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Stray field magnetic resonance tomography using ferromagnetic spheres

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Abstract

The methodology for obtaining two- and three-dimensional magnetic resonance images by using azimuthally symmetric dipolar magnetic fields from ferromagnetic spheres is described. We utilize the symmetric property of a geometric sphere in the presence of a large externally applied magnetic field to demonstrate that a complete two- or three-dimensional structured rendering of a sample can be obtained without the motion of the sample relative to the sphere. Sequential positioning of the integrated sample-sphere system in an external magnetic field at various angular orientations provides all the required imaging slices for successful computerized tomographic image reconstruction. The elimination of the requirement to scan the sample relative to the ferromagnetic tip in this imaging protocol is a potentially valuable simplification compared to previous scanning probe magnetic resonance imaging proposals. © 2006 Elsevier Inc. All rights reserved.

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There has been a steady advance in the field of magnetic resonance imaging (MRI) towards higher spatial resolution. The largest measurement challenges stem from weak signals typical of high-resolution magnetic resonance [1], and the limitation of available gradient field strengths from current carrying conductors. Following the original reports [2–4] of applying magnetic field gradients to samples in order to demonstrate magnetic resonance imaging of spatial spin distribution, improvements in conventional inductive detection [5,6] have resulted in spatial imaging resolution of approximately 1 μ m [7–10]. The attraction and intense research interest towards 3D MRI with even higher resolution is driven by the well-known advantages of MRI as a three-dimensional, non-invasive, multi-contrast, and chemically specific imaging tool [11,12].

The introduction of ferromagnetic structures for increased resolution in magnetic resonance imaging has opened additional avenues toward achieving higher resolution. Scaling considerations show that a miniaturized

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permanent magnet will produce higher fields than an electromagnet, and can be further scaled to a smaller size without any loss in field strength [13]. Miniaturization of permanent magnets also provides an increase in the magnetic field gradients while requiring no electrical power supply and no current leads. Finally, permanent magnets generate no heat and thus require no heat dissipation. This ability of ferromagnets to provide high magnetic field gradients that can in turn spatially resolve resonant spins has led Sidles to propose the magnetic resonance force microscope (MRFM) [14]. In this instrument, a microscopic magnetic particle on a mechanical cantilever acts as a source of imaging gradient fields as well as a force generator on the spins whose magnetic resonance the mechanical cantilever detects [15]. Magnetic resonance image can be obtained by mechanically scanning the tip in three dimensions over the sample.

In this article, we focus on the magnetic resonance imaging protocol that uses the interaction between a sample and the stray fields from a geometrically symmetric ferromagnetic sphere. We demonstrate that a two- or three-dimensional imaging of a sample can be obtained without the motion of the sample relative to the sphere. We believe that

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the use of such a sphere model is reasonable, as microscopic ferromagnetic spheres of high quality and shape uniformity have been successfully fabricated, manipulated, and integrated on sensors [16-19]. Our protocol is similar to the stray field magnetic resonance imaging (STRAFI) technique [20] where constant magnetic field gradients, on the order of 60 T/m, from superconducting magnets are used. Here, however, the nanometer scale ferromagnetic spheres can provide ultra-high magnetic field gradients on the order of $\sim 5 \times 10^6$ T/m for a 100 nm diameter Cobalt sphere. This means that they can in principle be utilized for three-dimensional magnetic resonance imaging with resolution reaching Angstrom levels. We emphasize, however, that our proposal is equally valid if larger size ferromagnetic spheres are utilized. They would provide lower imaging resolution, but would be more suitable for present capabilities in magnetic resonance detection.

We describe a two-dimensional imaging protocol first, before expanding this principle to the full three-dimensional method. Our model configuration is shown in Fig. 1a, where a sample with size of $\sim 1/10$ the size of the ferromagnetic sphere is positioned as shown. The sample can represent a biological sample $\sim 10 \,\mu\text{m}$ in size next to a sphere 100 µm in diameter, or at the extreme, a small molecule or protein of $\sim 10 \text{ nm}$ in size next to a 100 nm diameter ferromagnetic sphere. A large DC magnetic field $B_0 \sim 10$ Tesla (presently unavailable to us) is applied parallel to the z-direction, polarizing the spins of the sample as well as saturating the magnetization of the ferromagnetic sphere. A small radio frequency field B_1 is applied perpendicular to the large polarizing DC magnetic field B_0 . In the absence of the ferromagnetic sphere, the nuclear spins in the sample would experience the same externally applied field B_0 and therefore meet the magnetic resonance condition at the same magnetic resonance frequency $\omega_{\rm R}$. However, close to the ferromagnetic sphere, a large magnetic field gradient is present at the sample, and only certain spins of the sample satisfy the correct magnetic resonance condition at any given magnetic field and frequency:

$$\omega(\vec{r}) = \gamma |B(\vec{r})| \tag{1}$$

The magnetic field from the ferromagnetic sphere at point r in the sample has the following azimuthally symmetric dipolar form:

$$\vec{B}(\vec{r}) = \frac{3\vec{n}(\vec{m}\cdot\vec{n}) - \vec{m}}{\left|\vec{r}\right|^3}$$
(2)

where n is the unit vector that points from the center of the ferromagnetic sphere to the sample location, and m is the magnetic moment vector of the sphere. Since the external DC polarizing magnetic field B_0 is considered to be much larger than the field from the ferromagnetic sphere, only the z-component of the magnetic field from the ferromagnetic sphere, B_Z , needs to be considered [21–23] for imaging. For a ferromagnetic sphere, this z-component of the magnetic field has the azimuthally symmetric form:

Fig. 1. (a) Model configuration for two-dimensional magnetic resonance tomography using ferromagnetic spheres. Imaging contours of constant zcomponent of the magnetic field are perpendicular to the sphere surface and intersect the sample positioned at $\theta = 54.7^{\circ}$. (b) The imaging contours shown along the plane parallel to the two-dimensional sample surface. The magnetic resonance spectrum of the sample is a one-dimensional projection of the sample spin density. Sequential rotations by angle φ provide the required projections for the tomographic image reconstruction

$$B_Z(\vec{r}) = \frac{M_0}{|\vec{r}|^3} (3\cos^2\theta - 1)$$
(3)

where θ is the angle between the z-axis and the distance vector r, and M_0 is the magnitude of the saturation magnetic moment of the ferromagnetic sphere. Fig. 1(a) also shows the contours of constant values for the z-component of the magnetic field from the sphere, B_Z , along the x-zplane.

In contrast to the previous approaches [21–23], we propose to fix the sample directly on the sphere, as shown in Fig. 1(a), at an angular location where:

a

process.



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$$\frac{\partial B_Z(\vec{r})}{\partial r} = \frac{M_0}{\left|\vec{r}\right|^4} (3\cos^2\theta - 1) = 0 \tag{4}$$

At this angular orientation of $\theta = 54.7^{\circ}$, $B_Z \approx 0$, and the contours of constant *z*-component of the magnetic field B_Z from the ferromagnetic sphere are perpendicular to the sphere surface, so that the sample is intersected by approximately perpendicular imaging slices. In Fig. 1(b), the contours of constant *z*-component of the magnetic field from the ferromagnetic sphere are shown along the plane parallel to the two-dimensional sample surface. This view shows that the magnetic resonance spectrum of the two-dimensional sample (i.e., the configuration shown in Fig. 1) will be a one-dimensional projection of the sample spin density. This leads to the possibility of obtaining a computerized tomographic image [24–27] if multiple imaging slices from the dipolar field of the ferromagnetic sphere can be obtained at different angles, as we describe below.

The imaging slices at multiple angles required for the computerized tomographic image reconstruction process can be obtained from a configuration of Fig. 1 without the motion of the sample relative to the sphere. We come to this conclusion by considering what happens when the integrated sample/sphere system is jointly rotated by an angle φ around the $\theta = 54.7^{\circ}$ axis, as shown in Fig. 1. Although both the sample and the sphere are mechanically rotated by the same angle φ , the presence of a large polarizing magnetic field B_0 of ~10 Tesla along the z-axis ensures that the saturated magnetic moment of the ferromagnetic sphere remains oriented along the z-axis. As a result, the imaging contours of constant z-component of the magnetic field, B_Z , remain fixed in space. Therefore, rotating the fixed sample/sphere system at a uniform sequence of angles φ provides all of the required imaging slices for previously developed two-dimensional computerized tomography reconstruction algorithms [24-27].

We note that, depending on the instrumental constraints or preferences, the actual rotation of the integrated ferromagnetic-sphere/sample system shown in Fig. 1 could also be experimentally executed by multiple sequential rotations around the x- and y-axes, as shown in Fig. 2. As rotations do not commute, such sequential rotations around x- and y-axes would have to be carefully selected. For example, the rotation of the sample around $\theta_{\rm Y}$ and then around $\theta_{\rm X}$, shown in Fig. 2a, would result in the correct translation and rotation of the sample for proper tomographic slicing, while a single rotation around the z-axis, shown in Fig. 2b, would result in the correct translation but incorrect rotation of the sample for proper slicing by the contours of constant B_Z . Additionally, we restrict our sample size to a fraction of the ferromagnetic sphere dimension in order to maintain the slicing of the sample by approximately parallel contours of constant B_Z . We note that image reconstruction from non-parallel slices has been demonstrated in computerized tomography [26] and is mathematically justified [28,29].



Fig. 2. (a) Alternative procedure for proper image slicing by sequential rotations around the *y*- and *x*-axes. (b) A single rotation around the *z*-axis would result in an incorrect rotation of the sample for proper slicing by the magnetic field imaging contours.

To extend our methodology to the three-dimensional imaging case, we find it visually advantageous to represent the integrated sphere/sample system rotations (described in Figs. 1 and 2) in a stationary sample reference frame, as shown in Fig. 3. In this perspective, the same effect of image slicing as described in Figs. 1 and 2 can be employed. In this reference frame, the sample is fixed and located on top of the sphere, as shown in Fig. 3, while the ferromagnetic moment of the sphere is tilted away from the z-axis by $\theta = 54.7^{\circ}$ and rotated around the z-axis at a sequence of angles φ required for the tomographic image reconstruction process.

We now analyze the case of a three-dimensional sample mounted on a ferromagnetic sphere, as shown in Fig. 4. At the angular position of $\theta = 54.7^{\circ}$, as in the two-dimensional



Fig. 3. Stationary sample reference frame. The sample is fixed and located on top of the sphere while the magnetic moment of the sphere is tilted away from the *z*-axis by $\theta = 54.7^{\circ}$ and rotates around the *z*-axis at a sequence of angles φ .

imaging case, the sample is intersected by the planes of constant z-component of the magnetic field from the ferromagnetic sphere that are approximately perpendicular to the sphere surface. Consider now the rotation of the integrated sample/sphere system so that the angle $\varphi = 0$ is held fixed while the angle θ is sequentially reduced in value from $\theta = 54.7^{\circ}$ to $\theta = 0$. This results in the sequential slicing of the three-dimensional sample by the imaging planes that range from being approximately perpendicular to the sphere surface to being approximately parallel to the sphere surface, as Fig. 4(a) shows. Therefore, by rotating the sample/sphere system through several angular values that range from $\theta = 54.7^{\circ}$ to $\theta = 0$, all the required imaging slices are obtained for two-dimensional image reconstruction along the x-z plane where angle $\varphi = 0$. This protocol again relies on the principle that, although both the sample and the sphere are mechanically rotated by the angle θ , the large polarizing magnetic field along the z-axis ensures that the saturated magnetic moment of the ferromagnetic sphere remains oriented along the z-axis and the imaging contours remain fixed in space.

A three-dimensional imaging protocol follows directly from these principles as all of the slices needed for three-dimensional image reconstruction can be obtained by varying both angles φ and θ , as described in the stationary sample reference frame of Fig. 4(b). By sequentially varying the angles (θ , φ) of the ferromagnetic moment direction through all possible angular combinations from $\theta = 54.7^{\circ}$ to $\theta = 0^{\circ}$ and $\varphi = 0^{\circ}$ to $\varphi = 360^{\circ}$, as shown in Fig. 4(b), the sample will be intersected by imaging slices at all possible angular orientations. This is sufficient for a complete three-dimensional image reconstruction, although several



Fig. 4. (a) Rotation of the integrated sample/sphere system from $\theta = 54.7^{\circ}$ to $\theta = 0$ results in the sequential slicing of the three-dimensional sample by the imaging planes that range from being approximately perpendicular to the sphere surface to being approximately parallel to the sphere surface. (b) The stationary sample reference frame for the three-dimensional tomography. By sequentially varying of the angles (θ , φ) of the sphere magnetic moment direction through all angular combinations from $\theta = 54.7^{\circ}$ to $\theta = 0^{\circ}$ and $\varphi = 0^{\circ}$ to $\varphi = 360^{\circ}$, the sample is intersected by the imaging slices at all possible angular orientations.

points of interest need to be addressed regarding the image-reconstruction process.

It is apparent from Fig. 4(a) that the planes of constant z-component of the dipolar magnetic field B_Z from the ferromagnetic sphere are curved, non-parallel, and not equally spaced. This is not prohibitive for the image reconstruction procedure, as basic back-projection algorithms [24–27] can be used for obtaining a three-dimensional image of the sample. More specifically, for an angular orientation (θ , φ), a weighted value is assigned to each contour of constant z-component of the magnetic field B_Z from the magnetic resonance spectrum obtained at that angular orientation. The three-dimensional image reconstruction of the sample is then completed by repeating

the weighted value assignment procedure for all angular orientations (θ , φ). Although this procedure is sufficient for basic three-dimensional image reconstruction, this simple back-projection algorithm is known to produce starlike image artifacts, and is therefore not optimal. The less artifact-prone but more complicated filtered back-projection algorithms or, alternatively, the matrix-based iterative-reconstruction algorithms could be employed [26].

A second point of interest is the image resolution. It is apparent from the inspection of the contours of constant z-component of the magnetic field in Fig. 4(a) that the image resolution depends on the distance from the ferromagnetic sphere surface. Only two magnetic field gradient forms are of interest since there is no variation of the azimuthally symmetric contours of the constant z-component of the magnetic field with the change of angle φ . The variation of the imaging contours along the radial direction is described by Eq. (4), and the gradient of the imaging contours along the angular θ direction is:

$$\frac{1}{r}\frac{\partial B_Z(\vec{r})}{\partial \theta} = -\frac{M_0}{\left|\vec{r}\right|^4}(6\cos\theta\sin\theta) = -\frac{3M_0}{\left|\vec{r}\right|^4}\sin2\theta \tag{5}$$

Both gradients (Eqs. (4) and (5)) have an inverse radial dependence to the fourth power, which means that parts of the sample closer to the sphere will experience higher magnetic field gradients and therefore can in principle be imaged with higher resolution. This can also be deduced from Fig. 4(a). Strong dependence of the gradient fields on r in Eqs. (4) and (5) also explains why the use of the nanoscale ferromagnetic spheres is advantageous in potentially obtaining atomic resolution images from projections. We emphasize, again, that this technique is equally applicable to larger diameter ferromagnetic spheres and therefore to lower field gradients and imaging resolution in order to match the signal-to-noise capability of the potential detectors discussed bellow.

It is important to point out that in our imaging method it is not required to know a priori where the sample is located on the ferromagnetic sphere. If the ferromagnetic moment direction is sequentially varied through the angles (θ, φ) from $\theta = 0^{\circ}$ to $\theta = 180^{\circ}$ and $\varphi = 0^{\circ}$ to $\varphi = 360^{\circ}$, the sample will be intersected by the imaging slices at all possible angular orientations, and a three-dimensional image reconstruction through back-projection algorithms will reveal an image and the location of the sample on the ferromagnetic sphere.

In addition to understanding the imaging methodology and resolution, it is important to discuss the choice of experimental methods for sample/sphere positioning as well as magnetic resonance detection signal-to-noise considerations. Our protocol involves angular motion of the sample/sphere system around two rotational axes. Such sample positioning technology is well developed and routinely used in the STRAFI technique [20]. The sensitivity requirements depend on the desired imaging resolution. Micrometer scale magnetic resonance imaging resolution has been experimentally achieved using standard inductive coil detectors [7–9] that could be used to achieve similar resolution in our proposed technique. In addition to the conventional inductive detection, we also suggest that optical detection methods [30–32], micro-coils [33,34], superconducting quantum interference devices (SQUID) [35,36], Hall sensors[37,38], and superconducting resonators[39] are viable candidates to be implemented in this imaging method.

As compared to the scanning probe type cantilever detection [15], in our protocol the need for scanning the sample with respect to the ferromagnetic probe is eliminated, along with the potential problems of long term positioning drift between the sample and the ferromagnetic gradient source. It is also important to note that, with the elimination of the relative motion of the sphere with respect to the sample, the thermo-mechanical vibrations of the cantilever do not translate into relative thermal motion and therefore fluctuations of the magnetic fields and field gradients from the sphere at the sample location. The intrinsic thermal motion of the magnetic moment remains, however, and has to be carefully considered in the ferromagnetic sphere material selection [40].

Signal-to-noise ratio evaluations become challenging when potential atomic resolution, polarization, and intrinsic nuclear spin noise are evaluated. In most room temperature NMR experiments the fractional polarization is quite small, on the order of 10^{-5} , and improves at helium temperatures to values on the order of 10^{-3} . This will have important implications for potentially achieving atomic resolution ferromagnetic sphere tomography at those temperatures due to the significant spin noise background. The spin noise problem in magnetic resonance is a topic of ongoing investigations and still somewhat controversial [41]. Additionally, a single or few spin detection schemes will likely require new methodologies in the area of quantum measurement [42–45] that deviate significantly for the classical theory of magnetic resonance detection.

We have described a technique for magnetic resonance tomography using the dipolar magnetic fields from ferromagnetic spheres distinctly different from previous magnetic resonance scanning probe microscopy methods. In previous experimental schemes, the images are obtained by raster scanning a ferromagnetic probe over the sample in three dimensions, and de-convolving intensities from the obtained magnetic resonance spectra at each point [46,47]. In contrast, in the dipolar field magnetic resonance tomography scheme described in this article, the ferromagnetic sphere and the sample are fixed with respect to one another. We rely on the geometric symmetry of the sphere and on the principle that the ferromagnetic moment remains saturated and oriented along a large polarizing magnetic field despite the mechanical motion of the sphere. Angular positioning of the integrated sample/sphere system then provides all the required imaging slices for computerized tomographic image reconstruction. The elimination of the requirement of scanning the sample relative to a ferromagnetic tip in this new imaging protocol

could represent a valuable experimental simplification toward achieving higher MRI spatial resolution than is presently available.

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