

The Importance of MRI of the Wrist in Patients with Rheumatoid Arthritis

Filippo Del Grande, M.D., M.H.E.M.¹; Aaron J Flammang, MBA BSRT (MR)²; Abraham Padua RT (MR)²; John A. Carrino, M.D. M.P.H.¹

¹Johns Hopkins University School of Medicine, Russell H. Morgan Department of Radiology and Radiological Science, Baltimore, MD, USA

²Siemens Medical Solutions, USA

Introduction

Rheumatoid arthritis (RA) is a chronic inflammatory disease affecting the joints, mainly the metacarpo-phalangeal joints (MCP) and the carpal joints. MR imaging (MRI) is the only imaging modality that can directly detect the early findings of the disease such as bone marrow edema (BME) and synovitis [1]. Moreover MRI can detect tendosynovitis and bone erosions with greater sensitivity than in standard radiography. The treatments with anti-tumor necrosis factor (TNF) drugs are very expensive and carry potential severe infectious adverse effects. Therefore it is very

important to have an early and accurate diagnose of RA which is possible with MRI associated with clinical and laboratory findings. Moreover an early treatment with anti-rheumatic drugs seems to have a better disease outcome at 2 years [2].

We will review the MRI signs of RA such as BME, synovitis/tendosynovitis and bone erosions. The first three manifestation of the disease are included in the rheumatoid arthritis MRI scoring system (RAMRI) of the outcome measurement in Rheumatology clinical trials (OMER-ACT) [3]. We will not treat cartilage loss

that should be treated separately, due to its complexity and the presence of different promising emerging techniques.

MRI in patients with Rheumatoid arthritis

Bone marrow edema is a sign of early manifestation of RA and it shows low signal intensity ill defined areas on T1-weighted sequences and high signal intensity in fluid sensitive sequences such as STIR or T2 fat sat. Several interesting considerations are important regarding BME. First, BME follows the same pattern than bone erosions

suggesting that bone marrow edema is the precursor of bone erosions [4]. Second, BME is the most important prognostic factor for progression of the disease. The authors of one study [5] were able to correctly detect the disease progression in 82% of the cases with a sensitivity of 81% and a specificity of 82% using clinical (hand arthritis, morning stiffness) laboratory (positivity for RF) and imaging criteria (bone marrow edema score in MCP and carpal joints). Another 2-years randomized controlled trial [6] concluded that MRI bone marrow edema is the strongest predictor of progression of the disease in hand, wrist and foot. Third, according to Olech et al. [7] bone marrow edema is the most specific sign (among bone marrow edema, synovitis and bone erosions) for RA. The study [7] showed that BME was 65% sensitive and 82.5% specific for RA and, if the lunatum was not included in the scoring system, the sensitivity decrease to 62.5% but the specificity increase to 87.5%. BME was present in 15% of the control group subjects and, if the lunatum was excluded, in 12.5% of the subjects. Interestingly no one healthy subject presented bone marrow edema in the MCP joint.

Bone erosions represent “the focal loss of cortical and/or underlying trabecular bone” [8] and are better visualized in T1-weighted sequence as interruption of the cortical and trabecular bone. According to the RAMRI criterion bone erosions are defined as follows: “a sharply marginated bone lesion, with correct juxta-articular localization and typical signal characteristics, which is visible in two planes with a cortical break seen in at least one plane” [3]. MRI has a moderate sensitivity (61%) and a good specificity (93%) to detect bone erosions compared to CT as a gold standard whereas stan-

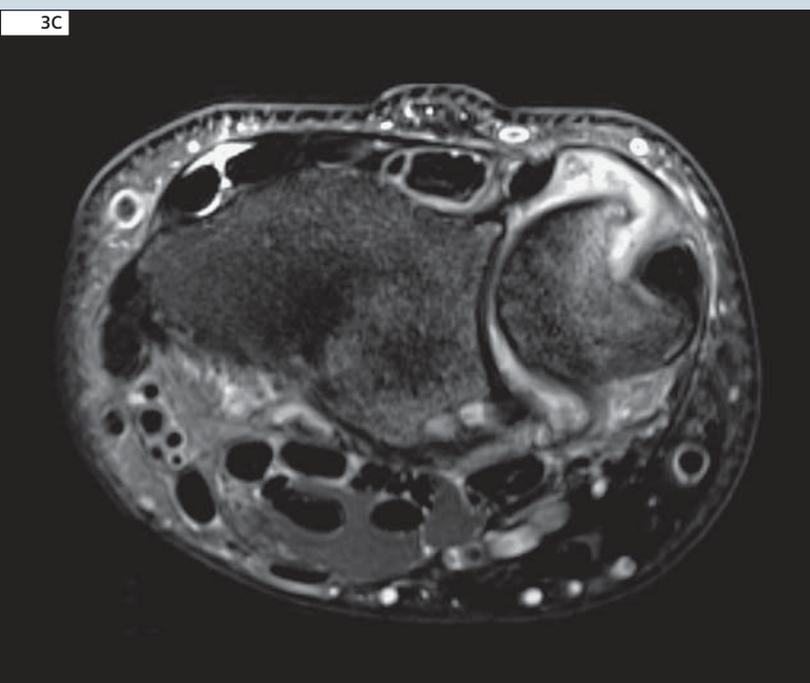
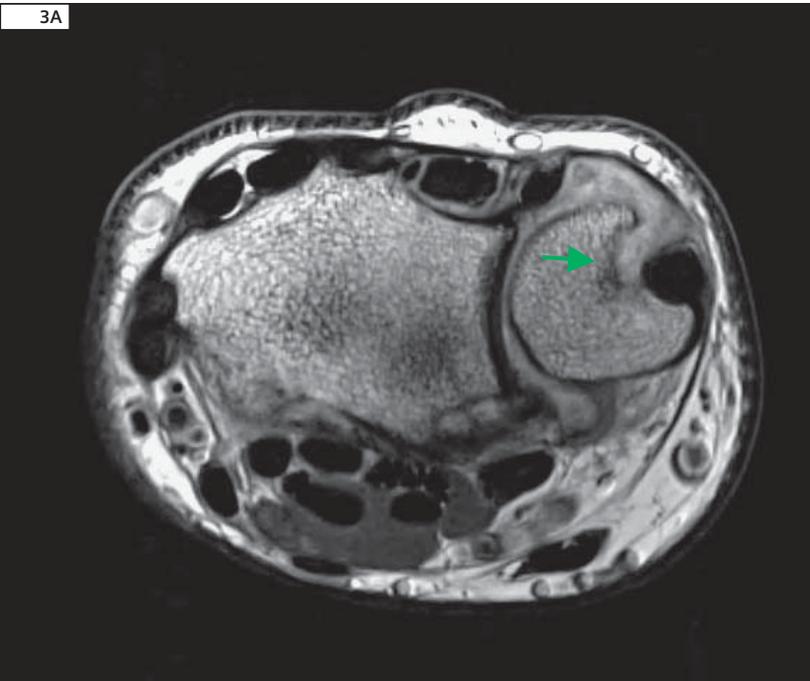
dard radiography has a sensitivity of 24% and a specificity of 99% [9]. In order to effectively diagnose bone erosions it is mandatory to run MRI protocols with thin slices (ideally 1 mm), high spatial resolution in two planes according to RAMRI score criteria [10]. Here 3 Tesla MRI could play an important role in the near future, due to its higher signal-to-noise ratio (SNR) and the potential to obtain high resolution isotropic 3D sequences.

The starting point of the disease process of RA seems to be the inflammation of the synovium which leads to high cellular inflammatory tissue, the pannus, which in turn is responsible for the cartilage and the bone destruction (bone erosion) [1, 11]. Synovium presents on MRI with low signal intensity on T1-weighted sequences and with high signal intensity on fluid sensitive sequences. Compared to free fluid the intensity signal of synovium is slightly higher on T1-weighted sequences and slightly lower on T2 fat set sequences.

Contrast-enhanced MRI is mandatory in RA patients in order to detect and quantify the inflammation synovium enhancement following the RAMRI score whereas detection and quantification of BME and bone erosion don't need the administration of Gadolinium. One interesting study of Agarwal et al. [12] used diffusion tensor imaging (DTI) in order to test an alternative method to evaluate the inflammation of the synovium in 18 patients and 6 volunteers. The principle of DTI is based on the isotropic or anisotropic movement of water molecules. The molecules are moving isotropic if they can move freely in all direction like in fluid whereas they are moving anisotropic if they are limited in the movement due to other components like in tissues. The degree and direction of diffusion can be described in a scale value from 0 to 1 [8]. In patients with RA the water molecules present a restricted motion due to the fractional anisotropy of the inflammatory fluid (effusion). According to the study these



2 Corresponding T1w contrast-media enhanced image to figure 1 is shown. Synovial enhancement as a consequence of acute inflammatory process is present in this case.



3 Synovial reaction and early signs of bone erosion of the ulnar styloid process. **3C**: Corresponding T2w image with fat saturation showing oedema within the bone and fluid collection as well as thickening of the synovia.

values are significantly altered in patients with RA compared to healthy individuals. DTI is by no doubt an interesting and promising technique to study the synovial inflammation with the great advantage of the absence of contrast media and the risks that are connected such as nephrogenic systemic fibrosis [13], acute adverse reactions and other, less important but common adverse events, such as extravasations. Moreover, according to the study, DTI has a similar sensitivity in detection synovial inflammation compared to conventional T1-weighted fast SE fat sat sequences. According to one recent study [14] even flexor tendosynovitis, that can be detected with MRI or sonography, has a strong predictive value for RA. The study analyzed 99 patients with unspecified

arthritis or suspected RA and negative standard X-rays. Moreover by adding RF or anti-cyclic citrullinated peptides (anti-CCP) the predictor was stronger with a sensitivity of 83% and a specificity of 63% [14].

Conclusion

MRI is an imaging modality that can detect RA in the early stadium. This information is mandatory to start appropriate therapies that are very expensive and that carry potential severe adverse reactions. Due to its higher SNR and the potential increase in spatial resolution, 3 Tesla MRI will probably play an important role to improve the detection of bone erosions and to implement new technique in the near future without the need of gadolinium administration.

Contact

Filippo Del Grande, M.D., M.H.E.M.
Research Fellow
Johns Hopkins Hospital
Musculoskeletal Radiology
Russell H. Morgan Department of
Radiology and Radiological Science
600 N. Wolfe Street
Phipps B-100
Baltimore, MD 21287
USA
Fdelgra1@jhmi.edu

John A. Carrino, M.D., M.P.H.
Associate Professor of Radiology and
Orthopaedic Surgery
Johns Hopkins University School of
Medicine
Section Chief, Musculoskeletal Radiology
Russell H. Morgan Department of
Radiology and Radiological Science
600 N. Wolfe Street
Phipps B-100
Baltimore, MD 21287
USA
carrino@jhmi.edu

References

- 1 Kosta PE, Voulgari PV, Zikou AK, Drosos AA, Argyropoulou MI. The usefulness of magnetic resonance imaging of the hand and wrist in very early rheumatoid arthritis. *Arthritis Res Ther* 2011 Jun 9;13(3):R84.
- 2 Lard LR, Visser H, Speyer I, vander Horst-Bruinsma IE, Zwinderman AH, Breedveld FC, et al. Early versus delayed treatment in patients with recent-onset rheumatoid arthritis: comparison of two cohorts who received different treatment strategies. *Am J Med* 2001 Oct 15;111(6):446-451.
- 3 Ostergaard M, Edmonds J, McQueen F, Peterfy C, Lassere M, Ejlberg B, et al. An introduction to the EULAR-OMERACT rheumatoid arthritis MRI reference image atlas. *Ann Rheum Dis* 2005 Feb;64 Suppl 1:i3-7.
- 4 Peterfy CG, Countryman P, Gabriele A, Shaw T, Anisfeld A, Tsuji W, et al. Magnetic resonance imaging in rheumatoid arthritis clinical trials: emerging patterns based on recent experience. *J Rheumatol* 2011 Sep;38(9):2023-2030.
- 5 Duer-Jensen A, Horslev-Petersen K, Hetland ML, Bak L, Ejlberg BJ, Hansen MS, et al. Bone edema on magnetic resonance imaging is an independent predictor of rheumatoid arthritis development in patients with early undifferentiated arthritis. *Arthritis Rheum* 2011 Aug;63(8):2192-2202.
- 6 Hetland ML, Ejlberg B, Horslev-Petersen K, Jacobsen S, Vestergaard A, Jurik AG, et al. MRI bone oedema is the strongest predictor of subsequent radiographic progression in early rheumatoid arthritis. Results from a 2-year randomised controlled trial (CIMESTRA). *Ann Rheum Dis* 2009 Mar;68(3):384-390.
- 7 Olech E, Crues JV, 3rd, Yocum DE, Merrill JT. Bone marrow edema is the most specific finding for rheumatoid arthritis (RA) on noncontrast magnetic resonance imaging of the hands and wrists: a comparison of patients with RA and healthy controls. *J Rheumatol* 2010 Feb;37(2):265-274.
- 8 Borrero CG, Mountz JM, Mountz JD. Emerging MRI methods in rheumatoid arthritis. *Nat Rev Rheumatol* 2011 Feb;7(2):85-95.
- 9 Dohn UM, Ejlberg BJ, Hasselquist M, Narvestad E, Moller J, Thomsen HS, et al. Detection of bone erosions in rheumatoid arthritis wrist joints with magnetic resonance imaging, computed tomography and radiography. *Arthritis Res Ther* 2008;10(1):R25.
- 10 Drape JL. MRI in rheumatoid arthritis: a dual emergency. *J Radiol* 2008 May;89(5 Pt 1):543-545.
- 11 Cyteval C. MR imaging of the hands in rheumatoid arthritis. *J Radiol* 2010 Jan;91(1 Pt 2):111-119.
- 12 Agarwal V, Kumar M, Singh JK, Rathore RK, Misra R, Gupta RK. Diffusion tensor anisotropy magnetic resonance imaging: a new tool to assess synovial inflammation. *Rheumatology (Oxford)* 2009 Apr;48(4):378-382.
- 13 Jalandhara N, Arora R, Batuman V. Nephrogenic systemic fibrosis and gadolinium-containing radiological contrast agents: an update. *Clin Pharmacol Ther* 2011 Jun;89(6):920-923.
- 14 Eshed I, Feist E, Althoff CE, Hamm B, Konen E, Burmester GR, et al. Tenosynovitis of the flexor tendons of the hand detected by MRI: an early indicator of rheumatoid arthritis. *Rheumatology (Oxford)* 2009 Aug;48(8):887-891.