

# ORIGINAL ARTICLE

## Effectiveness of MRI scans in patients with rheumatoid arthritis

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### ABSTRACT:

**Background:** Rheumatoid arthritis is a predominantly joint-based disease affecting approximately 1% of the world's population. Magnetic resonance imaging (MRI) has been shown to be a highly sensitive technique for the detection of inflammatory soft tissue proliferation, bone oedema and early erosions, and since the implementation of MRI into the clinical practice, numerous cross-sectional papers concerning the MRI-detectable features of RA have been published. Hence; we assessed the effectiveness of MRI scans in patients with rheumatoid arthritis. **Materials & methods:** 300 patients with RA who underwent clinical assessment with MRI. Synovitis was scored on a 0–3 scale at three different locations: radioulnar joint, radiocarpal joint and intercarpal–carpometacarpal joints (total maximum score 9). A score of 0 is normal, with no enhancement or enhancement up to the thickness of normal synovium, while the scores from 1 to 3 (mild, moderate, severe) refer to increments of one-third of the presumed maximum volume of enhancing tissue in the synovial compartment. Blood samples were collected at some time prior to the MRIs and the presence or absence of RF and serum levels of CRP and anti-CCP antibodies were determined. All the results were analyzed by SPSS software. Chi-square test was used for the assessment of level of significance. **Results:** Percentage of males in group 1 and group 2 was 28 and 24 percent respectively. Mean duration of disease in group 1 and group was 134 and 96 months respectively. Mean number of tender joints in group 1 and group 2 was 6.1 and 8.9 respectively. Significant results were obtained while comparing the mean duration of diseases and mean number of tender joints in group 1 and group 2 respectively. In patients with less than 3 years of diseases duration, in 8.5 percent of the patients in group 1, treatment was unchanged. **Conclusion:** MRI-detected inflammation contributes to the identification of unclassified arthritis patients who will develop RA. MRI-detected tenosynovitis was most helpful, and the accuracy was the highest in UA patients who presented with oligoarthritis. Furthermore, RA was unlikely to develop in UA patients with a normal MRI.

**Key words:** MRI, Rheumatoid arthritis, tenosynovitis.

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### INTRODUCTION

Rheumatoid arthritis is a predominantly joint-based disease affecting approximately 1% of the world's population. It is a chronic systemic autoimmune disorder that primarily affects the synovium and if left untreated leads to disorganization and destruction of the joints. In turn, joint destruction results in severe deformity and disability. Synovial hypertrophy and angiogenesis develop in the chronic phase of the condition, the hypertrophied synovium becoming locally invasive at the synovium-cartilage interface where it is thought to be responsible for causing bone erosions and subsequent joint destruction.<sup>1</sup>

Clinical course and progression of RA has been shown to be modified by biological treatment, mainly with anti-tumour necrosis factor (TNF)  $\alpha$  agents.<sup>1,2</sup> Disease remission in rheumatoid arthritis has been traditionally considered when there is no clinical or biochemical evidence of disease activity.<sup>3</sup> Rheumatoid arthritis (RA) is a chronic disease that if untreated results not only in pain, but also in progressive joint damage and functional decline. Up to 75% of the joint damage occurs within the

first 5 years of disease onset and continues throughout the course of the disease.<sup>4</sup>

Magnetic resonance imaging (MRI) has been shown to be a highly sensitive technique for the detection of inflammatory soft tissue proliferation, bone oedema and early erosions, and since the implementation of MRI into the clinical practice, numerous cross-sectional papers concerning the MRI-detectable features of RA have been published.<sup>5-7</sup>

The ability of MRI to provide additional and more sensitive information than clinical examination or conventional radiography is well established. MRI can identify bone erosions earlier than conventional radiography and can detect bone marrow edema and synovitis, which may be important precursors to erosive disease

MRI sensitively depicts inflammation; it visualizes synovitis, tenosynovitis and bone marrow oedema (BME). BME (also called osteitis in RA) is not depicted by other imaging modalities.

Hence; this study aimed to assess the diagnostic value of MRI in patients with early RA whose diagnosis cannot be made upon initial presentation.

**MATERIALS & METHODS**

The present study was conducted in the department of the radiodiagnosis. 300 patients with RA who underwent clinical assessment with MRI. Ethical approval was taken from the institutional ethical committee in written after explaining the entire research protocol. Clinical assessments were gathered and documented in structured data collection forms. The clinical and lab encounter abstracted was the encounter prior to the MRI being obtained. Coronal T1 and STIR MR images of the affected hands, wrists, or feet were performed with a low-field strength dedicated extremity unit. The extremities imaged were the dominant wrist and/or the most affected joint and/ or an extremity with questionable areas apparent on plain radiographs. The field of view for each sequence was 11 mm and the slice thickness was between 1.05 and 1.1 mm. Images was interpreted by one of four fellowship- trained musculoskeletal radiologists. Synovitis was scored on a 0–3 scale at three different locations: radioulnar joint, radiocarpal joint and intercarpal–carpometacarpal joints (total maximum score 9). A score of 0 is normal, with no enhancement or enhancement up to the thickness of normal synovium, while the scores from 1 to 3 (mild, moderate, severe) refer to increments of one-third of the

presumed maximum volume of enhancing tissue in the synovial compartment. Blood samples were collected at some time prior to the MRIs and the presence or absence of RF and serum levels of CRP and anti-CCP antibodies were determined. Anti-CCP antibodies were detected using ELISA with the second generation CCP test. ESR was also measured for each patient. RF and CCP were considered to be negative if they were less than 20 units. All the results were analyzed by SPSS software. Chi-square test was used for the assessment of level of significance.

**RESULTS**

**Graph 1** shows the demographic details of the patients. Mean age of the patient sin group 1 and group 2 was 53.2 and 50.1 years respectively. Percentage of males in group 1 and group 2 was 26 and 21 percent respectively. Mean duration of disease in group 1 and group was 141 and 99 months respectively. Mean number of tender joints in group 1 and group 2 was 7.5 and 10.1 respectively. **Table 1** shows p-value for the demographic details of the patients. Significant results were obtained while comparing the mean duration of diseases and mean number of tender joints in group 1 and group 2 respectively. **Graph 2** shows RA treatment status in patients after baseline MRI. In patients with less than 3 years of diseases duration, in 8.5 percent of the patients in group 1, treatment was unchanged.

**Table 1:** Baseline characteristics of all rheumatoid arthritis patients

Parameter	Group 1	Group 2	p-value
Mean age (years)	53.2	50.1	0.25
Males (%)	28	24	0.12
Mean duration of disease (months)	134	96	0.02*
Mean number of swollen joints	6.1	8.9	0.52
Mean number of tender joints	5.5	10.1	0.02*

\*: Significant

**Table 2:** Diagnosis at presentation (n=300)

RA	136
Unclassified arthritis	86
PsA or spondyloarthritis	9
Inflammatory OA	35
Reactive arthritis	6
Crystal arthropathy	5
RS3PE	8
SLE + MCTD	3
Other diagnoses	12

RS3PE: remitting seronegative symmetrical synovitis with pitting oedema.

**Table 3:** p-value for the RA treatment status in patients after baseline MRI

Parameter		Group 1	Group 2	p-value
Total patients	% of patients with unchanged treatment	18	92	0.01*
	% of patients with changes treatment	82	8	0.01*
Less than 3 year disease duration	% of patients with unchanged treatment	8.5	85.5	0.01*
	% of patients with changes treatment	91.5	14.5	0.01*
Less than 3 year disease duration	% of patients with unchanged treatment	24.5	75.5	0.01*
	% of patients with changes treatment	75.5	24.5	0.01*

## DISCUSSION

Early treatment requires early identification of RA. This is difficult if patients present with UA. It is inextricably linked to early recognition that the phenotype may not yet be completely matured; additional tests are therefore needed. When using the 2010 criteria, UA patients are mainly ACPA negative, as was also shown here. The regular predictors such as CRP and the number of swollen joints also have a limited predictive value. As it has been advocated that MRI-detected inflammation is valuable for the early identification of RA, this study aimed to assess the diagnostic value of MRI in patients with early RA whose diagnosis cannot be made upon initial presentation.<sup>8</sup> MRI is an important imaging technique that provides multiplanar images and is able to visualize a range of joint structures, including synovium, tendons, ligaments, bone, and cartilage. It does not use radiation, so it can be repeated as much as necessary, and allows longitudinal assessment. With the advances in sequence analysis software and lower costs, MRI is likely to become more accessible. MRI is recognized as the imaging technology of choice for visualization of the inflamed synovial membrane and bone edema. Furthermore, MRI has been shown to be a sensitive, non-invasive method for detection and quantification of bone erosions. Erosions are visible on MRI on average two years before they are visible on radiographs and may become consistently visualized on radiographs of the metacarpophalangeal (MCP) joints only when 20%-30% of the bone is eroded on MRI.<sup>9-11</sup>

In the present study, a relatively low level of clinical disease activity with a mean number of swollen or tender joints for both MRI-positive and MRI-negative patients was seen. Fox et al determined the impact of enhanced MRI on patient management in a group of patients referred for MRI by rheumatologists. The study included 48 patients with a mean age of 51 years. Significant management changes initially occurred in 79% of the positive and in 11% of the negative MR examinations with average follow-up of ~300 days. From the results, they concluded that enhanced MRI significantly altered clinical management in 50% of these patients with RA or suspected RA. Therefore, when the clinical picture in a patient with RA or suspected RA is unclear, enhanced MRI can provide useful guidance for treatment modifications.<sup>12</sup> Brown et al studied 107 RA patients receiving disease-modifying antirheumatic drug therapy who were judged by their consultant rheumatologist to be in remission and 17 normal control subjects. Patients underwent clinical, laboratory, functional, and quality of life assessments. The Disease Activity Score 28-joint assessment and the American College of Rheumatology remission criteria, together with strict clinical definitions of remission, were applied. Imaging of the hands and wrists using standardized acquisition and scoring techniques with conventional 1.5T magnetic resonance imaging (MRI) and ultrasonography (US) were performed. Irrespective of which clinical criteria were applied to determine remission, the majority of patients continued to have evidence of active inflammation, as

shown by findings on the imaging assessments. Most RA patients who satisfied the remission criteria with normal findings on clinical and laboratory studies had imaging-detected synovitis. This subclinical inflammation may explain the observed discrepancy between disease activity and outcome in RA. Imaging assessment may be necessary for the accurate evaluation of disease status and, in particular, for the definition of true remission.<sup>13</sup> Palosaari et al investigated if disease assessment by contrast-enhanced dynamic and static magnetic resonance imaging (MRI) and quantitative nanocolloid (NC) scintigraphy gives useful additional information in early rheumatoid arthritis (RA). The baseline MRI bone oedema score ( $\rho=0.67$ ), MRI synovitis score ( $\rho=0.57$ ), ESR ( $\rho=0.56$ ), CRP ( $\rho=0.48$ ), E-rate ( $\rho=0.47$ ) and (99m)Tc-NC uptake ( $\rho=0.45$ ) were related with the change in the MRI erosion score from baseline to 2 yrs ( $\rho=$  Spearman's correlation). In the multivariate logistic regression model, the bone marrow oedema score was the only baseline variable that predicted erosive progression at 2 yrs' follow-up (OR 4.2, 95% CI 1.3-13.8). The median (interquartile range) change in the erosion score from baseline to 2 yrs was 0 (0, 0) and 4 (2, 5) in the patients with ( $n=9$ ) and without ( $n=15$ ) a persistent clinical response over the 2 yrs, respectively ( $P=0.001$ ). The non-responders who presented with erosive progression from 1 yr to 2 yrs had higher MRI synovitis scores, bone oedema scores, E-rate and (99m)Tc-NC uptake at 1-yr follow-up than the non-responders without progressive bone damage. The degree of local synovial inflammation at baseline, evaluated by dynamic and static MRI and quantitative NC scintigraphy, is closely related to the progression of wrist joint erosions during the first 2 yrs of the disease. Furthermore, at follow-up, if no persistent clinical response is achieved, these imaging methods may help to predict future erosiveness and help in clinical therapeutic decision making. Inflammatory changes (synovitis and bone marrow edema) and destructive changes (bone erosion) were evaluated by magnetic resonance imaging (MRI) in patients with rheumatoid arthritis (RA), and their relations with disease activity were assessed during treatment with tumor necrosis factor (TNF) inhibitors. Ten patients with early active RA underwent MRI at 0 and 16 weeks of TNF-inhibitor treatment. The carpal bones of the dominant hand were evaluated by the outcome measures in rheumatology clinical trials MRI score for RA. After 16 weeks, the mean disease activity score (DAS 28) decreased significantly from 5.54 to 2.70, while the number of tender joints, number of swollen joints, and inflammatory parameters were also significantly improved. The mean synovitis and marrow edema scores determined by MRI showed a significant decrease from 6.1 to 2.2 and 12.8 to 6.2, respectively, while the annual bone-erosion progression score decreased from 12.6 to 2.0. Although synovitis persisted in some patients, imaging remission was achieved in two patients. In conclusion, TNF-inhibitor therapy achieved an early decrease of disease activity and MRI revealed

amelioration of joint destruction. The MRI score for RA is useful for assessing the early response to TNF inhibitors.<sup>14-16</sup>

Studies in past has suggested that MRI-detected tenosynovitis was associated with the development of RA in patients with unclassified arthritis (UA) independent of other inflammatory measures, including swollen joints and elevated CRP.

The negative predictive value for MRI tenosynovitis is high in RA, but its positive predictive value is limited.

MRI was found to be the most diagnostic medium in patients with UA presenting with oligoarthritis (ie, 2 to 4 swollen joints).

Patients with UA presenting with a normal MRI were unlikely to develop RA, suggesting that MRI can be used in the early diagnostic process of the disease.<sup>17-19</sup>

### CONCLUSION

From the above results, the authors conclude that useful information regarding the treatment therapy is provided by a single MRI done during the phase of treatment. MRI-detected inflammation contributes to the identification of unclassified arthritis patients who will develop RA. MRI-detected tenosynovitis was most helpful, and the accuracy was the highest in UA patients who presented with oligoarthritis. Furthermore, RA was unlikely to develop in UA patients with a normal MRI.

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