

Fat quantification in the liver

Bachir Taouli, M.D.

Assistant Professor, Department of Radiology
New York University Langone Medical Center
E-mail: bachir.taouli@nyumc.org

In this presentation, we will review:

1. The clinical relevance of diagnosis and quantification of liver fat
2. The available MR sequences for diagnosis of liver fat, and how these perform for liver fat quantification

Introduction-clinical relevance

Since the 1980s, obesity has become an epidemic in the United States (1-4). The prevalence of obesity was estimated to be 27% in 1999 (5). Overweight and obesity account for nearly 17% of all deaths in the US (6). Obesity is significantly associated with diabetes, insulin resistance, and nonalcoholic fatty liver disease (NAFLD). NAFLD is estimated to occur in 30 to 100% of obese adults (7), and includes a spectrum of liver abnormalities from steatosis to nonalcoholic steatohepatitis (NASH). NASH has a poorer prognosis compared with simple steatosis, leading to cirrhosis in up to 25% of cases (8), with the risk of liver failure, and hepatocellular carcinoma. NASH is diagnosed by the presence of inflammation and fibrosis, compared to simple steatosis.

Diagnosis and quantification of liver fat with MRI and Proton MR Spectroscopy (MRS)

Among the MRI methods used to date, the two-point Dixon method (in and out-of-phase imaging) (9) provides an accurate assessment of liver fat as shown in previous studies, with correlation coefficients of 0.86-0.98 (10-12). However, fat quantification measured with in- and out-of-phase imaging may be inaccurate when there is a high amount of liver fat (13) and concomitant iron deposition (14), and in those cases MRS should be used in conjunction with in- and out-of-phase imaging to resolve fat-water predominance. 3-point Dixon method (9) and the iterative decomposition of water and fat with echo asymmetry and least-squares estimation (IDEAL) method (15,16) have better accuracy for fat quantification in the liver, compared to 2-point Dixon method.

MRS is a potential noninvasive alternative to biopsy for assessing the degree of lipid accumulation within the liver (17-22). It allows in vivo evaluation of metabolites such as water and fat within a selected volume of interest. Signals from protons in different molecular groups have different resonant frequencies or different chemical shifts within a spectrum. Proton signals from water and fat are well separated with a chemical shift of 3.5 ppm or 225 Hz at 1.5T. MRS can be used to directly quantify fat fraction. Spectroscopic evaluation of liver FF may be achieved during a single breath-hold in portions of the liver, and requires the evaluation of the two dominant peaks within the unsuppressed spectrum, water at 4.7 ppm and lipid at 1.0-1.5 ppm. Livers containing increased lipid content demonstrate an increase in the lipid peak relative to normal liver. Quantitative analysis requires the correction for factors that affect signal intensity, such as signal decay from T2 relaxation and accounting for unsaturated lipids, as a small portion of the MR signal from these molecules overlaps with the water resonance at 4.7 ppm. Signal saturation from incomplete T1 relaxation is minimized by using long TR.

References

1. Mokdad AH, Ford ES, Bowman BA, Dietz WH, Vinicor F, Bales VS, Marks JS. Prevalence of obesity, diabetes, and obesity-related health risk factors, 2001. *Jama* 2003;289:76-79.
2. Hedley AA, Ogden CL, Johnson CL, Carroll MD, Curtin LR, Flegal KM. Prevalence of overweight and obesity among US children, adolescents, and adults, 1999-2002. *Jama* 2004;291:2847-2850.
3. Ogden CL, Carroll MD, Flegal KM. Epidemiologic trends in overweight and obesity. *Endocrinol Metab Clin North Am* 2003;32:741-760, vii.
4. Ogden CL, Fryar CD, Carroll MD, Flegal KM. Mean body weight, height, and body mass index, United States 1960-2002. *Adv Data* 2004;1-17.
5. Yanovski SZ, Yanovski JA. Obesity. *N Engl J Med* 2002;346:591-602.
6. Mokdad AH, Marks JS, Stroup DF, Gerberding JL. Actual causes of death in the United States, 2000. *Jama* 2004;291:1238-1245.
7. Luyckx FH, Desai C, Thiry A, Dewe W, Scheen AJ, Gielen JE, Lefebvre PJ. Liver abnormalities in severely obese subjects: effect of drastic weight loss after gastroplasty. *Int J Obes Relat Metab Disord* 1998;22:222-226.
8. Gholam PM, Kotler DP, Flancbaum LJ. Liver pathology in morbidly obese patients undergoing Roux-en-Y gastric bypass surgery. *Obes Surg* 2002;12:49-51.
9. Dixon WT. Simple proton spectroscopic imaging. *Radiology* 1984;153:189-194.
10. Levenson H, Greensite F, Hoefs J, Friloux L, Applegate G, Silva E, Kanel G, Buxton R. Fatty infiltration of the liver: quantification with phase-contrast MR imaging at 1.5 T vs biopsy. *AJR Am J Roentgenol* 1991;156:307-312.
11. Fishbein MH, Gardner KG, Potter CJ, Schmalbrock P, Smith MA. Introduction of fast MR imaging in the assessment of hepatic steatosis. *Magn Reson Imaging* 1997;15:287-293.
12. Rinella ME, McCarthy R, Thakrar K, Finn JP, Rao SM, Koffron AJ, Abecassis M, Blei AT. Dual-echo, chemical shift gradient-echo magnetic resonance imaging to quantify hepatic steatosis: Implications for living liver donation. *Liver Transpl* 2003;9:851-856.
13. Chang JS, Taouli B, Salibi N, Hecht EM, Chin DG, Lee VS. Opposed-phase MRI for fat quantification in fat-water phantoms with 1H MR spectroscopy to resolve ambiguity of fat or water dominance. *AJR Am J Roentgenol* 2006;187:103-106.
14. Westphalen AC, Qayyum A, Yeh BM, Merriman RB, Lee JA, Lamba A, Lu Y, Coakley FV. Liver fat: effect of hepatic iron deposition on evaluation with opposed-phase MR imaging. *Radiology* 2007;242:450-455.
15. Reeder SB, Wen Z, Yu H, Pineda AR, Gold GE, Markl M, Pelc NJ. Multicoil Dixon chemical species separation with an iterative least-squares estimation method. *Magn Reson Med* 2004;51:35-45.
16. Reeder SB, McKenzie CA, Pineda AR, Yu H, Shimakawa A, Brau AC, Hargreaves BA, Gold GE, Brittain JH. Water-fat separation with IDEAL gradient-echo imaging. *J Magn Reson Imaging* 2007;25:644-652.
17. Longo R, Ricci C, Masutti F, Vidimari R, Croce LS, Bercich L, Tiribelli C, Dalla Palma L. Fatty infiltration of the liver. Quantification by 1H localized magnetic resonance spectroscopy and comparison with computed tomography. *Invest Radiol* 1993;28:297-302.
18. Thomsen C, Becker U, Winkler K, Christoffersen P, Jensen M, Henriksen O. Quantification of liver fat using magnetic resonance spectroscopy. *Magn Reson Imaging* 1994;12:487-495.
19. Longo R, Pollesello P, Ricci C, Masutti F, Kvam BJ, Bercich L, Croce LS, Grigolato P, Paoletti S, de Bernard B, et al. Proton MR spectroscopy in quantitative in vivo determination of fat content in human liver steatosis. *J Magn Reson Imaging* 1995;5:281-285.
20. Tiikkainen M, Bergholm R, Vehkavaara S, Rissanen A, Hakkinen AM, Tamminen M, Teramo K, Yki-Jarvinen H. Effects of identical weight loss on body composition and features of insulin resistance in obese women with high and low liver fat content. *Diabetes* 2003;52:701-707.
21. Szczepaniak LS, Nurenberg P, Leonard D, Browning JD, Reingold JS, Grundy S, Hobbs HH, Dobbins RL. Magnetic resonance spectroscopy to measure hepatic triglyceride content: prevalence of hepatic steatosis in the general population. *Am J Physiol Endocrinol Metab* 2005;288:E462-468.
22. Thomas EL, Hamilton G, Patel N, O'Dwyer R, Dore CJ, Goldin RD, Bell JD, Taylor-Robinson SD. Hepatic triglyceride content and its relation to body adiposity: a magnetic resonance imaging and proton magnetic resonance spectroscopy study. *Gut* 2005;54:122-127.