

Predictors of DAPSA28 remission in patients with psoriatic arthritis initiating a first TNF-inhibitor: results from 13 European registries

Louise Linde^{*1,2}, Lykke M. Ørnbjerg^{*1,2}, Stylianos Georgiadis^{*1,2}, Simon H. Rasmussen¹, Ulf Lindström³, Johan Askling⁴, Brigitte Michelsen^{1,5,6}, Daniela Di Giuseppe⁷, Johan K. Wallman⁸, Bjorn Gudbjornsson^{9,10}, Thorvardur Jon Love^{10,11}, Dan C. Nordström¹², Timo Yli-Kerttula¹³, Lucie Nekvindová^{14,15}, Jiří Vencovský^{14,15}, Florenzo Iannone¹⁶, Alberto Cauli¹⁷, Anne Gitte Loft^{18,19}, Bente Glinthborg^{1,2,34}, Karin Laas²⁰, Ziga Rotar^{21,22}, Matija Tomšič^{21,22}, Gary J. Macfarlane²³, Burkhard Möller²⁴, Marleen van de Sande^{25,26}, Catalin Codreanu²⁷, Michael J. Nissen²⁸, Merih Birlik²⁹, Sukran Erten³⁰, Maria J. Santos^{31,32}, Elsa Vieira-Sousa³³, Merete L. Hetland^{1,34**} and Mikkel Østergaard^{1,34**}

* LL, LMØ and SG contributed equally to this manuscript.

** MLH and MØ contributed equally to this manuscript.

1. Copenhagen Center for Arthritis Research (COPECARE), Center for Rheumatology and Spine Diseases, Rigshospitalet, Glostrup, Denmark
2. DANBIO registry, Rigshospitalet, Glostrup, Denmark
3. Department of Rheumatology and Inflammation Research, Sahlgrenska Academy at University of Gothenburg, Gothenburg, Sweden
4. Department of Medicine Solna, Karolinska Institutet, Stockholm, Sweden
5. Center for treatment of Rheumatic and Musculoskeletal Diseases (REMEDY), Diakonhjemmet Hospital, Oslo, Norway
6. Research Unit, Sørlandet Hospital, Kristiansand, Norway
7. Clinical Epidemiology Division, Department of Medicine Solna, Karolinska Institutet, Stockholm, Sweden
8. Department of Clinical Sciences Lund, Rheumatology, Skåne University Hospital, Lund University, Lund, Sweden
9. Centre for Rheumatology Research, Landspítali, University Hospital, Reykjavik, Iceland
10. Faculty of Medicine, University of Iceland, Reykjavik, Iceland
11. Department for Science and Research, Landspítali University Hospital, Reykjavik, Iceland
12. Departments of Medicine and Rheumatology, Helsinki University Hospital, Helsinki, Finland
13. Department of Rheumatology, Satakunta Central Hospital, Rauma, Finland
14. Institute of Rheumatology, Prague, Czech Republic
15. Department of Rheumatology, First Faculty of Medicine, Charles University, Prague, Czech Republic
16. Rheumatology Unit, DiMePre-Jo, University of Bari, Bari, Italy
17. Rheumatology Unit, Department of Medical Sciences and Public Health, AOU and University of Cagliari, Monserrato, Italy
18. Department of Rheumatology, Aarhus University Hospital, Aarhus, Denmark
19. Department of Clinical Medicine, Aarhus University, Aarhus, Denmark

20. Department of Rheumatology, East-Tallinn Central Hospital, Tallinn, Estonia
21. Department of Rheumatology, University Medical Centre Ljubljana, Ljubljana, Slovenia
22. Faculty of Medicine, University of Ljubljana, Ljubljana, Slovenia
23. Aberdeen Centre for Arthritis and Musculoskeletal Health (Epidemiology Group), University of Aberdeen, Aberdeen, UK
24. Inselspital, University Hospital Bern, Department for Rheumatology and Immunology, on behalf of the Swiss Clinical Quality Management for Rheumatic Diseases, SCQM
25. Amsterdam UMC, University of Amsterdam, Department of Rheumatology & Clinical Immunology and Department of Experimental Immunology, Amsterdam Institute for Infection & Immunity, Amsterdam, the Netherlands
26. Amsterdam Rheumatology and immunology Center (ARC), Amsterdam, the Netherlands
27. Center for Rheumatic Diseases, University of Medicine and Pharmacy, Romanian Registry of Rheumatic Diseases, Bucharest, Romania
28. Department of Rheumatology, Geneva University Hospital, Geneva, Switzerland
29. Division of Rheumatology, Department of Internal Medicine, Dokuz Eylul University School of Medicine, Izmir, Turkey
30. Department of Rheumatology, Ankara Yıldırım Beyazıt University Ankara City Hospital, Ankara, Turkey
31. Serviço de Reumatologia, Hospital Garcia de Orta, Almada, Portugal
32. Reuma.pt, Sociedade Portuguesa de Reumatologia, Lisbon, Portugal
33. Department of Rheumatology, Hospital de Santa Maria, CHULN, Instituto Medicina Molecular, Faculdade de Medicina da Universidade de Lisboa, Centro Académico de Medicina de Lisboa, Lisbon, Portugal
34. Department of Clinical Medicine, University of Copenhagen, Copenhagen, Denmark

Corresponding author: Louise Linde

COPECARE, Rigshospitalet, Valdemar Hansens Vej 17, 2600 Glostrup, Denmark.

Email: louise.linde@regionh.dk

ORCID iD: 0000-0003-0863-1352

Abstract

Objectives

In bio-naïve patients with Psoriatic arthritis (PsA) initiating a Tumour Necrosis Factor inhibitor (TNFi), we aimed to identify baseline predictors of Disease Activity index for PsA in 28 joints (DAPSA28) remission (primary objective) and DAPSA28 moderate response at 6 months, as well as drug retention at 12 months across 13 European registries.

Methods

Baseline demographic and clinical characteristics were retrieved and the three outcomes investigated per registry and in pooled data, using logistic regression analyses on multiply imputed data. In the pooled cohort, selected predictors that were either consistently positive or negative across all three outcomes, were defined as common predictors.

Results

In the pooled cohort (n=13,369), six-month proportions of remission, moderate response and 12-month drug retention were 25%, 34% and 63% in patients with available data (n=6,954, n=5,275 and n=13,369, respectively). Baseline predictors of remission, moderate response and 12-month drug retention were identified, five common across all three outcomes. Odds ratios (95% confidence interval) for DAPSA28 remission were: age, per year: 0.97 (0.96-0.98); disease duration, years (< 2 years as reference): 2-3 years: 1.20 (0.89-1.60), 4-9 years: 1.42 (1.09-1.84), ≥10 years: 1.66 (1.26-2.20); men vs. women: 1.85 (1.54-2.23); CRP >10 vs. ≤ 10 mg/l: 1.52 (1.22-1.89) and one mm increase in patient fatigue score: 0.99 (0.98-0.99).

Conclusion

Baseline predictors of remission, response and adherence to TNFi were identified, of which five were common for all three outcomes, indicating that the predictors emerging from our pooled cohort may be considered generalisable from the country- to disease-level.

Keywords: Psoriatic arthritis, first TNF-inhibitor, predictors, DAPSA28, drug retention, real-world evidence

Key messages

- This real-world study across 13 European countries presents data on 13,369 psoriatic arthritis (PsA) patients.
- Baseline predictors of remission, response and drug-retention following treatment with a first TNFi were identified.
- Consistency of predictors across registries and treatment outcomes, suggests generalisability from the country- to disease-level.

INTRODUCTION

Tumor necrosis factor inhibitors (TNFi) have contributed to major improvements in clinical outcomes and quality of life for patients with psoriatic arthritis (PsA). However, many patients treated with TNFi fail to achieve the recommended treatment target of remission or, alternatively, low disease activity(1,2).

As the palette of treatment options continues to increase, understanding baseline determinants of a good response to TNFi is important for clinicians and patients in their shared decision-making. Several possible baseline predictors of treatment response in PsA have been investigated in individual countries or regions, including demographic, clinical, patient-reported and life-style characteristics, but no consistent pattern of predictors has emerged from the studies(3,4,13–17,5–12). Cross-country differences in baseline characteristics in PsA patients initiating TNFi treatment have been reported in a previous study from the EuroSpA collaboration(2), and such differences may have contributed to the inconsistencies in observed predictors of a treatment response across studies from individual countries.

In addition to differences in patient characteristics, a wide range of outcome measures has been applied(3,4,13–17,5–12), possibly reflecting the different views on how best to capture the full spectrum of PsA with its various clinical manifestations(18). In 2017, an international task force proposed the Disease Activity index for PSoriatic Arthritis (DAPSA)(19) for disease activity assessment in PsA(20). The DAPSA includes a 66/68 swollen/tender joint count, which, however, is not always performed in routine clinical settings. Therefore, the modified DAPSA28, based on a 28 joint count has been developed and compared to the original DAPSA and found valid (21). The

1
2
3
4 authors suggested that DAPSA28 might be an alternative if the full DAPSA was missing in registry
5 studies (21). While treatment responses according to DAPSA28 have been reported previously for
6 14,261 European patients with PsA initiating a TNFi (2), predictors of such a response using
7 DAPSA28 as outcome have not been investigated in a real-world cohort.
8
9
10

11
12
13 Thus, in this study of PsA patients starting their first TNFi, the primary aim was to identify baseline
14 predictors of DAPSA28 remission after 6 months' treatment. Secondary aims were to identify
15 baseline predictors of achieving DAPSA28 moderate response after 6 months and baseline
16 predictors of 12-month drug retention.
17
18
19
20
21

22 **METHODS**

23 *Data sources*

24
25
26 This study included secondary use of data on patients registered with a PsA diagnosis from 13
27 European registries: ATTRA (Czech Republic), DANBIO (Denmark), ROB-FIN (Finland),
28 ICEBIO (Iceland), GISEA (Italy), NOR-DMARD (Norway), Reuma.pt (Portugal), RRBR
29 (Romania), biorx.si (Slovenia), BIOBADASER (Spain), SRQ (Sweden), SCQM (Switzerland) and
30 TURKBIO (Turkey). In all registries, data are collected prospectively as part of routine clinical
31 practice. Based on a predefined study protocol, anonymised data were uploaded by individual
32 registries onto a secure central server.
33
34
35
36
37
38

39 *Patients and visits*

40
41
42 Patients were included if they had a registered clinical diagnosis of PsA, were aged ≥ 18 years at
43 diagnosis, and had initiated a first TNFi treatment at some point between diagnosis and 90 years of
44 age, with a start date between January 1st 2009 and December 31st 2018. The baseline visit was
45 defined as a registered visit within 30 days before to 30 days after the registered date of TNFi
46 treatment start (i.e., baseline date), with priority given to visits before treatment start. The 6-month
47 visit was defined as the one closest in time to 180 days within a range of 90 to 270 days after the
48 baseline date. Baseline patient characteristics included demography, clinical measures, treatment
49 and patient-reported outcomes (**table 1**).
50
51
52
53
54
55
56
57
58
59
60

Endpoints

The primary endpoint was DAPSA28 remission (i.e. DAPSA28 \leq 4) at 6 months on the first TNFi (21). Secondary endpoints were 1) DAPSA28 moderate response at 6 months (here defined as a 75% improvement from the baseline DAPSA28, similar to the corresponding response definition for the original DAPSA score, as no validated definition for DAPSA28 moderate response is available (22)) and 2) 12-month drug retention.

Patients with no available 6-month DAPSA28 data were classified as having achieved DAPSA28 remission and DAPSA28 moderate response, respectively, if they fulfilled both of the following two criteria: 1) they had stopped the TNFi before 6 months *and* no subsequent biological (b) or targeted synthetic (ts) disease-modifying anti-rheumatic drug (DMARD) was started within 6 months from the previous treatment start, *and* 2) if the clinician had stated “remission” as the reason for discontinuation (**figures 1a and b**). Patients who stopped the TNFi during the first 6 months due to lack of effect, were considered as *not* having achieved DAPSA28 remission or DAPSA28 moderate response. Patients discontinuing treatment due to AE, other reasons, or no stated reason, were not included in the analyses.

The 12-month drug retention was defined as the proportion of patients with a treatment duration \geq 52 weeks. Treatment duration was defined as the number of weeks between the registered date of treatment start and the registered stop date. If the same drug was restarted within 3 months of a registered stop date, and no other treatment was recorded in between, the treatment periods were considered as one. Switch to a biosimilar of the same drug was disregarded. A treatment without a registered stop date was assumed to have been discontinued if a new b or ts DMARD treatment was recorded in the registry, and the stop date was then defined as the date of next treatment start. If no new treatment had been registered, a stop date was entered 12 months after the last registered visit. In the remaining observations, the stop date was defined as the date of data extraction, date of death, or end of registry follow-up, whichever came first.

Ethics

All participating registries obtained necessary approvals from relevant authorities prior to data transfer to the EuroSpA coordinating center. This study was designed, implemented and reported in accordance with the Guidelines for Good Pharmacoepidemiology Practices (GPP) of the International Society for Pharmacoepidemiology (ISPE 2008), the STROBE (Strengthening the

1
2
3
4 Reporting of Observational Studies in Epidemiology) guidelines(23) and the ethical principles laid
5 down in the Declaration of Helsinki.

6 7 8 *Statistics*

9
10 The statistical approach used for the current study has previously been applied in a cohort of
11 patients with axial spondyloarthritis and is summarized below (24).

12
13 Descriptive analyses of the baseline patient characteristics were performed per registry, in the
14 pooled cohort, and additionally for patients with and without available data on DAPSA28 remission
15 and moderate response at 6 months (in the pooled cohort only).

16
17 Logistic regression analyses were used to identify baseline variables associated with the primary
18 and secondary endpoints. Regression models were applied separately per registry and in the pooled
19 cohort. Events-per-variable (EPV) was used to evaluate the sample size within the logistic
20 regression models. Likelihood ratio tests were used to assess all models. Results of the multivariate
21 models are presented as odds ratio (OR) with 95% or 85% confidence intervals (CI), see below.

22 23 24 25 26 27 28 29 *Independent variables*

30
31 Sex, smoking status (current vs. previous/never), use of concomitant conventional synthetic (cs)
32 DMARDs, C-reactive protein (CRP) (≤ 10 vs. >10 mg/l) and year of TNFi start (2009-2014 vs.
33 2015-2018) were included as categorical variables. Age at treatment start, time since diagnosis,
34 Body Mass Index (BMI), 28 tender and swollen joint counts, physician global score, Health
35 Assessment Questionnaire (HAQ)(25), patient pain and fatigue scores were included as continuous
36 variables. Age at diagnosis, erythrocyte sedimentation rate (ESR) and patient global score were not
37 included in the models as they were considered to represent an overlap with time since diagnosis,
38 CRP and patient pain and fatigue scores, respectively. For further details on independent variables,
39 see **tables 2-5**.

40 41 42 43 44 45 46 47 48 *Missing data*

49
50 Patients with no registration of concomitant csDMARDs were considered not using such drugs. For
51 all remaining independent baseline variables, multiple imputation by chained equations (MICE) was
52 applied in a pooled dataset containing all registries (30 imputed datasets).

Variable selection

Variable selection in multiply imputed data for each endpoint followed. First, variable selection was performed separately in each of the 30 imputed datasets; the final model included the predictors that appeared in at least half of the models. Once the set of predictors was selected, the model was fitted to all imputed datasets and the model estimates were pooled according to Rubin's rules(24,26).

Analyses in individual registries

To compare the selected predictors across registries, prediction models were first applied in each registry. A significance level of 0.157 was chosen due to small EPV values in some registries, corresponding to a 85% CI(27). The individual registry regression analyses were evaluated for consistency of selected predictors by visual inspection to determine if pooling of the data was feasible.

Analyses in the pooled cohort

The pooled dataset was split into a derivation cohort and a validation cohort for each of the three endpoints, ensuring that 50% of patients from each registry went into each cohort, respectively. Registries with $EPV \geq 1$ in the derivation cohort were pooled. Age, sex and registry were a priori forced into the models, and continuous variables were categorized if the assumption of linearity was violated. A significance level of 0.05 and a corresponding 95% CI was applied. Selected predictors that were either consistently positive or negative across all three outcomes, were defined as common predictors. The performance of the final multivariable models was evaluated in the validation cohorts by calculating the Area Under the Receiver Operating Curve (AUROC) (28).

Additional analyses

In addition, we assessed whether differences in per registry proportions for DAPSA28 remission, moderate response and drug retention impacted the identified predictors, by stratifying the pooled cohort into three ordered levels based on visual inspection of the distribution of the outcomes in the registries. Prediction models were applied to each stratum, adjusting for registry using a variable selection process similar to the analyses in individual registries.

Finally, as DAPSA is the gold standard in the assessment of PsA patients, we conducted a prediction analysis in a subset with available remission and response criteria based on 66/68 joint counts, i.e. applying DAPSA remission (≤ 4) and DAPSA moderate response (75% improvement

1
2
3
4 from baseline) as outcomes and substituting the 28 joint counts with 66/68 joint counts as predictors
5
6 (22). R version 4.1.0 was used for statistical analyses.
7

8 **Results**

9 *Cohorts*

10
11 Across the 13 registries, 13,369 PsA patients had started a first TNFi treatment during the study
12
13 period. Baseline patient characteristics by registry and pooled are shown in **table 1**, with
14
15 corresponding information on data availability in **Supplementary Table S1, available at**
16
17 ***Rheumatology online***. Numerical baseline differences between patients *with* versus *without* 6-
18
19 month follow-up data were only seen for concomitant csDMARD (**Supplementary Table S2,**
20
21 **available at *Rheumatology online***).
22

23 *DAPSA28 remission and moderate response*

24
25 Of the 13 registries, 11 collected data on DAPSA28 (n=11,333) (**table 1**). A total of 6,442 (57%)
26
27 patients had a DAPSA28 assessment at 6-month follow-up visit after initiating their first TNFi, with
28
29 1,713 (27%) of these having achieved DAPSA28 remission. Of the 4,891 (43%) patients with no
30
31 DAPSA28 assessment at 6 months, 512 were instead classified according to their discontinuation
32
33 reason prior to 6 months follow-up (**figure 1a**). In total, 1,723 of 6,954 patients (25%) were
34
35 classified as having achieved DAPSA28 remission at 6 months. Proportions of DAPSA28 remission
36
37 ranged from 18% to 34% across registries (**table 2**). Corresponding results for DAPSA28 moderate
38
39 response are presented in **figure 1b** and **table 3**.
40

41 *Drug retention*

42
43 All patients initiating a first TNFi were included in the drug retention analyses. Thereof, 8,461
44
45 (63%) were still on treatment at 12 months, with proportions ranging from 54% to 76% across
46
47 registries (**table 4**).
48

49 *Prediction analyses in individual registries*

50
51 Eleven registries fulfilled the EPV criteria and were eligible for prediction analyses of the primary
52
53 endpoint DAPSA28 remission at 6 months. Male sex was identified as a predictor in 9 registries
54
55 (positive in 8 and negative in 1), while negative predictors included older age at treatment start (9
56
57 registries), higher tender joint count (7 registries), and higher BMI, patient pain and fatigue scores
58
59 in 5 registries. The remaining baseline variables were found predictive in less than half of the
60

1
2
3
4 eligible registries in which the variable was available, see **table 2** and **Supplementary Table S3**,
5 **available at *Rheumatology* online**, for presentation of odds ratios (OR).
6
7

8
9 Eleven and 13 registries, respectively, were eligible for analyses of the secondary endpoints 6-
10 month DAPSA28 moderate response and 12-month drug retention. Higher swollen joint count was
11 identified as a positive predictor of DAPSA28 moderate response in 8 registries and CRP >10 mg/l
12 in 6 registries. Negative predictors included older age at treatment start (6 registries) and current
13 smoking, higher BMI and higher patient fatigue score (5 registries). Male sex and longer disease
14 duration were positive predictors of 12-month drug retention in 10 and 8 registries, respectively,
15 while TNFi start year 2015-2018 was a negative predictor in 10 registries. Concomitant csDMARD
16 was a positive predictor in 6 and a negative predictor in 1 registry. The remaining baseline variables
17 were found predictive in less than half of the registries in which the variable was available, see
18 **table 3-4** and **Supplementary Tables S4-S5**, **available at *Rheumatology* online**, for presentation
19 of ORs.
20
21
22
23
24
25
26
27

28 *Prediction analyses in the pooled cohort*

29
30 The consistency of predictors in the regression analyses per registry was found to justify pooling the
31 data (**tables 2-4**). Common baseline predictors across all three outcomes (6-month DAPSA28
32 remission/6-month DAPSA28 moderate response/12-month drug retention) in the derivation cohort
33 were: male sex, longer disease duration, higher CRP (positive predictors); older age at treatment
34 start, higher fatigue score (negative predictors) (**table 5**).
35
36
37
38

39
40 A higher pain score was a negative predictor of DAPSA28 remission and 12-month drug retention
41 but a positive predictor of DAPSA28 moderate response (**table 5**).
42

43
44 The performance of the final models as assessed by the Area under the Receiver Operating Curve
45 (AUROC) in the validation cohort was estimated to 0.75 (DAPSA28 remission), 0.73 (DAPSA28
46 moderate response) and 0.64 (12-month drug retention), i.e. the models were able to correctly
47 predict remission in 75%, moderate response in 73% and 12-month drug retention in 64% of
48 patients (**table 5**).
49
50
51

52
53 In the pooled analyses *stratified* according to the proportion of patients achieving DAPSA28
54 remission, DAPSA28 moderate response and 12-month drug retention, the common predictors
55 identified in the pooled *unstratified* analyses (positive: male sex, longer disease duration, higher
56 CRP; negative: older age at treatment start and higher patient fatigue score) were identified in at
57
58
59
60

1
2
3
4 least 2 of 3 strata across the three outcomes (**Supplementary Table S6, available at**
5 ***Rheumatology online***).

6
7
8 In the additional analyses with DAPSA remission and moderate response as outcomes, fewer data
9 were available compared to the DAPSA28 analyses (**Supplementary Tables S1 and S7, available**
10 **at *Rheumatology online***). Baseline differences between patients *with* versus *without* 6-month
11 follow-up DAPSA were comparable with those seen in the DAPSA28 analyses, as were the
12 predictors in the regression analyses per registry (data not shown). In the prediction models on
13 pooled data, we identified the same predictors as for DAPSA28. In addition, 66 swollen joint count
14 was a common positive predictor, which is in contrast to the DAPSA28 analyses, where 28 swollen
15 joint count was not identified as a common predictor (**Supplementary Table S7**).

22 **Discussion**

23
24 In this study, we identified five common baseline predictors of TNFi treatment response and
25 retention, for the first time applying the DAPSA28 as endpoint in a large scale prediction analysis
26 across 13 European countries through the EuroSpA collaboration.

27
28 The main findings were that male sex, longer disease duration and higher CRP were positive
29 predictors of DAPSA28 remission and DAPSA28 moderate response at 6 months and of drug
30 retention after 12 months, while older age at treatment start and a higher patient fatigue score were
31 negative predictors.

32
33 In the EuroSpA collaboration, we have previously shown how baseline characteristics and treatment
34 outcomes differ across European countries, possibly illustrating different prescription practices and
35 access to therapy(2). To analyse if cross-country differences might contribute to inconsistencies in
36 baseline predictors of treatment response across registries, we also stratified the pooled cohort by
37 the proportion of patients achieving DAPSA28 remission, moderate response and 12-month drug
38 retention, respectively, and identified baseline predictors for each stratum. We found that although
39 the identified baseline predictors across strata and endpoints were not identical to the per-registry
40 and unstratified pooled analyses, no major differences emerged. This suggests that despite the
41 known and unknown differences across the individual countries, pooling of the cohorts to allow
42 large scale analyses seems an acceptable approach. Thereby, the baseline predictors emerging from
43 our pooled analyses may be considered generalizable from the country- to disease-level.

1
2
3
4 We found that starting TNFi from 2015-2018 versus 2009-2014 reduced the chance of 12-month
5 drug retention. This observed decrease in treatment retention over time may be explained by the
6 emerging options for switching to another TNFi or a drug with a different mode of action, should
7 the treatment target not be met. In support of this argument, a recent study on time trends in
8 treatment response in European patients with PsA has indicated considerably longer drug retention
9 rates prior to 2009(29).
10
11
12
13
14

15 A major strength of this study was the availability of similar clinical variables from 13 different
16 European registries, allowing for the inclusion of the largest number of patients with PsA to date in
17 a thorough analysis of baseline predictors of treatment response to TNFi. In previous similar
18 studies, various outcome measures and baseline characteristics have been investigated, however few
19 consistent predictors have emerged across the studies(3–7,13,14,16,17). Similarly, a meta-analysis
20 from 2015 including 4034 patients with PsA identified several possible but no consistent predictors,
21 which was ascribed to variation in the study design and heterogeneity in the treatment response
22 measures used in the included studies(15).
23
24
25
26
27
28
29

30 In agreement with our findings, male sex has been suggested as a predictor for a good treatment
31 response in other studies of patients with PsA(4,6,9,10,13,14). Similarly, our study adds weight to
32 findings from previous smaller studies that have reported younger age at treatment start to be
33 associated with better treatment responses(9,10,30). On the other hand, we found a positive
34 association between longer disease duration at TNFi treatment start and both drug retention and
35 treatment response. The patients with longer disease duration in our cohort had earlier onset PsA,
36 which might also have contributed to the better outcomes, as there is evidence pointing towards a
37 more aggressive disease course in PsA with onset later in life(31). Smaller studies have reported
38 contradictory results regarding disease duration(16,17,32).
39
40
41
42
43
44
45

46 Higher CRP at baseline was, in our study, predictive of a good treatment response. In contrast,
47 although CRP was included in many previous studies, it only predicted a good treatment response
48 in a minority(3,9,12,17). Across those studies, the baseline level of inflammation, as assessed by the
49 CRP was generally low, and the room for improvement therefore limited, which may potentially
50 explain why this signal was not previously detected. It could also be an indication that many aspects
51 besides inflammation play a role in this heterogeneous disease entity.
52
53
54
55
56

57 Baseline patient pain and fatigue scores were consistently associated with all treatment outcomes in
58 our pooled cohort, with fatigue as a consistently negative predictor and pain as a negative predictor
59
60

1
2
3
4 of remission and drug retention but a positive predictor of DAPSA28 moderate response. Previous
5 smaller studies have not found any clear pattern of associations between patient scores and
6 treatment outcomes, but some have reported that worse scores at baseline predicted poorer
7 outcomes(4,5,9,11–17). There is emerging evidence suggesting that the fatigue and pain
8 experienced by patients may not be fully explained by the rheumatic disease. For example in a
9 study of fatigue in PsA, inflammation, disease duration and chronic pain only explained two thirds
10 of the experienced fatigue(33), and moreover, pain experienced by patients may be modulated by
11 the concept of pain catastrophizing, a negative cognitive–affective response to anticipated or actual
12 pain(34,35). Our findings may reflect such underlying mechanisms. Nevertheless, our findings
13 suggest that the patient perspective is important for predicting the success of therapies, however,
14 further investigation into the concepts of patient assessments is warranted.
15
16
17
18
19
20
21
22

23
24 Functional disability measured by HAQ has previously been associated with poor outcomes in
25 rheumatoid arthritis(36,37), but our results only showed a negative association with
26 remission/response and not with drug retention. We find that the setting may not have been suitable
27 for detecting such associations. For example, our patients have a relatively short disease duration
28 and a high HAQ score may thus partly reflect reversible disease activity. In addition, drug retention
29 is not a strictly clinical outcome measure and may be impacted by various factors not related to the
30 disease status itself, i.e. treatment guidelines, access to drug, etc.
31
32
33
34
35

36
37 Previously, other data on the use of csDMARDs in combination with TNFi suggested no additional
38 effect of combination therapy on treatment response, but a possible beneficial effect on treatment
39 retention(7,12,15,38–40). We have previously reported improved clinical response rates when
40 combining adalimumab and infliximab but not etanercept with a csDMARD in PsA(41). In the
41 current study, we were unable to replicate these findings as we analyzed TNFis as one group,
42 however, our findings are in agreement with previous studies regarding drug retention.
43
44
45
46

47
48 Cardiovascular risk factors, such as smoking and obesity, are overrepresented in patients with PsA
49 compared to the general population(42,43), but the role of such factors during treatment with TNFi
50 is unclear. In a few previous studies, smoking and obesity were associated with a poorer treatment
51 response(4,7,13), while others found no such effect(10,14,16). In our pooled cohort, smoking was a
52 negative predictor of DAPSA28 remission and drug retention but not associated with DAPSA28
53 moderate response. Smoking was, however, negatively associated with DAPSA28 moderate
54 response in half of the registries. Variation in smoking habits across countries in addition to
55
56
57
58
59
60

1
2
3
4 heterogeneity in the data collection, may play a role in the differences observed between the per
5 registry and pooled analyses. BMI showed a similar tendency in our data, in line with our recent
6 findings from a study on predictors of treatment response in axSpA(24).
7
8

9
10 Limitations to our study include its observational nature, which does not allow any causal
11 conclusions to be drawn, and the lack of an endorsed PsA data collection framework limits
12 generalizability of findings to this patient group. In addition, issues with data availability prompted
13 us to use DAPSA28 over DAPSA although the latter is the gold standard in assessing PsA. We
14 were, however, reassured in finding largely similar predictors in the subset of patients with
15 available DAPSA scores. Selection bias based on availability of the DAPSA28 outcome cannot be
16 ruled out, however, baseline characteristics for patients with and without available DAPSA28
17 scores at follow-up were largely similar, and we therefore consider our findings to be generalizable.
18
19

20
21 In addition, we have previously discussed other limitations including the unbalanced sizes of the
22 registries and missing data, which also apply to this study(24); moreover, we were not able to
23 include psoriasis and other relevant comorbidities in the prediction models due to a lack of good
24 quality data. Finally, we primarily investigated predictors of short- and medium-term outcomes,
25 which is a limited window for a disease like PsA with fluctuating disease activity over time. An aim
26 for future studies could be to investigate the maintenance of treatment responses within a longer
27 time-frame, including available visits regardless of prespecified time-windows.
28
29

30
31 The performance of the final models was found acceptable for DAPSA28 remission and DAPSA28
32 moderate response but poor for 12-month drug retention. This suggests that additional factors such
33 as e.g. socio-economic parameters, comorbidities and biomarkers (imaging and serological) are still
34 needed for better prediction of treatment retention and response.
35
36

37
38 In conclusion, baseline predictors of remission, response and drug retention in European patients
39 with PsA treated with a first TNFi were identified, five of which were common across the
40 outcomes. The consistency of predictors across registries and treatment outcomes, despite
41 heterogeneity in patient characteristics and treatment practices, indicate that the baseline predictors
42 emerging from our pooled analyses may be considered generalisable from the country- to disease-
43 level.
44
45
46
47
48
49
50
51
52
53
54
55
56
57
58
59
60

Acknowledgements

The EuroSpA Research Collaboration Network was financially supported by Novartis Pharma AG. Novartis had no influence on the data collection, statistical analyses, manuscript preparation or decision to submit the manuscript.

Funding: This work was supported by Novartis Pharma AG.

Conflicts of interest: Louise Linde, Lykke M. Ørnbjerg, Stylianos Georgiadis and Simon H. Rasmussen: research grants from Novartis; Johan Askling: PI for agreements between Karolinska Institutet and Abbvie, Astra-Zeneca, BMS, Eli Lilly, Galapagos, Janssen, MSD, Pfizer, Roche, Samsung Bioepis, Sanofi, and UCB; Brigitte Michelsen: research grant from Novartis; Johan K. Wallman: speaking fee from AbbVie, Amgen. Research support from AbbVie, Amgen, Eli Lilly, Novartis, Pfizer; Bjorn Gudbjornsson: consulting and/or Speaking fees from Amgen and Novartis; Dan C. Nordström: consulting and/or speaking fees from Abbvie, BMS, Lilly, MSD, Novartis, Pfizer, Roche and UCB; Jiří Vencovský: consulting and/or speaking fees from Abbvie, Biogen, Boehringer Ingelheim, Eli Lilly, Gilead, Merck Sharp and Dohme, Pfizer and UCB; Florenzo Iannone: consulting and/or speaking from Abbvie, Amgen, AstraZeneca, BMS, Galapagos, Janssen, Lilly, MSD, Novartis, Pfizer, Roche, UCB; Alberto Cauli: consulting and/or speaking fees from Abbvie, Amgen, Biogen, BMS, Galapagos, Eli Lilly, Janssen, Novartis, Pfizer, UCB; Anne Gitte Loft: Research Grant from Novartis, and speaking and/or consulting fees from AbbVie, Janssen, Lilly, MSD, Novartis, Pfizer, Roche, and UCB; Bente Glinthorg: Research grants: Pfizer, Abbvie, BMS; Karin Laas: consulting and/or speaking fees from Amgen, Johnson and Johnson and Novartis; Ziga Rotar: speaking or consultancy fees from Abbvie, Novartis, MSD, Medis, Biogen, Eli Lilly, Pfizer, Sanofi, Lek, Janssen; Matija Tomšič: consulting and/or speaking fees from Abbvie, Amgen, Biogen, Eli Lilly, Janssen, Medis, MSD, Novartis, Pfizer, Sanofi, Sandoz-Lek; Gary J. Macfarlane: research grant from GSK; Burkhard Möller: speaking fee from: Eli-Lilly, Janssen, Novartis, Pfizer. Grants from Amgen; Marleen van de Sande: research grant and/or consulting fee, and/or speaker fee from Eli Lilly, Novartis, UCB, Janssen, Abbvie; Catalin Codreanu: speaking and consultancy fees from AbbVie, Amgen, Boehringer Ingelheim, Ewopharma, Lilly, Novartis, Pfizer; Michael J. Nissen: consulting and/or speaking fees from AbbVie, Eli Lilly, Janssens, Novartis and Pfizer; Sukran Erten: speaking fees from Celltrion,

1
2
3
4 Pfizer, MSD, **Maria J. Santos**: speaker fees from Abbvie, AstraZeneca, Lilly, Novartis and Pfizer;
5
6 **Elsa Vieira-Sousa**: research grants from MSD, Pfizer, UCB. Speaker fees from Novartis, Abbvie,
7
8 MSD, Celgene, UCB; **Merete L. Hetland**: research grants from Abbvie, Biogen, BMS, Celltrion,
9
10 Eli Lilly, Janssen Biologics B.V, Lundbeck Fonden, MSD, Medac, Pfizer, Roche, Samsung
11
12 Biopies, Sandoz, Novartis and **Mikkel Østergaard**: research grants from Abbvie, BMS, Merck,
13
14 Celgene and Novartis, and speaker and/or consultancy fees from Abbvie, BMS, Boehringer-
15
16 Ingelheim, Celgene, Eli-Lilly, Hospira, Janssen, Merck, Novartis, Novo, Orion, Pfizer, Regeneron,
17
18 Roche, Sandoz, Sanofi and UCB. The remaining authors have declared no conflicts of interest.
19
20

21 **Data availability:** The data in this article was collected in the individual registries and made
22
23 available for secondary use through the EuroSpA Research Collaboration Network
24
25 [<https://eurospa.eu/#registries>]. Relevant patient level data may be made available on reasonable
26
27 request to the corresponding author, but will require approval from all contributing registries.
28
29
30
31
32
33
34
35
36
37
38
39
40
41
42
43
44
45
46
47
48
49
50
51
52
53
54
55
56
57
58
59
60

References

1. Gossec L, Baraliakos X, Kerschbaumer A, De Wit M, McInnes I, Dougados M, et al. EULAR recommendations for the management of psoriatic arthritis with pharmacological therapies: 2019 update. *Ann Rheum Dis*. 2020;79(6):700-712.
2. Brahe CH, Ørnbjerg LM, Jacobsson L, Nissen MJ, Kristianslund EK, Mann H, et al. Retention and response rates in 14 261 PsA patients starting TNF inhibitor treatment - Results from 12 countries in EuroSpA. *Rheumatol (United Kingdom)*. 2020;59(7):1640-1650.
3. Scrivo R, Giardino AM, Salvarani C, Foti R, Afeltra A, Viapiana O, et al. An observational prospective study on predictors of clinical response at six months in patients with active psoriatic arthritis treated with golimumab. *Clin Exp Rheumatol*. 2020;38(1):107-114.
4. Ogdie A, Palmer JL, Greenberg J, Curtis JR, Harrold LR, Solomon DH, et al. Predictors of achieving remission among patients with psoriatic arthritis initiating a tumor necrosis factor inhibitor. *J Rheumatol*. 2019;46(5):475-482.
5. Flouri ID, Markatseli TE, Boki KA, Papadopoulos I, Skopouli FN, Voulgari P V., et al. Comparative analysis and predictors of 10-year tumor necrosis factor inhibitors drug survival in patients with spondyloarthritis: First-year response predicts longterm drug persistence. *J Rheumatol*. 2018;45(6):785-794.
6. Iannone F, Lopriore S, Bucci R, Scioscia C, Anelli MG, Notarnicola A, et al. Two-year survival rates of anti-TNF- α therapy in psoriatic arthritis (PsA) patients with either polyarticular or oligoarticular PsA. *Scand J Rheumatol*. 2015;44(3):192-9.
7. Fagerli KM, Lie E, Van Der Heijde D, Heiberg MS, Lexberg ÅS, Rødevand E, et al. The role of methotrexate co-medication in TNF-inhibitor treatment in patients with psoriatic arthritis: Results from 440 patients included in the NOR-DMARD study. *Ann Rheum Dis*. 2014;73(1):132-7.
8. Chimenti MS, Perricone C, Graceffa D, Di Muzio G, Ballanti E, Guarino MD, et al. Complement system in psoriatic arthritis: A useful marker in response prediction and monitoring of anti-TNF treatment. *Clin Exp Rheumatol*. 2012;30(1):23-30.
9. Glinborg B, Åstergaard M, Dreyer L, Krogh NS, Tarp U, Hansen MS, et al. Treatment response, drug survival, and predictors thereof in 764 patients with psoriatic arthritis treated

- with anti-tumor necrosis factor α therapy: Results from the nationwide Danish DANBIO registry. *Arthritis and Rheumatism*. 2011;63(2):382-90.
10. Saad AA, Ashcroft DM, Watson KD, Symmons DPM, Noyce PR, Hyrich KL, et al. Efficacy and safety of anti-TNF therapies in psoriatic arthritis: an observational study from the British Society for Rheumatology Biologics Register. *Rheumatology (Oxford)*. 2010;49(4):697-705.
 11. Saber TP, Ng CT, Renard G, Lynch BM, Pontifex E, Walsh CAE, et al. Remission in psoriatic arthritis: Is it possible and how can it be predicted? *Arthritis Res Ther*. 2010;12(3):R94.
 12. Kristensen LE, Gülfe A, Saxne T, Geborek P. Efficacy and tolerability of anti-tumour necrosis factor therapy in psoriatic arthritis patients: Results from the South Swedish Arthritis Treatment Group register. *Ann Rheum Dis*. 2008;67(3):364-9.
 13. Vieira-Sousa E, Eusébio M, Ávila-Ribeiro P, Khmelinskii N, Cruz-Machado R, Rocha TM, et al. Real-world longterm effectiveness of tumor necrosis factor inhibitors in psoriatic arthritis patients from the rheumatic diseases portuguese register. *J Rheumatol*. 2020;47(5):690-700.
 14. Michelsen B, Sexton J, Wierød A, Bakland G, Rødevand E, Krøll F, et al. Four-year follow-up of inflammatory arthropathy patients treated with golimumab: Data from the observational multicentre NOR-DMARD study. *Semin Arthritis Rheum*. 2020;50(1):12-16.
 15. Maneiro JR, Souto A, Salgado E, Mera A, Gomez-Reino JJ. Predictors of response to TNF antagonists in patients with ankylosing spondylitis and psoriatic arthritis: Systematic review and meta-analysis. *RMD Open*. 2015 published online February 18 2015. doi: [10.1136/rmdopen-2014-000017](https://doi.org/10.1136/rmdopen-2014-000017).
 16. Iannone F, Santo L, Anelli MG, Bucci R, Semeraro A, Quarta L, et al. Golimumab in real-life settings: 2 Years drug survival and predictors of clinical outcomes in rheumatoid arthritis, spondyloarthritis, and psoriatic arthritis. *Semin Arthritis Rheum*. 2017;47(1):108-114.
 17. Eder L, Chandran V, Schentag CT, Shen H, Cook RJ, Gladman DD. Time and predictors of response to tumour necrosis factor-(alpha) blockers in psoriatic arthritis: An analysis of a longitudinal observational cohort. *Rheumatology*. 2010;49(7):1361-6.
 18. Mease PJ, Coates LC. Considerations for the definition of remission criteria in psoriatic

- 1
2
3
4 arthritis. *Seminars in Arthritis and Rheumatism*. 2018;47(6):786-796.
5
6
7 19. Schoels M, Aletaha D, Funovits J, Kavanaugh A, Baker D, Smolen JS. Application of the
8 DAREA/DAPSA score for assessment of disease activity in psoriatic arthritis. *Ann Rheum*
9 *Dis*. 2010;69(8):1441-7.
10
11
12 20. Smolen JS, Braun J, Dougados M, Emery P, FitzGerald O, Helliwell P, et al. Treating
13 spondyloarthritis, including ankylosing spondylitis and psoriatic arthritis, to target:
14 Recommendations of an international task force. *Ann Rheum Dis*. 2014;73(1):6-16.
15
16
17 21. Michelsen B, Sexton J, Smolen JS, Aletaha D, Krogh NS, Van Der Heijde D, et al. Can
18 disease activity in patients with psoriatic arthritis be adequately assessed by a modified
19 Disease Activity index for PSoriatic Arthritis (DAPSA) based on 28 joints? *Ann Rheum Dis*.
20 2018;77(12):1736-1741.
21
22
23 22. Schoels MM, Aletaha D, Alasti F, Smolen JS. Disease activity in psoriatic arthritis (PsA):
24 Defining remission and treatment success using the DAPSA score. *Ann Rheum Dis*.
25 2016;75(5):811-8.
26
27
28 23. Vandembroucke JP, Von Elm E, Altman DG, Gøtzsche PC, Mulrow CD, Pocock SJ, et al.
29 Strengthening the Reporting of Observational Studies in Epidemiology (STROBE):
30 Explanation and elaboration. *PLoS Med*. 2007;4(10):1628–54.
31
32
33 24. Ørnbjerg LM, Linde L, Georgiadis S, Rasmussen SH, Lindström U, Askling J, et al.
34 Predictors of ASDAS-CRP inactive disease in axial spondyloarthritis during treatment with
35 TNF-inhibitors: Data from the EuroSpA collaboration. *Semin Arthritis Rheum*. 2022;56. doi:
36 10.1016/j.semarthrit.2022.152081.
37
38
39 25. Fries JF, Spitz PW, Young DY. The dimensions of health outcomes: the health assessment
40 questionnaire, disability and pain scales. *J Rheumatol*. 1982;9(5):789–93.
41
42
43 26. Wood AM, White IR, Royston P. How should variable selection be performed with multiply
44 imputed data? *Stat Med*. 2008;27(17):3227-46.
45
46
47 27. Heinze G, Wallisch C, Dunkler D. Variable selection – A review and recommendations for
48 the practicing statistician. *Biometrical Journal*. 2018;60(3):431-49.
49
50
51 28. Hanley JA, McNeil BJ. The meaning and use of the area under a receiver operating
52 characteristic (ROC) curve. *Radiology*. 1982;143(1):29-36.
53
54
55
56
57
58
59
60

- 1
2
3
4 29. Christiansen SN, Ørnbjerg LM, Rasmussen SH, Loft AG, Askling J, Iannone F, et al.
5 European bio-naïve spondyloarthritis patients initiating TNFi: Time trends in baseline
6 characteristics, treatment retention and response. *Rheumatology (Oxford)* 2022;61(9):3799-
7 3807.
8
9
10
11 30. Iervolono S, Di Minno MND, Peluso R, Lofrano M, Russolillo A, Di Minno G, et al.
12 Predictors of early minimal disease activity in patients with psoriatic arthritis treated with
13 tumor necrosis factor- α blockers. *J Rheumatol.* 2012;39(3):568-73.
14
15
16
17 31. Fragoulis GE, Nikiphorou E, McInnes IB, Siebert S. Does Age Matter in Psoriatic Arthritis?
18 A Narrative Review. *J Rheumatol.* 2021;49(10):1085-1091.
19
20
21 32. Gratacós J, Casado E, Real J, Torre-Alonso JC. Prediction of major clinical response
22 (ACR50) to infliximab in psoriatic arthritis refractory to methotrexate. *Ann Rheum Dis.*
23 2007;66(4):493-7.
24
25
26
27 33. Skougaard M, Jørgensen TS, Rifbjerg-Madsen S, Coates LC, Egeberg A, Amris K, et al.
28 Relationship Between Fatigue and Inflammation, Disease Duration, and Chronic Pain in
29 Psoriatic Arthritis: An Observational DANBIO Registry Study. *J Rheumatol*
30 2020;47(4):548–52.
31
32
33
34 34. Edwards RR, Bingham CO, Bathon J, Haythornthwaite JA. Catastrophizing and pain in
35 arthritis, fibromyalgia, and other rheumatic diseases. *Arthritis Care Res (Hoboken)*
36 2006;55(2):325–32.
37
38
39
40 35. Hammer HB, Uhlig T, Kvien TK, Lampa J. Pain Catastrophizing, Subjective Outcomes, and
41 Inflammatory Assessments Including Ultrasound: Results From a Longitudinal Study of
42 Rheumatoid Arthritis Patients. *Arthritis Care Res* 2018;70(5):703–12.
43
44
45
46 36. Sokka T, Häkkinen A, Krishnan E, Hannonen P. Similar prediction of mortality by the health
47 assessment questionnaire in patients with rheumatoid arthritis and the general population.
48 *Ann Rheum Dis* 2004;63(5):494–7.
49
50
51 37. Young A, Dixey J, Kulinskaya E, Cox N, Davies P, Devlin J, et al. Which patients stop
52 working because of rheumatoid arthritis? Results of five years' follow up in 732 patients
53 from the early RA study (ERAS). *Ann Rheum Dis.* 2002;61(4):335-40.
54
55
56
57 38. Behrens F, Cañete JD, Olivieri I, Van Kuijk AW, McHugh N, Combe B. Tumour necrosis
58 factor inhibitor monotherapy vs combination with MTX in the treatment of PsA: A
59
60

- 1
2
3
4 systematic review of the literature. *Rheumatol (United Kingdom)*. 2014;54(5):915-26.
5
6
7 39. Mease PJ, Gladman DD, Collier DH, Ritchlin CT, Helliwell PS, Liu L, et al. Etanercept and
8 Methotrexate as Monotherapy or in Combination for Psoriatic Arthritis: Primary Results
9 From a Randomized, Controlled Phase III Trial. *Arthritis Rheumatol*. 2019;71(7):1112-1124.
10
11
12 40. George MD, Baker JF, Ogdie A. Comparative persistence of methotrexate and tumor
13 necrosis factor inhibitors in rheumatoid arthritis, psoriatic arthritis, and ankylosing
14 spondylitis. *J Rheumatol*. 2020;47(6):826-834.
15
16
17
18 41. Lindström U, Di Giuseppe D, Delcoigne B, Glintborg B, Möller B, Ciurea A, et al.
19 Effectiveness and treatment retention of TNF inhibitors when used as monotherapy versus
20 comedication with csDMARDs in 15 332 patients with psoriatic arthritis. Data from the
21 EuroSpA collaboration. *Ann Rheum Dis*. 2021;80(11):1410-1418.
22
23
24
25 42. Landgren AJ, Bilberg A, Eliasson B, Larsson I, Dehlin M, Jacobsson LTH, et al.
26 Cardiovascular risk factors are highly overrepresented in Swedish patients with psoriatic
27 arthritis compared with the general population. *Scand J Rheumatol*. 2020;49(3):195-199.
28
29
30
31 43. Kaine J, Song X, Kim G, Hur P, Palmer JB. Higher incidence rates of comorbidities in
32 patients with psoriatic arthritis compared with the general population using U.S.
33 Administrative claims data. *J Manag Care Spec Pharm*. 2019;25(1):122-132.
34
35
36
37
38
39
40
41
42
43
44
45
46
47
48
49
50
51
52
53
54
55
56
57
58
59
60

1
2
3
4 **Figure 1. Classification of patients starting their first TNF-inhibitor with regards to**
5 **DAPSA28 remission (A) and DAPSA28 moderate response (B) at six months.**
6
7

8 *Excluding Italy and Spain due to no available CRP; **according to the opinion of the clinician;
9 ***remission: n=1,723 (panel A)/response: n=1,803 (panel B); ****no remission: n=5,231 (panel
10 A)/no response: n=3,472 (panel B); *****including patients stopping TNFi *after* 6 months for all
11 reasons, patients stopping TNFi *within* 6 months for other reasons and patients continuing on TNFi
12 but without an assessment.
13
14
15
16

17
18 TNFi: Tumor Necrosis Factor alpha inhibitor; DAPSA28: Disease Activity index for PSoriatic
19 Arthritis in 28 joints.
20
21
22
23
24
25
26
27
28
29
30
31
32
33
34
35
36
37
38
39
40
41
42
43
44
45
46
47
48
49
50
51
52
53
54
55
56
57
58
59
60

Table 1. Baseline characteristics of PsA patients starting a first TNFi, pooled and stratified by registry.

Country	All	Czech Republic	Denmark	Finland	Iceland	Italy	Norway	Portugal	Romania	Slovenia	Spain	Sweden	Switzerland	Turkey
Registry	Pooled	ATTRA	DANBIO	ROB-FIN	ICEBIO	GISEA	NOR-DMARD	Reuma.pt	RRBR	Biorx.si	BIOBADASER	SRQ	SCQM	TURKBIO
Number of patients, n	13369	718	2090	234	306	1591	717	675	86	367	445	5225	628	287
Demography and diagnosis														
Age at treatment start, years	49 (40-58)	49 (40-57)	48 (39-56)	48 (40-56)	50 (39-59)	51 (42-59)	47 (39-57)	49 (40-57)	52 (47-61)	51 (43-57)	50 (40-57)	50 (40-59)	50 (40-58)	41 (34-51)
Age at diagnosis, years	43 (34-52)	40 (31-49)	43 (34-52)	40 (30-48)	43 (32-53)	45 (36-54)	41 (32-51)	42 (33-51)	47 (39-55)	43 (35-51)	45 (36-53)	43 (34-53)	44 (35-54)	36 (29-45)
Time since diagnosis, years	3 (1-8)	6 (2-12)	3 (1-7)	5 (2-11)	4 (1-9)	3 (1-7)	3 (1-9)	4 (2-8)	4 (2-6)	5 (2-10)	3 (1-7)	3 (1-8)	2 (1-6)	3 (1-7)
Men, n (%)	6385 (48%)	386 (54%)	928 (44%)	118 (50%)	126 (41%)	733 (46%)	345 (48%)	338 (50%)	37 (43%)	194 (53%)	227 (51%)	2552 (49%)	293 (47%)	108 (38%)
BMI, kg/m ²	27.0 (24.1-30.5)	28.1 (24.9-32.0)	27.2 (23.9-30.5)	27.8 (25.2-31.4)	30.1 (26.8-34.4)	26.2 (23.5-29.4)	NA	26.5 (24.0-29.4)	28.5 (25.5-31.8)	26.6 (23.8-29.7)	27.1 (24.2-30.7)	NA	26.5 (23.5-29.8)	28.1 (25.3-31.2)
Current smokers, n (%)	1865 (17%)	89 (16%)	582 (29%)	14 (12%)	26 (15%)	67 (8%)	131 (22%)	74 (16%)	4 (5%)	54 (15%)	98 (23%)	528 (12%)	127 (24%)	71 (26%)
Fulfilling the CASPAR criteria, n (%)	2497 (93%)	675 (95%)	284 (96%)	NA	47 (94%)	71 (96%)	NA	455 (89%)	79 (92%)	364 (99%)	NA	NA	502 (87%)	20 (87%)
Clinical measures														
Swollen joint count (28)	2 (0-5)	7 (3-10)	1 (0-3)	2 (1-5)	4 (2-6)	1 (0-3)	1 (0-3)	3 (1-6)	-	6 (3-9)	2 (1-4)	2 (0-5)	2 (0-4)	2 (0-4)
Swollen joint count (66)	3 (1-7)	9 (5-12)	3 (0-6)	3 (1-6)	-	1 (0-4)	NA	4 (1-8)	-	NA	NA	3 (1-6)	3 (1-6)	-
Tender joint count (28)	4 (1-9)	10 (5-13)	4 (1-8)	3 (1-6)	4 (2-6)	3 (1-8)	2 (1-6)	4 (2-9)	-	8 (4-12)	3 (1-6)	4 (2-8)	3 (1-7)	4 (1-8)
Tender joint count (68)	7 (3-12)	12 (8-19)	8 (4-14)	4 (2-9)	-	4 (2-10)	NA	7 (3-13)	-	NA	NA	6 (3-11)	6 (2-11)	-
CRP, mg/l	6 (3-14)	15 (6-28)	5 (2-12)	6 (3-13)	8 (3-15)	NA	5 (2-11)	8 (4-19)	-	7 (3-16)	NA	5 (2-12)	5 (2-10)	9 (3-17)

ESR, mm/hr	15 (7-29)	30 (17-45)	NA	14 (5-24)	NA	15 (8-30)	12 (6-22)	24 (11-42)	-	24 (12-40)	17 (7-34)	12 (6-24)	11 (6-20)	NA
Physician global score (mm)	40 (25-60)	65 (50-80)	25 (15-40)	38 (26-51)	56 (41-70)	50 (30-70)	30 (21-40)	50 (36-65)	-	60 (40-70)	NA	40 (30-50)	40 (30-60)	31 (20-62)
DAPSA28, units	25 (17-37)	41 (30-52)	23 (16-34)	21 (16-33)	28 (21-34)	NA	17 (12-26)	28 (19-40)	-	38 (26-51)	NA	24 (17-35)	19 (13-29)	26 (17-34)
DAPSA (original), units	25 (18-35)	36 (27-43)	26 (19-36)	21 (15-29)	-	NA	NA	26 (19-37)	-	NA	NA	23 (17-31)	21 (15-32)	-
DAS28-CRP, units	4.2 (3.3-5.0)	5.2 (4.6-5.8)	4.0 (3.1-4.8)	3.9 (3.2-4.7)	4.3 (3.9-4.9)	NA	3.5 (2.7-4.3)	4.3 (3.6-5.2)	-	5.0 (4.1-5.6)	NA	4.1 (3.3-4.8)	3.6 (2.7-4.5)	4.2 (3.2-4.9)
Treatment														
n (%)														
Infliximab	2251 (17%)	99 (14%)	576 (28%)	56 (24%)	188 (61%)	114 (7%)	91 (13%)	52 (8%)	8 (9%)	26 (7%)	39 (9%)	907 (17%)	64 (10%)	31 (11%)
Etanercept	4654 (35%)	126 (18%)	495 (24%)	60 (26%)	67 (22%)	657 (41%)	211 (29%)	270 (40%)	18 (21%)	63 (17%)	170 (38%)	2290 (44%)	147 (23%)	80 (28%)
Adalimumab	3987 (30%)	352 (49%)	626 (30%)	87 (37%)	9 (3%)	614 (39%)	87 (12%)	198 (29%)	41 (48%)	172 (47%)	132 (30%)	1312 (25%)	243 (39%)	114 (40%)
Certolizumab pegol	847 (6%)	47 (7%)	208 (10%)	6 (3%)	0 (0%)	28 (2%)	190 (26%)	12 (2%)	0 (0%)	31 (8%)	38 (9%)	248 (5%)	16 (3%)	23 (8%)
Golimumab	1630 (12%)	94 (13%)	185 (9%)	25 (11%)	42 (14%)	178 (11%)	138 (19%)	143 (21%)	19 (22%)	75 (20%)	66 (15%)	468 (9%)	158 (25%)	39 (14%)
TNFi start year*, n (%)														
2009-2014	7541 (56%)	344 (48%)	1231 (59%)	179 (76%)	144 (47%)	1254 (79%)	469 (65%)	336 (50%)	0 (0%)	219 (60%)	95 (21%)	2708 (52%)	452 (72%)	110 (38%)
2015-2018	5828 (44%)	374 (52%)	859 (41%)	55 (24%)	162 (53%)	337 (21%)	248 (35%)	339 (50%)	86 (100%)	148 (40%)	350 (79%)	2517 (48%)	176 (28%)	177 (62%)
Concomitant csDMARD (%)**	7832 (59%)	588 (82%)	1311 (63%)	190 (81%)	129 (42%)	916 (58%)	529 (74%)	463 (69%)	85 (99%)	285 (78%)	323 (73%)	2539 (49%)	361 (57%)	113 (39%)
Patient reported outcomes (PROs)														

Patient pain score (mm)	61 (42-77)	70 (50-80)	63 (43-78)	54 (36-72)	67 (50-78)	60 (50-80)	48 (29-65)	60 (48-80)	-	70 (56-80)	NA	61 (43-75)	60 (40-70)	75 (55-80)
Patient fatigue score (mm)	65 (41-80)	65 (50-80)	70 (50-84)	NA	70 (50-80)	NA	45 (15-70)	NA	NA	NA	NA	64 (41-78)	-	70 (50-75)
Patient global score (mm)	64 (45-80)	70 (58-80)	72 (52-87)	51 (31-70)	74 (54-85)	60 (50-80)	51 (31-70)	64 (48-80)	-	70 (60-80)	60 (50-80)	60 (42-75)	60 (40-80)	70 (54-75)
HAQ (units)	0.9 (0.5-1.4)	1.2 (0.9-1.6)	1.0 (0.6-1.5)	0.9 (0.5-1.4)	1.2 (0.8-1.5)	1.0 (0.4-1.5)	0.5 (0.2-0.9)	1.1 (0.5-1.5)	-	1.1 (0.5-1.6)	NA	0.9 (0.5-1.2)	0.8 (0.4-1.1)	0.8 (0.6-0.9)
Comorbidities and conditions associated with PsA														
Psoriasis	1904 (83%)	NA	378 (100%)	203 (87%)	NA	NA	NA	311 (61%)	41 (48%)	328 (89%)	-	NA	529 (89%)	90 (100%)
Uveitis	63 (3%)	NA	NA	10 (4%)	NA	NA	NA	1 (0%)	0 (0%)	6 (2%)	14 (3%)	NA	32 (5%)	NA
Inflammatory bowel disease	148 (8%)	NA	-	7 (3%)	NA	92 (100%)	NA	0 (0%)	0 (0%)	2 (1%)	NA	NA	22 (4%)	-
Cardiovascular disease	898 (26%)	262 (36%)	-	67 (29%)	NA	123 (100%)	108 (23%)	9 (2%)	43 (50%)	116 (32%)	18 (5%)	NA	108 (24%)	-
Diabetes	396 (12%)	57 (8%)	-	16 (7%)	NA	119 (100%)	27 (6%)	27 (5%)	14 (16%)	27 (7%)	34 (9%)	NA	27 (6%)	-
Kidney	92 (3%)	7 (1%)	NA	0 (0%)	NA	-	7 (1%)	8 (2%)	7 (8%)	4 (1%)	1 (0%)	NA	9 (3%)	NA
Data are as observed, median (interquartile range) or percentage. Percentages are calculated based on the number of patients with available data, unless stated otherwise. Cells are marked with "-" if based on <50 patients.														
NA: Not available; BMI: Body Mass Index; CASPAR: CIASsification criteria for Psoriatic ARthritis; CRP: C-reactive protein; ESR: erythrocyte sedimentation rate; DAPSA28: Disease Activity index for Psoriatic Arthritis in 28 joints; DAPSA (original): based on 66/68 joints; DAS28-CRP: disease activity score in 28 joints based on CRP; HLA-B27: Human Leukocyte Antigen subtypes B*2701-2759; TNFi: Tumor Necrosis Factor Inhibitor; csDMARD: conventional synthetic Disease Modifying Anti-Rheumatic Drug; HAQ: Health Assessment Questionnaire.														
*2009 was chosen as the first three biological Disease Modifying Anti-Rheumatic Drugs (bDMARDs) (adalimumab, etanercept and infliximab) from that year were all well-established treatment options across the European countries. 2015 was chosen as secukinumab was approved as the first non-TNFi bDMARD treatment option that year; **patients with no registration of concomitant use of csDMARDs were considered not using such drugs, all data are thus considered available.														

Table 2. Summary of predictors of DAPSA28 remission after 6 months of treatment with the first TNFi per registry* for registries with EPV per available independent variables ≥ 1 .

Country	Czech Republic	Denmark	Finland	Iceland	Norway	Portugal	Romania	Slovenia	Sweden	Switzerland	Turkey	Row sum**
Registry	ATTRA	DANBIO	ROB-FIN	ICEBIO	NOR-DMARD	Reuma.pt	RRBR	Biorx.si	SRQ	SCQM	TURKBIO	
Patients with DAPSA28 remission assessment, n	480	1496	113	177	546	383	82	287	3074	157	159	
DAPSA28 remission, n (%)	127 (27)	344 (23)	35 (31)	45 (25)	163 (30)	102 (27)	17 (21)	51 (18)	748 (24)	37 (24)	54 (34)	
EPV per available IVs	9.1	24.6	2.7	3.2	12.5	7.8	1.7	3.9	57.5	2.6	3.9	
Age at treatment start, years	-	-	-		-	-		-	-	-	-	9
Men	+	+	+	+	+	+	-		+		+	9
Time since diagnosis, years	+	+			+		+		+			5
BMI, kg/m ²	-	-			NA	-	-		NA	-		5
Current smokers		-								-		2
Concomitant csDMARD							constant					0
1 st TNFi start, year (2015-2018)***		+					constant			+		2
CRP>10 mg/l****	+	+					constant		+			3
Patient pain score, mm		-	-	-	-	-					+	6
Patient fatigue score, mm		-	NA		-	NA	NA	NA	-	-	-	5
Physician global score, mm	-		+						-			3

HAQ, units	-	-		-				-	-			5
Swollen joint count (28)					+				+			2
Tender joint count (28)		-	-	-	-			-	-	-		7
Sum of independent predictors*****	7	11	5	4	7	4	3	3	9	6	4	
Total number of available IVs*****	14	14	13	14	13	13	10	13	13	14	14	

Baseline variables that are identified as predictors in at least half of registries in which the variable is available are highlighted in bold.

DAPSA28: Disease Activity index for Psoriatic Arthritis in 28 joints; EPV: events-per-variable; IVs: independent variables; BMI: Body Mass Index; ; csDMARD: conventional synthetic Disease Modifying Anti-Rheumatic Drug; CRP: C-reactive protein; HAQ: Health Assessment Questionnaire; TNFi: Tumor Necrosis Factor Inhibitor; +: Odds Ratio (OR)>1; -: OR<1; constant: dichotomous variable, where only one category was available in the registry ; NA: variable not delivered by the registry; *Italy and Spain excluded due to no available CRP; **number of times a variable is selected as a predictor; ***TNFi initiation since January 1st 2009 was chosen as the start of data collection, as the first three bDMARDs (adalimumab, etanercept and infliximab) were then well-established treatment options across the European countries. 2015 was chosen as the separator between the time periods, as secukinumab was approved as the first non-TNFi bDMARD treatment option that year; ****the CRP cut-off was decided based on the various detection limits used across registries; *****sum of predictors selected per cohort; *****number of independent variables (after excluding NA and constant variables).

Table 3. Summary of predictors of DAPSA28 moderate response after 6 months of treatment with the first TNFi per registry* for registries with EPV per available independent variables ≥ 1 .

Country	Czech Republic	Denmark	Finland	Iceland	Norway	Portugal	Romania	Slovenia	Sweden	Switzerland	Turkey	Row sum**
Registry	ATTRA	DANBIO	ROB-FIN	ICEBIO	NOR-DMARD	Reuma.pt	RRBR	Biorx.si	SRQ	SCQM	TURKBIO	
Patients with DAPSA28 response assessment, n	462	1172	84	68	472	268	20	275	2205	116	133	
DAPSA28 moderate response, n (%)	265 (57)	317 (27)	34 (41)	13 (19)	143 (30)	106 (40)	11 (55)	124 (45)	711 (32)	17 (15)	62 (47)	
EPV per available IVs	15.5	22.6	2.6	0.9	11	8.2	1	9.5	54.7	1.2	4.4	
Age at treatment start, years	-	-	-		-			-	-			6
Men		+			+	+			+			4
Time since diagnosis, years	+	+	+						+			4
BMI, kg/m ²	-	-			NA	-		-	NA	-		5
Current smokers		-	-				constant	-	-	-		5
Concomitant csDMARD							constant		+			1
1 st TNFi start, year (2015-2018)***	+						constant					1
CRP>10 mg/l****	+	+			+	+	constant	+	+			6
Patient pain score, mm		+				-			+		+	4
Patient fatigue score, mm		-	NA		-	NA	NA	NA	-	-	-	5
Physician global score, mm	-					-		+				3

HAQ, units	-	-			-			-	-			5
Swollen joint count (28)	+	+			+	+	+		+	+	+	8
Tender joint count (28)		+						+				2
Sum of independent predictors*****	8	11	3	0	6	6	1	7	10	4	3	
Total number of available IVs*****	14	14	13	14	13	13	9	13	13	14	14	

Baseline variables that are selected as predictors in at least half of registries in which the variable is available are highlighted in bold.

DAPSA28: Disease Activity index for Psoriatic Arthritis in 28 joints; EPV: events-per-variable; IVs: independent variables; BMI: Body Mass Index; csDMARD: conventional synthetic Disease Modifying Anti-Rheumatic Drug; CRP: C-reactive protein; HAQ: Health Assessment Questionnaire; TNFi: Tumor Necrosis Factor Inhibitor; +: Odds Ratio (OR)>1; -: OR<1; constant: dichotomous variable, where only one category was available in the registry; NA: variable not delivered by the registry; *Italy and Spain excluded due to no available CRP; **number of times a variable is selected as a predictor; ***TNFi initiation since January 1st 2009 was chosen as the start of data collection, as the first three bDMARDs (adalimumab, etanercept and infliximab) were then well-established treatment options across the European countries. 2015 was chosen as the separator between the time periods, as secukinumab was approved as the first non-TNFi bDMARD treatment option that year; ****the CRP cut-off was decided based on the various detection limits used across registries; *****sum of predictors selected per cohort; *****number of independent variables (after excluding NA and constant variables).

Table 4. Summary of predictors of 12-month drug retention on the first TNFi per registry for registries with EPV per available independent variables ≥ 1 .

Country	Czech Republic	Denmark	Finland	Iceland	Italy	Norway	Portugal	Romania	Slovenia	Spain	Sweden	Switzerland	Turkey	Row sum*
Registry	ATTRA	DANBIO	ROB-FIN	ICEBIO	GISEA	NOR-DMARD	Reuma.pt	RRBR	Biorx.si	BIOBADASER	SRQ	SCQM	TURKBIO	
Number of patients	718	2090	234	306	1591	717	675	86	367	445	5225	628	287	
12-months drug retention, n (%)	504 (70)	1225 (59)	150 (64)	206 (67)	861 (54)	389 (54)	512 (76)	63 (73)	231 (63)	281 (63)	3468 (66)	387 (62)	184 (64)	
EPV per available IVs	15.3	61.8	6.5	7.1	60.9	25.2	12.5	1.9	10.5	18.2	135.2	17.2	7.4	
Age at treatment start, years					-	-			-	-	-		-	6
Men	+	+	+			+	+		+	+	+	+	+	10
Time since diagnosis, years	+	+		+	-		+		+	+	+	+		9
BMI, kg/m ²			+		+	NA			-		NA			3
Current smokers				-			-	+			-		-	5
Concomitant csDMARD	+	+	-			+	+			+		+		7
1 st TNFi start, year (2015-2018)**	-	-		-	-	-	-	constant	-	-		-	-	10
CRP>10 mg/l***		+			NA	+	+			NA	+		+	5
Patient pain score, mm			-	-		-	-			NA	-			5
Patient fatigue score, mm		-	NA		NA	+	NA	NA	NA	NA	-			3
Physician global score, mm	+									NA				1

HAQ, units										NA				0
Swollen joint count (28)	+				-						+			3
Tender joint count (28)	-	-									-	-	-	5
Sum of independent predictors****	7	7	4	4	5	7	7	1	5	5	9	5	6	
Total number of available IVs*****	14	14	13	14	12	13	13	12	13	9	13	14	14	

Baseline variables that are selected as predictors in at least half of registries in which the variable is available are highlighted in bold.

EPV: events-per-variable; IVs: independent variables; BMI: Body Mass Index; csDMARD: conventional synthetic Disease Modifying Anti-Rheumatic Drug; CRP: C-reactive protein; HAQ: Health Assessment Questionnaire; TNFi: Tumor Necrosis Factor Inhibitor; +: Odds Ratio (OR)>1; -: OR<1; constant: dichotomous variable, where only one category was available in the registry e; NA: variable not delivered by the registry.

*number of times a variable is selected as a predictor; **TNFi initiation since January 1st 2009 was chosen as the start of data collection, as the first three bDMARDs (adalimumab, etanercept and infliximab) were then well-established treatment options across the European countries. 2015 was chosen as the separator between the time periods, as secukinumab was approved as the first non-TNFi bDMARD treatment option that year; ***the CRP cut-off was decided based on the various detection limits used across registries; ****sum of predictors selected per cohort; *****number of independent variables (after excluding NA and constant variables).

Table 5. Univariate and final multivariate analyses for predicting DAPSA28 remission and DAPSA28 moderate response at 6 months and 12-month drug retention on the first TNFi in pooled data (derivation cohorts) for registries with EPV ≥ 1 .

	Prediction of DAPSA28 remission (n=3435)				Prediction of DAPSA28 moderate response (n=2537)				Prediction of 12-month drug retention (n=6642)			
	Univariate		Multivariate		Univariate		Multivariate		Univariate		Multivariate	
Patients achieving the outcome, n (%)	836 (24%)				860 (34%)				4170 (63%)			
	OR (95% CI)				OR (95% CI)				OR (95% CI)			
Age at treatment start, years	0.97	(0.97 - 0.98)	0.97	(0.96 - 0.98)	0.98	(0.97 - 0.99)			0.99	(0.99 - 1.00)	0.99	(0.99 - 1.00)
Men	2.43	(2.07 - 2.86)	1.85	(1.54 - 2.23)	1.96	(1.66 - 2.31)	1.71	(1.42 - 2.06)	1.66	(1.50 - 1.84)	1.47	(1.32 - 1.63)
Time since diagnosis, years	1.01	(1.00 - 1.02)			1.02	(1.01 - 1.04)	1.03	(1.01 - 1.04)	1.02	(1.01 - 1.03)		
BMI, kg/m ²	0.97	(0.94 - 0.99)	0.98	(0.95 - 1.00)	0.97	(0.95 - 0.99)	0.97	(0.95 - 0.99)	1.00	(0.98 - 1.01)		
Current smokers	0.69	(0.54 - 0.87)	0.74	(0.57 - 0.96)	0.82	(0.65 - 1.04)			0.73	(0.63 - 0.84)	0.77	(0.66 - 0.89)
Concomitant csDMARD	1.15	(0.98 - 1.35)			1.40	(1.17 - 1.68)	1.23	(1.01 - 1.50)	1.11	(1.00 - 1.22)		
1 st TNFi start, year (2015-2018)*	1.19	(1.01 - 1.39)			1.21	(1.02 - 1.42)			0.73	(0.66 - 0.81)	0.65	(0.58 - 0.72)
CRP>10 mg/l**	1.32	(1.09 - 1.58)	1.52	(1.22 - 1.89)	1.93	(1.62 - 2.29)	1.61	(1.33 - 1.95)	1.22	(1.07 - 1.39)	1.24	(1.08 - 1.43)
Patient pain score, mm	0.98	(0.97 - 0.98)			0.99	(0.99 - 1.00)	1.01	(1.00 - 1.01)	0.99	(0.99 - 0.99)	0.99	(0.99 - 1.00)

Patient fatigue score, mm	0.98	(0.97 - 0.98)	0.99	(0.98 - 0.99)	0.99	(0.98 - 0.99)	0.99	(0.98 - 0.99)	0.99	(0.99 - 0.99)	1.00	(0.99 - 1.00)
Physician global score, mm	0.99	(0.98 - 0.99)	0.99	(0.98 - 1.00)	1.01	(1.00 - 1.01)			1.00	(1.00 - 1.00)		
HAQ, units	0.32	(0.27 - 0.38)	0.57	(0.45 - 0.71)	0.73	(0.63 - 0.84)	0.75	(0.61 - 0.91)	0.79	(0.72 - 0.87)		
Swollen joint count (28)	0.97	(0.94 - 0.99)	1.05	(1.01 - 1.08)	1.08	(1.06 - 1.10)			1.00	(0.98 - 1.01)		
Tender joint count (28)	0.92	(0.90 - 0.93)			1.02	(1.00 - 1.03)			0.97	(0.96 - 0.98)	0.97	(0.96 - 0.99)
Age at treatment start, years (41-49)***							0.72	(0.56 - 0.92)				
Age at treatment start, years (50-57)							0.46	(0.36 - 0.60)				
Age at treatment start, years (58-84)							0.48	(0.37 - 0.63)				
Time since diagnosis, years (2nd quartile)***			1.20	(0.89 - 1.60)							1.12	(0.95 - 1.31)
Time since diagnosis, years (3rd quartile)			1.42	(1.09 - 1.84)							1.29	(1.11 - 1.50)
Time since diagnosis, years (4th quartile)			1.66	(1.26 - 2.20)							1.43	(1.21 - 1.69)
Patient pain score, mm (44-61)			0.64	(0.49 - 0.83)								
Patient pain score, mm (62-75)			0.77	(0.58 - 1.04)								
Patient pain score, mm (76-100)			0.80	(0.55 - 1.16)								
Swollen joint count (2-4)***							1.73	(1.38 - 2.16)				

Swollen joint count (5-28)								2.22	(1.74 - 2.84)				
Tender joint count (3-4)***			0.87	(0.66 - 1.15)									
Tender joint count (5-8)			0.60	(0.45 - 0.80)									
Tender joint count (9-28)			0.52	(0.36 - 0.74)									
AUROC (95% CI) ****			0.75 (0.73 - 0.77)				0.73 (0.70 - 0.75)				0.64 (0.62 - 0.65)		

Baseline variables that are common predictors across all outcomes are highlighted in bold. Registries with EPV ≥ 1 in derivation cohort, considering all independent variables, were included in all models (RRBR excluded from all analyses, ICEBIO and SCQM excluded from DAPSA28 response analyses).

DAPSA28: Disease Activity index for Psoriatic Arthritis in 28 joints; TNFi: Tumor Necrosis Factor Inhibitor; OR: odds ratio; 95CI: 95% confidence interval. BMI: Body Mass Index; csDMARD: conventional synthetic Disease Modifying Anti-Rheumatic Drug; CRP: C-reactive protein; HAQ: Health Assessment Questionnaire; AUROC: Area under the Receiver Operating Curve.

*TNFi initiation since January 1st 2009 was chosen as the start of data collection, as the first three bDMARDs (adalimumab, etanercept and infliximab) were then well-established treatment options across the European countries. 2015 was chosen as the separator between the time periods, as secukinumab was approved as the first non-TNFi bDMARD treatment option that year; **the CRP cut-off was decided based on the various detection limits used across registries; ***continuous independent variables were categorized if linearity assumption was violated. Cut-offs for time since diagnosis in DAPSA28 remission: 2nd quartile (2-3 yrs), 3rd quartile (4-9 yrs) and 4th quartile (10-56 yrs); 12-month drug retention: 2nd quartile (2 -3 yrs), 3rd quartile (4-8 yrs) and 4th quartile (9-56 yrs); ****AUROC was calculated in derivation cohort.

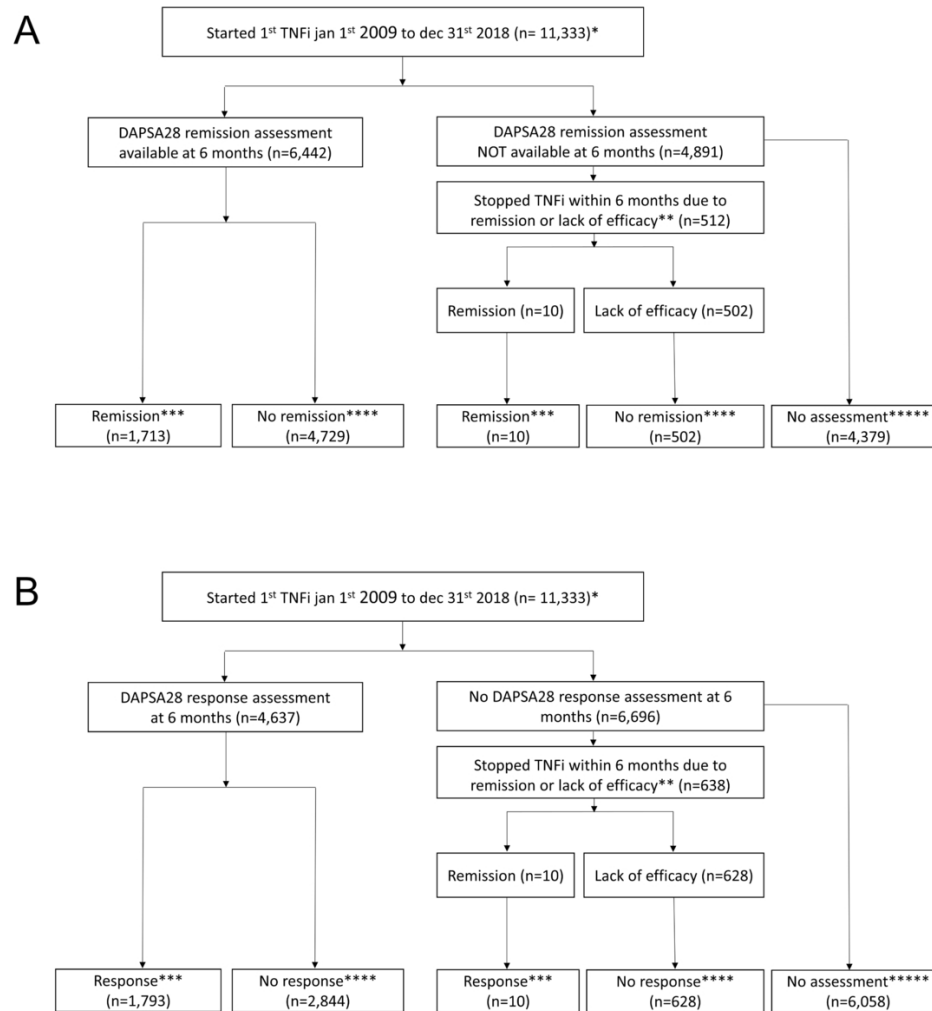


Figure 1

97x107mm (300 x 300 DPI)