## REVIEW

Rheumatic & Musculoskeletal Diseases

RMD

# Improving inflammatory arthritis management through tighter monitoring of patients and the use of innovative electronic tools

Piet van Riel,<sup>1,2</sup> Rieke Alten,<sup>3</sup> Bernard Combe,<sup>4</sup> Diana Abdulganieva,<sup>5</sup> Paola Bousquet,<sup>6</sup> Molly Courtenay,<sup>7</sup> Cinzia Curiale,<sup>8</sup> Antonio Gómez-Centeno,<sup>9</sup> Glenn Haugeberg,<sup>10</sup> Burkhard Leeb,<sup>11,12,13</sup> Kari Puolakka,<sup>14</sup> Angelo Ravelli,<sup>15</sup> Bernhard Rintelen,<sup>11,12</sup> Piercarlo Sarzi-Puttini<sup>16</sup>

## ABSTRACT

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For numbered affiliations see end of article.

#### **Correspondence to**

Professor Piet van Riel; piet.vanriel@radboudumc.nl

Treating to target by monitoring disease activity and adjusting therapy to attain remission or low disease activity has been shown to lead to improved outcomes in chronic rheumatic diseases such as rheumatoid arthritis and spondyloarthritis. Patient-reported outcomes, used in conjunction with clinical measures, add an important perspective of disease activity as perceived by the patient. Several validated PROs are available for inflammatory arthritis, and advances in electronic patient monitoring tools are helping patients with chronic diseases to self-monitor and assess their symptoms and health. Frequent patient monitoring could potentially lead to the early identification of disease flares or adverse events, early intervention for patients who may require treatment adaptation, and possibly reduced appointment frequency for those with stable disease. A literature search was conducted to evaluate the potential role of patient self-monitoring and innovative monitoring of tools in optimising disease control in inflammatory arthritis. Experience from the treatment of congestive heart failure, diabetes and hypertension shows improved outcomes with remote electronic self-monitoring by patients. In inflammatory arthritis, electronic self-monitoring has been shown to be feasible in patients despite manual disability and to be acceptable to older patients. Patients' self-assessment of disease activity using such methods correlates well with disease activity assessed by rheumatologists. This review also describes several remote monitoring tools that are being developed and used in inflammatory arthritis, offering the potential to improve disease management and reduce pressure on specialists.

## Key messages

## What is already known about this subject?

- Treating to target in chronic rheumatic diseases such as rheumatoid arthritis and spondyloarthritis by monitoring disease activity and adjusting therapy to attain remission or low disease activity has been shown to lead to improved outcomes.
- Patient-reported outcomes add the patient's perspective of disease activity to that of clinical measures.

#### What does this study add?

- Diseases outside of inflammatory arthritis show improved outcomes with remote self-monitoring by patients.
- Frequent remote patient monitoring could lead to early identification of disease flares, early intervention for patients requiring treatment adaptation or reduced appointment frequency for stable patients.

### How might this impact on clinical practice?

There are various remote monitoring tools for inflammatory arthritis in use or being developed with the potential to help improve disease management.

arthritis (RA) and spondyloarthritis (SpA), including ankylosing spondylitis (AS) and psoriatic arthritis (PsA). Integral to the principle of treating to target is that disease activity is measured on a regular basis and therapy is adjusted accordingly to achieve a target agreed by the physician and the patient.<sup>1 2</sup> Targeting low-disease activity or remission in the management of RA is part of the European League Against Rheumatism (EULAR) recommendations and, as has

A tight control or treat-to-target management strategy has become the standard of care for rheumatic diseases such as rheumatoid

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previously been widely reported (eg, the DREAM, TICORA and CAMERA studies), this has been shown to lead to improved outcomes.<sup>3–7</sup> In a recent study, patients with RA who achieved guideline-recommended low disease activity (Disease Activity Score (DAS)28-CRP <2.6) used fewer healthcare resources, including a 36-45% lower hospital admission rate (p<0.05), compared with patients who did not achieve target disease activity levels.<sup>8</sup> In outpatient clinics that monitor patients using outcome measures as standard practice, ~75% of patients with RA have been reported to be in remission or in low disease activity.<sup>9</sup> Furthermore, patients in remission who are used to modern technology have started to request the possibility of reporting their disease status by using their personal technology devices, for example, home PCs.

The treat-to-target strategy for RA was originally adopted from the treatment of hypertension and diabetes, where it resulted in considerable improvements in outcomes, however, with the difference that hypertension and diabetes are diseases not providing the patient with immediate alerts, namely with pain. Treat-to-target strategy in inflammatory arthritis usually uses clinical measurements of disease activity such as DAS28 or Bath Ankylosing Spondylitis Disease Activity Index (BASDAI). While EULAR recommends the use of composite indices to assess disease activity, the American College of Rheumatology (ACR) recommends both composite indices or patient-driven composite tools (PAS, PAS-II or Routine Assessment of Patient Index Data (RAPID) 3).<sup>10</sup> As such, in addition to clinical disease activity measures, the patient's perspective of disease state and burden is increasingly being recognised as an important consideration,<sup>11-13</sup> and the most recent ACR guidelines refer to the use of patient-reported outcomes (PROs) as activity measures.<sup>14</sup> The US Food and Drug Administration defines a PRO as: "any report of the status of a patient's health condition that comes directly from the patient, without interpretation of the patient's response by a clinician or anyone else. The outcome can be measured in absolute terms (eg, severity of a symptom, sign or state of a disease) or as a change from a previous measure".<sup>15</sup> PROs for inflammatory arthritis have been developed and validated to correlate closely with clinical measures of disease activity. PROs have been described as 'critical, relevant and complementary' in the context of the physician-patient interaction.<sup>16</sup> It is of fundamental importance for the treating physician to have PRO information before making decisions on treatment and/or new interventions.<sup>16</sup> Pincus *et al*<sup>17</sup> have shown that the combination of the three PRO measures from the seven ACR core data set measures is as informative as the ACR20 responses and DAS scores in distinguishing between placebo and effective treatment. The Rheumatoid Arthritis Disease Activity Index (RADAI)-5 PRO showed a similar outcome concerning remission rates as the Simple Disease Activity Index (SDAI)-remission and even has a higher sensitivity to indicate the 2011

EULAR/ACR Boolean remission criteria than the SDAI-remission criterion.<sup>18</sup> Thus, PROs could act as a surrogate for clinical measurement and enable remote monitoring of patients with RA.

In many other disease areas, such as diabetes, hypertension and congestive heart failure, patient selfmonitoring is a well-accepted and common practice in supporting a tight control strategy. For example, among patients with hypertension at high risk of cardiovascular disease, self-monitoring with self-titration of antihypertensive medication resulted in a 9.2 mm Hg lower systolic blood pressure at 12 months compared with usual care.<sup>19</sup> Advances in e-health tool technology are helping patients with chronic diseases to self-monitor and assess their symptoms and health, facilitating the incorporation of routine collection of PROs into clinical practice.<sup>20</sup>

## Need for frequent patient monitoring in inflammatory arthritis

The consequences of poor control of inflammatory arthritis include swollen and painful joints, irreversible joint damage, functional disability, decreased work productivity, sleep disturbance and a reduced ability of patients to lead a normal active life. In addition to advocating treat-to-target strategy, the EULAR treatment recommendations suggest that clinic visits should be scheduled every 1–3 months when treating rheumatic disease with biologics.<sup>3</sup> However, this frequency of visits may not always be possible due to specific barriers such as geographical and health-system-related constraints and, even in the case of high-quality care, patients' lives may not be predictable due to disease activity fluctuations.<sup>21</sup>

Frequent remote patient monitoring could potentially lead to the early identification of disease flares, prioritisation of patients who may require a treatment review, and possibly reduced appointment frequency for those with stable disease. Evidence has shown that fluctuations in disease activity do have a direct effect on the destruction of joints.<sup>22</sup> Therefore, after the newly diagnosed patient has reached a state of remission or low disease activity, there might be a potential benefit of remote monitoring carried out in between regular scheduled clinic visits to ensure that disease activity remains tightly controlled. Remote patient monitoring may also reduce the number of visits to the physician's office and be more convenient for many patients, especially those who are functionally incapacitated or who live far away from the nearest rheumatology clinic.<sup>23</sup>

There is evidence to support a correlation between higher patient engagement in their treatment and improved adherence to therapy.<sup>24</sup> Self-monitoring by patients is one method that can potentially increase engagement with their treatment. Self-monitoring may also lead to more consistent reporting in the long term, as outcomes are reported by the same person over time. While it is generally acknowledged that PROs are a subjective measurement, patients are best placed to provide evaluations of their pain and global estimates of well-being.

In summary, self-monitoring of PROs by patients may lead to improved disease control, potential early identification of disease flares and improved convenience for patients over clinic visits. Patients may also be more engaged in their treatment and improve their adherence to therapy.

## Aims

This review considers the role of PROs and patient selfmonitoring in inflammatory arthritis. A search of the literature was conducted to look at the potential of electronic patient monitoring tools to support this role. The authors' personal experiences of certain tools are included, as well as examples of electronic monitoring tools currently in use from disease areas other than inflammatory arthritis.

## LITERATURE SEARCH METHODOLOGY

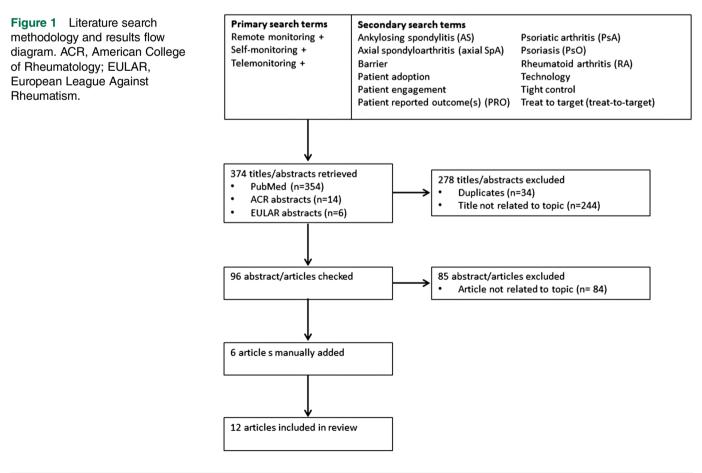
Patient self-monitoring and remote monitoring with innovative monitoring tools is a well-accepted and common practice in several chronic disease areas. A literature search was conducted to evaluate the potential role of patient self-monitoring and innovative monitoring tools in optimising disease control in inflammatory arthritis. Readers should note that this was a narrative review and the methods of the review are explained below.

The following databases were searched: PubMed (January 2000 to June 2015), accepted abstracts from ACR and EULAR annual congresses (ACR 2012–2014 and EULAR 2011–2015). Figure 1 shows the search terms used. Searches were performed using a combination of a single primary search term in conjunction with each secondary term (primary term AND secondary term). Relevance to the topic was determined by scanning the title and, where available, the abstract of the retrieved articles. Hits were collated and manually de-duplicated.

Examples that have been drawn from other disease areas were gathered from the authors' experience in order to add a wider healthcare context outside of rheumatology.

### LITERATURE SEARCH RESULTS

The literature search retrieved 374 titles/abstracts (354 titles from PubMed, 14 ACR abstracts, 6 EULAR abstracts), of which 278 were excluded by the initial scan and collation (34 duplicates, 244 titles not related to topic). The remaining 96 abstracts/articles were checked for relevance; 85 were excluded as not related to the topic, leaving 11 articles.<sup>20 23 25–33</sup> The findings of the initial literature search were reviewed and the authors suggested additional articles for inclusion that were not found as a result of search terms or human



error of the manual searching of databases. After manual addition of these 6 articles,<sup>8</sup>  $9^{-34-37}$  17 articles were selected for inclusion in the review (see online supplementary appendix A and figure 1). Key evidence from the literature search is summarised below.

## DISCUSSION—LITERATURE SEARCH FINDINGS

Innovative electronic remote monitoring and PRO reporting solutions could enable better data capture, easier incorporation of data into electronic medical records, and more frequent monitoring of disease activity in patients with RA between clinic visits.<sup>23</sup> Remote data collection offers the additional advantage of convenience to patients, especially those who are functionally incapacitated or who live far away from the nearest rheumatology clinic, as the data can be collected at home. Remote monitoring and reporting of PROs may facilitate a treat-to-target approach and help to achieve a low disease activity state or remission among patients with RA.<sup>23</sup> Results from a 2004 survey of 135 US physicians indicated that a large majority (83%) thought that remote patient monitoring would prove beneficial for the healthcare industry.<sup>33</sup> Their main concerns related to the privacy of medical information on the internet and the security of online transactions.

Specific data on the use of patient-led remote PRO monitoring tools in inflammatory arthritis appear to be relatively sparse based on the search criteria employed in the current review. This apparent knowledge gap suggests that research into the utility of electronic PRO reporting tools in inflammatory arthritis is warranted. Areas to be investigated include: any differences in longterm outcomes in patients remotely monitoring PROs; patient satisfaction after long-term use of such a remote monitoring tool; any difference in number of clinic visits or healthcare resource usage among patients using remote monitoring tools versus those not; any difference in cost of treatment; potential barriers to implementation of such tools.

#### Potential for remote monitoring of inflammatory diseases

Electronic remote monitoring tools for inflammatory rheumatic diseases offer additional data to support clinical decision-making, may improve the quality of care by effective patient communication and contribute to empowerment of patients.<sup>32</sup> The use of electronic remote monitoring tools to support tight control in such diseases is of great interest to rheumatologists, given the need for tight disease control to prevent symptoms, avoid joint damage and recognise complications early.

Technology for remote monitoring should be simple and practical to use. In addition, monitoring systems should be automated where possible in order to spare staff resources. There is a concern that use of IT applications by patients with RA may be limited by their age and manual disability. However, a recent study found that manual disability in patients with RA is not an obstacle for using mobile applications.<sup>26</sup> The mobile application for smartphones that was tested comprised a simple questionnaire over four screens. Fifteen patients with RA with an average age of  $63\pm10$  years completed the questionnaire twice, taking  $91\pm23$  s the first time and  $49\pm20$  s the second time. All patients agreed that the application was generally easy to use and intuitive, and that the mobile visual analogue scale was at least as easy to complete as in paper form.

A study in 153 patients with RA, systemic lupus erythematosus or SpA compared completion of standardised questionnaires using paper and pencil or electronically on a tablet PC.<sup>32</sup> The scores obtained by the two methods did not differ, and patients reported no major difficulties using the tablet PC. Most patients (62%) expressed a preference for using remote data entry in the future, while 7 (5%) patients felt uncomfortable with the tablet PC due to their rheumatic disease.

Disease activity measured by patients and reported with an electronic tool has been shown to correlate well with DAS28 results from a clinical examination.<sup>28</sup> A study of 51 patients with rheumatic disease reported a high correlation of 0.88 for DAS, with moderate correlation (0.63) for number of tender joints and a lower correlation (0.41) for number of swollen joints. In 37 (73%) patients, self-monitoring and the clinical examination by the physician resulted in an identical classification for low, moderate or high disease activity, with self-monitoring resulting in a higher classification in 12 (24%) cases and a lower classification in 2 (4%) cases.<sup>28</sup>

In another study, patients' self-assessment of disease activity (RAPID3 and 4) correlated strongly with that of rheumatologists (DAS44, Clinical Disease Activity Index (CDAI), SDAI).<sup>25</sup> Ninety patients with RA with a mean age of  $55\pm14$  years were educated to use a smartphone application for self-assessment, with weekly questionnaires to complete. Strong correlations were seen between patient and rheumatologist assessment of disease activity when comparing RAPID3 and DAS44 (R=0.60), CDAI (R=0.53) and SDAI (R=0.49), with similar correlations seen with RAPID4.<sup>25</sup>

## Use of electronic patient monitoring tools in other disease areas

A variety of electronic patient monitoring tools are already well accepted in other chronic disease areas (table 1). In cardiology and congestive heart failure, for example, patients undergoing cardiac resynchronisation therapy who were followed with quarterly in-office visits without a daily remote monitoring system had an 86% higher risk of delayed detection of adverse events, during a median follow-up of 7 months, than those who used remote monitoring.<sup>29</sup>

In diabetes management, a combined programme of automated telemonitoring, clinician notification and informal caregiver involvement was associated with consistent improvements in adherence to treatment,

areas				
Disease area	Participants	Intervention	Follow-up	Outcome
Cardiology/ congestive heart failure <sup>29</sup>	99 patients receiving cardiac resynchronisation therapy	Daily remote monitoring (RM) vs standard programme of in-office visits	7 months	Rate of detection of clinical adverse events was 23.8% in the RM group vs 48.7%; HR 0.14 (95% CI 0.06 to 0.37)
Diabetes <sup>30</sup>	301 patients with type 2 diabetes	Automated telemonitoring, clinician notification and informal caregiver involvement	3–6 months	Significant improvements over time in long-term medication non-adherence, physical functioning, depressive symptoms and diabetes-related distress (all p<0.001). Significant improvements in patient-reported frequency of weekly medication adherence, self-monitored blood glucose (SMBG) performance, checking feet and abnormal SMBG readings
Hypertension <sup>31</sup>	778 patients taking antihypertensive drugs	Usual care vs usual care with home blood pressure monitor (BPM) vs web-based pharmacist care with home BPM	12 months	55% of patients in the pharmacist-care group vs 37% in the usual care with home BPM group had BP <140/90 mm Hg. Home BPM accounted for 30.3% of the intervention effect, secure electronic messaging for 96%, and medication intensification for 29.3%

Table 1	Summary of evidence of impact of remote patient monitoring tools on patient outcomes across various disease
areas	

diabetes self-management behaviours, physical functioning and psychological distress.<sup>30</sup> A study on a remote monitoring tool in diabetes found that the ability to raise an automatic alert in case of measurements below or above certain limits offered a sense of security, and treating physicians were able to follow the therapeutic course in an easy and timely manner. Furthermore, the remote nature of the monitoring may be especially favourable for elderly, sometimes immobile patients.<sup>27</sup>

Another example comes from the treatment of hypertension. A study evaluated the role of home monitoring, communication with pharmacists, medication intensification, medication adherence and lifestyle factors in contributing to the effectiveness of an intervention to improve blood pressure control in patients with uncontrolled essential hypertension.<sup>31</sup> Study arms analysed were usual care with а home blood pressure monitor and pharmacist-assisted care with a home blood pressure monitor delivered via a patient website. At 12 months follow-up, patients in the web-based pharmacist care group were more likely to have a blood pressure below 140/90 mm Hg compared with patients in the group with home blood pressure monitors only (55% vs 37%; p=0.001). The effect of web-based pharmacist care on improved blood pressure control was explained in part through a combination of home blood pressure monitoring, secure messaging and antihypertensive medication intensification.<sup>31</sup>

# DISCUSSION—REMOTE MONITORING TOOLS FOR INFLAMMATORY ARTHRITIS

Further to the results of the literature search discussed above, several of the current authors have personal experience with remote monitoring tools being developed for use in inflammatory arthritis. These tools are described below and summarised in table 2. They represent only a sample of the existing tools; many rheumatology registries also make use of web-based tools.

## GoTreatIT Rheuma (Norway)

The GoTreatIT tool (http://www.diagraphit.com) was developed as a hospital computer system for patient monitoring in clinical practice. The tool incorporates disease measures (all in Norwegian and English) and PRO tools (most of them available in more than 20 languages). It is currently used in 13 hospital centres and by 3 private practising rheumatologists in Norway, and other centres have plans to use it. The tool is used for data collection to the national arthritis registry (NorArtritt). Furthermore, GoTreatIT is also used by more than 10 rheumatology centres in Finland and used for data collection to the Finnish arthritis registry (ROB-FIN). It has been used in a cross-sectional study reporting similar disease burdens in RA, PsA and axial SpA, to compare disease status and treatment in RA between Norway and Finland, and to explore the

ТооІ	Disease	PROs/disease activity measures available	Platform	Automatic alerts for healthcare professionals	Patient's ability to view results	Data security
iMonitor(http://www.pfizer.co.uk/content/ medical-and-educational-goods-and -services-megs-imonitor)	RA PsA AS	BASFI BASDAI HAQ Pt-DAS28 RADAI5 RAID RAPID3	PC Tablet Smartphone	✓	✓	✓
GoTreatIT (http://www.diagraphit.com)	RA PsA Axial SpA	DAS DAS28 BASDAI/ASDAS Patient reported joint pain HAQ MHAQ MDHAQ VAS pain fatigue QUEST RA questions PROMIS20 RAID BASFI/BASG	PC Tablet (mobile phones soon to be supported)	✓ (Alerts for patients when a report is due is under development)	1	•
Sanoïa (http://www.sanoia.com)	RA PsA AS	HAQ RAID RAPID3 ASAS NSAID ASAS QOL ASAS HI BASDAI BASFI	PC Tablet Smartphone	1	1	1
Andar (http://www.sanoia.com)	RA	RAPID3 DAS28 SDAI CDAI	PC	✓	1	1

Table 2 Some examples of remote monitoring tools available for inflammatory arthritis, based on authors' experience

AS, ankylosing spondylitis; ASAS HI, Assessment of SpondyloArthritis international Society Health Index; ASAS NSAID, Assessment of SpondyloArthritis international Society Nonsteroidal Antiinflammatory Drug; ASAS QoL, Assessment of SpondyloArthritis international Society Quality of Life; ASDAS, Ankylosing Spondylitis Disease Activity Score; Axial SpA, spondyloarthritis; BASFI, Bath Ankylosing Spondylitis Functional Index; BASG, Bath Ankylosing Spondylitis Global assessment; HAQ, Health Assessment Questionnaire; MDHAQ, Multidimensional Health Assessment Questionnaire; MHAQ, Modified Health Assessment Questionnaire; PROMIS20, Patient-Reported Outcomes Measurement Information System20; Pt-DAS28, Patient Derived Disease Activity Score28; QUEST RA, Quantitative Patient Questionnaires in Standard Monitoring of Patients with Rheumatoid Arthritis; RA, rheumatoid arthritis; RAID, Rheumatoid Arthritis Impact of Disease; PROs, patient-reported outcomes; PsA, psoriatic arthritis; VAS, Visual Analog Scale.

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change in disease status and treatment in patients with RA in a 10-year period at an outpatient clinic.

A technical solution called GoTreatIT Web has recently been developed which allows the patient to report their disease status via the internet directly into the hospital system using secure transfer of information to the hospital server. The self-reported data become immediately visible for healthcare personal at the outpatient clinic. A 2012 presentation at the EULAR congress reported clinical workflow efficiencies with use of the tool, by combining patient monitoring and registry data collection in a single workflow.

## Sanoïa (France)

Sanoïa (http://www.sanoia.com), launched in 2010, provides online secure health records that allow patients to track and store their own health data. It is available in 14 languages on PCs, tablet computers and smartphones. Forms such as BASDAI, Health Assessment Questionnaire, ASAS-QoL, RAPID3 and treatment trackers are available, and the patient can generate and print reports and graphical summaries. The patient decides whether to allow the physician to see their data. In September 2015, 4695 patients with RA were registered, and patients with AS and PsA started using the tool. A randomised controlled trial is underway to evaluate the effect of the tool on the quality of patient–doctor interactions (http://www.clinicaltrials. gov/ct2/show/NCT02200068).

## Andar (Spain)

Andar (http://www.proyectoandar.com) is a standardised monitoring tool in which the patient completes the RAPID3 questionnaire and clinical and laboratory measurements can be added by the healthcare professional, allowing calculation of composite indices (DAS28, SDAI, CDAI). Initially, this was developed as a paper-based questionnaire that patients completed before each clinic visit. It has now been developed as a web-based tool. Patients determine their own treatment targets and can view the evaluations. Physicians add blood test results, and nurses decide whether patients need urgent visits on the basis of monthly alerts.

## iMonitor

iMonitor is a web-based software application that allows patients to report information about their disease state for RA, PsA and AS. It can be accessed by PC, tablet or smartphone. Data are protected during storage and transmission and are encrypted using a PIN code entered by the user. Physicians can choose from a selection of PROs and set individual treatment targets and thresholds for each patient. The physician can then review PRO results entered by patients before an appointment, and real-time monitoring keeps them up to date with their patient's condition. Physicians receive alerts when established thresholds are not met or if PROs are not completed on time. Those patients with poor disease control can be prioritised, contacted and reviewed, as needed.  $^{\rm 37}$ 

## Patient groups most likely to benefit

Certain patients may particularly benefit from the use of remote monitoring tools. For example, patients with early RA who are most likely to benefit from a treat-to-target strategy may be the first candidates to adopt such tools. Others who may be suitable include patients with a high technological understanding, those with high engagement with their own disease management, those with barriers to frequent clinic visits (eg, poor mobility or great distance from the clinic), and those at high risk of flare or with a high need for monitoring (eg, patients whose disease activity fluctuates greatly between clinic visits). In addition, patients with stable disease may also be a target group for use of remote monitoring tools which allow them to report a stable condition without needing to attend a clinic for assessment.

## **DISCUSSION—FUTURE PERSPECTIVE**

As the cultural trend of moving towards digital monitoring and record keeping in healthcare develops, we anticipate that further work to develop the current and future range of remote PRO monitoring tools will continue. Our current review and search criteria highlighted a low number of published articles specifically relating to remote PRO reporting tools. While we recognise the limits of our search, there is a need for greater interest and research in the potential benefits of these tools.

## CONCLUSIONS

A treat-to-target strategy targeting low disease activity or remission in the management of RA is the standard of care and has been shown to lead to better outcomes. Remote monitoring and reporting of PROs may facilitate a treat-to-target approach and help to achieve a low disease activity state or remission among patients with RA.<sup>23</sup> PROs used in conjunction with rheumatologist-led disease activity monitoring may add an important perspective on disease activity, as it is perceived by the patient. Several validated PROs exist for inflammatory arthritis.

There is an unmet need for more frequent patient monitoring in chronic inflammatory arthritis to improve disease management and potentially to reduce pressure on specialists, as well as to achieve a better understanding of the disease course, which should be considered as more than just the linear path between two consecutive observation points. Evidence from several disease areas suggests that electronic tools that allow patients to give feed back on their disease may be beneficial. Innovative electronic tools that allow more frequent monitoring have the potential to improve disease management and may be more widely adopted in the future. Multiplatform availability of electronic monitoring devices is an important consideration in encouraging the widest usage possible. Innovative electronic tools, such as iMonitor, GoTreatIT, Sanoïa and Andar, may help to support physician time management, to reduce the burden on clinic time, and to prioritise patients who may need further attention.

### Author affiliations

<sup>1</sup>Radboud University Medical Centre, Radboud Institute for Health Sciences, IQ healthcare, Nijmegen, The Netherlands

<sup>2</sup>Department of Rheumatology, Bernhoven, Uden, The Netherlands

<sup>3</sup>Department Internal Medicine, Rheumatology, Schlosspark Klinik University Medicine Berlin, Berlin, Germany

<sup>4</sup>Département de Rhumatologie, Hôpital Lapeyronie, Montpellier Université, Montpellier, France

<sup>5</sup>Department of Internal Medicine, Kazan State Medical University, Kazan, Russia

<sup>6</sup>Pfizer Regional Medical Team, Rome, Italy

<sup>7</sup>School of Healthcare Sciences, College of Biomedical and Life Sciences, Cardiff University, Wales, UK

<sup>8</sup>Pfizer, Rome, Italy

<sup>9</sup>Department of Rheumatology, Parc Taulí Hospital Universitari, Sabadell, Spain

<sup>10</sup>Department of Rheumatology, Martina Hansens Hospital, Bærum, Norway
<sup>11</sup>Landesklinikum Stockerau, Center for Rheumatology Lower Austria, Stockerau, Austria

<sup>12</sup>Karl Landsteiner Institute for Clinical Rheumatology, Stockerau, Austria
<sup>13</sup>Department for Rheumatology and Immunology, Medical University of Graz, Graz, Austria

<sup>14</sup>Department of Medicine, South Karelia Central Hospital, Lappeenranta, Finland

<sup>15</sup>University of Genoa and Istituto Giannina Gaslini, Genoa, Italy

<sup>16</sup>Department of Internal Medicine, University of Milan, Milano, Italy

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