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(54) **SYSTEM AND METHOD FOR METABOLIC MR IMAGING OF A HYPERPOLARIZED AGENT**

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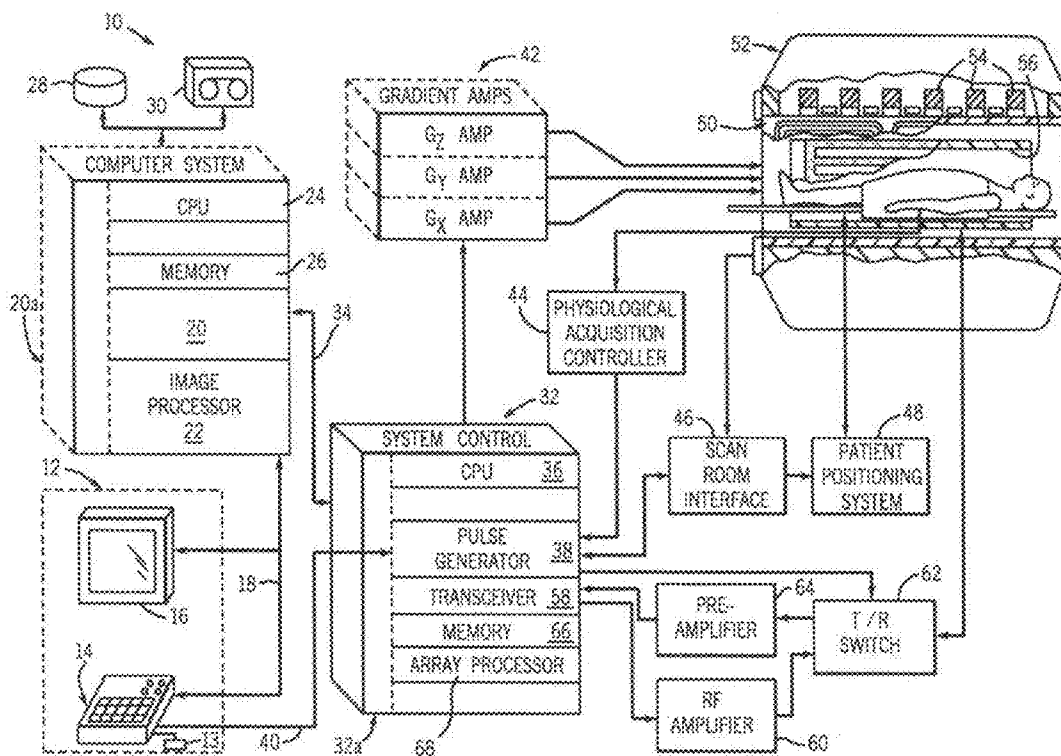
(57) **ABSTRACT**

A system and method for metabolic MR imaging of a hyperpolarized agent includes exciting a single metabolic species of a hyperpolarized agent injected into a subject of interest. MR signals are acquired from the excited single metabolic species and an image is reconstructed from the acquired MR signals.
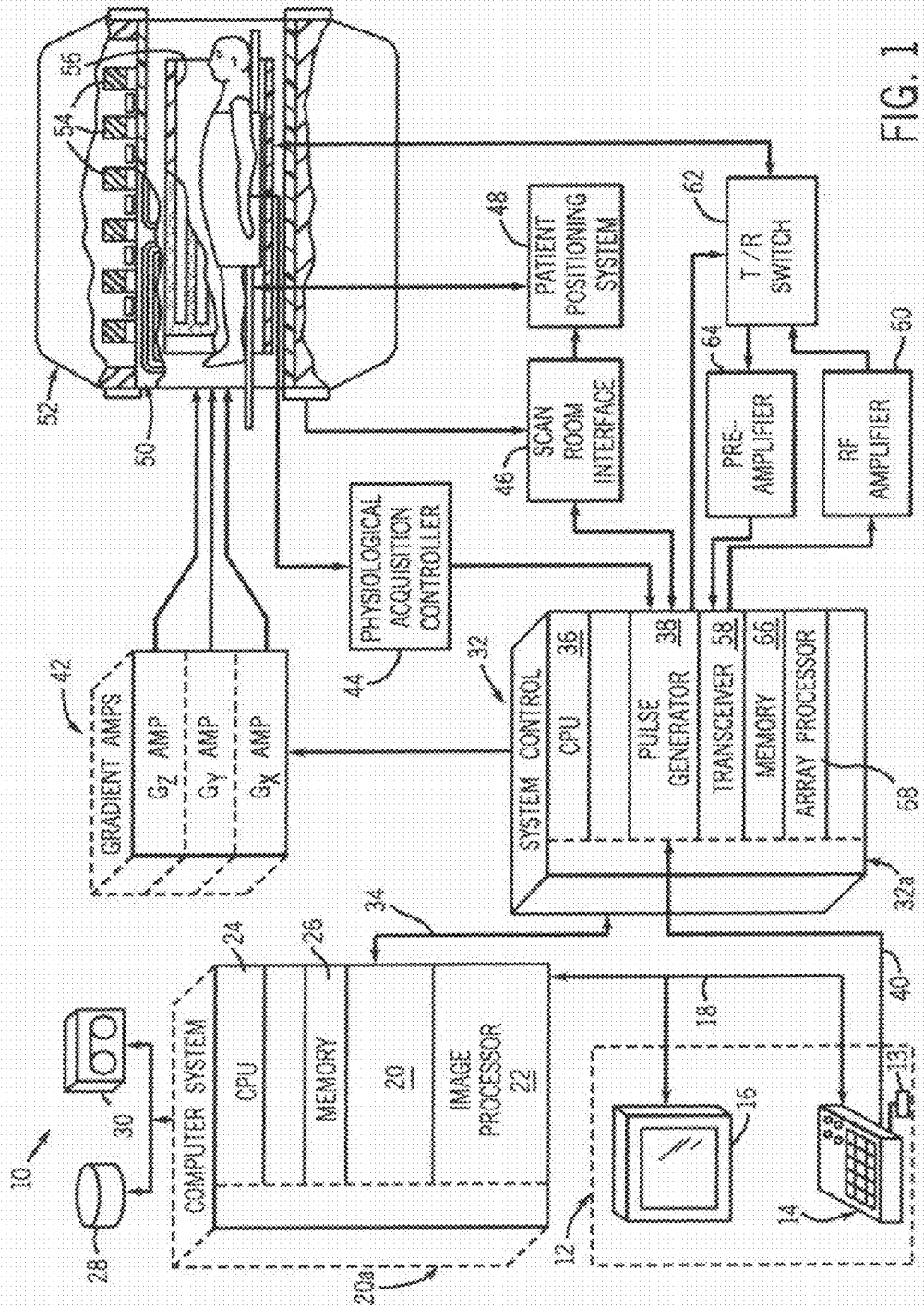
FIG. 1

FIG. 2

FIG. 3

Flip angle 1 deg.

88

88

90

90

90

Signal (Mxy)

Phaseshift during one TR

FIG. 4
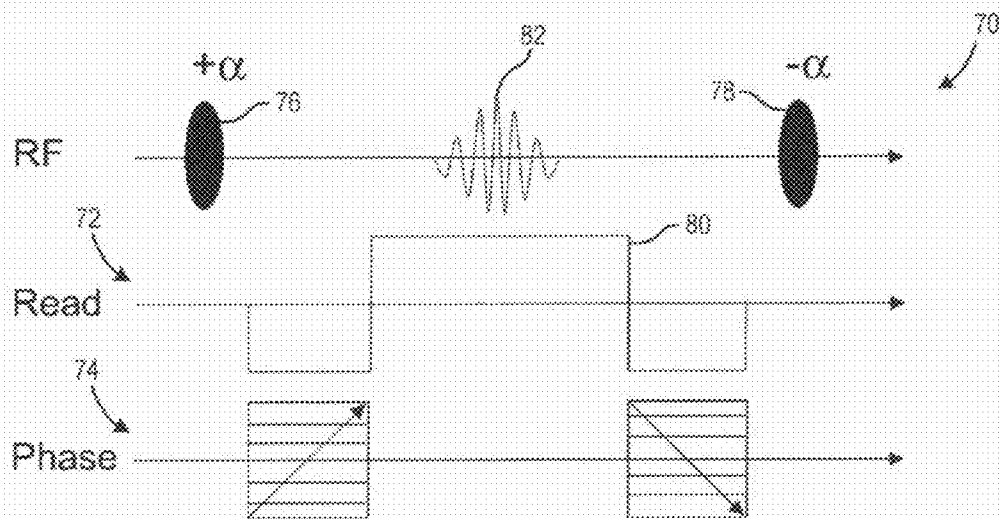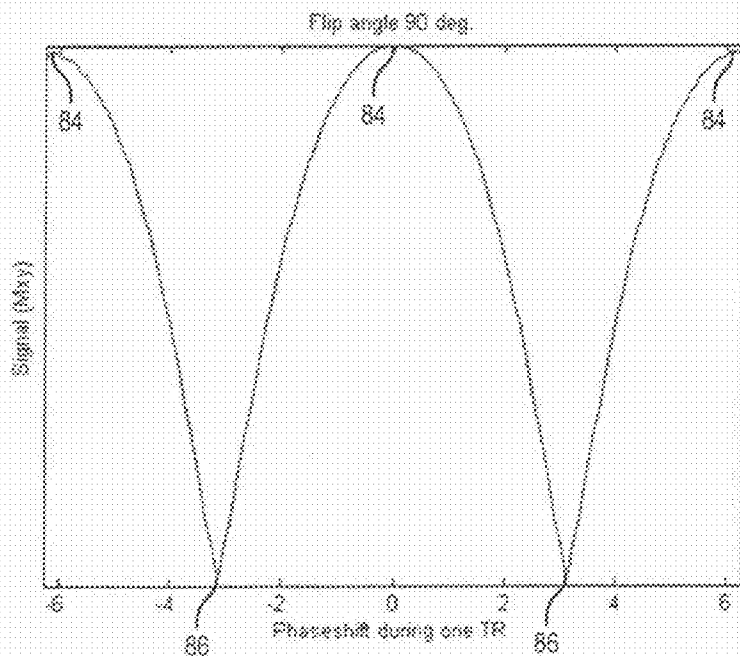
94

94

Flip angle 1 deg.

92

Longitudinal magnetization (Mz)

Phaseshift during one TR

FIG. 5

FIG. 6

FIG. 7

FIG. 8

FIG. 9

FIG. 10

FIG. 11

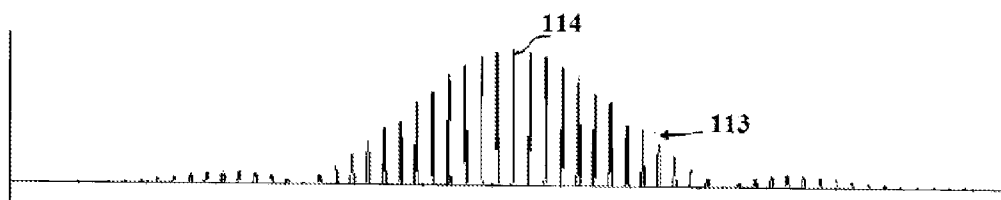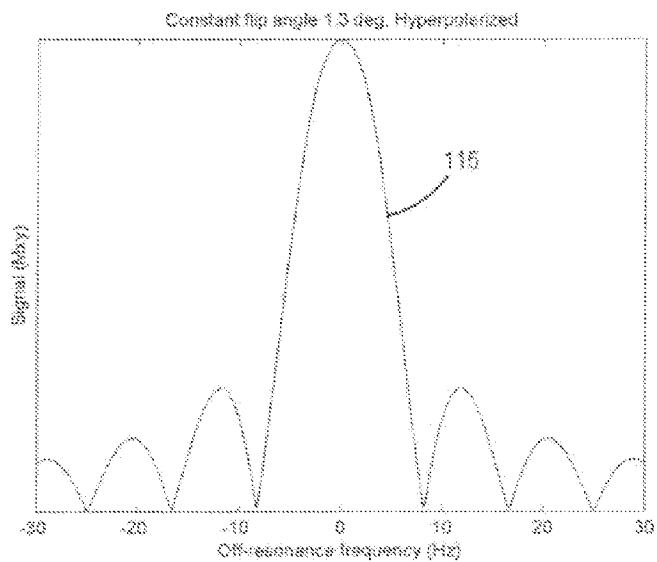

FIG. 12

FIG. 13

FIG. 14

DETERMINE LOW FLIP ANGLE
MODULATION SCHEME ⟋122

120

ADJUST EXCITATION PROFILE ⟋124

INTRODUCE HYPERPOLARIZED
CONTRAST AGENT ⟋136

EXCITE DESIRED METABOLIC
SPECIES ⟋128

ADJUST EXCITATION
136 ⟋ PROFILE / MODULATION
SCHEME

ACQUIRE SIGNALS ⟋130

Y   EXCITE DIFFERENT
METABOLIC SPECIES
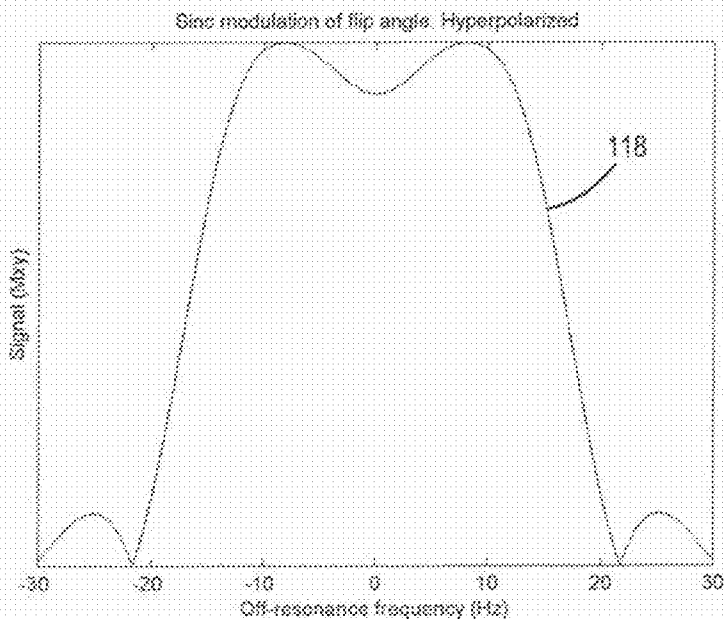? ⟋132

134

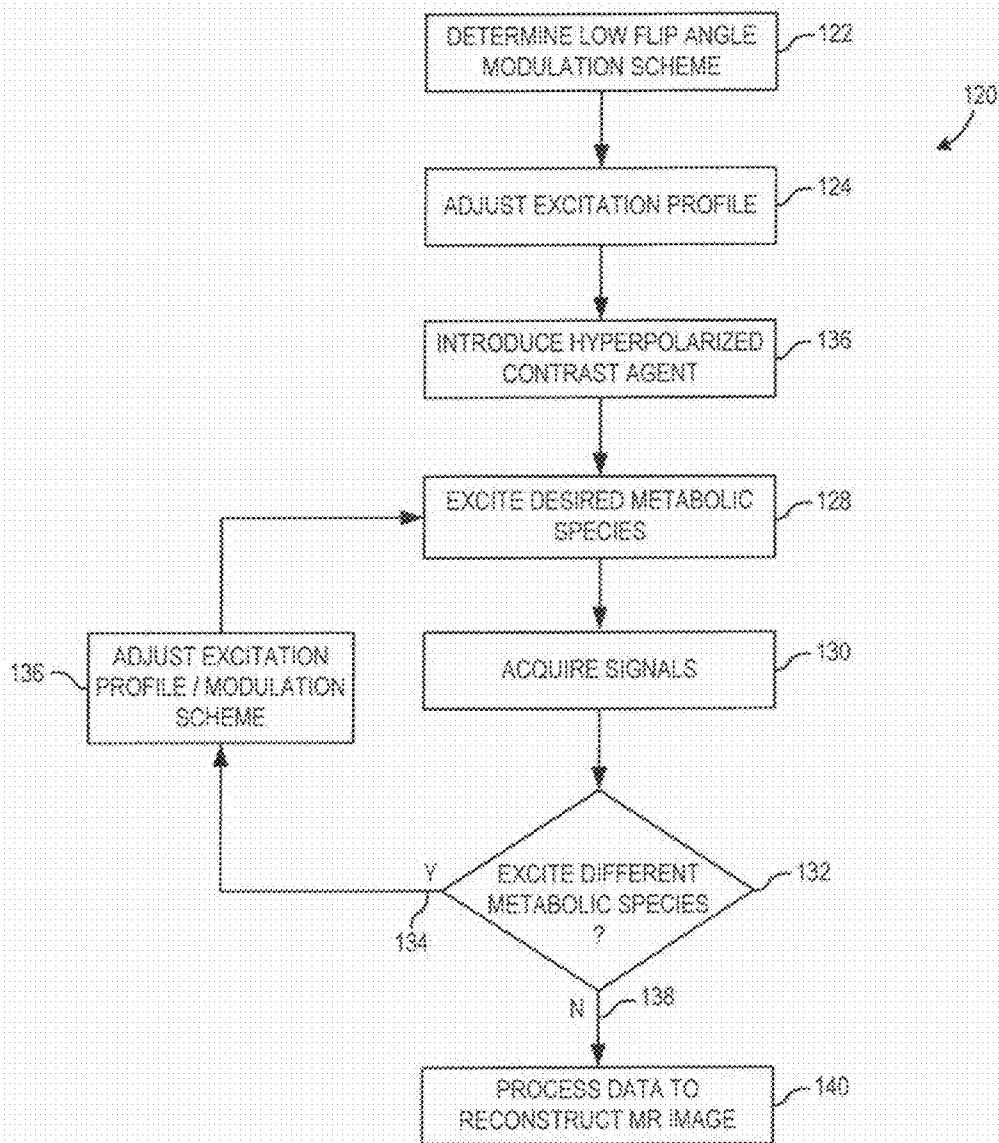N ⟋ 138

PROCESS DATA TO
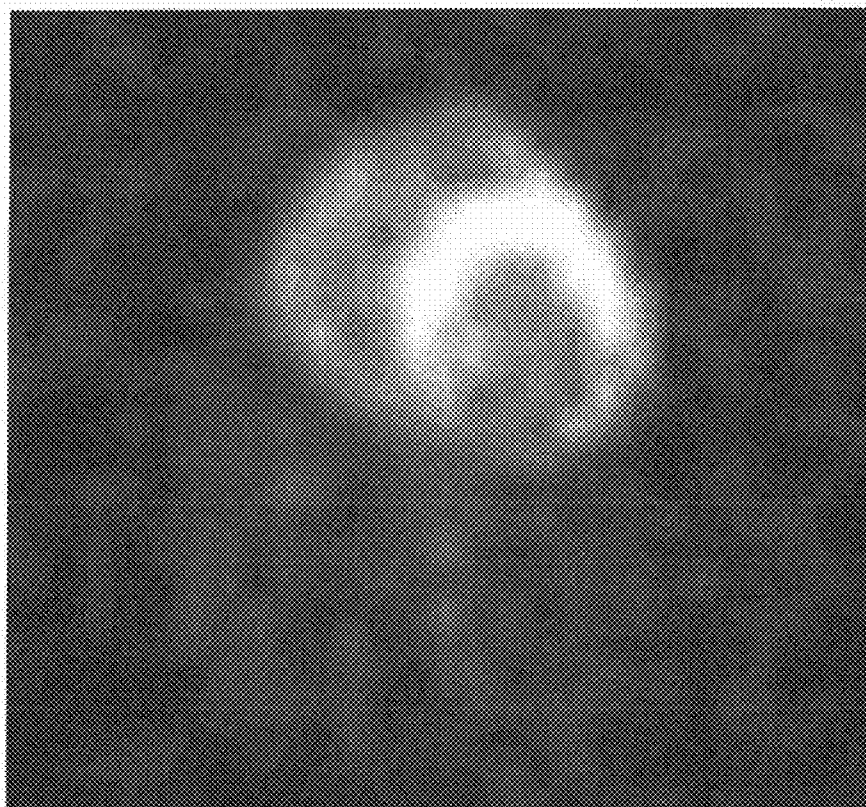RECONSTRUCT MR IMAGE ⟋140

FIG. 15

FIG. 16

SYSTEM AND METHOD FOR METABOLIC MR IMAGING OF A HYPERPOLARIZED AGENT

CROSS-REFERENCE TO RELATED APPLICATION

[0001] The present application is a continuation of and claims priority to U.S. patent application Ser. No. 11/623,277 filed Jan. 15, 2007, the disclosure of which is incorporated herein.

BACKGROUND OF THE INVENTION

[0002] The present invention relates generally to MR imaging and, more particularly, to metabolic MR imaging of a hyperpolarized agent.

[0003] When a substance such as human tissue is subjected to a uniform magnetic field (polarizing field B_0), the individual magnetic moments of the spins in the tissue attempt to align with this polarizing field, but precess about it in random order at their characteristic Larmor frequency. If the substance, or tissue, is subjected to a magnetic field (excitation field B_1) which is in the x-y plane and which frequency is near the Larmor frequency, the net aligned moment, or "longitudinal magnetization", M_Z, may be rotated, or "tipped", into the x-y plane to produce a net transverse magnetic moment M_t. A signal is generated by the excited spins after the excitation signal B_1 is terminated and this signal may be received and processed to form an image.

[0004] When utilizing these signals to produce images, magnetic field gradients (G_x, G_y, and G_z) are employed. Typically, the region to be imaged is scanned by a sequence of measurement cycles in which these gradients vary according to the particular localization method being used. The resulting set of received NMR signals are digitized and processed to reconstruct the image using one of many well known reconstruction techniques. It is desirable that the imaging process, from data acquisition to reconstruction, be performed as quickly as possible for improved patient comfort and throughput.

[0005] For some procedures and investigations, it is also desirable for MR images to display spectral information in addition to spatial information. One known method for acquiring MR signals and reconstructing MR images containing both spatial and spectral information is "chemical shift imaging" (CSI). CSI has been employed to monitor metabolic and other internal processes of patients, including imaging hyperpolarized substances such as ^{13}C labeled contrast agents and metabolites thereof. However, after injection of the hyperpolarized agent, imaging is a challenging task. The hyperpolarization of the agent has a limited lifetime, and imaging must be done rapidly. For example, typical T1 lifetimes of hyperpolarized agents are on the order of a few minutes in vivo. Furthermore, the RF excitations of the pulse sequence may destroy the hyperpolarization irreversibly.

[0006] The CSI method has some drawbacks which limit available signal-to-noise ratio, and thus image quality. For example, CSI tends to acquire data slowly, considering the short lifetimes of the increased magnetization of hyperpolarized substances. In addition, CSI typically exposes the imaging subject to a large number of RF excitations. These properties are especially unfavorable for a hyperpolarized agent because the hyperpolarized agent magnetization has a limited lifetime and is destroyed by the RF excitations of the CSI

sequence. As a consequence, the available magnetization cannot be fully utilized by the CSI method, and the signal-to-noise ratio (SNR) is thus reduced.

[0007] Additionally, MR procedures which require very fast, or periodic data acquisition, such as cardiac imaging or metabolic imaging of the heart, are difficult to perform with CSI sequences because CSI can take more than 15 seconds for a 16×16 matrix, whereas cardiac and related metabolic imaging should be completed within a few heartbeats or a few seconds.

[0008] It would therefore be desirable to have a system and method capable of exciting and imaging a metabolic species of a hyperpolarized agent without affecting magnetization of metabolic species at other frequencies.

BRIEF DESCRIPTION OF THE INVENTION

[0009] Embodiments of the present invention provide a system and method of MR that overcome the aforementioned drawbacks. A single metabolic species of a hyperpolarized agent injected into a subject of interest is excited, and MR signals from the excited single metabolic species are acquired. An image is reconstructed from the acquired MR signals.

[0010] Therefore, in accordance with one aspect of the present invention, an MRI apparatus includes an MRI assembly having a plurality of gradient coils positioned about a bore of a magnet to impress a polarizing magnetic field. An RF transceiver system and an RF switch are controlled by a pulse module to transmit and receive RF signals to and from an RF coil assembly to acquire MR images. The MRI apparatus also includes a system controller coupled to the MRI assembly, the system controller configured to cause the RF coil assembly to excite a single metabolic species of a hyperpolarized agent injected into a subject of interest. The system controller further causes the RF transceiver system to acquire MR signals from the excited single metabolic species and reconstruct an image from the acquired MR signals.

[0011] In accordance with another aspect of the invention, a method of hyperpolarized agent MR imaging includes injecting a hyperpolarized agent into a subject of interest and exciting a first metabolic species of the hyperpolarized agent. The method also includes acquiring MR signals from the excited first metabolic species and reconstructing an image from the acquired MR signals.

[0012] According to a further embodiment of the invention, a computer readable storage medium includes a computer program stored thereon comprising instructions which when executed by a computer, causes the computer to modulate a plurality of flip angle train RF pulses of a pulse sequence. The instructions further cause the computer to acquire MR data from the plurality of molecules and generate an image from the MR data.

[0013] Various other features and advantages will be made apparent from the following detailed description and the drawings.

BRIEF DESCRIPTION OF THE DRAWINGS

[0014] The drawings illustrate embodiments presently contemplated for carrying out the invention.

[0015] In the drawings:

[0016] FIG. 1 is a schematic block diagram of an MR imaging system for use with embodiments of the present invention.

[0017] FIG. 2 is a pulse sequence diagram of a trueFISP sequence with the slice gradient omitted.

[0018] FIG. 3 is a graph of the signal of the trueFISP sequence of FIG. 2 as a function of phase accumulation between two successive RF pulses with a constant flip angle of 90°.

[0019] FIG. 4 is a graph of the signal of the trueFISP sequence of FIG. 2 as a function of phase accumulation between two successive RF pulses with a constant flip angle of 1°.

[0020] FIG. 5 is a graph of the longitudinal magnetization of the trueFISP sequence of FIG. 2 as a function of phase accumulation between two successive RF pulses with a constant flip angle of 1°.

[0021] FIG. 6 is a modulation envelope of an RF pulse amplitude in accordance with an embodiment of the present invention.

[0022] FIG. 7 is a graph of trueFISP signals from hyperpolarized spins as function of phase shift per TR and RF pulse number.

[0023] FIG. 8 is another modulation envelope of the RF pulse amplitude in accordance with an embodiment of the present invention.

[0024] FIG. 9 is a pulse sequence diagram incorporating the pulse sequence modulation envelope of FIG. 8 in accordance with an embodiment of the present invention.

[0025] FIG. 10 is another modulation envelope of the RF pulse amplitude in accordance with an embodiment of the present invention.

[0026] FIG. 11 is a pulse sequence diagram incorporating the pulse sequence modulation envelope of FIG. 10 in accordance with an embodiment of the present invention.

[0027] FIG. 12 is a graph of frequency selectivity of a trueFISP sequence incorporating a constant amplitude of the RF pulses in accordance with an embodiment of the present invention.

[0028] FIG. 13 is a graph of frequency selectivity of a trueFISP sequence incorporating a Gaussian modulation envelope of the RF pulse amplitude in accordance with an embodiment of the present invention.

[0029] FIG. 14 is a graph of frequency selectivity of a trueFISP sequence incorporating a sinc modulation envelope of the RF pulse amplitude in accordance with an embodiment of the present invention.

[0030] FIG. 15 is a flowchart setting forth the steps of a technique in accordance with an embodiment of the present invention.

[0031] FIG. 16 is an exemplary image acquired in accordance with an embodiment of the present invention.

DETAILED DESCRIPTION OF THE INVENTION

[0032] Referring to FIG. 1, the major components of a preferred magnetic resonance imaging (MRI) system 10 that may incorporate embodiments of the present invention are shown. The operation of the system is controlled from an operator console 12 which includes a keyboard or other input device 13, a control panel 14, and a display screen 16. The console 12 communicates through a link 18 with a separate computer system 20 that enables an operator to control the production and display of images on the display screen 16. The computer system 20 includes a number of modules which communicate with each other through a backplane 20a. These include an image processor module 22, a CPU module 24 and a memory module 26, known in the art as a frame buffer for

storing image data arrays. The computer system 20 is linked to disk storage 28 and removable storage 30 for storage of image data and programs, and communicates with a separate system control 32 through a high speed serial link 34. The input device 13 can include a mouse, joystick, keyboard, track ball, touch activated screen, light wand, voice control, or any similar or equivalent input device, and may be used for interactive geometry prescription.

[0033] The system control 32 includes a set of modules connected together by a backplane 32a. These include a CPU module 36 and a pulse generator module 38 which connects to the operator console 12 through a serial link 40. It is through link 40 that the system control 32 receives commands from the operator to indicate the scan sequence that is to be performed. The pulse generator module 38 operates the system components to carry out the desired scan sequence and produces data which indicates the timing, strength and shape of the RF pulses produced, and the timing and length of the data acquisition window. The pulse generator module 38 connects to a set of gradient amplifiers 42, to indicate the timing and shape of the gradient pulses that are produced during the scan. The pulse generator module 38 can also receive patient data from a physiological acquisition controller 44 that receives signals from a number of different sensors connected to the patient, such as ECG signals from electrodes attached to the patient. And finally, the pulse generator module 38 connects to a scan room interface circuit 46 which receives signals from various sensors associated with the condition of the patient and the magnet system. It is also through the scan room interface circuit 46 that a patient positioning system 48 receives commands to move the patient to the desired position for the scan.

[0034] The gradient waveforms produced by the pulse generator module 38 are applied to the gradient amplifier system 42 having Gx, Gy, and Gz amplifiers. Each gradient amplifier excites a corresponding physical gradient coil in a gradient coil assembly generally designated 50 to produce the magnetic field gradients used for spatially encoding acquired signals. The gradient coil assembly 50 forms part of a magnet assembly 52 which includes a polarizing magnet 54 and a whole-body RF coil 56. A transceiver module 58 in the system control 32 produces pulses which are amplified by an RF amplifier 60 and coupled to the RF coil 56 by a transmit/receive switch 62. The resulting signals emitted by the excited nuclei in the patient may be sensed by the same RF coil 56 and coupled through the transmit/receive switch 62 to a preamplifier 64. The amplified MR signals are demodulated, filtered, and digitized in the receiver section of the transceiver 58. The transmit/receive switch 62 is controlled by a signal from the pulse generator module 38 to electrically connect the RF amplifier 60 to the coil 56 during the transmit mode and to connect the preamplifier 64 to the coil 56 during the receive mode. The transmit/receive switch 62 can also enable a separate RF coil (for example, a surface coil) to be used in either transmit or receive mode.

[0035] The MR signals picked up by the RF coil 56 are digitized by the transceiver module 58 and transferred to a memory module 66 in the system control 32. A scan is complete when an array of raw k-space data has been acquired in the memory module 66. This raw k-space data is rearranged into separate k-space data arrays for each image to be reconstructed, and each of these is input to an array processor 68 which operates to Fourier transform the data into an array of image data. This image data is conveyed through the serial

FIGS. 12-14 will cover a broader frequency range. Furthermore, calculations were made for a sequence with 96 phase-encoding steps and a total flip angle of 120°. The signal responses shown in FIGS. 12-14 are shown after 48 RF pulses, i.e., at the center of the k-space, with a linear phase-encoding scheme.

[0048] FIG. 12 shows that, with constant RF amplitude at a flip angle of 1.3°, the usable frequency range for a reasonably uniform signal intensity is approximately 5 Hz, or approximately $\frac{1}{3}$ ppm at 1.5 T. FIG. 13 shows that, with Gaussian amplitude modulation, the usable frequency range for a reasonable uniform signal intensity is approximately double the usable frequency range shown in FIG. 12. FIG. 14 shows that, with a sinc function modulation, the usable frequency range for a reasonably uniform signal intensity is approximately 30 Hz, or approximately 2 ppm, at 1.5 T.

[0049] As shown in FIG. 14, imaging of a single hyperpolarized agent metabolite should be possible without additional shimming actions given that other metabolites of interest are sufficiently far away from the excited frequency range. Further optimization of the excitation profiles shown in FIGS. 12-14 can be achieved by adjusting the modulation profile with the respect to the position in the k-space, e.g., adjusting the maximum of the modulation profile such that it does not coincide with the passage of the k-space center.

[0050] Referring now to FIG. 15, a hyperpolarized agent imaging technique 120 in accordance with one embodiment of the present invention is shown. The technique 120 begins with determining, at block 122, a low flip angle modulation scheme for the RF pulse sequence of a modified trueFISP pulse sequence. Determination of the low flip angle modulation scheme includes basing the determination on a desired frequency selectivity as described above as well as on the shimming, or the B_0 homogeneity. The low flip angle modulation scheme is based on a constant amplitude envelope, a Gaussian modulation envelope, a sinc modulation envelope, or the like. Additionally, determination of the low flip angle modulation scheme may include a consideration of the hyperpolarized contrast agent. In a preferred embodiment, the hyperpolarized contrast agent includes ^{13}C nuclei; however, it is contemplated that other hyperpolarized contrast agents may be used, such as ^{14}N, ^{31}P, ^{19}F, and ^{23}Na nuclei, other NMR relevant nuclei.

[0051] Following the determination at block 122 of the low flip angle modulation scheme, an excitation profile of the modified trueFISP pulse sequence is adjusted at block 124 for excitation of a single metabolic species of the hyperpolarized contrast agent. For example, a hyperpolarized ^{13}C-pyruvate may be injected into an imaging subject, and the excitation profile may be adjusted to excite ^{13}C-bicarbonate nuclei. The hyperpolarized contrast agent is then introduced into the imaging subject at block 126. Next, the modified trueFISP pulse sequence excites the desired metabolic species at block 128. This excitation may be delayed a specific time period after introduction of the agent to allow for perfusion into tissues, or for the agent to reach an organ of diagnostic interest. Alternatively, a period of delay may correspond to an amount of time for the contrast agent to be metabolized. Signals are then acquired from the excited metabolic species at block 130.

[0052] Technique 120 then determines at block 132 whether to excite a different metabolic species for acquiring signals therefrom. If so 134, then the excitation profile is adjusted at block 136 to excite the nuclei for another metabo-

lite. The modulation scheme of the RF pulse sequence may also be adjusted, if desired, such that the modulation scheme for one metabolite is distinct from the modulation scheme for another metabolite. Technique 120 then excites and acquires signals from the other metabolite as described above in blocks 128 and 130. If not 138, MR images are reconstructed for the acquired signal data at block 140. In a preferred embodiment, an image is reconstructed for each metabolite acquired. However, it is contemplated that a combined image may be reconstructed for all metabolites acquired.

[0053] FIG. 16 shows an exemplary in vivo trueFISP image of a pig heart, acquired with a constant flip angle of 2°. Hyperpolarized ^{13}C-pyruvate was injected intravenously, and images were acquired with the excitation profile adjusted to the resonance frequency of ^{13}C-bicarbonate, which is 160 Hz apart from the resonance frequency of pyruvate. FIG. 16 shows the sum of five consecutive images, acquired with 9-Hz frequency separation around the bicarbonate resonance.

[0054] Therefore, in accordance with one embodiment of the present invention, an MRI apparatus includes an MRI assembly having a plurality of gradient coils positioned about a bore of a magnet to impress a polarizing magnetic field. An RF transceiver system and an RF switch are controlled by a pulse module to transmit and receive RF signals to and from an RF coil assembly to acquire MR images. The MRI apparatus also includes a system controller coupled to the MRI assembly, the system controller configured to cause the RF coil assembly to excite a single metabolic species of a hyperpolarized agent injected into a subject of interest. The system controller further causes the RF transceiver system to acquire MR signals from the excited single metabolic species and reconstruct an image from the acquired MR signals.

[0055] In accordance with another embodiment of the invention, a method of hyperpolarized agent MR imaging includes injecting a hyperpolarized agent into a subject of interest and exciting a first metabolic species of the hyperpolarized agent. The method also includes acquiring MR signals from the excited first metabolic species and reconstructing an image from the acquired MR signals.

[0056] According to a further embodiment of the invention, a computer readable storage medium includes a computer program stored thereon comprising instructions which when executed by a computer, causes the computer to modulate a plurality of flip angle train RF pulses of a pulse sequence. The instructions further cause the computer to acquire MR data from the plurality of molecules and generate an image from the MR data.

[0057] The present invention has been described in terms of the preferred embodiment, and it is recognized that equivalents, alternatives, and modifications, aside from those expressly stated, are possible and within the scope of the appending claims.

What is claimed is:

1. An MRI apparatus comprising:

a magnetic resonance imaging (MRI) assembly having a plurality of gradient coils positioned about a bore of a magnet to impress a polarizing magnetic field and an RF transceiver system and an RF switch controlled by a pulse module to transmit RF signals to an RF coil assembly to acquire MR images; and

a system controller coupled to the MRI assembly, the system controller configured to:
  cause the RF coil assembly to excite a single metabolic species of a hyperpolarized agent injected into a subject of interest;
  cause the RF transceiver system to acquire MR signals from the excited single metabolic species; and
  reconstruct an image from the acquired MR signals.

2. The MRI apparatus of claim 1 wherein the excitation of the single metabolic species excites the single metabolic species only.

3. The MRI apparatus of claim 1 wherein the excited single metabolic species comprises a metabolite of the hyperpolarized agent injected into the subject of interest.

4. The MRI apparatus of claim 1 wherein the system controller is further configured to repeat the excitation and the acquisition for each single metabolic species of the hyperpolarized agent.

5. The MRI apparatus of claim 1 wherein the single metabolic species comprises a single metabolic species of 13C.

6. A method of hyperpolarized agent MR imaging comprising the steps of:
  injecting a hyperpolarized agent into a subject of interest;
  exciting only a first metabolic species of the hyperpolarized agent;
  acquiring MR signals from the excited first metabolic species; and
  reconstructing an image from the acquired MR signals.

7. The method of claim 6 wherein the excitation of only the first metabolic species does not excite other metabolic species of the hyperpolarized agent.

8. The method of claim 6 wherein the excitation of the first metabolic species comprises a metabolite of the hyperpolarized agent injected into the subject of interest.

9. The method of claim 6 wherein the step of exciting comprises:
  modifying a flip angle train according to a first low angle modulation pattern; and
  applying a pulse sequence having the modified flip angle train to the first metabolic species.

10. The method of claim 9 wherein the step of modifying comprises modifying the flip angle train according to a constant amplitude low angle modulation pattern having amplitudes less than 2 degrees.

11. The method of claim 9 wherein the step of modifying comprises modifying the flip angle train according to a Gaussian modulated low angle modulation pattern.

12. The method of claim 9 wherein the step of modifying comprises modifying the flip angle train according to a sinc function modulated low angle modulation pattern.

13. The method of claim 9 further comprising the steps of:
  exciting only a second metabolic species of the hyperpolarized agent; and
  acquiring MR signals from the excited second metabolic species.

14. The method of claim 13 wherein the step of exciting the second metabolic species comprises:
  modifying the flip angle train according to a second low angle modulation pattern distinct from the first low angle modulation pattern; and
  applying a pulse sequence having the modified flip angle train according to the second low angle modulation pattern to the second metabolic species.

15. A non-transitory computer readable storage medium having stored thereon a computer program comprising instructions which when executed by a computer, causes the computer to:
  modulate a plurality of flip angle train RF pulses of a pulse sequence;
  excite, with the pulse sequence, a plurality of molecules of a single hyperpolarized metabolite within an object;
  acquire MR data from the plurality of molecules; and
  generate an image from the MR data.

16. The computer readable storage medium of claim 15 wherein the instructions, in exciting of the plurality of molecules of the single hyperpolarized metabolite with the pulse sequence, do not excite molecules of other hyperpolarized metabolites within the object.

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