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Author manuscript

*Magn Reson Imaging*. Author manuscript; available in PMC 2018 May 01.

Published in final edited form as:

*Magn Reson Imaging*. 2017 May ; 38: 63–70. doi:10.1016/j.mri.2016.12.018.

## A Simple Analytic Method for Estimating T<sub>2</sub> in the Knee from DESS

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### Abstract

**Purpose**—To introduce a simple analytical formula for estimating T<sub>2</sub> from a single Double-Echo in Steady-State (DESS) scan.

**Methods**—Extended Phase Graph (EPG) modeling was used to develop a straightforward linear approximation of the relationship between the two DESS signals, enabling accurate T<sub>2</sub> estimation from one DESS scan. Simulations were performed to demonstrate cancellation of different echo pathways to validate this simple model. The resulting analytic formula was compared to previous methods for T<sub>2</sub> estimation using DESS and fast spin-echo scans in agar phantoms and knee cartilage in three volunteers and three patients. The DESS approach allows 3D (256×256×44) T<sub>2</sub>-mapping with fat suppression in scan times of 3–4 minutes.

**Results**—The simulations demonstrated that the model approximates the true signal very well. If the T<sub>1</sub> is within 20% of the assumed T<sub>1</sub>, the T<sub>2</sub> estimation error was shown to be less than 5% for typical scans. The inherent residual error in the model was demonstrated to be small both due to signal decay and opposing signal contributions. The estimated T<sub>2</sub> from the linear relationship agrees well with reference scans, both for the phantoms and in vivo. The method resulted in less underestimation of T<sub>2</sub> than previous single-scan approaches, with processing times 60 times faster than using a numerical fit.

**Conclusion**—A simplified relationship between the two DESS signals allows for rapid 3D T<sub>2</sub> quantification with DESS that is accurate, yet also simple. The simplicity of the method allows for immediate T<sub>2</sub> estimation in cartilage during the MRI examination.

### Keywords

DESS; T<sub>2</sub>; Cartilage; Osteoarthritis

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## 1. Introduction

The Double-Echo Steady-State (DESS) sequence offers distortion-free, SNR-efficient 3D-imaging with good contrast flexibility [1–3]. DESS is a gradient-spoiled steady-state sequence, collecting two echoes per repetition (TR), with the spoiler gradient separating the two echoes. The two signals, often labeled  $S_1$  (before the spoiler) and  $S_2$  (after the spoiler), have different contrast. The  $S_1$  signal is often said to have  $T_1/T_2$  weighting and  $S_2$  to have a mixture of  $T_2$  and diffusion weighting, but in reality both the contrasts are quite complicated functions of the extrinsic scan parameters (TR, TE, flip angle  $\alpha$ , gradient amplitude  $G$  and gradient duration  $\tau$ ) as well as the intrinsic tissue parameters ( $T_1$ ,  $T_2$ , and diffusivity). While DESS has been used for various anatomies such as the breast [4,5] and the prostate [6,7], it has found particularly widespread use for musculoskeletal imaging, where for example it has been used in the Osteoarthritis Initiative (OAI) to determine Osteoarthritis (OA) progression [8].

To improve OA detection, DESS has been used for quantitative imaging of  $T_2$  and apparent diffusion coefficient (ADC) [9–12]. These have been found to correlate with collagen network organization and proteoglycan concentration, respectively [13,14]. A number of signal models have been developed to quantitatively describe both of the DESS signals and their dependency on tissue parameters. These include both closed-form and non-closed-form analytical methods as well as numerical methods [15–20].

Some methods model the DESS signals completely and can be considered fully accurate for a single  $T_1$ ,  $T_2$ , and diffusivity in a voxel [16,17]. However, owing to their complexity, using these models for parameter estimation typically requires solving them with a numerical search over a predetermined solution set. This inhibits immediate, automatic creation of parameter maps right at the scanner, which would be of great value for clinical scans. Furthermore, the method in ref. [16], which describes the magnetization with a continued fraction, does not allow for pulsed gradients. For these reasons, approximate models are often used to estimate the desired parameter in spite of known errors. For example, in ref. [9], the following model, originally suggested in ref. [1], was used to estimate  $T_2$  from a single DESS scan:

$$\frac{S_2}{S_1} = e^{-\frac{2(TR-TE)}{T_2}} \quad (1)$$

This relationship essentially describes a spin echo relationship between  $S_2$  and  $S_1$  in consecutive TRs. Although this approach may offer useful results in spite of errors in  $T_2$ , practical  $T_2$  mapping would benefit from a comparably simple analytical formula that is even more accurate.

In this work, we introduce a new model that can be used for  $T_2$  estimation from a single DESS scan. Our analysis demonstrates that higher-order echo pathways can often be ignored, based on pathway cancellation effects and signal decay, leading to a simple and accurate linear relationship between the signals from two DESS echoes. We validate the overall approach using simulations, phantom scans and in vivo knee scans.

## 2. Theory

### 2.1 Derivation of Simplified Signal Ratio

A simple yet accurate linear approximation of the relationship between the two DESS echoes can be derived by tracing the components that contribute to the magnetization, starting at  $S_2$ , until we arrive at  $S_1$ . We use the Extended Phase Graph (EPG) formalism, which decomposes magnetization into a basis of positively and negatively dephased transverse states (coefficients  $F_0, F_1, F_2, \dots$ ) as well as sinusoidally varying longitudinal states (coefficients  $Z_0, Z_1, Z_2, \dots$ ) [20,21]. The procedure is explained graphically in Fig. 1a–c, using similar graphical structures as those introduced by Scheffler et al. [18]. As explained by Weigel et al. [17,21], in the EPG formalism, the magnetization can be represented by a matrix of coefficients ( $F_n$  and  $Z_n$ ) that multiply transverse states with  $n$  cycles of positive or negative dephasing, and longitudinal states with  $n$  cycles of sinusoidal variation:

$$M = \begin{bmatrix} F_0 & F_1 & F_2 & \dots \\ F_0^* & F_{-1}^* & F_{-2}^* & \dots \\ Z_0 & Z_1 & Z_2 & \dots \end{bmatrix}, \quad (2)$$

where an asterisk (\*) denotes a complex conjugate. The observable signal is the DC transverse term  $F_0$ . Relaxation and diffusion over time are represented by scaling of the elements ( $T_1$  relaxation is represented by an additive factor to the DC longitudinal term,  $Z_0$ ) and the effect of a gradient is represented by shifting the transverse coefficients  $F_n$ .

The effects of  $T_1/T_2$  decay, as well as diffusion, on state  $n$ , can be described by the scaling terms  $\tilde{E}_{2,n} = e^{-\frac{TR}{T_2} - [(n^2 + n + \frac{1}{2})TR - \frac{\tau}{6}] \Delta k^2 D}$  and  $\tilde{E}_{1,n} = e^{-\frac{TR}{T_1} - n^2 TR \Delta k^2 D}$  (assuming the gradient in the center of the TR) [17]. The effect of an RF pulse with flip angle  $\alpha$  and phase  $\phi$  is represented by matrix multiplication with the matrix

$$R(\alpha, \phi) = \begin{bmatrix} \cos^2 \frac{\alpha}{2} & e^{2i\phi} \sin^2 \frac{\alpha}{2} & -ie^{i\phi} \sin \alpha \\ e^{-2i\phi} \sin^2 \frac{\alpha}{2} & \cos^2 \frac{\alpha}{2} & ie^{-i\phi} \sin \alpha \\ -\frac{i}{2} e^{-i\phi} \sin \alpha & \frac{i}{2} e^{i\phi} \sin \alpha & \cos \alpha \end{bmatrix} \quad (3)$$

Here, TR and TE represent the repetition and echo time, respectively,  $\alpha$  is the flip angle,  $D$  is the diffusivity, and the dephasing per unit length induced by the unbalanced gradient is denoted by  $k = \gamma G \tau$ , where  $G$  and  $\tau$  are the spoiler amplitude and duration, respectively, and  $\gamma$  is the gyromagnetic ratio. We label the states immediately before/after the RF pulse with  $-/+$  superscripts, respectively. To determine a relationship between the two DESS signals, we look at the echo pathway of the observable signal immediately before the pulse,  $F_0^-$ , and write down the components that contributed to it. We repeat this until we reach the observable signal immediately after the pulse,  $F_0^+$  (Fig. 1). As justified in Section 2.2, we will ignore signal contributions from states that have experienced two cycles of dephasing or

more, as was done for the  $S_2$  signal in ref. [22]. We will assume  $\phi=90^\circ$ , resulting in real valued states, without loss of generality.

$$\begin{aligned} F_0^- &= \tilde{E}_{2,-1} F_{-1}^+ = \tilde{E}_{2,-1} \left( -\sin^2 \frac{\alpha}{2} F_1^- + \cos^2 \frac{\alpha}{2} F_{-1}^- - \sin \alpha Z_1^- \right) \\ &= -\tilde{E}_{2,-1} \left( \sin^2 \frac{\alpha}{2} \tilde{E}_{2,0} F_0^+ - \cos^2 \frac{\alpha}{2} \tilde{E}_{2,-2} F_{-2}^+ + \sin \alpha \tilde{E}_{1,1} Z_1^+ \right) \\ &\approx -\tilde{E}_{2,-1} \left( \sin^2 \frac{\alpha}{2} \tilde{E}_{2,0} F_0^+ + \sin \alpha \tilde{E}_{1,1} Z_1^+ \right) \end{aligned} \quad (4a)$$

$$\begin{aligned} Z_1^+ &= \frac{1}{2} \sin \alpha F_1^- + \frac{1}{2} \sin \alpha F_{-1}^- + \cos \alpha Z_1^- \\ &= \frac{1}{2} \sin \alpha \tilde{E}_{2,0} F_0^+ + \frac{1}{2} \sin \alpha \tilde{E}_{2,-2} F_{-2}^- + \cos \alpha \tilde{E}_{1,1} Z_1^- \\ &\approx \frac{1}{2} \sin \alpha \tilde{E}_{2,0} F_0^+ + \cos \alpha \tilde{E}_{1,1} Z_1^- \\ &\Rightarrow Z_1^+ \approx \frac{1}{2} \frac{\sin \alpha \tilde{E}_{2,0}}{1 - \cos \alpha \tilde{E}_{1,1}} F_0^+ \end{aligned} \quad (4b)$$

Combining Eqs. (4a–b) and using some algebra and trigonometric identities yields

$$\frac{F_0^-}{F_0^+} \approx -\tilde{E}_{2,-1} \tilde{E}_{2,0} \sin^2 \left( \frac{\alpha}{2} \right) \frac{1 + \tilde{E}_{1,1}}{1 - \cos \alpha \tilde{E}_{1,1}} \quad (5)$$

Using the stated expressions for the decay factors as well as the fact that the observed signal magnitudes are  $S_1 = F_0 e^{-\frac{TE}{T_2}}$  and  $S_2 = -\tilde{E}_{2,-1} F_{-1} e^{\frac{TE}{T_2}} e^{-\frac{TE}{T_2}'}$  (where we have used that the two signals are out of phase and the second signal experiences  $T_2'$  rephasing), we arrive at the following expression:

$$\frac{S_2}{S_1} = e^{-\frac{2(TR-TE)}{T_2} - (TR - \frac{\tau}{3}) \Delta k^2 D} \sin^2 \left( \frac{\alpha}{2} \right) \left( \frac{1 + e^{-\frac{TR}{T_1} - TR \Delta k^2 D}}{1 - \cos \alpha e^{-\frac{TR}{T_1} - TR \Delta k^2 D}} \right) \quad (6)$$

This relationship, which assumes the echoes are acquired symmetrically during the TR (with the same duration from the RF pulse to  $S_1$  as from  $S_2$  to the subsequent RF pulse), provides a simple analytic estimate of  $T_2$  given an estimate of  $T_1$  and either minimal diffusion sensitivity or an estimate of diffusivity.

In the case of low spoiling ( $k \approx 0$ ), the model reduces to the following relationship:

$$\frac{S_2}{S_1} = e^{-\frac{2(TR-TE)}{T_2}} \sin^2 \left( \frac{\alpha}{2} \right) \left( \frac{1 + e^{-\frac{TR}{T_1}}}{1 - \cos \alpha e^{-\frac{TR}{T_1}}} \right) \quad (7)$$

For  $\alpha = 90^\circ$ , this further reduces to

$$\frac{S_2}{S_1} = e^{-\frac{2(TR-TE)}{T_2}} \frac{1 + e^{-\frac{TR}{T_1}}}{2} \quad (8)$$

In the case of very long  $T_1$  Eq. (7) becomes independent of flip angle and results in Eq. (1). This equation was presented by Bruder et al. and has been used by Welsch et al. for  $T_2$  estimation in cartilage [1,9]. However, this relationship results in errors unless the flip angle is close to  $90^\circ$  or  $TR/T_1$  is very small. The relationship also assumes weak spoiling so that diffusion effects may be ignored. Eq. (6) better approximates the signal over a large range of  $T_1$ , flip angle  $\alpha$ , and (zeroth) spoiler gradient moments. This is demonstrated in Fig. 2a, which shows the approximated signal from Eqs. (1) and (6), as well as highly accurate numerical computation of the signal using EPG matrices. The signal ratio is modeled with a typical  $T_1$  value of 1.2 s for cartilage [23], assuming both weak and strong spoiling (with spoiler moments of 0.001 and 156.6 mT/m•ms, respectively), as well as a long  $T_1$  value of 5 s for the weakly spoiled case, and a TR of 22.5 ms. Other parameters were TE = 6.5 ms,  $T_2 = 40$  ms, gradient duration  $\tau = 3.4$  ms, and ADC =  $1.6 \cdot 10^{-3}$  mm/s. In order for Eq. (1) to be valid with smaller flip angles (about  $20^\circ$ – $60^\circ$ ),  $T_1$  clearly needs to be much larger than TR.

## 2.2 Cancellation of higher-order echo pathways

The model in Eq. (6) performs well for two reasons. The equation neglects the contribution of echo pathways between  $S_1$  and  $S_2$  that have spent more than two TRs in the transverse plane. This is partly justified by simple signal decay – the more time that the magnetization spends in the transverse plane, the more it will be attenuated both by  $T_2$  relaxation and diffusion. However, a more important mechanism is the cancellation of echo pathways due to opposing phase, independent of relaxation. It is easy to show that all contributions to  $S_2$  that have experienced two transverse TRs will have negative phase relative to  $S_1$ . However, this does not hold for higher order echo pathways. For example, if we denote TRs where the magnetization starts in the  $+/-$   $n^{\text{th}}$  transverse state or the  $n^{\text{th}}$  longitudinal state by  $F_{+/-n}$  and  $Z_n$ , then the pathways  $F_0 F_1 F_{-2} F_{-1}$  and  $F_0 Z_1 F_1 F_{-2} F_{-1}$  will have opposite sign. By accumulating the contributions of all such pathways, the positive and negative contributions (relative to  $S_1$ ) of higher-order pathways can be modeled and compared. For example, by including pathways with two and four transverse TRs, Eq. (6) can be extended to

$$\frac{S_2}{S_1} = B_{2-} - B_{4+} + B_{4-} \quad (9)$$

where  $B_{2-}$  is the two-TR component given by Eq. (6), and  $B_{4+}$  and  $B_{4-}$  are the four-TR components with the same and opposite phase to  $S_1$ , respectively, given by

$$B_{4+} = 2CDEF \quad (10)$$

$$B_{4-} = C(D^2 + E^2)F \quad (11)$$

where

$$C = e^{-\frac{4TR-2TE}{T_2} - (6TR - \frac{2r}{3})\Delta k^2 D} \quad (12)$$

$$D = \cos^2\left(\frac{\alpha}{2}\right) \quad (13)$$

$$E = \frac{\frac{1}{2}\sin^2\alpha e^{-\frac{TR}{T_1} - TR\Delta k^2 D}}{1 - \cos\alpha e^{-\frac{TR}{T_1} - TR\Delta k^2 D}} \quad (14)$$

$$F = \sin^2\left(\frac{\alpha}{2}\right) + \frac{\frac{1}{2}\sin^2\alpha e^{-\frac{TR}{T_1} - 4TR\Delta k^2 D}}{1 - \cos\alpha e^{-\frac{TR}{T_1} - 4TR\Delta k^2 D}} \quad (15)$$

Fig. 2b displays the components  $B_{n+}$  and  $B_{n-}$  for  $n = 2-12$  over a range of flip angles with the same parameters as the green curve in Fig. 2a. The higher-order components with positive and negative phase have very similar magnitude. Therefore, they will have a negligible effect on the final answer, particularly for flip angles above  $20^\circ$  or short  $T_2$ . This is demonstrated in Fig. 2c. This leads to Eq. (6) simulating the signal ratio  $S_2/S_1$  very accurately, even in cases where there is very little  $T_2$  decay, such as in fluid. Fig. 2d shows the same simulation as Fig. 2c, but for synovial fluid at 3T, having  $T_1 = 3.6$  s and  $T_2 = 0.77$  s [23]. The higher-order pathways contribute more to the signal ratio in this case, especially for low flip angles, but Eq. (6) nonetheless approximates the true signal well for flip angles above  $20^\circ$ . The approximation of neglecting higher-order states was also used by Buxton [22]. In that work, only the  $S_2$  signal was modeled. If  $S_1$  in Eq. (6) is assumed to have no transverse pathway contributions, the two models agree, as further explained in the Discussion.

### 3. Methods

All scans were performed on a 3T 750 scanner (GE Healthcare, Waukesha, WI, USA). Informed consent was obtained from all subjects in accordance with the Institutional Review Board protocol at our institution. The volunteer scans used a 16-channel receive coil wrapped around the knee (GEM Flex by NeoCoil, Waukesha, WI, USA), the phantom scans used a single-channel transmit-receive wrist coil (GE BC-10 by Mayo Clinic, Rochester

MN, USA), and the patient scans used an 8-channel transmit-receive knee coil (Knee Array Coil by Invivo, Gainesville, FL, USA).

### 3.1 Phantom scans

Three agar phantoms were scanned using DESS with scan parameters  $TR=13.4$  ms,  $TE=3.9$  ms, voxel size  $0.47 \times 0.47 \times 3.0$  mm<sup>3</sup>, spoiler moment  $7.83$  mT/m•ms and duration  $\tau=3.4$  ms, and flip angles  $\alpha = 10^\circ, 15^\circ, 20^\circ, 30^\circ, 40^\circ,$  and  $50^\circ$ . Fast spin-echo (FSE) scans with echo times  $TE = 17, 26, 34, 43, 51, 60,$  and  $68$  ms,  $TR = 1000$  ms and voxel size  $0.47 \times 0.47 \times 6.0$  mm<sup>3</sup> were performed and fitted to a monoexponential curve to obtain a reference  $T_2$ . The  $T_2$  value was estimated from the DESS scans using Eqs. (1) and (7) and compared to the value resulting from the reference FSE scans. For Eq. (7), reference  $T_1$  values known to be  $T_{1-1} = 2000$  ms,  $T_{1-2} = 1700$  ms, and  $T_{1-3} = 1600$  ms, measured with a 3TI MP-RAGE technique [24], and a flip angle measured with the Bloch-Siegert method [25], were used for processing each phantom.

### 3.2 Volunteer scans

To test Eq. (7) in vivo, a sagittal DESS scan was performed in the knee of three healthy volunteers. The scans used a spoiler with a moment of  $15.66$  mT/m•ms and duration  $3.4$  ms. Other parameters were  $\alpha=25^\circ$ ,  $TR=22.5$  ms,  $TE=6.5$  ms, slice thickness  $3.0$  mm. A spectrally selective RF pulse was used. The  $T_2$  of the cartilage was again estimated using the linear approximations of Eqs. (1) and (7). Reference FSE scans were acquired with  $TE = 9, 18, 27, 36, 45, 54, 63,$  and  $72$  ms and  $TR = 1000$  ms. Both scans were run with  $FOV = 13\text{cm} \times 13\text{cm}$  and acquisition matrix  $384 \times 384$ , which was automatically interpolated to  $512 \times 512$  by the scanner. For  $T_2$  quantification using Eq. (7),  $T_1=1.2$  s, typical for cartilage at 3T [23], and the prescribed flip angle of  $\alpha=25^\circ$  were used. Eq. (7) has low sensitivity to  $T_1$ , so small errors in the  $T_1$  assumption should not cause large errors in estimated  $T_2$ . Furthermore,  $T_1$  in knee cartilage does not change much whereas  $T_2$  is sensitive to degeneration. For comparison, a  $T_2$  map using a full numerical fit was produced from the DESS scan as well. This involved computing a dictionary of signal ratios for  $T_2$  ranging between  $10$ – $100$  ms (with a step size of  $1$  ms) using large EPG matrices (with  $6$  states) and then choosing the dictionary entry that best fitted each pixel. The total scan time was  $6:26$  min for FSE (acquiring  $6$  slices) and  $5:11$  min for DESS (acquiring  $36$  slices). In slices where both scans displayed clear anterior and posterior regions of the femoral cartilage, the resulting  $T_2$  maps in those two regions were divided into deep and superficial regions, resulting in  $4$  regions per slice. This was possible in  $10$  slices, resulting in a total of  $40$  regions. The root-mean-squared (RMS) difference between the results from Eq. (7) and the FSE results was computed and compared to the difference between the results from Eq. (1) and the reference scans. The  $T_2$  maps were computed in Matlab (The MathWorks, Natick, Ma) on a Linux terminal with  $128$  GB of RAM and  $32$   $2.6$  GHz CPUs.

### 3.3 Patient scans

As an initial demonstration of the value of Eq. (7) for routine diagnostic imaging, we selected data from three patients from a larger group scanned with DESS in a clinical setting (spoiler moment =  $31.32$  mT/m•ms, spoiler duration =  $3.4$  ms,  $\alpha=20^\circ$ ,  $TR=20.4$  ms,  $TE=6.4$  ms,  $FOV = 16\text{cm} \times 16\text{cm}$ , acquisition matrix  $416 \times 512$ ,  $80$  slices with thickness of  $1.6\text{mm}$

interpolated to 0.8 mm (160 slices), 2× parallel imaging in the phase encode direction, scan time = 5:00).  $T_2$  maps were constructed automatically at the scanner. A proton-density (PD) weighted scan was acquired for comparison.

## 4. Results

### 4.1 Phantom scans

The results from the phantom scans are shown in Fig. 3. The measured reference  $T_2$  values were  $T_{2-1} = 62$  ms,  $T_{2-2} = 44$  ms, and  $T_{2-3} = 31$  ms for the three phantoms. Eq. (1) underestimates  $T_2$  compared to the reference FSE estimate, and this becomes more pronounced for smaller flip angles. The estimate from Eq. (7) agrees very well with the reference scans for most flip angles ( $20^\circ$  and higher), but gives a substantial overestimate in the case of a large  $T_2$  and  $10^\circ$  or  $15^\circ$  flip angles. However, the error is much smaller than the error from Eq. (1).

### 4.2 Volunteer scans

Fig. 4 shows the  $T_2$  estimates from the DESS scan, using Eqs. (1) and (7), as well as the results from the reference FSE scans and a full numerical fit. The maps from Eq. (7) are more visually similar to the reference scans than those of Eq. (1) (Fig. 4a–c). The RMS difference between the estimates of Eq. (7) and FSE was 3.7 ms, while for Eq. (1) it was 9.6 ms. Plotting the data points from Eq. (7) against the FSE data points and drawing a best fit line that crossed the origin yielded a slope of 0.96, while for Eq. (1) it was 0.77 (Fig. 4f). This demonstrates that Eq. (7) gave  $T_2$  estimates that better agreed with FSE estimates than Eq. (1). The full numerical fit gives very similar results to those obtained by Eq. (7), as shown both by its  $T_2$  map and by a difference map (scaled by 10), while taking around 60× longer to produce, with an average processing time of 577 s per DESS data set ( $36 \times 512 \times 512$  data points) compared to 9 s for Eq. (7).

### 4.3 Patient scans

Fig. 5 shows the scan of a patient with a history of knee injuries and osteochondral defects, visible on the DESS images. Fibrocartilage formation is present in the damaged hyaline cartilage region and in the surrounding subchondral bone. The corresponding region in the DESS  $T_2$  map shows a focal decrease in  $T_2$  values. In another patient (not shown), the availability of the DESS  $T_2$  map made it possible to visualize an oblique meniscal tear in the anterior horn of the medial meniscus, which was challenging to delineate with only the morphological images. In a third patient (not shown), a  $T_2$  map acquired with DESS indicated tendinopathy where the patellar tendon had increased  $T_2$  values at its insertion site, which was also relatively challenging to visualize with only the morphological DESS images.

## 5. Discussion

The DESS sequence allows 3D estimation of  $T_2$  in cartilage with high SNR efficiency. The signal expressions are complicated, often necessitating imprecise simplifications or time-consuming numerical modeling of the signals. This study explores a linear relationship



between the two DESS signals that can be used for accurately estimating  $T_2$  from a single scan. We have focused on applying the method in cartilage, but the proposed method could also be used in various other anatomies. This could include  $T_2$  mapping in breast cancer patients, which to date is impractical due to the time consuming nature of spin-echo-based  $T_2$  measurements [26]. Also, this methodology has been used to perform relaxometry of short- $T_2$  tissues in the knee [27].

The proposed method can be viewed as an extension to Eq. (1), used in ref. [9], that accounts for  $T_1$ , flip angle, and diffusion. The expression results from applying a similar approximation to that used by Buxton [22], ignoring echo pathways that have spent more than two TRs in the transverse plane. It is important to note, however, that ref. [22] only dealt with sequences acquiring the second echo  $S_2$ . This required making the assumption that magnetization is transverse for only two TRs from the start of the sequence. In this work, the two-transverse-TR assumption is only applied *between* the echoes  $S_1$  and  $S_2$ , which is a much weaker assumption, resulting in good performance in spite of a simpler model. Effectively, the model of ref. [22] can be viewed as the model proposed here in the special case of the  $S_1$  signal having no transverse history, i.e., being the RF-spoiled or SPGR signal. If  $S_1$  in Eq. (6) is replaced with the well-known expression for the RF-spoiled signal, the two models agree (resulting in the same expression for  $S_2$ ).

The scan time for DESS used in this study was about 5 minutes. The patient scans were acquired at a higher resolution than the volunteer scans, but using parallel imaging, the scan time remained at 5 minutes at a cost in SNR. Scan time could be reduced by using lower resolution, higher readout bandwidth, fewer slices, or shorter RF pulses. However, in our experience, a 5 minute scan time is acceptable, especially considering that these 3D scans provide both morphological and quantitative information. Potentially, DESS could replace sagittal FSE scans that are widely used in many current protocols, but this needs to be studied further.

As mentioned, using the linear approximation of Eq. (7) for estimating  $T_2$  from a single DESS scan requires an assumed  $T_1$  value. This should not lead to large estimation errors, since the method has little sensitivity to  $T_1$ , which is also not expected to change much in cartilage. For example, using the same scan parameters as for the middle curve in Fig. 2a, with the same assumption of  $T_1 = 1.2$  s, yields  $T_2$  estimates of 38.8 ms and 41.8 ms ( $-3.0\%$  and  $+4.5\%$ ) when the real  $T_1$  is 0.96 s and 1.44 s (i.e., the actual  $T_1$  is 20% lower or higher than the assumed value), respectively. Both values are relatively close to the actual value of 40.0 ms. A correct assumption of  $T_1 = 1.2$  s yields  $T_2 = 40.6$  ms ( $+1.5\%$ ), a slight overestimate due to the approximation made. Similarly, sensitivity to  $B_1$  results in estimates of 36.9 ms and 43.1 ms ( $-7.75\%$  and  $+7.75\%$ ) when the actual flip angle  $\alpha$  is  $20^\circ$  and  $30^\circ$  (i.e., 20% lower or higher than the assumed value of  $25^\circ$ ). The  $B_1$  sensitivity could be mitigated by acquiring a  $B_1$  map along with the DESS scan. The sensitivity of the  $T_2$  estimate to errors in the  $T_1$  and  $\alpha$  assumptions is further demonstrated in Fig. 6. It should also be noted that assumptions of  $T_1$  and flip angle will always be needed when estimating  $T_2$  from a single DESS scan, regardless of the model used.

The proposed method can also be used in combination with other methods for quantitative DESS imaging. Cheng et al. proposed a multi-echo DESS sequence for imaging  $T_2$  and  $T_2^*$  that measures  $S_1$  and  $S_2$  at different echo times and uses the fact that  $T_2'$  causes dephasing of  $S_1$  and rephasing of  $S_2$  [28]. Put together, the two methods could potentially estimate other parameters such as  $T_1$  or flip angle. Also, it has previously been noted by Bieri et al. that the ratio  $(S_{2H}S_{1L})/(S_{1H}S_{2L})$ , where H and L denote scans with a large and small ( $k \approx 0$ ) spoiler gradient moment, respectively, has good diffusion sensitivity but low sensitivity to  $T_1$  and  $T_2$  [11]. This is consistent with the expression derived from Eq. (6):

$$\frac{S_{2H}S_{1L}}{S_{1H}S_{2L}} = e^{-(TR-\frac{\tau}{3})\Delta k^2 D} \frac{1 + e^{-\frac{TR}{T_1}} (e^{-TR\Delta k^2 D} - \cos\alpha) - e^{-\frac{2TR}{T_1}} \cos\alpha e^{-TR\Delta k^2 D}}{1 + e^{-\frac{TR}{T_1}} (1 - \cos\alpha e^{-TR\Delta k^2 D}) - e^{-\frac{2TR}{T_1}} \cos\alpha e^{-TR\Delta k^2 D}} \quad (16)$$

This relationship is independent of  $T_2$ , and largely insensitive to  $T_1$ .

In conclusion, we have demonstrated a simplified expression for the ratio between the DESS signals, providing good  $T_2$  estimation. The expression disregards higher-order echo pathways, an assumption we have shown to be valid due to both decay and cancellation of such pathways. The expression takes into account signal dependency to  $T_1$  and flip angle, and we have shown that the method works well by assuming a  $T_1$  typical for the tissue and using the prescribed flip angle in calculations.

## Acknowledgments

This study was supported by National Institutes of Health (NIH) P41 EB015891, R01 AR063643, and R01 EB002524 and by GE Healthcare.

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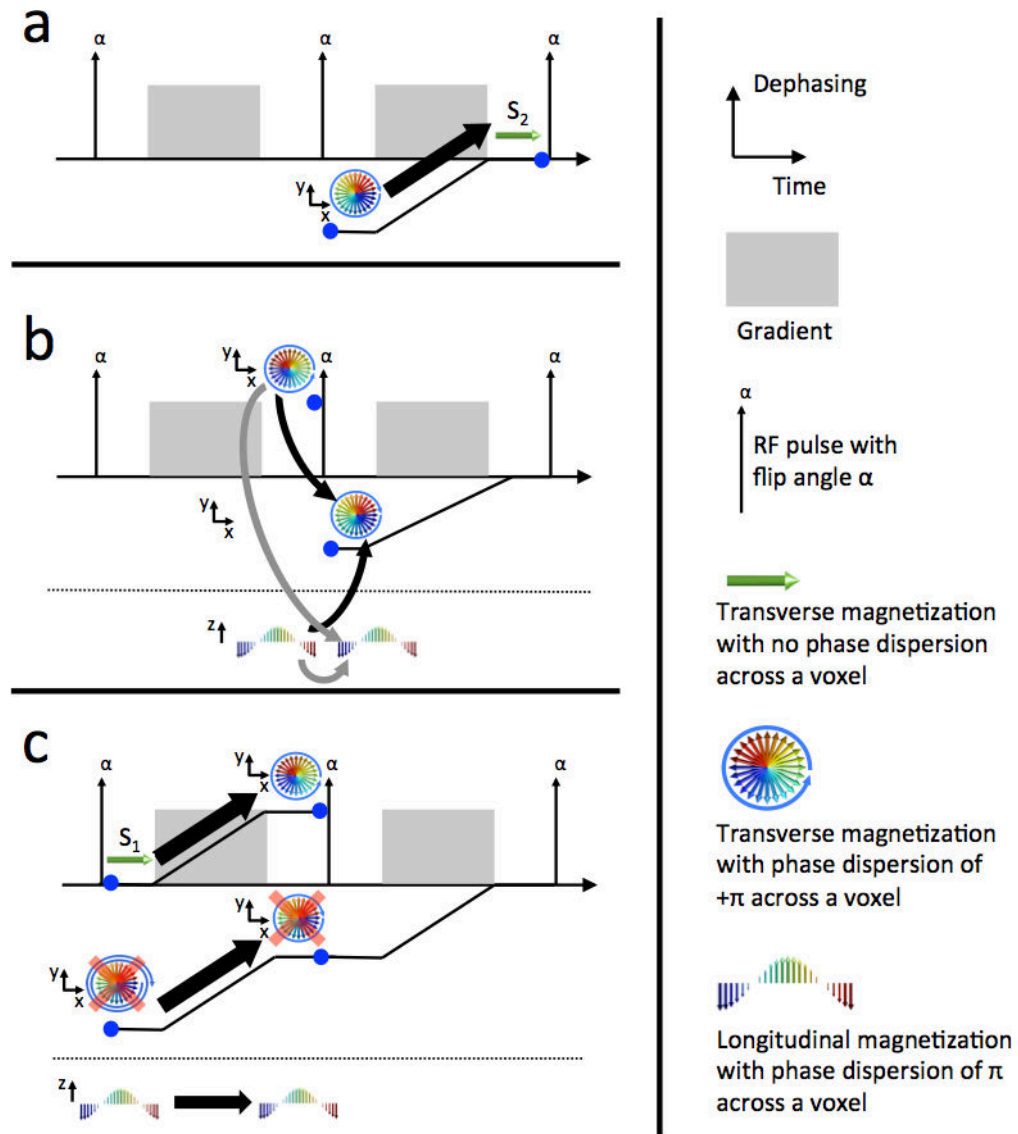
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### Highlights

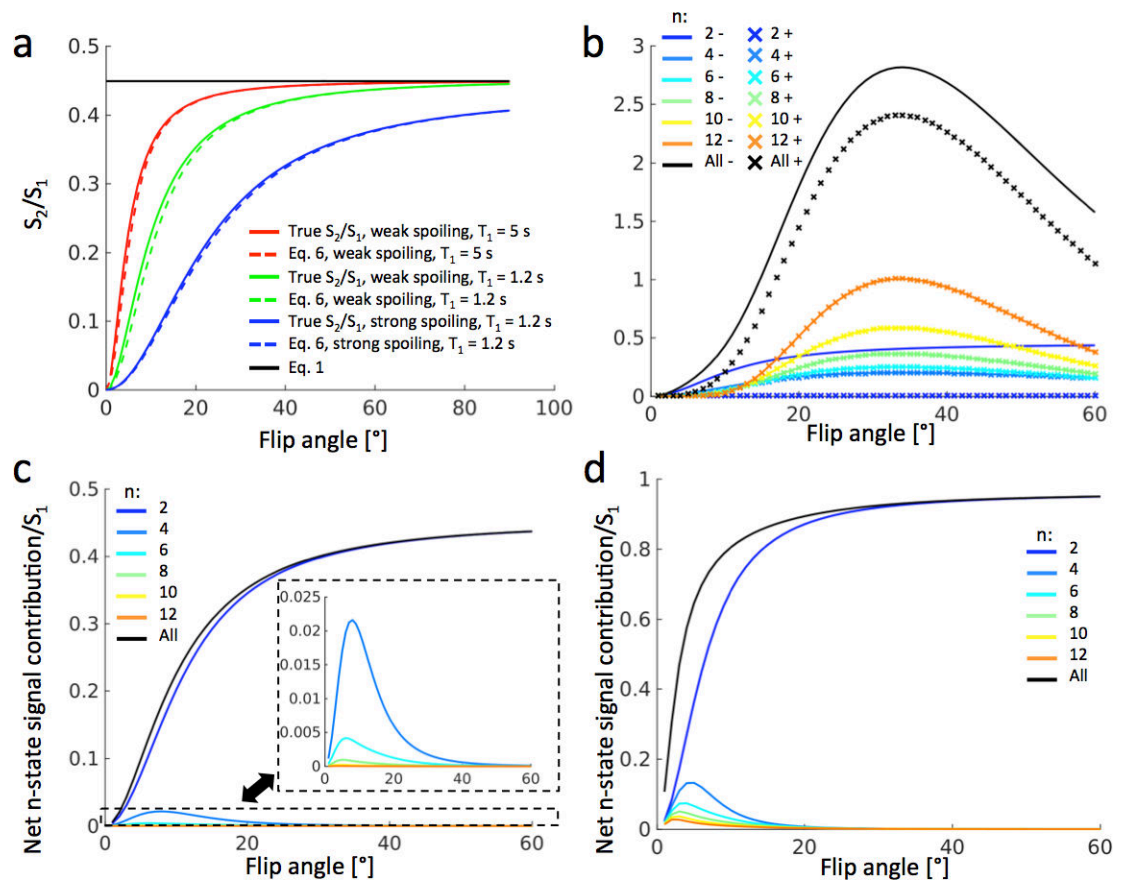
- A simple method for estimating T2 from a single DESS scan is proposed
- The method gave good results in simulations and in phantoms and in vivo scans
- Easy implementation and fast processing allow for near-instant T2 maps at scanner



**Figure 1.**

Diagrams showing dephasing across a voxel as a function of time using graphics similar to ref. [18]. In each panel, we consider the magnetization at the time of the right blue dot and examine the components from the time of the left blue dot that contributed to it. Graphical representations are shown of transverse and longitudinal states (above and below the dotted line, respectively). The accrued phase of the transverse magnetization is represented by the distance of the black line from the time axis. (a) The state that contributes to the measurable signal ( $S_2$ ) before the RF pulse is the “negatively dephased” signal following the previous RF pulse. (b) Two relevant paths before the RF pulse contribute to the state in panel a [21], shown with black arrows. These are transverse and longitudinal states that have a net dephasing from one gradient. They also contribute to the longitudinal state following the RF pulse (gray arrows). (c) The measurable signal following the previous RF pulse ( $S_1$ ), before dephasing by the gradient, contributes to the “positively dephased” transverse pathway from panel b. This is the desired endpoint so the pathway is not retraced further backward. The

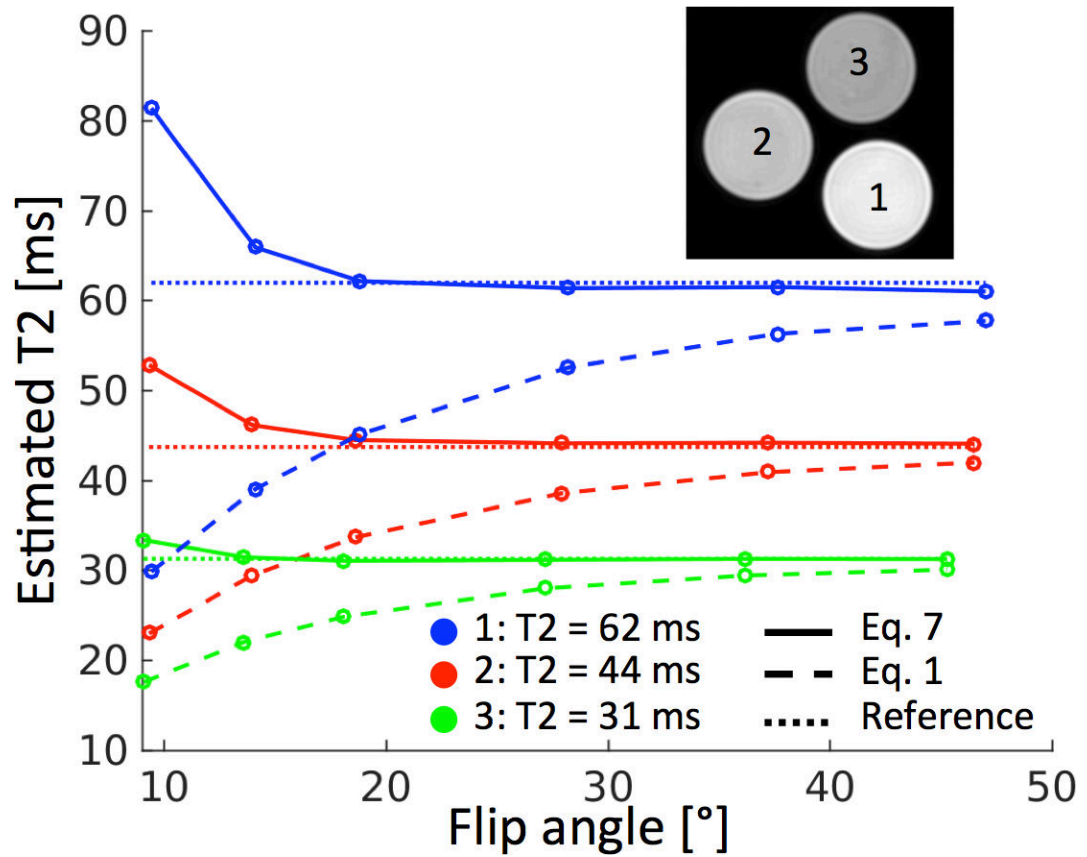
longitudinal pathway is not dephased by the gradient. The contribution of a third transverse pathway has been ignored since it comes from a doubly dephased signal, which can be neglected (shown with red x's). This gives a recursive relationship between S1 and S2 that can be easily solved, resulting in Eq. (6).



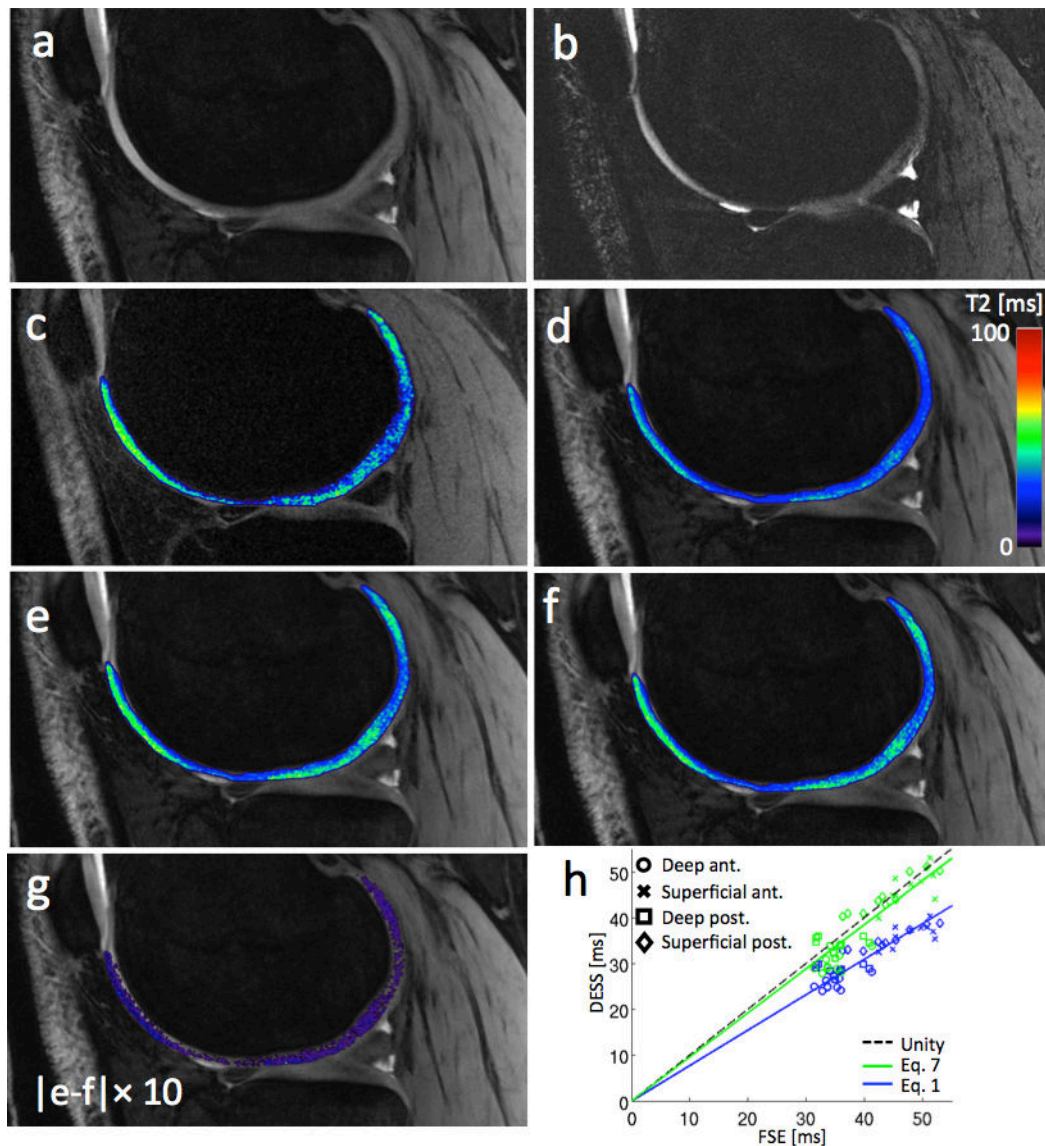
**Figure 2.**

(a) Comparison of proposed model of Eq. (6) (dashed) with complete simulations (colored solids) and with the simple exponential model of Eq. (1) (black solid) in a tissue with  $T_2 = 40$  ms. The blue (bottom) curves correspond to strong spoiling (gradient moment 156.6 mT/m•ms) and the green (middle) curves to weak spoiling (gradient moment 0.001 mT/m•ms), both with  $T_1 = 1.2$  s. The red (top) curves correspond to weak spoiling and  $T_1 = 5$  s. The green curves represent DESS scans that can be used for  $T_2$  estimation in cartilage. (b) Positive (dashed) and negative (solid) contributions to  $S_2/S_1$  from pathways spending 2–12 TRs in the transverse plane. The black curves represent the sum of the different pathways. For pathways of order 6 and above, the positive (x) curves overlap with the negative (solid) curves. (c) Net contributions from the pathways in panel b shows that pathways with more than 2 TRs in the transverse plane contribute minimally. (d) The same simulations as in panel c, performed for synovial fluid at 3T ( $T_1 = 3.6$  s,  $T_2 = 0.77$  s), showing that the proposed model (blue signal) approximates the true signal (black curve) well for flip angles above 20°, even for long  $T_1$  and  $T_2$ .



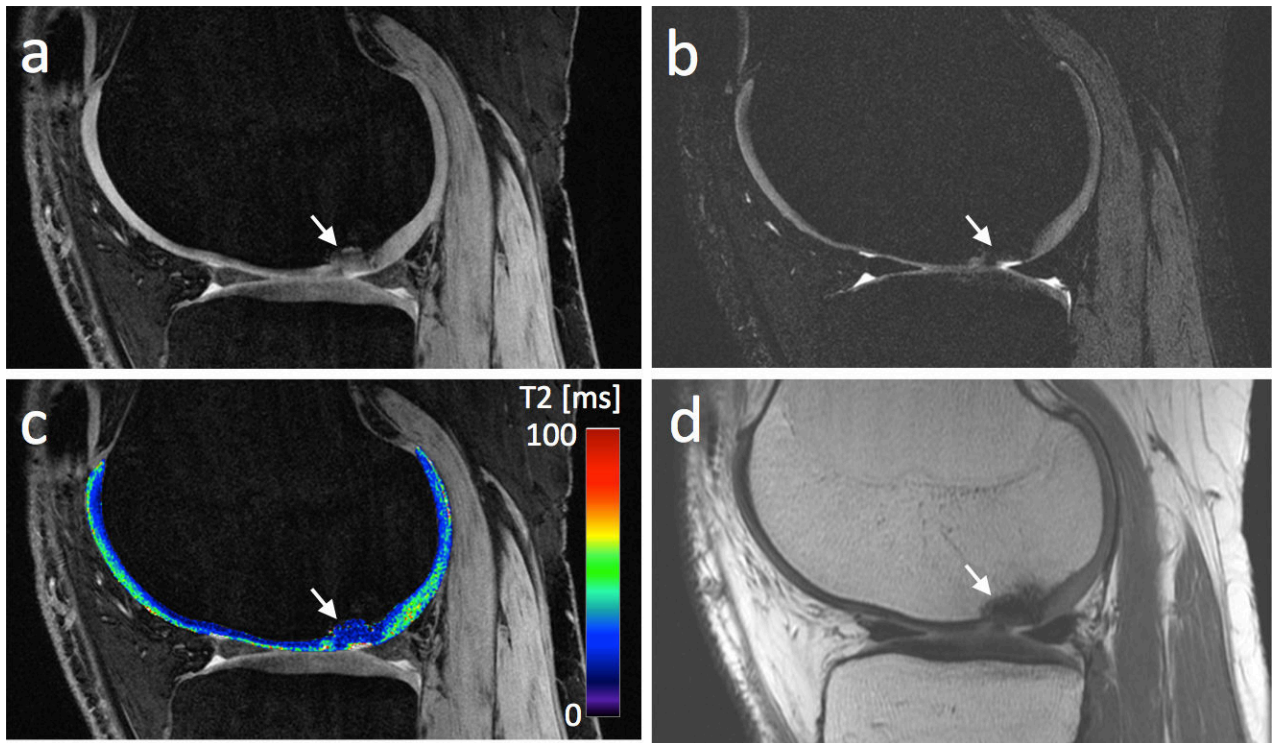


**Figure 3.** Results from estimating T<sub>2</sub> from DESS using Eq. (1) (dashed) and Eq. (7) (solid) as well as from reference FSE scans (dotted). For all phantoms, Eq. (7) clearly agrees better with the reference value than Eq. (1), and for flip angles of 20° or higher, Eq. (7) agrees very well with the reference scans.



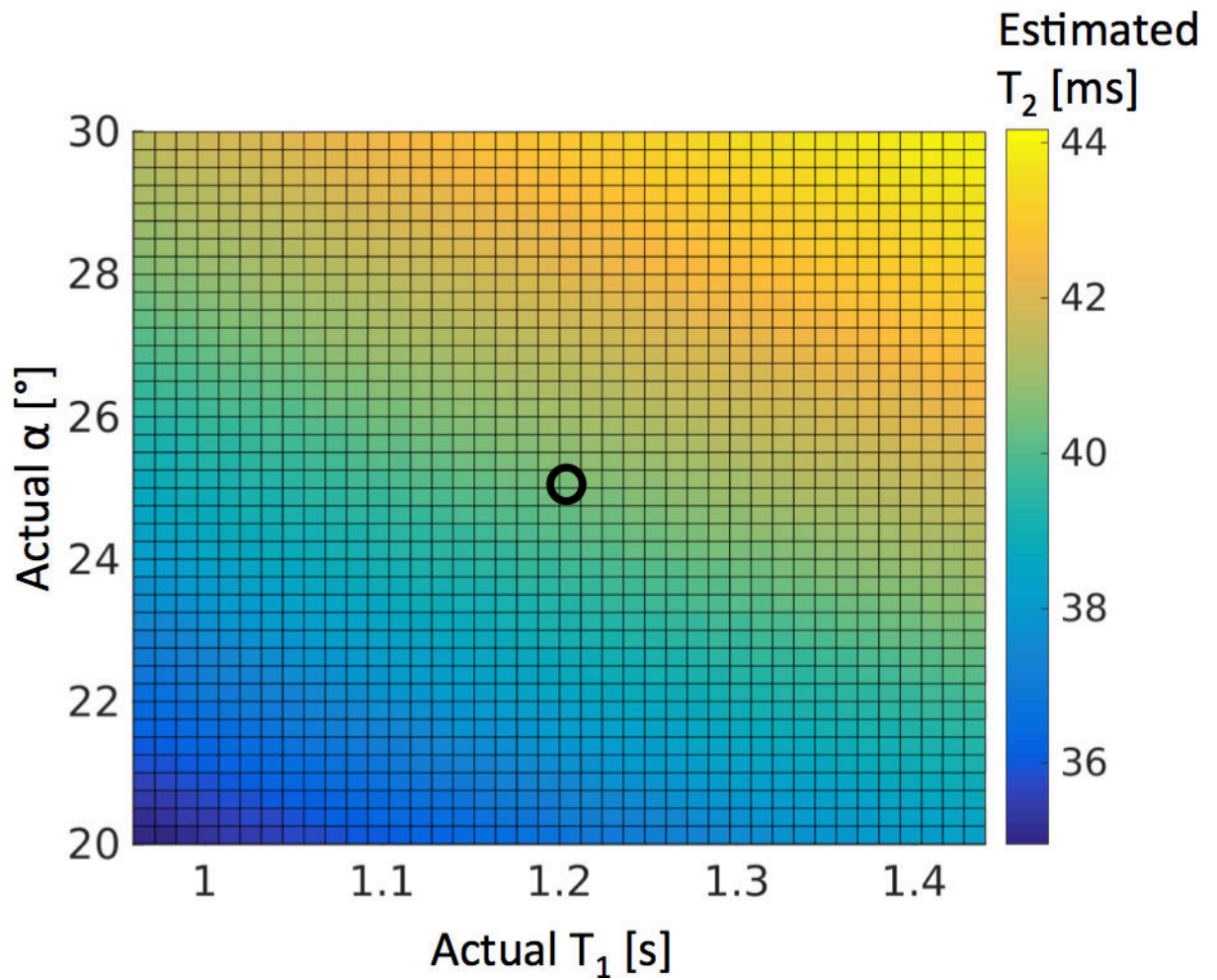
**Figure 4.**

(a) The first echo ( $S_1$ ) from a sample sagittal DESS scan. (b) The second echo ( $S_2$ ) from the same DESS scan. The windowing level is not the same as in panel a, in order to show the cartilage. (c) A sample  $T_2$  map from FSE scans of the subject in panels a–b. (d)  $T_2$  map of articular cartilage from a the DESS scan in panels a–b using Eq. (1). (e)  $T_2$  map using Eq. (7). (f)  $T_2$  map using a full numerical fit. The map looks very similar to the one in panel e, but took about  $60\times$  longer to produce. (g) A map showing the absolute difference between the maps in panels e and f, multiplied by 10. The difference in the cartilage is small, mostly 1 ms or less. Zero difference appears as transparent. (h) The  $T_2$  estimates from 4 regions of femoral cartilage in a total of 10 slices from 3 subjects. The trend line for Eq. (7) (green) better agrees with the FSE scans.



**Figure 5.**

(a) The first echo of a DESS scan of a patient with a chondral lesion (white arrow). (b) The second echo of the DESS scan. Panels a and b both show cartilage signal heterogeneity in the central femoral condyle (note the different windowing settings). (c) The  $T_2$  maps resulting from applying Eq. (7) to the data in panels a–b. (d) A PD weighted scan, acquired for reference, shows low signal in the lesion.



**Figure 6.**

The sensitivity of the  $T_2$  estimation method in Eq. (7) to errors in the assumption of  $T_1$  and  $B_1$ . The true  $T_2$  is 40 ms. The flip angle  $\alpha$  and  $T_1$  are assumed to be  $25^\circ$  and 1.2 s, but actually vary from these values by  $\pm 20\%$ . Other scan parameters were the same as in Fig. 2a. The black circle shows the point where the assumptions of  $\alpha$  and  $T_1$  are correct.