Liver Fat Quantification Using MRI for Assessment of NAFLD/NASH

Applications in Clinical Studies
Antaros Medical and ProSciento scientists have conducted many studies assessing liver lipid and glycogen homeostasis with interventions such as diet, surgery, as well as pharmaceutical interventions, including SGLT-2, GLP-1, and ARB. For these studies, a consistent and precise quantification of liver fat is essential, especially when assessing efficacy and safety endpoints for new therapies for metabolic diseases such as type 2 diabetes mellitus, insulin resistance, obesity and NAFLD/NASH.

High Precision Measurement
Magnetic Resonance Imaging (MRI) based methods that allow collection of fat and water specific signals are considered the most accurate non-invasive assessments of liver fat. Ultrasonography (US) and Computed Tomography (CT) measure liver fat using indirect properties. Moreover, MRI and Magnetic Resonance Spectroscopy (MRS) can both be utilized. MRI uses a model of the water and lipid signal and allows coverage of the entire liver and high spatial resolution, while MRS provides high spectral resolution from a few voxels. The MRI-based methods with full liver coverage and high spatial resolution allow specific analysis of the liver parenchyma, which is of importance for precision as it avoids signals from large vessels and bile ducts. Total method precision using repeated imaging (test-retest) and analysis in ten healthy volunteers is 0.1 percentage points (mean of the ten standard deviations). Antaros Medical also quantifies liver volume in all studies to allow assessment of changes in the non-lipid component (e.g. glycogen) of the liver. This allows estimation of total liver fat content in addition to the liver fat concentration.

Tolerability and Scanner Compatibility
The imaging protocols that Antaros Medical uses for the liver fat measurement only require a single breath hold, which is well tolerated by patients. We previously installed the protocols on all major clinical MRI vendors and field strengths, allowing maximum MRI site compatibility and subject inclusion in multi-center trials.

Subjects and Preparation
Subjects are always screened for standard contraindications for MRI. Fasted state (>4h) and standardized time of the day is preferred to minimize variability.
Procedure – Data Collection

Body coil or surface coils are used depending on the MRI system. The MRI protocol is a Proton Density Fat Fraction (PDFF) technique utilizing a spoiled 3D 6-echo gradient echo in the axial plane covering the entire liver. Image data is coded, exported and uploaded together with a scan log to our core lab for quality control and analysis.

Procedure – Image Analysis

Depending on the MRI system, we use built-in solutions (provided by the manufacturers) for water-fat image reconstruction or our own published reconstruction algorithm.9 The methods apply multi-peak lipid spectral models and simultaneous quantification and correction for T2*. In addition, special care is taken to reduce effects of phase errors (eddy currents).10,11 The liver fat value is the median value of all voxels in the identified volume of interest, excluding vessels and bile ducts. Quality control is performed in all steps of the process.

Advantages of Antaros Medical’s Liver Fat Quantification Using MRI

- High precision due to whole liver coverage with high spatial resolution analysis avoiding vessels and ducts leading to improved probability of detecting significant drug effects
- Accurate liver volume assessment included as a proxy for potential drug-induced effects on glycogen
- Integration with other liver morphology and functional measurements, such as diffusion, elastography and gadoxetate imaging for multiparametric assessment of NAFLD/NASH, as well as measurements of body composition17 avoiding additional investigations
- Already used in over 20 clinical studies with more than 3,000 PDFF assessments

References


About ProSciento’s CRI

Fact Sheets

Each fact sheet describes a select technique from our wide scope of complex methodologies for metabolic clinical research. Contact us for fact sheets highlighting additional techniques.

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