles²⁴.

Bare and PAAMA coated MNPs were characterized by VSM. Curves between uncoated and coated MNPs are superimposed, and give same diameters and polydispersities before and after coating. This shows that the coating has no influence of the magnetic behavior of MNPs against applied magnetic field.

The values of r_1 and r_2 for PAAMA coated MNPs at 20 MHz are compared with those of non-coated MNPs (figure 5). The diameter considered here is the maghemite diameter without the polymer corona. The values are clearly superimposed for the r_2 while for the r_1 for PAAMA coated MNPs are slightly below the r_1 for non-coated MNPs. It can be noted that the effect of coating on the r_1 and r_2 at high frequency is subject to high discrepancies of partially and uncontrolled MNPs aggregation phenomena²⁵. Hence, the r_1 profiles is more able to reveal the coating effect.

The r_1 profiles have been recorded for PAAMA coated MNPs and plotted in figure 6. The evolution of the curves with the MNPs diameter is very similar to the curves for non-coated MNPs. In order to better reveal the effect of the PAAMA coating, the profile for non-coated and PAAMA coated MNPs have been superimposed for four sizes, 5.6, 7.7, 10.0 and 12.7 nm in figure 7. Several observations can be made when the MNPs are coated: (a) the value of the plateau at low frequency is smaller; (b) the hollow before the bump is less pronounced; (c) the position of the hollow is shifted toward low frequency; (d) the position of the bump is not shifted; (e) the bump maximum is affected and (f) the profiles tend to merge at high frequency.

To illustrate these effects, we have also plotted in figure 8 the ratio A between the maximum of r_1 ($r_1^{\max}$) and the r_1 at 10 kHz (the plateau value r_1^{plateau}) and the ratio B

between the minimum of r_I before the bump (r_I^{deep}) and the r_I^{plateau} . From figure 8 it is clear that the height associated to the Curie term in the models²⁶ is increased with the coating. It can be also notice that the difference between the ratio A for non-coated and coated MNPs is roughly constant with the maghemite diameter. The striking point is that the position of the bump is not affected in the same time. The models predict a combined modification of the bump height and its position^{14,23}. As illustrated by Kruk et al.²⁷ in maghemite MNPs dispersed in decaline and in toluene, a slowdown of the solvent dynamics lead to both the decrease of the bump and the shift of its position toward low frequencies. In the latter case, two solvents of different viscosities were used. In the case of coated MNPs, only the dynamics of water inside the polymer corona and in the very close vicinity is modified (along with the dynamics of the whole MNPs). To modelize the relaxivity of polymer coated MNPs it is therefore necessary to reconsider the diffusion propagator of water with a two steps environments. On figure 8, we have also plotted the ratio B between the minimum of r_I before the bump (r_I^{deep}) and the r_I^{plateau} . Here again a clear feature can be observed.

The MNPs in the models are impenetrable sphere and the diffusion of water is homogenous whatever the distance to the MNPs. In the case of polymer coated MNPs there is a corona around the MNPs where the diffusion of water is hindered. The effect of the modification of water diffusion in the vicinity of MNPs has been underlined by Ye et al.²⁸ who compared the relaxivities of magnetite coated by silica or capped by cetyltrimethyl ammonium bromide.

Activation energy

The relaxivity r_1 and r_2 at 20 MHz have been measured at different temperatures ranging from 10 to 40°C. In this range, the relaxivities exhibit an Arrhenius behavior and it was possible to determine an activation energy E_a . It must be underlined that at 20 MHz the relaxivity is ruled by the Curie term. The latter is related to the characteristic time $\tau_D = d^2/D$ of the water diffusion around the MNPs (d is the diameter of the MNPs and D is the self-diffusion coefficient of water). The energy E_a values obtained are in the same range of E_a for bulk water (19 kJ/mol)²⁹. The results are presented in figure 9 as function of the maghemite size. For the non-coated MNPs, the first striking point is the difference of behaviour in the E_a variations for r_1 and for r_2 . For r_1 E_a is constant for small MNPs approximately up to 10nm, and then decreases. The behavior is opposite for r_2 E_a as the latter increases up to 10 nm and then is roughly constant. It must be stressed also that these variations are very different from those observed for MNPs with high magnetic anisotropy. In the case of cobalt ferrite MNPs E_a of both the r_1 and the r_2 decreases. These differences are very interesting because MNPs with high (cobalt ferrite) and low (maghemite) magnetic anisotropy energy have very different r_1 profiles, but above 20 MHz, their r_1 and r_2 values are very similar. Therefore, the only measurement of E_a for r_1 and the r_2 relaxivities using a tabletop relaxometer is not sufficient to investigate the magnetic anisotropy effects. The measurement of E_a of r_1 and r_2 thus opens an opportunity for such studies and furthermore for studies on the tuning of the anisotropy properties.

The coating influences slightly the r_1 and the r_2 activation energy. For small size MNPs, E_a is increased by the coating while no effect is observed for larger maghemite (>8 nm). Greater effect was expected as the E_a , at this frequency, is dominated by the Curie term,

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3 i.e. correlated with the τ_D time. Hence the polymer corona seems to affect the
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5 relaxivities properties only for small maghemite MNPs.
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9 To analyze more precisely the impact of the coating on the activation energy, we can
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11 evaluate the role of surface to volume ratio, that is proportional to the inverse of the
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13 particle diameter. Indeed, we should be able to highlight the role of the coating on the
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15 relaxation process. Thus, we plotted the ratio between activation energies E_a (measure
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17 either on T_1 or T_2) of coated and uncoated particles versus the inverse of the particles
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19 diameter (figure 10).
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24 There are clearly some dispersions in measurements, but it can be seen that the
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26 activation energy ratio measured on T_1 values are mainly larger than unity, and
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28 increases with the inverse of the diameter. The activation energy ratio values obtained
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30 from T_2 measurement shows an increase with increasing value of $1/D$ as well, as
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32 observed with E_a values measured from T_1 . These results show that the coating on the
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34 nanoparticles play a significant role in the relaxation process of water molecules in the
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36 polymer corona compared to bare MNPs.
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39 40 41 **Conclusion**

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43 The influences of the size, the sample polydispersity and the polymer coating of
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45 maghemite nanoparticles on NMR relaxivities have been investigated. In agreement
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47 with previous studies, r_1 and r_2 increase with the MNPs diameter. The NMR relaxation
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49 profile better reveal the influence of the size with a shift of the r_1 maximum toward low
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51 frequency. It is shown the importance of controlling the size of the sample as in addition
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53 to higher discrepancy in r_1 and r_2 values, the NMR relaxation profile is significantly
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55 changed with a general broadening and an important shift of the r_1 maximum toward
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low frequency. Polymer coating influences the relaxivities of maghemite MNPs as shown by the NMR relaxation profiles. The most striking and surprising feature is that the height of the bump is modified while its position is unchanged. The activation energy of T_1 and T_2 have been measured at 20 MHz for non-coated and coated maghemite MNPs. This activation energy E_a exhibits an interesting dependence on the magnetic properties as its variation with MNPs size is clearly different for maghemite and cobalt ferrite MNPs, which magnetic anisotropy energy is ten time higher in the latter case than in the former case. The E_a of maghemite MNPs is influenced by the polymer coating only for small size MNPs (<8 nm).

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Figure 1: TEM image of the bare maghemite nanoparticles before sorting and after sorting process.

300x207mm (72 x 72 DPI)

Figure 2: scheme of the bare maghemite nanoparticles (left) and PAAMA coated MNPs (right).

309x120mm (72 x 72 DPI)

Figure 3: effect of the polydispersity σ on r_1 and r_2 : $\sigma < 0.2$ (blue squares), $\sigma \approx 0.24$ (empty circles), $\sigma \approx 0.3$ (green triangles), $\sigma \approx 0.35$ (black diamonds), and $\sigma > 0.4$ (red disks).

366x141mm (72 x 72 DPI)

Figure 4: r1 NMRD profile for two samples of the same medium size but with very different polydispersity: d = 10.7 nm with $\sigma = 0.17$ (circle) and d = 10.8 nm $\sigma = 0.53$ (triangle).

262x201mm (72 x 72 DPI)

Figure 5: r_1 and r_2 relaxivities of uncoated (black symbol) and coated (empty symbol) maghemite as a function of the diameter (for $\sigma \leq 0.2$).

367x149mm (72 x 72 DPI)

Figure 6: r_1 profile of maghemite uncoated MNPs (A) and PAAMA-coated MNPs (B) for diameters of the maghemite core (polymer corona not included) ranging from 4.5 to 11.2 nm.

368x141mm (72 x 72 DPI)

Figure 7: r_1 profile of maghemite MNPs non coated (black symbol) and PAAMA coated (empty symbol) for several diameters of the maghemite core (polymer corona not included): 5.6, 7.7, 10 and 12.7 nm.

254x213mm (72 x 72 DPI)

Figure 8: ratio A between the maximum of r_1 (r_{1max}) and the r_1 at 10 kHz (the plateau value $r_{1plateau}$) and the ratio B between the minimum of r_1 before the bump (r_{1deep}) and the $r_{1plateau}$ as a function of the maghemite diameter for uncoated (black disks) and coated (empty circles) MNPs.

361x155mm (72 x 72 DPI)

Figure 9: Activation energy E_a of T1 (left) and T2 (right) E_a for non-coated (black symbol) and coated (empty symbol) maghemite as a function of the diameter.

368x147mm (72 x 72 DPI)

Figure 10: ratio of activation energy E_a of T1 (left) and T2 (right) E_a between non-coated and coated maghemite as a function of the inverse of the diameter. Lines are linear fits.

366x143mm (72 x 72 DPI)

NMR relaxivity on coated and non-coated size-sorted maghemite nanoparticles

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Figures

Figure 1: TEM image of the bare maghemite nanoparticles before sorting and after sorting process.

Figure 2: scheme of the bare maghemite nanoparticles (left) and PAAMA coated MNPs (right).

Figure 3: effect of the polydispersity σ on r_1 and r_2 : $\sigma < 0.2$ (blue squares), $\sigma \approx 0.24$ (empty circles), $\sigma \approx 0.3$ (green triangles), $\sigma \approx 0.35$ (black diamonds), and $\sigma > 0.4$ (red disks).

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Figure 8: ratio A between the maximum of r_I ($r_I^{\max}$) and the r_I at 10 kHz (the plateau value r_I^{plateau}) and the ratio B between the minimum of r_I before the bump (r_I^{deep}) and the r_I^{plateau} as a function of the maghemite diameter for uncoated (black disks) and coated (empty circles) MNPs.

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