**Very fast 2D proton MR spectroscopic imaging of the in vivo human brain at 3 Tesla with high spatial resolution using the SSFP based sequence "spectroscopic FAST"**

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**Introduction**

In the past, several methods for fast 2D and 3D ¹H MR spectroscopic imaging (MRSI) derived from their corresponding fast MRI methods have been proposed to map the distribution of brain metabolites. Besides echo planar MRSI and spiral MRSI [1-3], various pulse sequences based on the condition of steady state free precession (SSFP) are very promising because of their high signal-to-noise ratio per unit measurement time (SNRt) and their short minimum measurement time (Tmin) [4]. To date, only the SSFP based sequences spectroscopic CE-FAST (spCE-FAST) [5] and spectroscopic Missing-Pulse SSFP (spMP-SSFP) [6] have been applied to the in vivo human brain at 3 Tesla. Their efficient lipid suppression is essential for in vivo measurements to prevent lipid signal contamination of intracranial voxels by voxels from the scalp. In this work, for the first time the SSFP based sequence “spectroscopic FAST” (spFAST) [4] is used for fast 2D ¹H MRSI of the human brain at 3 Tesla. Except for spectroscopic TRUE-FISP [7] with which only single resonances are detectable, spFAST yields the highest possible SNRt among the SSFP based sequences. Therefore, smaller voxel sizes are accessible and thus a higher spatial resolution is achieved within a reasonable measurement time. Although in spFAST the signal intensity of lipids is much higher than the intensity of brain metabolites, contamination of voxels inside the brain by extracranial lipid signals is minimized by spectral-spatial RF excitation pulses and an elliptical k-space sampling scheme with subsequent spatial filtering. Furthermore, a large number of phase encoding steps ensures a better spatial localization of lipid signals in the scalp.

**Materials and methods**

The spFAST sequence is shown in Fig.1. After RF excitation with spectral-spatial composite pulses [8] (1-τ-1-τ-6-τ-1-τ-6) spectral minima appear at 1/τ, τ=2ms which are simultaneously slice and chemical shift selective (for both water and lipid suppression) and subsequent phase encoding gradients, an FID-like signal is acquired. Afterwards, the phase encoding gradients are compensated for by phase rewinding gradients to maintain the steady state. Prior to the next spectral-spatial composite pulse, the echo-like signal is spoiled by using 2ms gradients with an amplitude of 25 mTm to avoid interference of the signals.

In Fig.2 the relative signal intensities of metabolites with uncoupled spins, water and lipids are simulated in dependence on the flip angle. Assuming (T1=1400 ms, T2=210 ms) for metabolites, (T1=1200 ms, T2=90 ms) for water and (T1=210 ms, T2=65 ms) for lipids as typical in vivo relaxation times at 3 Tesla, the transverse magnetization of lipids is about twice the maximum signal intensity of metabolites which makes an efficient lipid suppression more difficult.

The spFAST sequence was implemented on a 3 Tesla Magnetom Allegra head scanner (Siemens Medical Solutions, Erlangen, Germany) and was applied to phantoms and healthy volunteers. A CP head coil was used for both RF transmission and signal reception. Further measurement parameters were: repetition time (TR) = 72 ms, duration of spectral-spatial composite pulse = 12.4 ms, excitation flip angle α = 20°, field of view (FOV) = 190×220 mm², slice thickness = 20 mm, acquisition bandwidth = 5 kHz, 256 complex data points, elliptical k-space sampling and subsequent spatial filtering with a Hamming function. We used a different number of phase encoding (PE) steps in three consecutive measurements to evaluate different spatial resolutions with regard to lipid signal localization and SNR:

1) PE steps: 28x32, nominal voxel size: 0.93 cm³, real voxel size: 2.02 cm³ due to elliptical k-space sampling and the spatial filtering, measurement time: 48 sec.

2) PE steps: 42x48, nominal voxel size: 0.41 cm³, real voxel size: 0.83 cm³ due to elliptical k-space sampling and the spatial filtering, measurement time: 1 min 49 sec.

3) PE steps: 66x64, nominal voxel size: 0.23 cm³, real voxel size: 0.47 cm³ due to elliptical k-space sampling and spatial filtering, measurement time: 3 min 13 sec.

Postprocessing consisted of spectral apodization with a sine-bell function, zero-filling from 256 to 1024 data points, Fourier transformation and phase correction.

**Results**

A representative spectrum of measurement 2 is depicted in Fig.4 which originates from the voxel marked in the reference image in Fig.5. The signals are assigned to N-acetyl-aspartate (NAA), total creatine (Cr), total choline (Cho) and myo-inositol (Ins). Residual lipid signals at around 1.5 ppm that contaminate voxels inside the brain are minimized to or below the level of the NAA signal throughout the whole brain. Metabolic images of NAA created by peak area integration are displayed in Fig.5 and demonstrate the different spatial resolutions of 2.02 cm³ (Fig.5a), 0.83 cm³ (Fig.5b) and 0.47 cm³ (Fig.5c). The signal-to-noise-ratio of the NAA peak was determined to ~50 for measurement 1, to ~30 for measurement 2 and to ~25 for measurement 3 (signal maximum / standard deviation of noise in a signal-free chemical shift region).

**Discussion and conclusions**

In spite of the higher relative intensity of lipid signals with respect to the intensity of metabolite signals as shown in the simulation, signal bleeding of extracranial lipid signals into voxels inside the brain is efficiently minimized by spectral-spatial composite pulses, elliptical k-space sampling and spatial filtering, the former two at the cost of larger voxel sizes. Even the “small” number of PE steps of measurement 1 (28x32) is sufficient for an adequate localization of lipid signals in the scalp.

Lipid signals were further reduced by the choice of the excitation flip angle of 20°: while the loss of signal intensity for metabolites with uncoupled spins compared to an optimal flip angle of 30° is only 5%, the signal loss of lipids is 21% compared to α = 20°. Further investigations are necessary to empirically adjust TR and the excitation flip angle for maximum SNR, of metabolites with coupled spins. However, the optimization is hampered because simulation programs for signals of coupled spins are not available yet and signal intensities strongly depend on T1 and T2 values which may vary in vivo. The short measurement time could be further reduced by an EPI-like readout as done in Ref. 9 for spCE-FAST. In conclusion, the spFAST pulse sequence was successfully implemented at 3 Tesla and provides 2D metabolic maps with a spatial resolution of 0.47 cm³ in about 3 minutes with a high SNR.

**References**