

# Clinical Evaluation of Fast T2-Corrected MR Spectroscopy Compared to Multi-Point 3D Dixon for Hepatic Lipid and Iron Quantification

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**Target Audience:** Translational Scientists, Body Radiologists, Sequence Developers

**Background:** Advanced MR imaging, using multi-point Dixon reconstruction, allows 3-dimensional assessment of hepatic fat fraction (FF) and local R2\*, which is a surrogate for iron concentration (1-5). In addition, a high-speed, T2-corrected single-voxel spectroscopy (HISTO-MRS) approach has been developed to interrogate hepatic water and lipid compartments, and to elucidate FF and R2 values of these components in a single breath hold (6). Thus far, this fast single breath-hold HISTO-MRS has not been formally evaluated in the clinical environment, nor compared to current 3D multi-point Dixon imaging in terms of lipid and iron estimation.

**Purpose:** 1) To evaluate breath hold HISTO-MRS in a routine clinical environment, in consecutive patients; 2) To correlate hepatic FF from HISTO-MRS and 3D multi-point Dixon at 1.5T; and 3) to compare HISTO-R2 measures with R2\* for sensitivity to iron content.

**Methods:** This investigation was approved by the IRB, was HIPAA-compliant, and all participants signed informed consent prior to imaging. All imaging was performed on a Siemens 1.5T Aera system using an 18-channel phased-array body coil. Thirty-nine consecutive patients arriving for routine abdominal MRI over a two week period were recruited for this study. The exclusion criteria consisted of standard MR contraindications. Each patient received two additional breath hold acquisitions: 1) single-voxel HISTO-MRS; and 2) 6-point 3D GRE (3D Dixon FQ). The HISTO-MRS voxel was prescribed using available T2-weighted single-shot localizer acquisitions. The voxel was placed in a region free of major hepatic vessels, usually in the right lobe. The MRS parameters were: STEAM sequence; TR/TM = 3000ms/10ms; 5 TEs = {12, 24, 36, 48, 72}ms; voxel = 20-30mm<sup>3</sup>; 1 signal average; time = 15sec. The 3D Dixon FQ sequence consisted of 80 slices prescribed to cover the full liver volume. Other parameters included: TR = 9.1ms; 6 TEs = 1.2/2.5/3.7/5.0/6.3/7.6ms; matrix = 256 x 154; FOV = 400-420mm; phase FOV = 75%; slice thickness = 3mm; BW = 1085Hz/px; time = 18sec. Post-processing for both techniques were performed automatically at the scanner console with inline reconstruction. For HISTO-MRS, the integral signal area of both water and total lipid (~1.3 ± 0.8ppm) spectra were automatically tabulated at each TE, whereby T2 of both components (R2<sub>water</sub> and R2<sub>lipid</sub>) were quantified using a non-linear least squares algorithm, and a goodness-of-fit measure (r<sup>2</sup>). The fat fraction (FF<sub>HISTO</sub>) was calculated from the ratio of T2-corrected lipid integral to T2-corrected lipid + water integral (1). Using a multi-step fitting approach, the 3D Dixon FQ data was processed automatically by fitting the multi-echo data to the magnitude of the complex signal model that included the coefficients and frequency components for a seven-peak fat model (5). Volumetric T2\*-corrected fat fraction (FF<sub>DIXON</sub>) maps were calculated from the fitting, along with R2\* maps. **Analysis:** Mean and standard deviation measurements of FF<sub>DIXON</sub> and R2\* were performed in Clear Canvas Workstation 2.0 (Toronto, CA) from region-of-interests (ROIs) in approximate region of the HISTO-MRS voxel. Agreement of FF<sub>HISTO</sub> and FF<sub>DIXON</sub> was performed with Bland-Altman analysis, with measures of bias and 95% confidence limits. Since R2\* has been shown to correlate with hepatic iron content (7), Pearson correlation was performed against both R2<sub>water</sub> and R2<sub>lipid</sub>. Significance was set to p<0.05.

**Results:** All patients tolerated the additional MR acquisitions. 1/39 patients were excluded due to large metal artifact, 3/39 3D Dixon FQ cases experienced fat-water “swap”, leading to uncalculated R2\* maps, while 1/39 HISTO-MRS case suffered from poor T2 fit (r<sup>2</sup><0.6) due to severe iron deposition. Therefore, 34 patients were included in the analysis. For low FF<sub>HISTO</sub> (< 6.0%, n=22), R2<sub>lipid</sub> could not be reliably calculated due to poor lipid SNR. Figure 1 shows representative HISTO-MRS spectra and corresponding FF<sub>DIXON</sub> and R2\* maps in one patient. Figure 2 shows the Bland-Altman plot comparing FF<sub>HISTO</sub> and FF<sub>DIXON</sub>. The bias was -0.5%, and 95% confidence was ±4.8%, which represents a significant correlation (r=0.99, p<0.01), despite 1 outlier at high FF (red circle in Fig. 2). R2<sub>water</sub> measurement from HISTO-MRS correlated significantly with R2\* (r=0.87, p<0.01), while R2<sub>lipid</sub> did not correlate (r=0.12, p=0.5), and remained relatively constant in patients with elevated FF (R2<sub>lipid</sub> = 20.5 ± 4.1s<sup>-1</sup>, FF > 8.0%, n=7).

**Discussion:** With automatic post-processing, HISTO-MRS was efficiently performed in a routine clinical setting. Significant agreement in FF was found between HISTO-MRS and 3D Dixon FQ. From the behavior of R2<sub>water</sub> and R2<sub>lipid</sub>, the results also confirm that hepatic iron content affects R2<sub>water</sub> compartment more significantly than R2<sub>lipid</sub>. Limitations of the current HISTO-MRS include poor estimation of severe iron deposition (due to long initial TE) and unreliable R2<sub>lipid</sub> for low FF<sub>HISTO</sub>.

**Conclusions:** Both HISTO-MRS and 3D Dixon FQ are viable clinical methods for the estimation of hepatic fat fraction (FF) and iron content (R2\* and R2<sub>water</sub>).

**References:** 1. Chebrolu et al. Magn Reson Med 2010;63:849-857; 2. Hernando et al. Magn Reson Med 2010;63:79-90; 3. Bydder et al. Magn Reson Imaging 2008;26:347-359; 4. Yu et al. Magn Reson Med 2011;66:199-206; 5. Zhong et al. Magn Reson Med; in press; 6. Pineda et al. Radiology 2009;252:568-576; 7. Wood et al. Blood 2005;106:1460-1465.

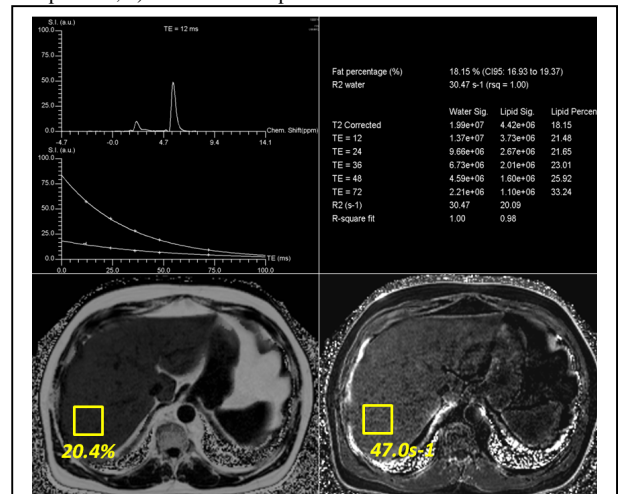


Figure 1. HISTO report (top) and Dixon FQ maps (bottom)

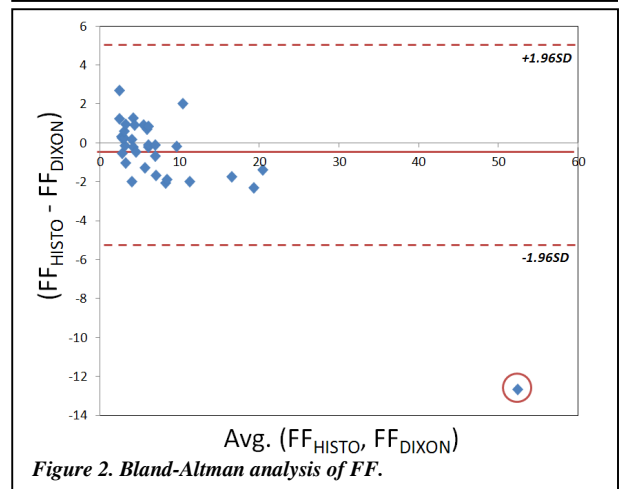


Figure 2. Bland-Altman analysis of FF.