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CLINICAL SCIENCE

Characteristics associated with poor COVID-19 outcomes in people with psoriasis, psoriatic arthritis and axial spondyloarthritis: data from the COVID-19 PsoProtect and Global Rheumatology Alliance physician-reported registries

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ABSTRACT

Objectives To investigate factors associated with severe COVID-19 in people with psoriasis (PsO), psoriatic arthritis (PsA) and axial spondyloarthritis (axSpA).

Methods Demographic data, clinical characteristics and COVID-19 outcome severity of adults with PsO, PsA and axSpA were obtained from two international physician-reported registries. A three-point ordinal COVID-19 severity scale was defined: no hospitalisation, hospitalisation (and no death) and death. ORs were estimated using multivariable ordinal logistic regression.

Results Of 5045 cases, 18.3% had PsO, 45.5% PsA and 36.3% axSpA. Most (83.6%) were not hospitalised, 14.6% were hospitalised and 1.8% died. Older age was non-linearly associated with COVID-19 severity. Male sex (OR 1.54, 95% CI 1.30 to 1.83), cardiovascular, respiratory, renal, metabolic and cancer comorbidities (ORs 1.25–2.89), moderate/high disease activity and/or glucocorticoid use (ORs 1.39–2.23, vs remission/low disease activity and no glucocorticoids) were associated with increased odds of severe COVID-19. Later pandemic time periods (ORs 0.42–0.52, vs until 15 June 2020), PsO (OR 0.49, 95% CI 0.37 to 0.65, vs PsA) and baseline exposure to TNFi, IL17i and IL-23i/IL-12+23i (OR 0.57, 95% CI 0.44 to 0.73; OR 0.62, 95% CI 0.45 to 0.87; OR 0.67, 95% CI 0.45 to 0.98; respectively; vs no disease-modifying

antirheumatic drug) were associated with reduced odds of severe COVID-19.

Conclusion Older age, male sex, comorbidity burden, higher disease activity and glucocorticoid intake were associated with more severe COVID-19. Later pandemic time periods, PsO and exposure to TNFi, IL17i and IL-23i/IL-12+23i were associated with less severe COVID-19. These findings will enable risk stratification and inform management decisions for patients with PsO, PsA and axSpA during COVID-19 waves or similar future respiratory pandemics.

INTRODUCTION

The COVID-19 pandemic has significantly impacted people with immune-mediated inflammatory diseases (IMIDs), particularly those taking immunomodulatory drugs such as biological or targeted synthetic disease-modifying antirheumatic drugs (b/tsDMARDs).^{1–4} While risk factors for severe COVID-19 outcomes have been demonstrated in both registry-based and population-based studies, for people with IMIDs collectively and for specific diseases such as rheumatoid arthritis, relevant risk factor data are limited for axial spondyloarthritis (axSpA) and psoriatic disease (including psoriasis without arthritis (PsO) and psoriatic arthritis

WHAT IS ALREADY KNOWN ON THIS TOPIC

- ⇒ Factors associated with severe COVID-19 outcomes have been demonstrated in both registry-based and population-based studies for people with immune-mediated inflammatory diseases (IMiDs) collectively and for specific IMiDs.
- ⇒ However, relevant risk factor data are limited for axial spondyloarthritis (axSpA) and psoriatic disease (including psoriasis without arthritis (PsO) and psoriatic arthritis (PsA)), a group of conditions that shares pathophysiological mechanisms and approved treatments, particularly targeted therapies.

WHAT THIS STUDY ADDS

- ⇒ Older age, male sex, comorbidity burden, higher disease activity and glucocorticoid intake were associated with more severe COVID-19.
- ⇒ Later pandemic time periods, PsO and exposure to TNFi, IL17i and IL-23i/IL-12+23i were associated with less severe COVID-19.

HOW THIS STUDY MIGHT AFFECT RESEARCH, PRACTICE OR POLICY

- ⇒ The findings from this study will enable risk stratification for patients with PsO, PsA and axSpA.
- ⇒ These findings will inform the development of tailored management strategies and evidence-based recommendations for patients with PsO, PsA and axSpA.

(PsA)).^{5–13} The association of specific classes of b/tsDMARDs commonly used in this population, including IL-17 inhibitors (IL17i) and IL-23 or IL-12/23 inhibitors (IL-23i/IL-12+23i), with COVID-19 outcomes has not been well studied. Improved understanding of the risks associated with exposure to these medications in this population will address knowledge gaps as we continue to navigate COVID-19 risks in the postvaccination era.

We used data from the COVID-19 Global Rheumatology Alliance (C19-GRA) and the Psoriasis Patient Registry for Outcomes, Therapy and Epidemiology of COVID-19 Infection (PsoProtect) physician-reported registries to evaluate the associations of baseline characteristics, including different classes of b/tsDMARDs, with COVID-19 severity in people with PsO, PsA and axSpA.

METHODS**Data source**

The C19-GRA physician-reported observational registry launched on 24 March 2020. Patients are eligible for inclusion if they have both a pre-existing rheumatic disease and SARS-CoV-2 infection. PsoProtect is a physician-reported observational registry launched on 27 March 2020. Patients are eligible for inclusion if they have both pre-existing PsO and SARS-CoV-2 infection. For both registries, data are entered voluntarily into the data entry systems by rheumatologists/dermatologists or under the supervision of rheumatologists/dermatologists. In Argentina, Brazil, France, Germany, Italy, Portugal and Sweden, C19-GRA data are transferred from national registries; in all other countries, data are entered directly into the registries' data entry systems. Countries were categorised according to the six WHO regions (www.who.int); the 'Americas' was further divided into north and south. Further details of the registries

have been described elsewhere.^{13–18} We used data collected on or before 25 October 2021.

COVID-19 reporting and primary outcome of interest

Both confirmed and presumptive cases of COVID-19 were reported. For analysis, patients were subsequently categorised into (1) confirmed or high likelihood of COVID-19 (chest imaging (CT or chest X-ray) showing bilateral infiltrates and/or symptoms after close contact with a known laboratory-confirmed COVID-19 positive patient) or (2) presumptive cases based on symptoms alone.

The primary outcome of interest of this study was COVID-19 outcome, assessed by use of an ordinal COVID-19 severity scale with three mutually exclusive categories: (1) no hospitalisation and no death; (2) hospitalisation, but no death and (3) death. 'Baseline characteristics' refer to demographic or clinical characteristics at the time of COVID-19 symptom onset (or diagnosis if asymptomatic).

IMiD treatment prior to COVID-19

Medications used to treat the IMiD prior to COVID-19 diagnosis were categorised into groups. Immunomodulatory drugs (conventional synthetic (cs)/biological (b)/targeted synthetic (ts) DMARDs) were distinguished from the PsO-specific non-biological systemic agent acitretin as well as from non-steroidal anti-inflammatory drugs (NSAIDs) and glucocorticoids (GC). csDMARDs included antimalarials, cyclosporine, leflunomide, methotrexate and sulfasalazine. bDMARDs included TNFi (eg, adalimumab, certolizumab, etanercept, golimumab, infliximab and TNFi biosimilars), IL-17i (eg, brodalumab, ixekizumab and secukinumab), IL-12/23i (ustekinumab) and IL-23i (eg, guselkumab, risankizumab and tildrakizumab). tsDMARDs included apremilast and JAKi (eg, baricitinib, tofacitinib and upadacitinib). IL-23i and IL-12/23i were combined in the same group for data analysis (IL-23i/IL-12+23i). Regarding NSAIDs, we asked physicians to report if at the time of COVID-19 symptom onset (or diagnosis if asymptomatic), the patient was taking NSAIDs, without specifying a minimal duration of a continuous treatment with NSAIDs. We chose no current DMARD use as the reference group after considering the groups' sample size and internal validity to be used as comparator for exposure to the various IMiD treatments. For more details regarding the choice of DMARD reference category, refer to online supplemental methods.

Statistical analyses

Descriptive tables were produced for the whole cohort and by diagnostic group (PsO, PsA and axSpA, as defined by the reporting healthcare professional). All patients with confirmed or presumptive COVID-19 were included in the primary analysis.

Independent associations between demographic and disease features and the ordinal COVID-19 outcome were estimated by multivariable ordinal logistic regression using the proportional odds model and were reported as OR and 95% CIs. In ordinal regression analysis, the effect size of a categorical predictor gives the change in log odds of being at least one level higher on the ordinal COVID-19 severity scale compared with the reference category of the predictor variable, while for a continuous predictor, it gives the change in odds of being one level higher on the ordinal COVID-19 severity scale for a unit increase in the continuous predictor. More details about assumptions of the proportional odds model are provided in online supplemental methods.

Factors potentially associated with the COVID-19 outcome considered in the models were age, sex, smoking habits (ever, unknown/missing, never), pandemic calendar period (until 15 June 2020, 16 June 2020 to 31 December 2020, 1 January 2021 and later), key comorbidities (chronic obstructive pulmonary disease (COPD) or asthma, other chronic lung disease, chronic kidney disease (CKD), hypertension, other cardiovascular disease (CVD), obesity, diabetes, cancer), IMiD diagnostic category, IMiD disease activity as per physician's global assessment (remission/low vs moderate/high), DMARD treatment prior to COVID-19 diagnosis, GC use and NSAID use.

For patients classified as having more than one IMiD or being treated with more than one of the medications of interest, we created a hierarchy based on clinical expertise to categorise patients. This way, non-overlapping (mutually exclusive) categories are obtained, allowing a clear reference group for interpretation of the regression models, and avoiding collinearities. Patients labelled as having both PsA and axSpA were counted as PsA patients. Patients receiving multiple csDMARDs were grouped according to the following hierarchy: cyclosporine>sulfasalazine>leflunomide>methotrexate>antimalarials, where 'A>B' means 'A has priority over B'. Patients receiving a b/tsDMARD and additionally a csDMARD were considered in the model solely in the b/tsDMARD group (ie, b/tsDMARD>csDMARD).

We tested four two-way additive interactions in the models: hypertension and CVD; obesity and diabetes; cancer and smoking habits; and disease activity and prednisolone-equivalent GC use. Online supplemental methods provide more details regarding statistical interactions.

To account for heterogeneity between participating countries regarding healthcare systems and infection dynamics, countries were considered as random effects in the regression analyses. To appropriately estimate the well-established non-linear effect of age on the outcome of SARS-CoV-2 infection, we included restricted cubic splines in the regression models. Four knots were chosen for most analyses, while three knots were chosen for the outcome mortality and the disease-specific analyses due to the limited effective sample size.¹⁹

Missing data were handled using multiple imputation; results of the logistic regression analyses for 10 imputed datasets were pooled by Rubin's rules. As disease activity was missing for all patients entered from France in the C19-GRA registry, country-level life expectancy was used in the imputation model to explain potential structural differences in disease activity between countries not accounted for in the patient-level data (data from 2018, source: <http://hdr.undp.org/>). For more details regarding excluded patients and handling of missing data, refer to online supplemental methods.

IMiDs differ regarding the DMARDs approved for their treatment. To explore the impact of this heterogeneity on the associations of interest, in addition to the primary analysis with all patients, diagnostic categories were defined, and stratified secondary analyses were undertaken separately for patients with PsO, PsA and axSpA.

The following sensitivity analyses were also performed to examine the robustness of our findings: (1) analysis limited to patients with confirmed or highly likely COVID-19; (2) analysis using the alternative binary outcome 'hospitalisation'; (3) analysis using the alternative binary outcome 'death'. In the model using death as dependent variable, comorbidities were analysed as an independent binary variable (3 or more comorbidities vs less than 3), to minimise the risk of overfitting. Data were considered statistically significant for p values < 0.05. All analyses were conducted in SAS (V.9.4) and R (V.4.0.4).

RESULTS

Study sample and baseline characteristics

The study population included 5045 cases, of which 921 (18.3%) were patients with PsO, 2293 (45.5%) with PsA, and 1831 (36.3%) with axSpA. Overall, the mean age was 50 years (SD 13.5), just over half were male (51.7%) and most were from Europe (77.5%) (table 1). Cases were reported fairly equally across the three pandemic time periods. Most cases had disease (IMiDs) in remission or minimal/low disease activity (82.7%). About half had no key comorbidities reported (52.9%). Of those with comorbidities, the most reported were hypertension (26.5%) and obesity (21.1%). Any csDMARD use was reported in 30.3%, with methotrexate as the most common (23.4%). Only 5.6% reported using sulfasalazine. bDMARD use was reported in 65.7% (TNFi 45.6%, IL17i 12.1%, IL-23i/IL-12+23i 8.1%). Only 1.2% reported JAKi use. Baseline GC use was reported in only 7.3% (4.6%, 0–7.5 mg/day and 1.4%, >7.5 mg/day) and NSAID use in 24%.

When stratified by condition (table 1), the main notable differences were that individuals with PsA were older (mean 53.2 years vs 46.9 years in axSpA and 48.4 years in PsO), a higher proportion of those with PsA had hypertension (32.8% vs 21.2% in axSpA and 21.3% in PsO) and a higher proportion of those with PsO were obese (29.2% vs 23% in PsA and 14.5% in axSpA). csDMARDs were most used among individuals with PsA (46.6% vs 19.3% in axSpA and 11.6% in PsO) while bDMARDs were most used among individuals with axSpA (73.6% vs 58.5% in PsA and 68.1% in PsO). Baseline GC usage was low overall but differed notably between the groups, with almost none in PsO (0.7%) vs 10.7% in PsA and 6.3% in axSpA. There was no difference across disease groups with regard to disease activity.

When stratified by medication group (online supplemental table 1), patients not taking DMARDs were slightly younger (mean 49.9 years) than patients taking DMARDs (range from 50 to 56.2 years, depending on the DMARD group) except for IL-17i/IL-23i/IL12+23i (mean 49.9 years) and TNFi (mean 48.3 years). Moreover, patients not taking DMARDs were slightly less often in remission/low disease activity (71.%) than patients taking DMARDs (range from 80.5% to 86.1%, depending on the DMARD group) except for JAKi (65.3% in remission/low disease activity).

COVID-19 outcomes

Baseline characteristics of the study population stratified by COVID-19 outcome are shown in online supplemental table 2. Most patients (4220, 83.6%) were not hospitalised, 736 (14.6%) were hospitalised and 89 (1.8%) died. The frequency of hospitalisation (without death) and death were slightly higher in PsA (17.1% and 2.2%, respectively), compared with axSpA (12.5% and 1.4%, respectively) and PsO (12.5% and 1.3%, respectively) (table 1).

Associations of baseline characteristics with COVID-19 severity

The results of the primary multivariable ordinal logistic regression model are shown in table 2 and the relationship between age and probability of hospitalisation and death is shown in figure 1.

Age was associated with COVID-19 severity in a non-linear way (stronger association for older age groups). Hypertension without CVD (OR 1.25, 95% CI 1.01 to 1.55), CVD without hypertension (OR 1.87, 95% CI 1.21 to 2.90), COPD or asthma (OR 1.75, 95% CI 1.33 to 2.31), other lung disease (OR 2.54, 95% CI 1.64 to 3.93), CKD (OR 2.32, 95% CI 1.50 to 3.58),

Table 1 Baseline characteristics of the study population (total and stratified by immune-mediated inflammatory disease diagnosis)

Parameter	Psoriatic arthritis	Axial spondyloarthritis	Psoriasis (without arthritis)	Total
N	2293	1831	921	5045
General				
Age (years)	53.2 (12.8)	46.9 (13.4)	48.4 (13.6)	50 (13.5)
≤30 years	110 (4.8)	211 (11.5)	100 (10.9)	421 (8.3)
31–50 years	785 (34.2)	900 (49.2)	400 (43.4)	2085 (41.3)
51–65 years	1032 (45)	567 (31)	336 (36.5)	1935 (38.4)
66–75 years	280 (12.2)	109 (6)	59 (6.4)	448 (8.9)
>75 years	86 (3.8)	44 (2.4)	26 (2.8)	156 (3.1)
Male sex	1053 (45.9)	996 (54.4)	557 (60.5)	2606 (51.7)
Ever-smoker	566 (33.2) (N=1705) (Missing=588)	313 (23.9) (N=1311) (Missing=520)	334 (36.3) (N=921) (Missing=0)	1213 (30.8) (N=3937) (Missing=1108)
Regions				
African Region	5 (0.2)	5 (0.3)	4 (0.4)	14 (0.3)
Eastern Mediterranean Region	34 (1.5)	31 (1.7)	7 (0.8)	72 (1.4)
European Region	1707 (74.4)	1396 (76.2)	805 (87.4)	3908 (77.5)
North American Region	454 (19.8)	221 (12.1)	53 (5.8)	728 (14.4)
South American Region	62 (2.7)	159 (8.7)	46 (5)	267 (5.3)
South-East Asian Region	10 (0.4)	8 (0.4)	3 (0.3)	21 (0.4)
Western Pacific Region	21 (0.9)	11 (0.6)	3 (0.3)	35 (0.7)
Time period				
Until 15 June 2020	744 (32.4)	564 (30.8)	417 (45.3)	1725 (34.2)
From 16 June 2020 to 31 December 2020	1043 (45.5)	880 (48.1)	340 (36.9)	2263 (44.9)
1 January 2021 and later	506 (22.1)	387 (21.1)	164 (17.8)	1057 (21)
Ordinal outcome				
Not hospitalised, no death	1850 (80.7)	1576 (86.1)	794 (86.2)	4220 (83.6)
Hospitalised, no death	392 (17.1)	229 (12.5)	115 (12.5)	736 (14.6)
Death	51 (2.2)	26 (1.4)	12 (1.3)	89 (1.8)
Disease activity	(N=2014) (Missing=279)	(N=1288) (Missing=532)	(N=920) (Missing=1)	N=4233 (Missing=812)
Remission/low disease activity	1670 (82.9)	1090 (83.9)	740 (80.4)	3500 (82.7)
Moderate/high disease activity	344 (17.1)	209 (16.1)	180 (19.6)	733 (17.3)
Comorbidities	(N=2266) (Missing=27)	(N=1809) (Missing=22)	(N=921) (Missing=0)	N=4996 (Missing=49)
Hypertension	744 (32.8)	383 (21.2)	196 (21.3)	1323 (26.5)
Cardiovascular disease	179 (7.9)	88 (4.9)	71 (7.7)	338 (6.8)
COPD or asthma	185 (8.2)	114 (6.3)	62 (6.7)	361 (7)
Other chronic lung disease	47 (2.1)	30 (1.7)	22 (2.4)	99 (2)
Chronic kidney disease	60 (2.6)	24 (1.3)	17 (1.8)	101 (2)
Diabetes	316 (13.9)	126 (7)	120 (13)	562 (11.2)
Cancer	65 (2.9)	26 (1.4)	24 (2.6)	115 (2.3)
Obesity	522 (23)	262 (14.5)	269 (29.2)	1053 (21.1)
No of comorbidities	1 (1.2)	0.6 (1)	1 (1.2)	0.8 (1.1)
No comorbidity	1077 (47.5)	1124 (62.1)	442 (48)	2643 (52.9)
1 comorbidity	573 (25.3)	418 (23.1)	248 (26.9)	1239 (24.8)
2 comorbidities	350 (15.4)	181 (10)	127 (13.8)	658 (13.2)
≥3 comorbidities	266 (11.7)	86 (4.8)	104 (11.3)	456 (9.1)
DMARDs (monotherapy or combination therapy)				
csDMARDs	1068 (46.6)	353 (19.3)	107 (11.6)	1528 (30.3)
Antimalarials	30 (1.3)	7 (0.4)	0	37 (0.7)
Methotrexate	889 (38.8)	194 (10.6)	100 (10.9)	1183 (23.4)
Leflunomide	94 (4.1)	13 (0.7)	0	107 (2.1)
Sulfasalazine	119 (5.2)	164 (9)	0	283 (5.6)
Cyclosporine	11 (0.5)	0	8 (0.9)	19 (0.4)
bDMARDs	1341 (58.5)	1347 (73.6)	627 (68.1)	3315 (65.7)
TNF inhibitors	895 (39)	1176 (64.2)	227 (24.6)	2298 (45.6)

Continued

Table 1 Continued

Parameter	Psoriatic arthritis	Axial spondyloarthritis	Psoriasis (without arthritis)	Total
N	2293	1831	921	5045
IL-17 inhibitors	301 (13.1)	164 (9)	145 (15.7)	610 (12.1)
IL-23/IL-12+23 inhibitors	145 (6.3)	7 (0.4)	255 (27.7)	407 (8.1)
tsDMARDs	111 (4.8)	11 (0.6)	19 (2.1)	141 (2.8)
JAK inhibitors	51 (2.2)	11 (0.6)	0	62 (1.2)
Apremilast	60 (2.6)	0	19 (2.1)	79 (1.6)
No DMARD treatment	234 (10.2)	309 (16.9)	179 (19.4)	722 (14.3)
Other therapies				
Glucocorticoids (#)	241 (10.7) (N=2242) (Missing=51)	110 (6.3) (N=1760) (Missing=71)	6 (0.7) (N=921) (Missing=0)	357 (7.3) (N=4923) (Missing=122)
0 mg/day <glucocorticoids ≤7.5 mg/day	167 (7.5) (N=2215) (Missing=78)	54 (3.1) (N=1724) (Missing=107)	3 (0.3) (N=919) (Missing=2)	224 (4.6) (N=4858) (Missing=187)
Glucocorticoids >7.5 mg/day	46 (2.1) (N=2215) (Missing=78)	19 (1.1) (N=1724) (Missing=107)	1 (0.1) (N=919) (Missing=2)	66 (1.4) (N=4858) (Missing=187)
NSAIDs	512 (24.5) (N=2094) (Missing=199)	553 (35.5) (N=1559) (Missing=272)	32 (3.5) (N=921) (Missing=0)	1097 (24) (N=4574) (Missing=471)
Acitretin	3 (0.1)	0	26 (2.8)	29 (0.6)

Data are N (column %) for categorical variables or mean (SD) for continuous variables. Table includes patients diagnosed with psoriasis without arthritis, psoriatic arthritis or axial spondyloarthritis, with a non-missing ordinal outcome and non-missing values for age, sex, time period and DMARDs. Further, patients receiving multiple b/tsDMARDs or DMARDs not typical for the three diagnoses were excluded, as well as patients labelled as having additional inflammatory rheumatic diseases (529 patients excluded in total). Data refer to patients with non-missing values for the respective variable; total N for patients with non-missing values is given in parentheses for variables with missing values; the total number of missing values is also given in parenthesis, for the applicable variables. (#) Includes patients with a missing glucocorticoid dosage. bDMARD, biological disease-modifying anti-rheumatic drugs; BMI, body mass index; COPD, chronic obstructive pulmonary disease; csDMARD, conventional synthetic DMARD; DMARD, disease-modifying antirheumatic drug; IL, interleukin; JAK, Janus kinase; N, number; NSAID, non-steroidal anti-inflammatory drugs; TNF, tumour necrosis factor; tsDMARD, targeted synthetic DMARD.

cancer in patients with missing data on smoking (OR 2.89, 95% CI 1.19 to 6.97), obesity without diabetes (OR 1.35, 95% CI 1.07 to 1.70), diabetes without obesity (OR 1.84, 95% CI 1.38 to 2.45), and coexistence of obesity and diabetes (OR 1.89, 95% CI 1.34 to 2.68) were associated with greater odds of worse COVID-19 severity compared with referents without each condition. Male sex was associated with 1.54 times greater odds of worse COVID-19 severity compared with female sex (95% CI 1.30 to 1.83). Moderate/high disease activity (with or without GC use) and remission/low disease activity (with GC use) were associated with higher odds of worse COVID-19 outcomes compared with being in remission/low disease activity without GC use (OR ranging from 1.39 to 2.23). Later pandemic time periods were associated with lower odds of worse COVID-19 severity compared with the baseline period of March 2020–15 June 2020 (OR 0.42, 95% CI 0.34 to 0.51 for 16 June 2020–31 December 2020; OR 0.52, 95% CI 0.41 to 0.67 for 1 January 2021 and later). Compared with PsA, PsO was associated with less COVID-19 severity (OR 0.49, 95% CI 0.37 to 0.65). For medication classes, none were associated with higher odds of COVID-19 severity. TNFi, IL17i and IL-23i/IL-12+23i all demonstrated reduced odds of severe COVID-19 outcomes (OR 0.57, 95% CI 0.44 to 0.73; OR 0.62, 95% CI 0.45 to 0.87; OR 0.67, 95% CI 0.45 to 0.98, respectively). Finally, NSAID use compared with no use of NSAIDs was associated with lower odds of severe COVID-19 outcomes (OR 0.77, 95% CI 0.60 to 0.98).

Stratified analyses

When stratified by condition, results were similar to the primary model (online supplemental table 3) and online supplemental

figures 1-3) with the following notable exceptions: hypertension alone and CVD alone were only significantly associated with the COVID-19 severity outcome among those with axSpA (OR 1.49, 95% CI 1.01 to 2.19; and OR 2.77, 95% CI 1.25 to 6.13; respectively) whereas COPD and asthma were associated with the COVID-19 severity outcome only among those with PsA (OR 1.95, 95% CI 1.34 to 2.82). The association of IL-23i/IL-12+23i with less severe COVID-19 outcomes was only statistically significant among those with PsO (OR 0.43, 95% CI 0.23 to 0.82); however, IL-23i/IL-12+23i were not used among patients with axSpA (not efficacious/licensed for this indication) and numbers were lower for PsA.

Sensitivity analyses

The results of sensitivity analyses are shown in online supplemental tables 4-6 and online supplemental figures 4-6. When restricting the analysis to confirmed COVID-19 cases (n=4176), multivariable model results were consistent with the primary model. Results were also similar to the primary model for the binary outcome of hospitalisation.

For the binary outcome of death, male sex (OR 2.00, 95% CI 1.22 to 3.26), having three or more comorbidities (OR 3.34, 95% CI 1.98 to 5.63) and baseline GC use (OR 1.91, 95% CI 1.002 to 3.64) remained associated with the outcome of interest. In this model, TNFi and IL17i continued to demonstrate reduced odds of severe COVID-19 outcomes (OR 0.50, 95% CI 0.26 to 0.98 and OR 0.11, 95% CI 0.02 to 0.51; respectively). However, sulfasalazine use (OR 2.64, 95% CI 1.13 to 6.17) and JAKi use (OR 7.49, 95% CI 2.61 to 21.47) were associated with greater odds of severe COVID-19 outcomes in this model.

Table 2 Multivariable ordinal logistic regression analysis of factors associated with COVID-19 severity (primary model, all patients)

N total		5045		
N deaths/hospitalisations without death/neither		89/736/4220		
	N deaths/hospitalisations without death/neither	OR	95% CI	
Male sex (vs female)	57/419/2130	1.54	1.30	1.83
Pandemic time period				
Until 15 June 2020	45/395/1285	1	(Reference)	
16 June 2020–31 December 2020	28/217/2018	0.42	0.34	0.51
1 January 2021 and later	16/124/917	0.52	0.41	0.67
Comorbidities				
Hypertension alone (vs no hypertension, no CVD)	28/242/847	1.25	1.01	1.55
CVD alone (vs no hypertension, no CVD)	7/38/76	1.87	1.21	2.90
CVD and hypertension (vs no hypertension, no CVD)	21/63/136	1.41	0.98	2.02
COPD or asthma	21/87/257	1.75	1.33	2.31
Other lung disease	11/34/55	2.54	1.64	3.93
Chronic kidney disease	14/42/46	2.32	1.50	3.58
Obesity alone (vs no obesity, no diabetes)	11/138/676	1.35	1.07	1.70
Diabetes mellitus alone (vs no obesity, no diabetes)	15/102/212	1.84	1.38	2.45
Obesity and diabetes mellitus (vs no obesity, no diabetes)	14/61/162	1.89	1.34	2.68
Cancer and known smoking habits (vs no cancer, never smoked)	4/29/57	1.13	0.68	1.88
Cancer and unknown smoking habits (vs no cancer, never smoked)	5/7/13	2.89	1.19	6.97
No cancer and ever smoked or unknown smoking habits (vs no cancer, never smoked)	38/281/1933	0.87	0.71	1.06
Rheumatic disease				
Psoriatic arthritis	51/392/1850	1	(Reference)	
Axial spondyloarthritis	26/229/1576	1.07	0.86	1.33
Psoriasis (without arthritis)	12/115/794	0.49	0.37	0.65
Medication				
No DMARD therapy	21/128/573	1	(Reference)	
Antimalarials	0/4/14	1.08	0.30	3.84
Methotrexate	13/133/449	1.03	0.76	1.40
Leflunomide	2/16/42	1.08	0.55	2.11
Sulfasalazine	12/32/136	1.41	0.91	2.17
Cyclosporine	0/1/18	0.31	0.04	2.47

Continued

Table 2 Continued

N total		5045		
N deaths/hospitalisations without death/neither		89/736/4220		
	N deaths/hospitalisations without death/neither	OR	95% CI	
TNF inhibitors	24/248/2026	0.57	0.44	0.73
IL-17 inhibitors	2/88/520	0.62	0.45	0.87
IL-23/IL-12+23 inhibitors	5/58/344	0.67	0.45	0.98
JAK inhibitors	7/10/45	1.58	0.83	3.01
Apremilast	3/18/58	1.04	0.57	1.91
Disease activity (DA) and glucocorticoids (GCs)				
Remission/low DA, no GCs	57/516/3358	1	(Reference)	
Remission/low DA, GCs	13/59/177	1.97	1.39	2.79
Moderate/high DA, no GCs	15/130/604	1.39	1.09	1.76
Moderate/high DA, GCs	5/32/82	2.23	1.39	3.58
NSAIDs	16/151/1072	0.77	0.60	0.98

Results for ordinal mixed effects logistic regression analysis in all patients (primary model). Shown are fixed effects, random effects for country are not shown. Missing values are imputed via multiple imputation, patient numbers may thus be rounded. The model was additionally adjusted for age employing four-knot restricted cubic splines. Significant associations highlighted in bold.

bDMARD, biological disease-modifying antirheumatic drug; BMI, body mass index; COPD, chronic obstructive pulmonary disease; csDMARD, conventional synthetic DMARD; CVD, cardiovascular disease; DMARD, disease-modifying antirheumatic drug; IL, interleukin; JAK, Janus kinase; N, number; NSAID, non-steroidal antiinflammatory drug; TNF, tumour necrosis factor; tsDMARD, targeted synthetic DMARD.

Discussion

In this registry-based study of individuals with PsO, PsA and axSpA with SARS-CoV-2 infection, we found that known risk factors for the general population (older age, the presence of comorbidities) and for IMIDs overall (higher disease activity, higher baseline GC usage) were associated with more severe COVID-19 outcomes. In addition, a diagnosis of COVID-19 in a later time period during the pandemic was associated with lower disease severity compared with early 2020. Consistent with previous studies, baseline TNFi use was associated with lower odds for severe COVID-19 outcomes; we also found that IL17i and IL-23i/IL-12+23i use had similar associations with lower odds for severe COVID-19 outcomes.

The findings of our study reiterate known risk factors in both the general population and among people with IMIDs: older age, male sex and presence of comorbidities, specifically cardiometabolic and pulmonary conditions, were associated with more severe COVID-19 outcomes.^{5,6} Our findings that disease activity and GC usage at baseline have an additive interaction are consistent with prior findings in the C19-GRA registry.²⁰

In this study, baseline use of TNFi was associated with lower odds of severe COVID-19 outcomes. This was previously shown in the C19-GRA registry,⁷ in a combined rheumatic disease, inflammatory bowel disease (IBD) and PsO analysis,⁸ and in a US-based administrative claims database study among individuals with RA.²¹ Mechanistic plausibility for trialling TNFi therapies for COVID-19 treatment has been discussed in the literature.^{22,23} These therapies neutralise TNF, a major cytokine

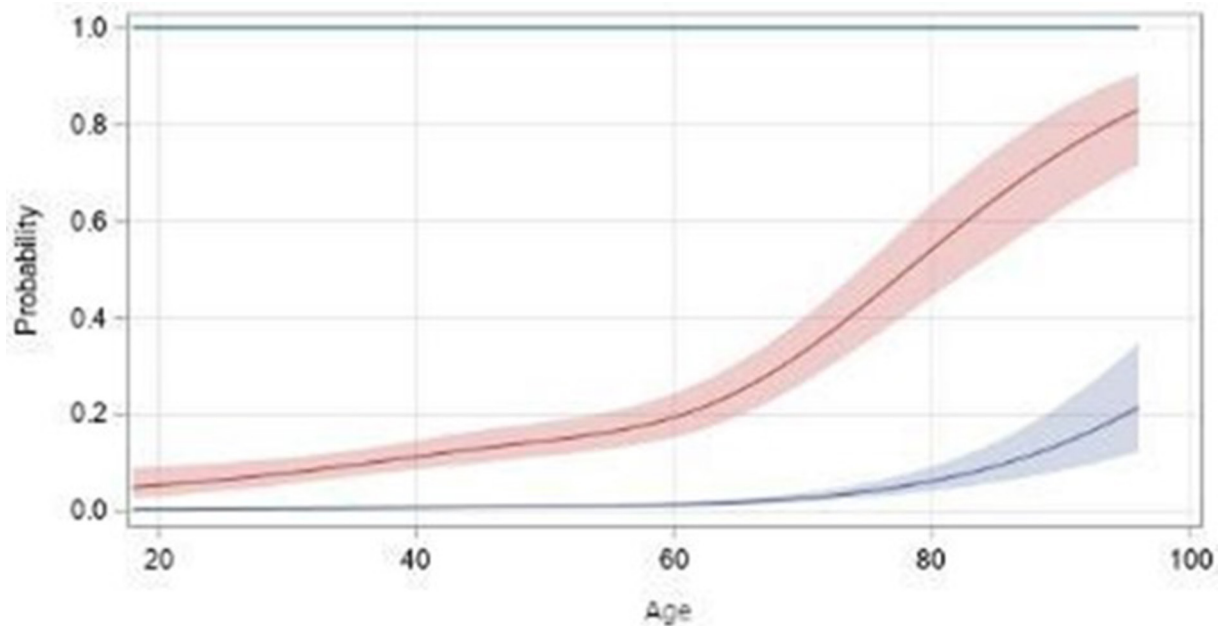


Figure 1 Relationship between age and probability of hospitalisation (red) and death (blue) estimated by four-knot restricted cubic splines, with 95% CIs (primary model, ordinal outcome, all patients).

in the excess inflammatory phase of COVID-19, and several trials are ongoing. A recent preprint announced results of a large randomised, placebo-controlled clinical trial led by the National Institutes of Health showing that treating adults hospitalised with COVID-19 with infliximab (a TNFi) did not significantly shorten time to recovery but did improve 14-day clinical status and substantially reduced 28-day mortality compared with standard of care²⁴—the peer-reviewed publication is awaited.

We also demonstrated that using IL17i and IL-23i/IL-12+23i was also associated with lower odds of severe COVID-19 outcomes. Prior population-level data from Israel and the UK have shown that the use of IL-17i was not associated with worse COVID-19 outcomes.^{25 26} At the same time, case reports and case series have also suggested that IL-17 and IL-23 inhibition may not have a negative effect on the course of COVID-19,^{27–29} though further inference on whether exposure to these medications might be associated with better COVID-19 outcomes is limited. IL-17 may play a pathogenic role in acute respiratory distress syndrome and lung inflammation associated with severe COVID-19. Patients with COVID-19 who experience pulmonary complications have increased and activated Th17 cell populations, and lung damage and hyperinflammation are linked to these patients' increased Th17 cell responses.^{30 31} The anti-IL-17 monoclonal antibody netakimab improved survival in a small clinical trial in patients with COVID-19; it decreased lung lesion volume and the need for oxygen support.³² However, in another study, netakimab therapy improved some clinical parameters and decreased C reactive protein levels, but it had no effect on the need for mechanical ventilation or patient survival in COVID-19 patients.³³ Suppressing inflammation via a variety of mechanisms has been shown to improve COVID-19 outcomes in people with severe disease (ie, GC, IL-6i, JAKi, maybe TNFi). Whether IL-17 will also have a role remains to be determined and requires further study.^{34 35} Importantly, in our study, we report associations and therefore we caution against interpreting our estimates causally, as the possibility of selection bias and unmeasured confounding cannot be excluded.

Apremilast was not associated with the severity of COVID-19 in patients with PsO/PsA. Although the number of patients taking apremilast was low, these data are important because they add to limited previous evidence of a favourable safety profile of apremilast on COVID-19 severity in patients with these conditions.^{36–38}

The finding that baseline NSAID use was associated with less COVID-19 severity is interesting but should be interpreted with caution. NSAID use is particularly prone to reporting bias, and inconsistencies in reporting might have resulted from the fact that we did not specify a minimal duration of a continuous treatment with NSAID and did not use a standardised questionnaire to collect NSAID data (eg, type of NSAID, dose and duration of treatment). General population studies in the UK and Denmark have not found associations between NSAID use and COVID-19-related hospitalisation or death.^{39–41} In our study, this association was seen particularly in individuals with axSpA and may be related to milder disease and/or well controlled of disease activity; confounding by indication cannot be excluded.

Finally, the results of one sensitivity analysis indicated that use of sulfasalazine and JAKi were associated with higher odds of death (binary outcome) due to COVID-19, though there were no associations with the ordinal COVID-19 severity outcome or with hospitalisation (binary outcome). In the C-19 GRA registry, we previously found an association of sulfasalazine use with worse COVID-19 outcome,⁷ a finding which was also seen in initial analyses of the Surveillance Epidemiology of Coronavirus Under Research Exclusion (IBD) database⁴² though later analyses were null.⁴³ While there are biologically plausible effects of sulfasalazine on SARS-CoV-2 viral entry,⁴⁴ our results may be due to residual confounding. The association of JAKi usage with COVID-19 outcomes is consistent with findings from some studies focused on people with RA.^{9 45} However, results from this sensitivity analysis should be interpreted with caution as the proportion of patients on JAKi was low (and no patients with PsO were taking this medication) and the respective 95% CI was wide.

Our study has several strengths, including the international nature of the combined registries, the large sample size and the granularity of information regarding IMID medications and disease activity. Our study also has limitations. First, the C19-GRA and PsoProtect registries were dependent on voluntary provider entry of cases, and there may be bias towards cases with more severe COVID-19 and those on DMARD therapy, as mostly secondary care clinicians were submitting cases. As such, proportions of events in our study sample should not be interpreted as incidence rates. Second, while we tried to mitigate the impacts of selection bias and confounding by indication, it is possible that our results may still be biased. However, we performed a series of sensitivity analyses to confirm the robustness of our findings, including restricting to a sample of confirmed cases of COVID-19, and our results were consistent across these additional analyses. Third, although we were able to adjust for several potential confounders in our models, there may still be residual unmeasured confounding. We did not have data available on disease duration or prior medication use, apart from what was reported at the time of COVID-19 diagnosis. Finally, vaccination status was not available for the patients in this dataset; however, the model adjustment for pandemic calendar period used in this study may act as a surrogate for vaccination status.

In conclusion, more severe COVID-19 outcomes in PsO, PsA and axSpA are largely associated with age, comorbidities, active disease and GC use. None of the bDMARDs typically used in PsO, PsA and axSpA, including TNFi, IL-17i and IL-23i/IL-12+23i, were associated with severe COVID-19 outcomes, and no biologics-specific differences were found. Our findings will help clinicians, scientific societies and policy makers worldwide develop tailored management strategies for patients with PsO, PsA and axSpA during COVID-19 waves or similar future respiratory pandemics.

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Correction notice This article has been corrected since it published Online First. The first authorship statement has been added.

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Appendix 2. Psoriasis Patient Registry for Outcomes, Therapy and Epidemiology of COVID-19 Infection (PsoProtect) Consortium.

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Freya Meynell
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SUPPLEMENTARY TABLES AND FIGURES

Supplementary table 1. Baseline characteristics of the study population (total, and stratified by medication group)							
Parameter	No DMARDs	csDMARDs only	TNFi	IL-17i/IL-23i/IL12+23i	JAKi	APREMILAST	TOTAL
N	722	867	2298	1017	62	79	5045
General							
Age [years]	49.9 (15.2)	54 (13.4)	48.3 (13.0)	49.9 (12.8)	54.3 (12.5)	56.2 (11.9)	50 (13.5)
≤ 30 years	81 (11.2)	36 (4.2)	226 (9.8)	72 (7.1)	3 (4.8)	3 (3.8)	421 (8.3)
31 – 50 years	281 (38.9)	291 (33.6)	1044 (45.4)	433 (42.6)	19 (30.6)	17 (21.5)	2085 (41.3)
51 – 65 years	252 (34.9)	377 (43.5)	836 (36.4)	402 (39.5)	24 (38.7)	44 (55.7)	1935 (38.4)
66 - 75 years	70 (9.7)	111 (12.8)	152 (6.6)	88 (8.7)	16 (25.8)	11 (13.9)	448 (8.9)
> 75 years	38 (5.3)	52 (6)	40 (1.7)	22 (2.2)	0	4 (5.1)	156 (3.1)
Male sex	357 (49.4)	385 (44.4)	1267 (55.1)	546 (53.7)	20 (32.3)	31 (39.2)	2606 (51.7)
Ever smoker	159 (29.6) (N=538) (NA=184)	196 (31.2) (N=628) (NA=239)	517 (29.1) (N=1777) (NA=521)	308 (35.0) (N=879) (NA=138)	9 (19.6) (N=46) (NA=16)	24 (34.8) (N=69) (NA=10)	1213 (30.8) (N=3937) (NA=1108)
Regions							
African Region	4 (0.6)	6 (0.7)	3 (0.1)	1 (0.1)	0	0	14 (0.3)
Eastern Mediterr. Region	19 (2.6)	17 (2)	27 (1.2)	7 (0.7)	2 (3.2)	0	72 (1.4)
European Region	550 (76.2)	679 (78.3)	1769 (77)	822 (80.8)	35 (56.5)	53 (67.1)	3908 (77.5)
North American Region	102 (14.1)	106 (12.2)	343 (14.9)	128 (12.6)	24 (38.7)	25 (31.6)	728 (14.4)
South American Region	29 (4.0)	38 (4.4)	146 (6.4)	53 (5.2)	1 (1.6)	0	267 (5.3)
South-East Asian Region	5 (0.7)	11 (1.3)	3 (0.1)	1 (0.1)	0	1 (1.3)	21 (0.4)
Western Pacific Region	13 (1.8)	10 (1.2)	7 (0.3)	5 (0.5)	0	0	35 (0.7)
Time period							
Until June 15, 2020	253 (35.0)	304 (35.1)	756 (32.9)	357 (35.1)	20 (32.3)	35 (44.3)	1725 (34.2)
June 16 to Dec 31, 2020	297 (41.1)	367 (42.3)	1073 (46.7)	469 (46.1)	27 (43.5)	30 (38)	2263 (44.9)
January 1, 2021 or later	172 (23.8)	196 (22.6)	469 (20.4)	191 (18.8)	15 (24.2)	14 (17.7)	1057 (21)
Immune-mediated inflammatory diseases							
SpA	309 (42.8)	164 (18.9)	1176 (51.2)	171 (16.8)	11 (17.7)	0	1831 (36.3)
PsA	234 (32.4)	607 (70.0)	895 (38.9)	446 (43.9)	51 (82.3)	60 (75.9)	2293 (45.5)
Psoriasis (no arthritis)	179 (24.8)	96 (11.1)	227 (9.9)	400 (39.3)	0	19 (24.1)	921 (18.3)
Disease activity							
	N=606 (NA=116)	N=762 (NA=105)	N=1821 (NA=477)	N=918 (NA=99)	N=49 (NA=13)	N=77 (NA=2)	N=4233 (NA=812)
Remission, min./low DA	432 (71.3)	625 (82)	1567 (86.1)	782 (85.2)	32 (65.3)	62 (80.5)	3500 (82.7)
Moderate, sev./high DA	174 (28.7)	137 (18)	254 (13.9)	136 (14.8)	17 (34.7)	15 (19.5)	733 (17.3)
Ordinal outcome							
Not hospitalized/dead	573 (79.4)	654 (75.4)	2026 (88.2)	864 (85)	45 (72.6)	58 (73.4)	4220 (83.7)
Hospitalized, no death	128 (17.7)	186 (21.5)	248 (10.8)	146 (14.4)	10 (16.1)	18 (22.8)	736 (14.6)
Death	21 (2.9)	27 (3.1)	24 (1)	7 (0.7)	7 (11.3)	3 (3.8)	89 (1.8)
Comorbidities							
	N=714 (NA=8)	N=854 (NA=13)	N=2281 (NA=17)	N=1008 (NA=9)	N=62 (NA=0)	N=77 (NA=2)	N=4996 (NA=49)

Hypertension	183 (25.6)	263 (30.8)	555 (24.3)	285 (28.3)	13 (21.0)	24 (31.2)	1323 (26.5)
Cardiovascular disease	56 (7.8)	84 (9.8)	109 (4.8)	81 (8.0)	0	8 (10.4)	338 (6.8)
COPD or asthma	52 (7.3)	58 (6.8)	130 (5.7)	102 (10.1)	12 (19.4)	7 (9.1)	361 (7.2)
Other lung disease	15 (2.1)	13 (1.5)	37 (1.6)	27 (2.7)	4 (6.5)	3 (3.9)	99 (2.0)
Chronic kidney disease	14 (2.0)	23 (2.7)	36 (1.6)	23 (2.3)	1 (1.6)	4 (5.2)	101 (2.0)
Obesity (BMI >= 30)	137 (19.2)	171 (20.0)	426 (18.7)	289 (28.7)	12 (19.4)	18 (23.4)	562 (11.2)
Diabetes	81 (11.3)	104 (12.2)	227 (10.0)	133 (13.2)	7 (11.3)	10 (13.0)	115 (2.3)
Cancer	31 (4.3)	24 (2.8)	20 (0.9)	28 (2.8)	1 (1.6)	11 (14.3)	1053 (21.1)
number of comorbidities	0.8 (1.1)	0.9 (1.2)	0.7 (1.0)	1 (1.3)	0.8 (1.0)	1.3 (1.4)	0.8 (1.1)
No comorbidity	371 (52.0)	436 (51.1)	1313 (57.6)	461 (45.7)	33 (53.2)	29 (37.7)	2643 (52.9)
One comorbidity	187 (26.2)	224 (26.2)	524 (23)	271 (26.9)	14 (22.6)	19 (24.7)	1239 (24.8)
Two comorbidities	98 (13.7)	107 (12.5)	281 (12.3)	144 (14.3)	12 (19.4)	16 (20.8)	658 (13.2)
≥ 3 comorbidities	58 (8.1)	87 (10.2)	163 (7.1)	132 (13.1)	3 (4.8)	13 (16.9)	456 (9.1)
csDMARD therapy							
csDMARDs	0	867 (100)	487 (21.2)	138 (13.6)	22 (35.5)	14 (17.7)	1528 (30.3)
Methotrexate	0	649 (74.9)	388 (16.9)	119 (11.7)	18 (29.0)	9 (11.4)	1183 (23.4)
Leflunomide	0	64 (7.4)	30 (1.3)	7 (0.7)	3 (4.8)	3 (3.8)	107 (2.1)
Sulfasalazine	0	181 (20.9)	86 (3.7)	12 (1.2)	3 (4.8)	1 (1.3)	283 (5.6)
Antimalarials	0	27 (3.1)	8 (0.4)	1 (0.1)	0	1 (1.3)	37 (0.7)
Cyclosporine	0	14 (1.6)	2 (0.1)	3 (0.3)	0	0	19 (0.4)
Further therapies							
Glucocorticoids (#)	52 (7.4) (N=698) (NA=24)	112 (13.1) (N=858) (NA=9)	121 (5.4) (N=2229) (NA=69)	57 (5.7) (N=1004) (NA=13)	12 (20.7) (N=58) (NA=4)	3 (3.9) (N=76) (NA=3)	357 (7.3) (N=4923) (NA=122)
0 mg/d < GCs <= 7.5 mg/d	31 (4.5) (N=691) (NA=31)	73 (8.6) (N=847) (NA=20)	75 (3.4) (N=2198) (NA=100)	38 (3.8) (N=991) (NA=26)	5 (9.1) (N=55) (NA=7)	2 (2.6) (N=76) (NA=3)	224 (4.6) (N=4858) (NA=187)
GCs > 7.5 mg/d	14 (2.0) (N=691) (NA=31)	27 (3.2) (N=847) (NA=20)	14 (0.6) (N=2198) (NA=100)	6 (0.6) (N=991) (NA=26)	4 (7.3) (N=55) (NA=7)	1 (1.3) (N=76) (NA=3)	66 (1.4) (N=4858) (NA=187)
NSAIDs	214 (31.9) (N=671) (NA=51)	199 (24.8) (N=802) (NA=65)	480 (23.9) (N=2012) (NA=286)	175 (18.2) (N=961) (NA=56)	14 (25.0) (N=56) (NA=6)	15 (20.8) (N=72) (NA=7)	1097 (24.0) (N=4574) (NA=471)
Acitretin	25 (3.5)	1 (0.1)	2 (0.1)	0	1 (1.6)	0	29 (0.6)

Data are N (column %) for categorical variables or mean (SD) for continuous variables. Table includes patients diagnosed with psoriasis, psoriatic arthritis or axial spondyloarthritis, with a non-missing ordinal outcome and non-missing values for age, sex, and disease-modifying anti-rheumatic drugs (DMARDs). Further, patients receiving DMARDs not typical for the three diagnoses were excluded, as well as patients with some additional rheumatic diseases (529 patients excluded in total). Data refers to patients with non-missing values for the respective variable; total N for patients with non-missing values is given in parentheses for variables with missing values; the total number of missing values is also given in parenthesis, for the applicable variables. (#) Includes patients with a missing glucocorticoid dosage.

bDMARD, biological disease-modifying anti-rheumatic drugs; BMI, body mass index; csDMARD, conventional synthetic disease-modifying anti-rheumatic drugs; DMARD, disease-modifying anti-rheumatic drugs; IL, interleukin; JAK, Janus kinase; N, number; NSAID, non-steroidal anti-inflammatory drugs; TNF; tumour necrosis factor; tsDMARD, targeted synthetic disease-modifying anti-rheumatic drugs.

Supplementary table 2. Baseline characteristics of the study population (total, and stratified by COVID-19 outcome)				
Parameter	Not hospitalised, no death	Hospitalised, no death	Death	Total
N	4220	736	89	5045
General				
Age [years]	48.4 (12.8)	57.3 (13.5)	67.7 (13.3)	50 (13.5)
≤ 30 years	406 (9.6)	15 (2)	0	421 (8.3)
31 – 50 years	1869 (44.3)	207 (28.1)	9 (10.1)	2085 (41.3)
51 – 65 years	1588 (37.6)	317 (43.1)	30 (33.7)	1935 (38.4)
66 - 75 years	297 (7)	129 (17.5)	22 (24.7)	448 (8.9)
> 75 years	60 (1.4)	68 (9.2)	28 (31.5)	156 (3.1)
Male sex	2130 (50.5)	419 (56.9)	57 (64)	2606 (51.7)
Ever smoker	999 (30.8) (N=3246) (Missing=974)	194 (31.0) (N=626) (Missing=110)	20 (30.8) (N=65) (Missing=24)	1213 (30.8) (N=3937) (Missing=1108)
Regions				
African Region	9 (0.2)	5 (0.7)	0	14 (0.3)
Eastern Mediterranean Region	59 (1.4)	13 (1.8)	0	72 (1.4)
European Region	3300 (78.2)	550 (74.7)	58 (65.2)	3908 (77.5)
North American Region	609 (14.4)	98 (13.3)	21 (23.6)	728 (14.4)
South American Region	221 (5.2)	36 (4.9)	10 (11.2)	267 (5.3)
South-East Asian Region	8 (0.2)	13 (1.8)	0	21 (0.4)
Western Pacific Region	14 (0.3)	21 (2.9)	0	35 (0.7)
Pandemic time period				
Until June 15, 2020	1285 (30.5)	395 (53.7)	45 (50.6)	1725 (34.2)
From June 16 to December 31, 2020	2018 (47.8)	217 (29.5)	28 (31.5)	2263 (44.9)
January 1, 2021, and later	917 (21.7)	124 (16.8)	16 (18)	1057 (21)
Immune-mediated inflammatory disease				
Axial spondyloarthritis	1576 (37.3)	229 (31.1)	26 (29.2)	1831 (36.3)
Psoriatic arthritis	1850 (43.8)	392 (53.3)	51 (57.3)	2293 (45.5)
Psoriasis (without arthritis)	794 (18.8)	115 (15.6)	12 (13.5)	921 (18.3)
Disease activity				
	N=3572 (Missing=648)	N=589 (Missing=147)	N=72 (Missing=17)	N=4233 (Missing=812)
Remission/low disease activity	2986 (83.6)	458 (77.8)	56 (77.8)	3500 (82.7)
Moderate/high disease activity	586 (16.4)	131 (22.2)	16 (22.2)	733 (17.3)
Comorbidities				
	N=4178 (Missing=42)	N=730 (Missing=6)	N=88 (Missing=1)	N=4996 (Missing=49)
Hypertension	973 (23.3)	302 (41.4)	48 (54.5)	1323 (26.5)
Cardiovascular disease	210 (5)	100 (13.7)	28 (31.8)	338 (6.8)
COPD or asthma	254 (6.1)	86 (11.8)	21 (23.9)	361 (7)
Other chronic lung disease	54 (1.3)	34 (4.7)	11 (12.5)	99 (2)
Chronic kidney disease	46 (1.1)	41 (5.6)	14 (15.9)	101 (2)
Diabetes	371 (8.9)	162 (22.2)	29 (33)	562 (11.2)
Cancer	70 (1.7)	36 (4.9)	9 (10.2)	115 (2.3)
Obesity	830 (19.9)	198 (27.1)	25 (28.4)	1053 (21.1)
Number of comorbidities	0.7 (1)	1.4 (1.4)	2.3 (1.8)	0.8 (1.1)
No comorbidity	2397 (57.4)	230 (31.5)	16 (18.2)	2643 (52.9)
One comorbidity	1004 (24)	217 (29.7)	18 (20.5)	1239 (24.8)
Two comorbidities	494 (11.8)	146 (20)	18 (20.5)	658 (13.2)
≥ 3 comorbidities	283 (6.8)	137 (18.8)	36 (40.9)	456 (9.1)
DMARDs				

(monotherapy or combination therapy)				
csDMARDs	1185 (28.1)	309 (42)	34 (38.2)	1528 (30.3)
Antimalarials	30 (0.7)	7 (1)	0	37 (0.7)
Methotrexate	915 (21.7)	246 (33.4)	22 (24.7)	1183 (23.4)
Leflunomide	82 (1.9)	22 (3)	3 (3.4)	107 (2.1)
Sulfasalazine	210 (5)	60 (8.2)	13 (14.6)	283 (5.6)
Cyclosporine	18 (0.4)	1 (0.1)	0	19 (0.4)
bDMARDs	2890 (68.5)	394 (53.5)	31 (34.8)	3315 (65.7)
TNF inhibitors	2026 (48)	248 (33.7)	24 (27)	2298 (45.6)
IL-17 inhibitors	520 (12.3)	88 (12)	2 (2.2)	610 (12.1)
IL-23/IL-12+23 inhibitors	344 (8.2)	58 (7.9)	5 (5.6)	407 (8.1)
tsDMARDs	103 (2.4)	28 (3.8)	10 (11.2)	141 (2.8)
JAK inhibitors	45 (1.1)	10 (1.4)	7 (7.9)	62 (1.2)
Apremilast	58 (1.4)	18 (2.4)	3 (3.4)	79 (1.6)
No DMARD treatment	573 (13.6)	128 (17.4)	21 (23.6)	722 (14.3)
Other therapies				
Glucocorticoids (#)	251 (6.1) (N=4116) (Missing=104)	89 (12.3) (N=722) (Missing=14)	17 (20) (N=85) (Missing=4)	357 (7.3) (N=4923) (Missing=122)
0 mg/d < Glucocorticoids <= 7.5 mg/d	165 (4) (N=4078) (Missing=142)	48 (6.9) (N=697) (Missing=39)	11 (13.3) (N=83) (Missing=6)	224 (4.6) (N=4858) (Missing=187)
Glucocorticoids > 7.5 mg/d	46 (1.1) (N=4078) (Missing=142)	16 (2.3) (N=697) (Missing=39)	4 (4.8) (N=83) (Missing=6)	66 (1.4) (N=4858) (Missing=187)
NSAIDs	948 (24.8) (N=3824) (Missing=396)	137 (20.3) (N=674) (Missing=62)	12 (15.8) (N=76) (Missing=13)	1097 (24) (N=4574) (Missing=471)
Acitretin	22 (0.5)	5 (0.7)	2 (2.2)	29 (0.6)

Data are N (column %) for categorical variables or mean (SD) for continuous variables. Data refers to patients with non-missing values for the respective variable; total N for patients with non-missing values is given in parentheses for variables with missing values; the total number of missing values is also given in parenthesis, for the applicable variables. (#) Includes patients with a missing glucocorticoid dosage.

bDMARD, biological disease-modifying anti-rheumatic drugs; BMI, body mass index; csDMARD, conventional synthetic disease-modifying anti-rheumatic drugs; DMARD, disease-modifying anti-rheumatic drugs; IL, interleukin; JAK, Janus kinase; N, number; NSAID, non-steroidal anti-inflammatory drugs; TNF; tumour necrosis factor; tsDMARD, targeted synthetic disease-modifying anti-rheumatic drugs.

Supplementary Table 3. Multivariable ordinal logistic regression analysis of factors associated with COVID-19 severity, stratified by IMID (PsA, axSpA and PsO)												
	Psoriatic arthritis				Axial spondyloarthritis				Psoriasis (without arthritis)			
N total	2293				1818				912			
N deaths / hospitalizations without death / other	51 / 392 / 1850				26 / 229 / 1563				12 / 115 / 785			
	N deaths / hospitalizations without death / neither	OR	95% CI		N deaths / hospitalizations without death / neither	OR	95% CI		N deaths / hospitalizations without death / neither	OR	95% CI	
Male sex (vs. female)	26 / 205 / 822	1.50	1.18	1.91	21 / 132 / 837	1.41	1.04	1.93	10 / 82 / 458	2.12	1.32	3.42
Pandemic time period												
Until June 15 th , 2020	24 / 207 / 513	1	[Reference]		13 / 123 / 423	1	[Reference]		8 / 65 / 338	1	[Reference]	
June 16, 2020 – December 31, 2020	16 / 117 / 910	0.41	0.31	0.55	11 / 71 / 792	0.36	0.26	0.52	1 / 29 / 308	0.52	0.31	0.88
January 1, 2021, and later	11 / 68 / 427	0.53	0.37	0.75	2 / 35 / 348	0.35	0.22	0.56	3 / 21 / 139	0.89	0.49	1.63
Comorbidities												
Hypertension alone (vs. no hypertension, no CVD)	18 / 141 / 466	1.06	0.79	1.41	8 / 68 / 255	1.49	1.01	2.19	2 / 34 / 120	1.69	0.97	2.95
CVD alone (vs. no hypertension, no CVD)	2 / 18 / 31	1.46	0.76	2.83	3 / 11 / 22	2.77	1.25	6.13	2 / 9 / 22	1.71	0.67	4.36
CVD and hypertension (vs. no hypertension, no CVD)	11 / 37 / 81	1.07	0.66	1.74	5 / 15 / 33	1.72	0.82	3.62	5 / 11 / 22	2.00	0.83	4.83
COPD or asthma	14 / 49 / 124	1.95	1.34	2.82	5 / 23 / 86	1.40	0.81	2.40	2 / 14 / 46	2.02	0.94	4.35
Other chronic lung disease	3 / 21 / 23	2.41	1.28	4.55	5 / 7 / 18	2.75	1.18	6.40	3 / 6 / 13	2.94	1.11	7.78
Chronic kidney disease	9 / 26 / 26	2.56	1.46	4.48	3 / 11 / 10	3.04	1.22	7.54	2 / 5 / 10	1.10	0.34	3.54
Obesity alone (vs. no obesity, no diabetes)	9 / 75 / 306	1.40	1.02	1.94	1 / 33 / 189	1.37	0.87	2.15	1 / 30 / 175	1.31	0.77	2.26
Diabetes mellitus alone (vs. no obesity, no diabetes)	6 / 60 / 115	1.85	1.25	2.74	5 / 23 / 60	1.54	0.88	2.70	4 / 19 / 36	3.33	1.68	6.61

Obesity and diabetes mellitus (vs. no obesity, no diabetes)	10 / 35 / 93	1.85	1.18	2.92	3 / 9 / 27	1.82	0.79	4.21	1 / 17 / 42	2.00	0.93	4.30
Cancer and known smoking habits (vs. no cancer, never smoked)	2 / 19 / 30	1.20	0.62	2.33	0 / 4 / 11	1.01	0.26	3.89	2 / 6 / 16	1.28	0.45	3.60
Cancer and unknown smoking habits (vs. no cancer, never smoked)	2 / 4 / 8	2.74	0.86	8.74	3 / 3 / 5	2.42	0.59	9.92		n.a.		
No cancer and ever smoked or unknown smoking habits (vs. no cancer, never smoked)	24 / 155 / 934	0.83	0.63	1.09	11 / 83 / 716	0.88	0.59	1.32	3 / 43 / 272	0.98		
Medication												
No DMARD therapy	8 / 49 / 177	1	[Reference]		8 / 47 / 254	1	[Reference]		5 / 32 / 141	1	[Reference]	
Antimalarials	0 / 3 / 12	0.78	0.17	3.45	0 / 1 / 2	1.96	0.13	30.73		n.a.		
Methotrexate	11 / 101 / 342	0.97	0.64	1.47	2 / 14 / 35	1.70	0.80	3.61	0 / 18 / 72	0.77	0.37	1.58
Leflunomide	2 / 16 / 36	1.24	0.60	2.59		n.a.				n.a.		
Sulfasalazine	7 / 15 / 54	1.86	0.98	3.54	5 / 17 / 82	1.25	0.66	2.37				
Cyclosporine	0 / 1 / 10	0.61	0.07	5.13		n.a.						
TNF inhibitors	11 / 104 / 780	0.47	0.32	0.71	10 / 128 / 1038	0.75	0.49	1.13	3 / 16 / 207	0.38	0.19	0.75
IL-17 inhibitors	1 / 49 / 251	0.56	0.34	0.90	1 / 21 / 142	0.71	0.38	1.33	0 / 18 / 127	0.53	0.25	1.12
IL-23/IL-12+23 inhibitors	3 / 31 / 111	0.76	0.43	1.34		n.a.			2 / 27 / 225	0.43	0.23	0.82
JAK inhibitors	7 / 9 / 35	1.65	0.80	3.42	0 / 1 / 10	0.52	0.06	4.72		n.a.		
Apremilast	1 / 14 / 45	0.91	0.43	1.91		n.a.			2 / 4 / 13	1.14	0.34	3.76
Disease activity (DA) and glucocorticoids (GCs)												
Remission/low DA, no GCs	30 / 273 / 1431	1	[Reference]		18 / 159 / 1274	1	[Reference]		9 / 83 / 64	1	[Reference]	
Remission/low DA, GCs	9 / 35 / 121	1.63	1.02	2.59	4 / 21 / 53	2.54	1.41	4.57	0 / 2 / 3	2.18	0.24	20.25
Moderate/high DA, no GCs	8 / 63 / 241	1.44	1.01	2.04	3 / 37 / 213	1.60	0.99	2.59	3 / 30 / 141	1.11	0.65	1.90

Moderate/high DA, GCs	4 / 21 / 57	2.02	1.12	3.64	2 / 11 / 23	2.85	1.25	6.52		n.a.		
NSAIDs	8 / 75 / 479	0.75	0.55	1.02	6 / 69 / 565	0.72	0.48	1.09	2 / 7 / 23	1.69	0.65	4.40

Results for ordinal mixed effects logistic regression analysis in patients with specific disease entities. Shown are fixed effects, random effects for country are not shown. Missing values are imputed via multiple imputation, patient numbers may thus be rounded. The model was additionally adjusted for age employing five-knot restricted cubic splines. Significant associations highlighted in bold.

axSpA, axial spondyloarthritis; bDMARD, biological disease-modifying anti-rheumatic drugs; BMI, body mass index; csDMARD, conventional synthetic disease-modifying anti-rheumatic drugs; DMARD, disease-modifying anti-rheumatic drugs; IL, interleukin; JAK, Janus kinase; N, number; NSAID, non-steroidal anti-inflammatory drugs; PsA, psoriatic arthritis; PsO, psoriasis without arthritis; TNF; tumour necrosis factor; tsDMARD, targeted synthetic disease-modifying anti-rheumatic drugs.

Supplementary Table 4. Sensitivity analysis – Multivariable ordinal logistic regression analysis of factors associated with COVID-19 severity (confirmed cases)				
N deaths/hospitalisations/neither	75 / 680 / 3421			
	N deaths/hospitalizations/neither	OR	95% CI	
Male sex (vs. female)	47 / 387 / 1733	1.48	1.23	1.78
Pandemic time period				
Until June 15 th , 2020	36 / 361 / 739	1	[Reference]	
June 16, 2020 – December 31, 2020	24 / 203 / 1858	0.28	0.23	0.35
January 1, 2021, and later	15 / 116 / 824	0.36	0.28	0.47
Comorbidities				
Hypertension alone (vs. no hypertension, no CVD)	21 / 228 / 678	1.32	1.05	1.66
CVD alone (vs. no hypertension, no CVD)	6 / 35 / 62	1.84	1.14	2.97
CVD and hypertension (vs. no hypertension, no CVD)	18 / 55 / 117	1.28	0.86	1.90
COPD or asthma	19 / 78 / 202	1.83	1.35	2.47
Other chronic lung disease	10 / 34 / 47	2.35	1.48	3.74
Chronic kidney disease	11 / 38 / 36	2.39	1.48	3.87
Obesity alone (vs. no obesity, no diabetes)	9 / 130 / 554	1.36	1.06	1.74
Diabetes mellitus alone (vs. no obesity, no diabetes)	14 / 97 / 171	1.91	1.40	2.60
Obesity and diabetes mellitus (vs. no obesity, no diabetes)	10 / 53 / 135	1.65	1.13	2.42
Cancer and known smoking habits (vs. no cancer, never smoked)	3 / 28 / 47	1.09	0.63	1.88
Cancer and unknown smoking habits (vs. no cancer, never smoked)	4 / 6 / 11	2.80	1.04	7.51
No cancer and ever smoked or unknown smoking habits (vs. no cancer, never smoked)	32 / 256 / 1662	0.84	0.67	1.04
Rheumatic disease				
Psoriatic arthritis	43 / 354 / 1535	1	[Reference]	
Axial spondyloarthritis	21 / 215 / 1308	1.08	0.86	1.36
Psoriasis (without arthritis)	11 / 111 / 578	0.55	0.41	0.74
Medication				
No DMARD therapy	20 / 124 / 492	1	[Reference]	
Antimalarials	0 / 4 / 13	1.09	0.29	4.05
Methotrexate	12 / 122 / 371	1.00	0.73	1.39
Leflunomide	1 / 13 / 36	0.79	0.37	1.72
Sulfasalazine	10 / 32 / 113	1.54	0.97	2.44
Cyclosporine	0 / 1 / 12	0.41	0.05	3.46
TNF inhibitors	18 / 226 / 1628	0.60	0.46	0.79
IL-17 inhibitors	1 / 80 / 412	0.67	0.47	0.96

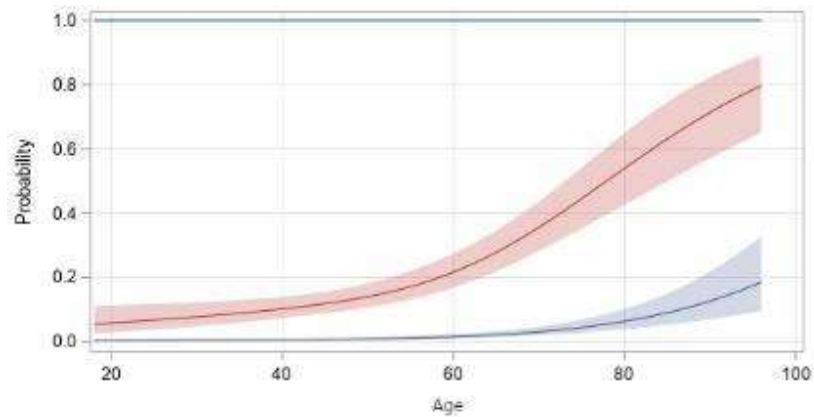
IL-23/IL-12+23 inhibitors	3 / 53 / 264	0.66	0.43	0.9999
JAK inhibitors	7 / 8 / 41	1.43	0.71	2.87
Apremilast	3 / 17 / 42	1.34	0.69	2.59
Disease activity (DA) and glucocorticoids (GCs)				
Remission/low DA, no GCs	49 / 477 / 2748	1	[Reference]	
Remission/low DA, GCs	10 / 57 / 156	1.78	1.22	2.60
Moderate/high DA, no GCs	12 / 119 / 449	1.46	1.12	1.90
Moderate/high DA, GCs	4 / 27 / 68	1.81	1.06	3.08
NSAIDs	13 / 138 / 878	0.76	0.59	0.97
<p>Results for ordinal mixed effects logistic regression analysis in patients with a confirmed COVID-19 diagnosis. Shown are fixed effects, random effects for country are not shown. Missing values are imputed via multiple imputation, patient numbers may thus be rounded. The model was additionally adjusted for age employing four-knot restricted cubic splines. Significant associations highlighted in bold.</p> <p>bDMARD, biological disease-modifying anti-rheumatic drugs; BMI, body mass index; csDMARD, conventional synthetic disease-modifying anti-rheumatic drugs; DMARD, disease-modifying anti-rheumatic drugs; IL, interleukin; JAK, Janus kinase; N, number; NSAID, non-steroidal anti-inflammatory drugs; TNF, tumour necrosis factor; tsDMARD, targeted synthetic disease-modifying anti-rheumatic drugs.</p>				

Supplementary Table 5. Sensitivity analysis – Multivariable binary logistic regression analysis of factors associated with COVID-19 severity (outcome hospitalisation)				
N hospitalizations /other	837 / 4220			
	N hospitalizations/other	OR	95% CI	
Male sex (vs. female)	479 / 2130	1.50	1.26	1.79
Pandemic time period				
Until June 15 th , 2020	442 / 1285	1	[Reference]	
June 16, 2020 – December 31, 2020	250 / 2018	0.40	0.33	0.49
January 1, 2021, and later	145 / 917	0.51	0.40	0.66
Comorbidities				
Hypertension alone (vs. no hypertension, no CVD)	272 / 847	1.31	1.05	1.62
CVD alone (vs. no hypertension, no CVD)	45 / 76	1.92	1.22	3.02
CVD and hypertension (vs. no hypertension, no CVD)	87 / 136	1.36	0.93	1.98
COPD or asthma	111 / 257	1.69	1.27	2.25
Other chronic lung disease	45 / 55	2.39	1.49	3.82
Chronic kidney disease	58 / 46	2.29	1.44	3.66
Obesity alone (vs. no obesity, no diabetes)	153 / 676	1.38	1.09	1.74
Diabetes mellitus alone (vs. no obesity, no diabetes)	117 / 212	1.92	1.43	2.58
Obesity and diabetes mellitus (vs. no obesity, no diabetes)	75 / 162	1.75	1.22	2.51
Cancer and known smoking habits (vs. no cancer, never smoked)	33 / 57	1.19	0.70	2.02
Cancer and unknown smoking habits (vs. no cancer, never smoked)	13 / 13	2.06	0.80	5.33
No cancer and ever smoked or unknown smoking habits (vs. no cancer, never smoked)	330 / 1933	0.87	0.71	1.06
Rheumatic disease				
Psoriatic arthritis	450 / 1850	1	[Reference]	
Axial spondyloarthritis	260 / 1576	1.08	0.87	1.35
Psoriasis (without arthritis)	127 / 794	0.47	0.36	0.63
Medication				
No DMARD therapy	151 / 573	1	[Reference]	
Antimalarials	4 / 14	1.20	0.33	4.39
Methotrexate	49 / 449	1.12	0.82	1.53
Leflunomide	18 / 42	1.13	0.56	2.27
Sulfasalazine	44 / 136	1.25	0.79	1.97
Cyclosporine	1 / 18	0.31	0.04	2.46
TNF inhibitors	278 / 2026	0.59	0.46	0.77
IL-17 inhibitors	91 / 520	0.69	0.49	0.96
IL-23/IL-12+23 inhibitors	63 / 344	0.74	0.50	1.10

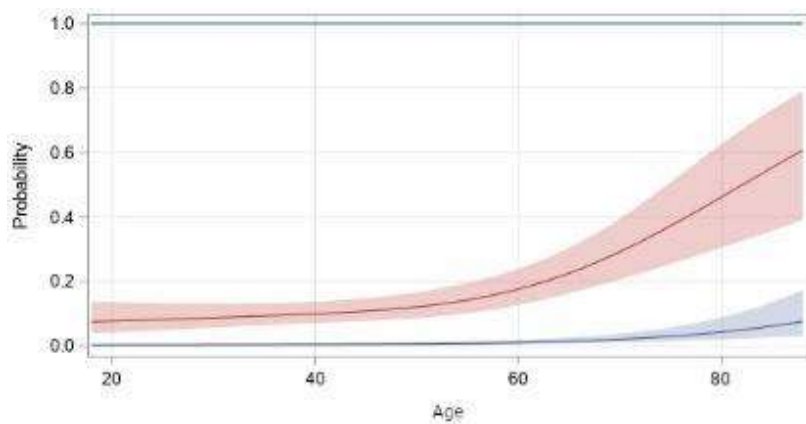
JAK inhibitors	17 / 45	1.34	0.68	2.65
Apremilast	21 / 58	1.07	0.57	2.00
Disease activity (DA) and glucocorticoids (GCs)				
Remission/low DA, no GCs	580 / 3358	1	[Reference]	
Remission/low DA, GCs	74 / 177	1.97	1.38	2.82
Moderate/high DA, no GCs	145 / 504	1.36	1.07	1.74
Moderate/high DA, GCs	38 / 82	2.34	1.43	3.81
NSAIDs	171 / 1072	0.78	0.61	0.99
<p>Results for binary mixed effects logistic regression analysis in all patients, for the outcome of hospitalization. Shown are fixed effects, random effects for country are not shown. Missing values are imputed via multiple imputation, patient numbers may thus be rounded. The model was additionally adjusted for age employing four-knot restricted cubic splines. Significant associations highlighted in bold.</p> <p>bDMARD, biological disease-modifying anti-rheumatic drugs; BMI, body mass index; csDMARD, conventional synthetic disease-modifying anti-rheumatic drugs; DMARD, disease-modifying anti-rheumatic drugs; IL, interleukin; JAK, Janus kinase; N, number; NSAID, non-steroidal anti-inflammatory drugs; TNF; tumour necrosis factor; tsDMARD, targeted synthetic disease-modifying anti-rheumatic drugs.</p>				

Supplementary Table 6. Sensitivity analysis – Multivariable binary logistic regression analysis of factors associated with COVID-19 severity (outcome death)				
N deaths / survivors	89 / 5070			
	N deaths/ survivors	OR	95% CI	
Male sex (vs. female)	57 / 2605	2.00	1.22	3.26
Pandemic time period				
Until June 15 th , 2020	45 / 1723	1	[Reference]	
June 16, 2020, and later	44 / 3347	0.67	0.41	1.10
Comorbidities				
Three or more comorbidities (vs. 0 – 2 comorbidities)	36 / 435	3.34	1.98	5.63
Smoking habits				
Ever smoked (vs. never smoked)	20/ 1209	0.76	0.41	1.39
Unknown smoking habits (vs. never smoked)	24 / 1141	1.99	0.95	4.16
Medication				
No DMARD therapy	21 / 714	1	[Reference]	
Methotrexate	13 / 596	0.57	0.26	1.24
Leflunomide	2 / 61	0.58	0.11	3.03
Sulfasalazine	12 / 171	2.64	1.13	6.17
TNF inhibitors	24 / 2360	0.50	0.26	0.98
IL-17 inhibitors	2 / 625	0.11	0.02	0.51
IL-23/IL-12+23 inhibitors	5 / 403	0.39	0.13	1.14
JAK inhibitors	7 / 58	7.49	2.61	21.47
Apremilast	3 / 82	1.05	0.25	4.40
Disease activity (DA) and glucocorticoids (GCs)				
Moderate/high DA (vs. remission/low DA)	20 / 857	1.12	0.60	2.06
GC therapy (vs. no GC therapy)	18 / 352	1.91	1.002	3.64
NSAIDs	16 / 1252	0.81	0.42	1.54
Results for binary mixed effects logistic regression analysis in all patients, for the outcome death. Shown are fixed effects, random effects for country are not shown. Missing values are imputed via multiple imputation, patient numbers may thus be rounded. The model was additionally adjusted for age employing three-knot restricted cubic splines. Significant associations highlighted in bold.				
bDMARD, biological disease-modifying anti-rheumatic drugs; BMI, body mass index; csDMARD, conventional synthetic disease-modifying anti-rheumatic drugs; DMARD, disease-modifying anti-rheumatic drugs; IL, interleukin; JAK, Janus kinase; N, number; NSAID, non-steroidal anti-inflammatory drugs; TNF; tumour necrosis factor; tsDMARD, targeted synthetic disease-modifying anti-rheumatic drugs.				

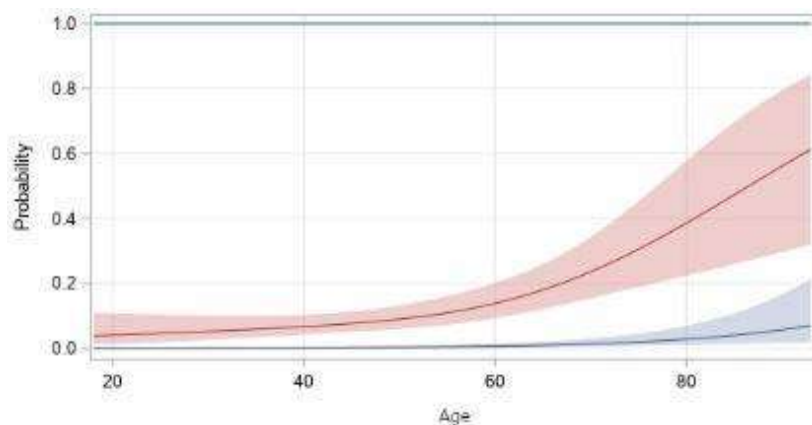
Supplementary figure 1. Relationship between age and probability of hospitalisation (red) and death (blue) in patients with psoriatic arthritis (disease-specific model), estimated by three-knot restricted cubic splines, with 95% confidence intervals.



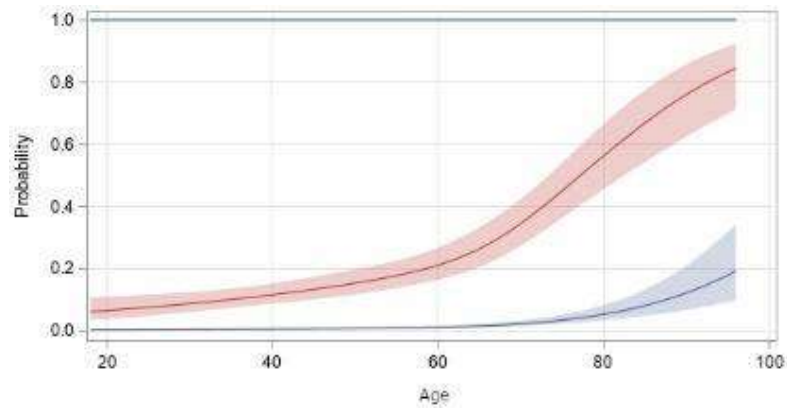
Supplementary figure 2. Relationship between age and probability of hospitalisation (red) and death (blue) in patients with axial spondyloarthritis (disease-specific model), estimated by three-knot restricted cubic splines, with 95% confidence intervals.



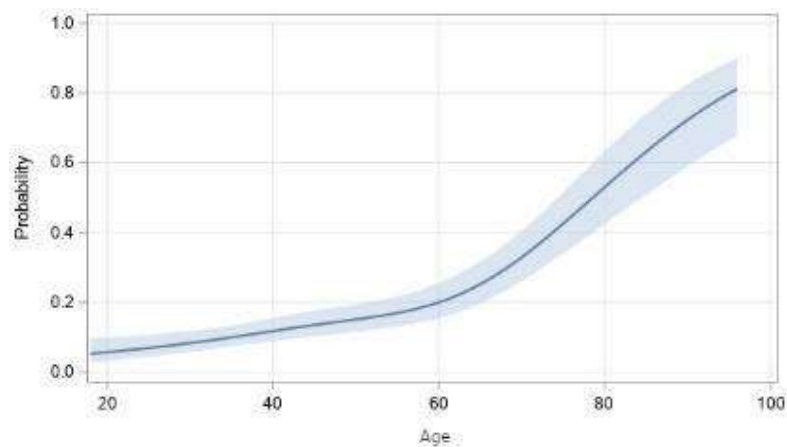
Supplementary figure 3. Relationship between age and probability of hospitalisation (red) and death (blue) in patients with psoriasis without arthritis (disease-specific model), estimated by three-knot restricted cubic splines, with 95% confidence intervals.



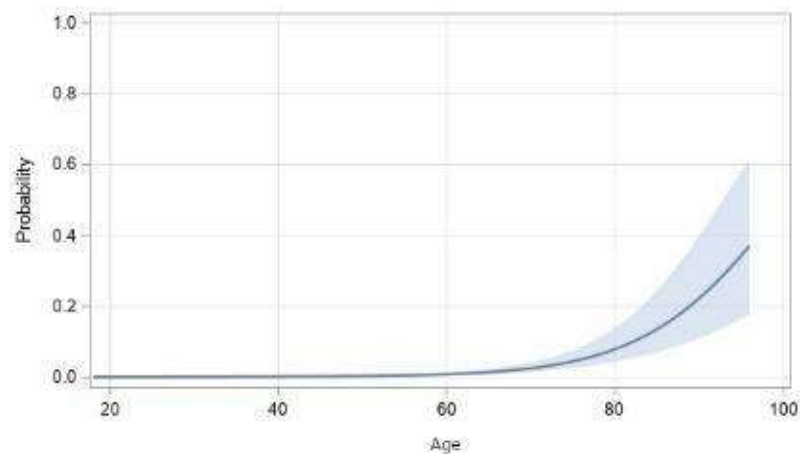
Supplementary figure 4. Relationship between age and probability of hospitalisation (red) and death (blue) estimated by four-knot restricted cubic splines, with 95% confidence intervals, for one imputation stratum (model with confirmed cases only).



Supplementary figure 5. Relationship between age and probability of hospitalisation estimated by four-knot restricted cubic splines, with 95% confidence interval (binary outcome model: hospitalisation).



Supplementary figure 6. Relationship between age and probability of death estimated by three-knot restricted cubic splines, with 95% confidence interval, for one imputation stratum (binary outcome model: death).



SUPPLEMENTARY METHODS

Choice of reference category for medication exposure

Regarding the choice of the reference category for disease-modifying antirheumatic drugs (DMARDs), the most populated categories and therefore possible choices were TNF inhibitors (TNFi), methotrexate, and absence of DMARD therapy. While the TNFi group was the largest group overall, patients receiving TNFi were younger, had less comorbidities, smoked less, took steroids less often, and had better disease control than the overall average, and therefore this group was felt to be inappropriate as a reference group. The proportion of patients receiving methotrexate, on the other hand, was much higher among psoriatic arthritis (PsA) patients compared to patients with axial spondyloarthritis (axSpA) or psoriasis without arthritis (PsO). Moreover, in spondyloarthritis patients with purely axial disease, conventional synthetic DMARDs like methotrexate are ineffective and rarely prescribed. Thus, absence of DMARD therapy was adopted as the medication reference group for DMARDs.

Assumptions of the proportional odds model

In the proportional odds model, the odds ratio for the events “hospitalized or deceased” vs. “neither hospitalized nor deceased” is assumed to be equal to the odds ratio for the events “deceased” vs. “not deceased (hospitalized or not hospitalized)”. This is the odds ratio reported as results for the model. Potential deviations from this assumption were assessed graphically by plotting the stratified means for the levels of the ordinal outcome together with the expected values given that the proportional odds assumption holds,[1] without detecting deviations of concern overall (data not shown).

Statistical interactions

Four two-way interactions were modeled in an additive sense[2]:

1. Between hypertension and cardiovascular disease (CVD): hypertension alone, CVD alone, hypertension combined with CVD; vs. no hypertension and no CVD.
2. Between obesity and diabetes: obesity alone, diabetes alone, obesity combined with diabetes; vs. no obesity and no diabetes.
3. Between smoking status and cancer: cancer and known smoking habits, cancer unknown smoking habits, no cancer and ever smoked or unknown smoking habits; vs. no cancer and never smoked. The interaction between smoking status and cancer was modelled in a non-standard way due to the group of patients

with missing information on smoking habits presenting an OR exceeding the ORs for patients that had never or ever smoked in preliminary analyses, suggesting it was more appropriate to consider it as a separate subgroup within the interaction.

4. Between disease activity and prednisolone-equivalent glucocorticoid (GC) use: remission/low disease activity and GC use, moderate/high disease activity and no GC use, moderate/high disease activity and GC use; vs. remission/low disease activity and no GC use.

Patients excluded and handling of missing data

Patients under the age of 18, patients with missing primary outcome as well as missing values for age, sex, pandemic time period, DMARD exposure, patients treated with more than one biological DMARD, patients treated with DMARDs not typically used or licensed for PsO, PsA or axSpA, PsA (abatacept, B-cell inhibitors, IL-1 and IL-6 inhibitors), and patients diagnosed with multiple inflammatory rheumatic diseases (except for Sjögren's syndrome, and if not receiving B-cell therapies, azathioprine, mycophenolate, ciclosporin, cyclophosphamide or tacrolimus) were excluded from the analysis (529 patients excluded in total).

Further, for some analyses, medications without patients at all or medications with all patients falling into the same outcome category had to be excluded for statistical reasons in some of the secondary and sensitivity analyses (leflunomide, cyclosporine, IL-23 inhibitors, and apremilast, for the analyses focusing on patients with axSpA; antimalarials, leflunomide, sulfasalazine, cyclosporine, JAK inhibitors, and glucocorticoids in the presence of moderate/high disease activity, for the analysis focusing on patients with PsO; antimalarials and cyclosporine, for the analysis using the binary outcome mortality). Missing values for comorbidities, glucocorticoid therapy, disease activity and NSAIDs were derived by multiple imputation using full conditional specification.[3]

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Fedra Irazoque	Private Practice
Genevieve Katigbak	Private Practice
Gilbert Kepecs	Private Practice
Greta Reyes-Cordero	Private Practice
Jaimie Russell	Private Practice
Jessica Chapman	Private Practice
Joseph Huffstutter	Private Practice
Kara Long	Private Practice
Khurram Abbass	Private Practice
Lauren Kaufman	Private Practice
Marco Martinez Martinez	Private Practice

MONICA MACIAS-PALACIOS	Private Practice
NAGA PRABU	Private Practice
Paloma De Abreu Trigueros	Private Practice
Raheem Kherani	Private Practice
Randall Beyl	Private Practice
Sarah Middleton	Private Practice
Sharath Kumar	Private Practice
Susan Barrett	Private Practice
Viktoria Pavlova	Private Practice
Yanira Yinde	Private Practice
Samir Patel	Queen Elizabeth Hospital Woolwich
Alexandra Balbir-Gurman	Rambam Rheumatology Institute, Haifa
Bruno Ferreira	Rede Sarah de Hospitais de Rabilitação
Kathleen Anthony	Rheumatology and Osteoporosis Specialists
Richard Stern	Rheumatology Associastes
Lilliam Miranda	Rheumatology Center INC
Shabbir Chikani	Rheumatology centre, HCG hospital
Ann Clarke	Richmond Road Diagnostic And Treatment Center
Ammar Haikal	Riverside Medical Group
Michael Guma	Riverside Medical Group
Sushama Mody	Riverside Medical Group
Diana O'Kane	RNHRD at Royal United Hospital Bath
Sheila O'Reilly	Royal Derby Hospital
HUMAID AL WAHSHI	Royal Hospital
Nasra Al-Adhoubi	Royal Hospital
Jenny Tyler	Royal United hospital
Jennifer Tyler	Royal United Hospital, Bath
Vivien Hsu	Rutgers Robert Wood Johnson
Mária Oetterová	Safarik University hospital, Kosice
Audrey Low	Salford Royal NHS FT
Beverley Harrison	Salford Royal NHS FT

Guillermo Pons-Estel	Sanatorio Parque
Mariana Peixoto Guimaraes Ubirajara e Silva de Souza	Santa Casa of Belo Horizonte
Carolina Zorzanelli Costa	Santa Casa of Vitoria
Veronika Sharp	Santa Clara Valley Medical Center
Laura Groseanu	Sf Maria Clinical Hospital, Bucharest
Daric Mueller	Shores Rheumatology PC
Lillian Barra	SJHC London
Tom Appleton	SJHC London
South African Rheumatism and Arthritis Association	South Africa
Laura Chadwick	St Helens & Knowsley NHS Foundation Trust
Richard Conway	St James' Hospital, Dublin
Janet Pope	St. Joseph's Health Care
Maria del Carmen Hernandez	Star Medica Centro
Cecília Resende Brunow Bazzo	State University of Londrina
Steven Feldman	Steven Feldman MD PA
Anna Sabová	Súkromná Reumatologická Ambulancia, Vranov nad Topľou
Talal Ali Al Lawati	Sultan Qaboos University Hospital
Rachael Flood	Tallaght University Hospital
MOOIKHIN HNG	Tareum
Arundathi Jayatilleke	Temple University Hospital
Simeon Grazio	The teaching hospital sisters of charity
Keiko Koshiba	Toho University Omori Medical Center
Masashi Funada	Toho University Omori Medical Center
Midori Sato	Tokyo Medical Center
Fukumi NAKAMURA	Tokyo Metropolitan Bokutoh General Hospital
Kazuhisa Yokota	Tokyo Metropolitan Health and Hospitals Corporation, Ebara Hosipital
Lingli Dong	Tongji Hospital
A. Silvia Ross	Triangle Arthritis & Rheumatology
John FitzGerald	UCLA
Tanaz Kermani	UCLA
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Alojzija Hocevar	UMC Ljubljana
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Su-Ann Yeoh	University College London Hospital
Maria Oetterova	University Hospital
Nicola Gullick	University Hospital
Emőke Šteňová	University Hospital Bratislava
Marko Barešić	University Hospital Center Zagreb
Ivan Padjen	University Hospital Centre Zagreb
Melanie-Ivana Čulo	University Hospital Dubrava, Zagreb
Wilson Bautista Molano	University Hospital Fundación Santa Fe de Bogotá
James Pilcher	University Hospital Lewisham
Kristina Kovačević Stranski	University Hospital Osijek
Lubica Capova	University Hospital, Bratislava
Zelmira Macejova	University Hospital, Košice
Jarosław Nowakowski	University Hospital, Krakow
Takashi Kida	University Hospital, Kyoto Prefectural University of Medicine
SAUDATU ISSAKA	University Hospitals
Licia Maria Henrique da Mota	University of Bras
JoAnn Zell	University of Colorado
Christine Peschken	University of Manitoba
Angelito Flora	University of the Philippines, Philippine General Hospital
Evelyn Salido	University of the Philippines, Philippine General Hospital
Geraldine Zamora	University of the Philippines, Philippine General Hospital
Alison Bays	University of Washington, Seattle
David Karp	UT Southwestern Medical Center
Ezzati Fatemeh	UT Southwestern Medical Center
Guillermo Quiceno	UT Southwestern Medical Center
Kathryn Dao	UT Southwestern Medical Center
APARNA DAS	UTMB

Lauro Quintanilla

Vicki Quincey

Deborah Parks

Kelly Weselman

Fabiane Shizue Sakai

Katie Williams

Kirsty Devine

Lenny Geurts-van Bon

Sarah Goglin

Virginia Medica

Waikato Hospital

Washington University Div of Rheumatology

Wellstar Kennestone Hospital

Windmills Hospital

York District Hospital

York/Scarborough Hospitals

Ziekenhuisgroep Twente

Zuckerberg San Francisco General Hospital

Appendix 2. Psoriasis Patient Registry for Outcomes, Therapy and Epidemiology of COVID-19 Infection (PsoProtect) Consortium.

Name

A van geest
Aadarsh Shah
AC de Waal
Alba Catala
Alberto Barea
Alberto Romero Maté
Alekya Singapore
Alexandra Paolino
Alexandra Vincent
Alice Mwale
Alison Sears
Amy de la Breteque
Amy Foulkes
Ana Brasileiro
Ana Maria Morales Callaghan
Ana Martinez
Andrea Carugno
Andrea Chiricozzi
Andrea Conti
Andrew DeCrescenzo
Andrew Pink
Angela Braeger
Anke Piekar
Ann Jones
Ann Sergeant
Anna Baran

Anne Groeneveld
Annette Essex
Anthony Bewley
Antoine Fauconneau
Antony Raharja
Aparna Sinha
Areti Makrygeorgou
Arias Martín Nadia
Astrid van Huizen
Beata Fabos
Beatriz Pérez Suárez
Benhadou Farida
Birgitta Wilson Claréus
Bola Coker
Canelle Mazaud
Caoimhe Fahy
Caoimhe M. R. Fahy
Carla Tubau Prims
Carmen Bugarin Diz
Caroline Campbell
Carolyn Martin
Carrie Davis
Catherine Holden
Catherine Motosko
Catherine Quinlan
Catriona Maybury
Cesar Gonzalez
Charlotte Barclay
Chifari Angelo
Choon Siew Eng

Cid Yazigi Sabbag
Claudia de la Cruz
Claudia Guebenlian Bakerdjian
Claudio Greco
Cristina Echeverria
Cristina Mariela Echeverría
Dagmara Samselska
Danang Tri Wahyudi
Daniela Armijo
Danielle Brassard
Daryl Teo
David Fairhurst
Deanna Cummings
Deepti Kolli
Denis Jullien
Denise Peeters
Descamps Vincent
Diana Patricia Ruiz Genao
Diana Ruiz Genao
Eileen Parry
Elaine Agius
Eleanor Henderson
Elena Hawryluk
Eliseo Martínez-García
Elizabeth Molina
Elizabeth Stewart
Ellie Henderson
Elzbieta Klujso
Emily Brown
Emily Dwyer

Emmanuel Mahe
Emmanuel Toni
Emmerson Gale Vista
Emmylou Casanova
Enikö Sonkoly
Enrique Loayza
Erin Kamp
Esteban Daudén
Esther Balogh
F Aubin
Felicity Edwards
Ferial Ismail
Fernando Valenzuela
Fikki Orekoya
Florentina-Silvia Delli
Francesca Capon
Freya Meynell
Gabriel Magariños
Gabrielle Becher
Gabrielle Key
Gaelle Quereux
Gaurav Dhawan
Gemma Keogh
Georgi Popov
Georgie King
Girard Celine
Gloria Aparicio
Gordana Krnjevic Pezic
Graham Johnston
Gustavo Anibal Cardozo

H. El Khattabi
Haleema Alfailakawi
Hazel Oon
Hazel Rooney
Helen McAteer
Helene Aubert
Hervé Bachelez
Hoseah Waweru
Ian Pearson
Ignacio Yanguas
Iman Kotb
Ines Barbosa
Isabella Tosi
J Charles
Jack French
James John Murphy
Jamie Weisman
Jennifer Elias
Jennifer Soung
Jenny Carolan
Jenny Hughes
Jill Ramsay
Jing Husaini
JM. Ortiz
Jo Lambert
Joanne Topliffe
Joel M Gelfand
John Newsham
Jonathan Ng
Jose-Manuel Carrascosa

Joseph J Schwartz
Julia Bowman
Julie Charles
Juul van de Reek
Karina Jackson
Karolina Vorčáková
Katarzyna Gryś
Katherine Poirier
Kathryn Kerisit, MD
Kayleigh J Mason
Keith Wu
Kimberly Anne G. Ednalino
Kirsty Wynes
Kyle Eash
Lars Iversen
Laura Speck
Laure Mery-Bossard
Lauren Booker
Leah Hoggan
leandro perrotat
Leila Asfour
Leila Kattach
Leontien de Graaf
Lesort Cécile
Lian van der Gang
Lieve Meuleman
Lim Jin Huang
Linda Lawson
Linda McMahan
Line Kibsgaard

Lisa Kirby
Lisa van der Rijst
Liv Eidsmo
Lluís Rusiñol Batlle
Lone Skov
Lorraine Gribben
Lucy Moorhead
Luigi Naldi
Luis Puig
Lyndsey Florence
M.M. Bakker
Mahira Hamdy El Sayed
Mahmood Abubakar
Malcolm Rustin
Manel Velasco
Mangiarotti Germán
Manisha Panchal
Manja Bloem
Manpreet Lakhan
Manuel Dario Franco
Mara Maccarone
Margot Common
Maria Fernanda Lui
Marie Dedroog
Marie-Eve Fortier
Marie-Louise Svensson
Mariusz Sikora
Mark Vandaele
Maruska Marovt
Masanori Okuse

Matthias Schmuth
Melanie Bruton
Melanie Claridge
Melanie westmoreland
Melissa Sweeney
Michela Magnano
Mireille van Baar
Miriam Saposnik
Mohamed EL-Komy
Musumeci Maria Letizia
N Beneton
Nick J Reynolds
Nick Reynolds
Nora Noemí Kogan
Omid Zargari
Pablo De Caso
Pamela Campbell
Paola Di Meglio
Paolo Gisondi
Paula Bourren
Paula luna
Paulo Varela
Penny Nash
Peter Foley
Peter Holló
Peter Jenkin
Phan Céline
Philip Hampton
Phyllis Spuls
Pia Tookey

Piergiacomo Calzavara-Pinton
Portia Goldsmith
Pter Holló
Rachel Bak
Radhika Patel
Raquel Rivera Diaz
Rebecca Rose
Reinhart Speeckaert
Ricardo Romiti
Richard Warren
Richard Woolf
Rogelio Mercado
Rohima Khatun
Rolland Gyulai
Romana Ceovic
Romana Machackova
Ronald Vender
Rosa Andres Ejarque
Rosa Taberner
Rosalie Juch
Russell cohen md faad
Sandy McBride
Sara Cacciapuoti
Sarah Drummond
Sarah Kirk
Sarah McCusker
Saskia Reeken
Selva Anahí Gutiérrez Yañez
Shanti Ayob
Shrita Shinton

Silvia Pérez Barrio
Simina Stefanescu
Sinan Doğan
Sinéad M Langan
Sophia Mohme
Sophia Strong-Sheldrake
Sophie Wanten
Stefano Piaserico
Stephanie Ball
Stephanie Ogden
Susan Ann Lloyd
Susan Hall
Susan Woollett
Susana Armesto Alonso
Susannah Hoey
Taku Suzuki
Tatiana del Río
Tatiana Pecova
Teena Mackenzie
Telgdy Enikő
Teresa Tsakok
Thomas Beaulieu
Tiago Torres
Ting Seng Tang
Tom Hillary
Tomas Kampe
Toomas Talme
Tracy Brown
Tracy Smith
Tran Hong Truong

Trupti Desai
Victoria Brown
Victoria King
Vito Di Lernia
Wisam Alwan
Ya-Hsin Wang
Yena Kim
Zafeiriou Efterpi
zahira koreja
Zaida Troyano
Zeeshaan Hasan
Zeljko Mijuskovic
Zenas Yiu
Zeynep Topkarci

SUPPLEMENTARY TABLES AND FIGURES

Supplementary table 1. Baseline characteristics of the study population (total, and stratified by medication group)							
Parameter	No DMARDs	csDMARDs only	TNFi	IL-17i/IL-23i/IL12+23i	JAKi	APREMILAST	TOTAL
N	722	867	2298	1017	62	79	5045
General							
Age [years]	49.9 (15.2)	54 (13.4)	48.3 (13.0)	49.9 (12.8)	54.3 (12.5)	56.2 (11.9)	50 (13.5)
≤ 30 years	81 (11.2)	36 (4.2)	226 (9.8)	72 (7.1)	3 (4.8)	3 (3.8)	421 (8.3)
31 – 50 years	281 (38.9)	291 (33.6)	1044 (45.4)	433 (42.6)	19 (30.6)	17 (21.5)	2085 (41.3)
51 – 65 years	252 (34.9)	377 (43.5)	836 (36.4)	402 (39.5)	24 (38.7)	44 (55.7)	1935 (38.4)
66 - 75 years	70 (9.7)	111 (12.8)	152 (6.6)	88 (8.7)	16 (25.8)	11 (13.9)	448 (8.9)
> 75 years	38 (5.3)	52 (6)	40 (1.7)	22 (2.2)	0	4 (5.1)	156 (3.1)
Male sex	357 (49.4)	385 (44.4)	1267 (55.1)	546 (53.7)	20 (32.3)	31 (39.2)	2606 (51.7)
Ever smoker	159 (29.6) (N=538) (NA=184)	196 (31.2) (N=628) (NA=239)	517 (29.1) (N=1777) (NA=521)	308 (35.0) (N=879) (NA=138)	9 (19.6) (N=46) (NA=16)	24 (34.8) (N=69) (NA=10)	1213 (30.8) (N=3937) (NA=1108)
Regions							
African Region	4 (0.6)	6 (0.7)	3 (0.1)	1 (0.1)	0	0	14 (0.3)
Eastern Mediterr. Region	19 (2.6)	17 (2)	27 (1.2)	7 (0.7)	2 (3.2)	0	72 (1.4)
European Region	550 (76.2)	679 (78.3)	1769 (77)	822 (80.8)	35 (56.5)	53 (67.1)	3908 (77.5)
North American Region	102 (14.1)	106 (12.2)	343 (14.9)	128 (12.6)	24 (38.7)	25 (31.6)	728 (14.4)
South American Region	29 (4.0)	38 (4.4)	146 (6.4)	53 (5.2)	1 (1.6)	0	267 (5.3)
South-East Asian Region	5 (0.7)	11 (1.3)	3 (0.1)	1 (0.1)	0	1 (1.3)	21 (0.4)
Western Pacific Region	13 (1.8)	10 (1.2)	7 (0.3)	5 (0.5)	0	0	35 (0.7)
Time period							
Until June 15, 2020	253 (35.0)	304 (35.1)	756 (32.9)	357 (35.1)	20 (32.3)	35 (44.3)	1725 (34.2)
June 16 to Dec 31, 2020	297 (41.1)	367 (42.3)	1073 (46.7)	469 (46.1)	27 (43.5)	30 (38)	2263 (44.9)
January 1, 2021 or later	172 (23.8)	196 (22.6)	469 (20.4)	191 (18.8)	15 (24.2)	14 (17.7)	1057 (21)
Immune-mediated inflammatory diseases							
SpA	309 (42.8)	164 (18.9)	1176 (51.2)	171 (16.8)	11 (17.7)	0	1831 (36.3)
PsA	234 (32.4)	607 (70.0)	895 (38.9)	446 (43.9)	51 (82.3)	60 (75.9)	2293 (45.5)
Psoriasis (no arthritis)	179 (24.8)	96 (11.1)	227 (9.9)	400 (39.3)	0	19 (24.1)	921 (18.3)
Disease activity							
	N=606 (NA=116)	N=762 (NA=105)	N=1821 (NA=477)	N=918 (NA=99)	N=49 (NA=13)	N=77 (NA=2)	N=4233 (NA=812)
Remission, min./low DA	432 (71.3)	625 (82)	1567 (86.1)	782 (85.2)	32 (65.3)	62 (80.5)	3500 (82.7)
Moderate, sev./high DA	174 (28.7)	137 (18)	254 (13.9)	136 (14.8)	17 (34.7)	15 (19.5)	733 (17.3)
Ordinal outcome							
Not hospitalized/dead	573 (79.4)	654 (75.4)	2026 (88.2)	864 (85)	45 (72.6)	58 (73.4)	4220 (83.7)
Hospitalized, no death	128 (17.7)	186 (21.5)	248 (10.8)	146 (14.4)	10 (16.1)	18 (22.8)	736 (14.6)
Death	21 (2.9)	27 (3.1)	24 (1)	7 (0.7)	7 (11.3)	3 (3.8)	89 (1.8)
Comorbidities							
	N=714 (NA=8)	N=854 (NA=13)	N=2281 (NA=17)	N=1008 (NA=9)	N=62 (NA=0)	N=77 (NA=2)	N=4996 (NA=49)

Hypertension	183 (25.6)	263 (30.8)	555 (24.3)	285 (28.3)	13 (21.0)	24 (31.2)	1323 (26.5)
Cardiovascular disease	56 (7.8)	84 (9.8)	109 (4.8)	81 (8.0)	0	8 (10.4)	338 (6.8)
COPD or asthma	52 (7.3)	58 (6.8)	130 (5.7)	102 (10.1)	12 (19.4)	7 (9.1)	361 (7.2)
Other lung disease	15 (2.1)	13 (1.5)	37 (1.6)	27 (2.7)	4 (6.5)	3 (3.9)	99 (2.0)
Chronic kidney disease	14 (2.0)	23 (2.7)	36 (1.6)	23 (2.3)	1 (1.6)	4 (5.2)	101 (2.0)
Obesity (BMI >= 30)	137 (19.2)	171 (20.0)	426 (18.7)	289 (28.7)	12 (19.4)	18 (23.4)	562 (11.2)
Diabetes	81 (11.3)	104 (12.2)	227 (10.0)	133 (13.2)	7 (11.3)	10 (13.0)	115 (2.3)
Cancer	31 (4.3)	24 (2.8)	20 (0.9)	28 (2.8)	1 (1.6)	11 (14.3)	1053 (21.1)
number of comorbidities	0.8 (1.1)	0.9 (1.2)	0.7 (1.0)	1 (1.3)	0.8 (1.0)	1.3 (1.4)	0.8 (1.1)
No comorbidity	371 (52.0)	436 (51.1)	1313 (57.6)	461 (45.7)	33 (53.2)	29 (37.7)	2643 (52.9)
One comorbidity	187 (26.2)	224 (26.2)	524 (23)	271 (26.9)	14 (22.6)	19 (24.7)	1239 (24.8)
Two comorbidities	98 (13.7)	107 (12.5)	281 (12.3)	144 (14.3)	12 (19.4)	16 (20.8)	658 (13.2)
≥ 3 comorbidities	58 (8.1)	87 (10.2)	163 (7.1)	132 (13.1)	3 (4.8)	13 (16.9)	456 (9.1)
csDMARD therapy							
csDMARDs	0	867 (100)	487 (21.2)	138 (13.6)	22 (35.5)	14 (17.7)	1528 (30.3)
Methotrexate	0	649 (74.9)	388 (16.9)	119 (11.7)	18 (29.0)	9 (11.4)	1183 (23.4)
Leflunomide	0	64 (7.4)	30 (1.3)	7 (0.7)	3 (4.8)	3 (3.8)	107 (2.1)
Sulfasalazine	0	181 (20.9)	86 (3.7)	12 (1.2)	3 (4.8)	1 (1.3)	283 (5.6)
Antimalarials	0	27 (3.1)	8 (0.4)	1 (0.1)	0	1 (1.3)	37 (0.7)
Cyclosporine	0	14 (1.6)	2 (0.1)	3 (0.3)	0	0	19 (0.4)
Further therapies							
Glucocorticoids (#)	52 (7.4) (N=698) (NA=24)	112 (13.1) (N=858) (NA=9)	121 (5.4) (N=2229) (NA=69)	57 (5.7) (N=1004) (NA=13)	12 (20.7) (N=58) (NA=4)	3 (3.9) (N=76) (NA=3)	357 (7.3) (N=4923) (NA=122)
0 mg/d < GCs <= 7.5 mg/d	31 (4.5) (N=691) (NA=31)	73 (8.6) (N=847) (NA=20)	75 (3.4) (N=2198) (NA=100)	38 (3.8) (N=991) (NA=26)	5 (9.1) (N=55) (NA=7)	2 (2.6) (N=76) (NA=3)	224 (4.6) (N=4858) (NA=187)
GCs > 7.5 mg/d	14 (2.0) (N=691) (NA=31)	27 (3.2) (N=847) (NA=20)	14 (0.6) (N=2198) (NA=100)	6 (0.6) (N=991) (NA=26)	4 (7.3) (N=55) (NA=7)	1 (1.3) (N=76) (NA=3)	66 (1.4) (N=4858) (NA=187)
NSAIDs	214 (31.9) (N=671) (NA=51)	199 (24.8) (N=802) (NA=65)	480 (23.9) (N=2012) (NA=286)	175 (18.2) (N=961) (NA=56)	14 (25.0) (N=56) (NA=6)	15 (20.8) (N=72) (NA=7)	1097 (24.0) (N=4574) (NA=471)
Acitretin	25 (3.5)	1 (0.1)	2 (0.1)	0	1 (1.6)	0	29 (0.6)

Data are N (column %) for categorical variables or mean (SD) for continuous variables. Table includes patients diagnosed with psoriasis, psoriatic arthritis or axial spondyloarthritis, with a non-missing ordinal outcome and non-missing values for age, sex, and disease-modifying anti-rheumatic drugs (DMARDs). Further, patients receiving DMARDs not typical for the three diagnoses were excluded, as well as patients with some additional rheumatic diseases (529 patients excluded in total). Data refers to patients with non-missing values for the respective variable; total N for patients with non-missing values is given in parentheses for variables with missing values; the total number of missing values is also given in parenthesis, for the applicable variables. (#) Includes patients with a missing glucocorticoid dosage.

bDMARD, biological disease-modifying anti-rheumatic drugs; BMI, body mass index; csDMARD, conventional synthetic disease-modifying anti-rheumatic drugs; DMARD, disease-modifying anti-rheumatic drugs; IL, interleukin; JAK, Janus kinase; N, number; NSAID, non-steroidal anti-inflammatory drugs; TNF; tumour necrosis factor; tsDMARD, targeted synthetic disease-modifying anti-rheumatic drugs.

Supplementary table 2. Baseline characteristics of the study population (total, and stratified by COVID-19 outcome)				
Parameter	Not hospitalised, no death	Hospitalised, no death	Death	Total
N	4220	736	89	5045
General				
Age [years]	48.4 (12.8)	57.3 (13.5)	67.7 (13.3)	50 (13.5)
≤ 30 years	406 (9.6)	15 (2)	0	421 (8.3)
31 – 50 years	1869 (44.3)	207 (28.1)	9 (10.1)	2085 (41.3)
51 – 65 years	1588 (37.6)	317 (43.1)	30 (33.7)	1935 (38.4)
66 - 75 years	297 (7)	129 (17.5)	22 (24.7)	448 (8.9)
> 75 years	60 (1.4)	68 (9.2)	28 (31.5)	156 (3.1)
Male sex	2130 (50.5)	419 (56.9)	57 (64)	2606 (51.7)
Ever smoker	999 (30.8) (N=3246) (Missing=974)	194 (31.0) (N=626) (Missing=110)	20 (30.8) (N=65) (Missing=24)	1213 (30.8) (N=3937) (Missing=1108)
Regions				
African Region	9 (0.2)	5 (0.7)	0	14 (0.3)
Eastern Mediterranean Region	59 (1.4)	13 (1.8)	0	72 (1.4)
European Region	3300 (78.2)	550 (74.7)	58 (65.2)	3908 (77.5)
North American Region	609 (14.4)	98 (13.3)	21 (23.6)	728 (14.4)
South American Region	221 (5.2)	36 (4.9)	10 (11.2)	267 (5.3)
South-East Asian Region	8 (0.2)	13 (1.8)	0	21 (0.4)
Western Pacific Region	14 (0.3)	21 (2.9)	0	35 (0.7)
Pandemic time period				
Until June 15, 2020	1285 (30.5)	395 (53.7)	45 (50.6)	1725 (34.2)
From June 16 to December 31, 2020	2018 (47.8)	217 (29.5)	28 (31.5)	2263 (44.9)
January 1, 2021, and later	917 (21.7)	124 (16.8)	16 (18)	1057 (21)
Immune-mediated inflammatory disease				
Axial spondyloarthritis	1576 (37.3)	229 (31.1)	26 (29.2)	1831 (36.3)
Psoriatic arthritis	1850 (43.8)	392 (53.3)	51 (57.3)	2293 (45.5)
Psoriasis (without arthritis)	794 (18.8)	115 (15.6)	12 (13.5)	921 (18.3)
Disease activity				
	N=3572 (Missing=648)	N=589 (Missing=147)	N=72 (Missing=17)	N=4233 (Missing=812)
Remission/low disease activity	2986 (83.6)	458 (77.8)	56 (77.8)	3500 (82.7)
Moderate/high disease activity	586 (16.4)	131 (22.2)	16 (22.2)	733 (17.3)
Comorbidities				
	N=4178 (Missing=42)	N=730 (Missing=6)	N=88 (Missing=1)	N=4996 (Missing=49)
Hypertension	973 (23.3)	302 (41.4)	48 (54.5)	1323 (26.5)
Cardiovascular disease	210 (5)	100 (13.7)	28 (31.8)	338 (6.8)
COPD or asthma	254 (6.1)	86 (11.8)	21 (23.9)	361 (7)
Other chronic lung disease	54 (1.3)	34 (4.7)	11 (12.5)	99 (2)
Chronic kidney disease	46 (1.1)	41 (5.6)	14 (15.9)	101 (2)
Diabetes	371 (8.9)	162 (22.2)	29 (33)	562 (11.2)
Cancer	70 (1.7)	36 (4.9)	9 (10.2)	115 (2.3)
Obesity	830 (19.9)	198 (27.1)	25 (28.4)	1053 (21.1)
Number of comorbidities	0.7 (1)	1.4 (1.4)	2.3 (1.8)	0.8 (1.1)
No comorbidity	2397 (57.4)	230 (31.5)	16 (18.2)	2643 (52.9)
One comorbidity	1004 (24)	217 (29.7)	18 (20.5)	1239 (24.8)
Two comorbidities	494 (11.8)	146 (20)	18 (20.5)	658 (13.2)
≥ 3 comorbidities	283 (6.8)	137 (18.8)	36 (40.9)	456 (9.1)
DMARDs				

(monotherapy or combination therapy)				
csDMARDs	1185 (28.1)	309 (42)	34 (38.2)	1528 (30.3)
Antimalarials	30 (0.7)	7 (1)	0	37 (0.7)
Methotrexate	915 (21.7)	246 (33.4)	22 (24.7)	1183 (23.4)
Leflunomide	82 (1.9)	22 (3)	3 (3.4)	107 (2.1)
Sulfasalazine	210 (5)	60 (8.2)	13 (14.6)	283 (5.6)
Cyclosporine	18 (0.4)	1 (0.1)	0	19 (0.4)
bDMARDs	2890 (68.5)	394 (53.5)	31 (34.8)	3315 (65.7)
TNF inhibitors	2026 (48)	248 (33.7)	24 (27)	2298 (45.6)
IL-17 inhibitors	520 (12.3)	88 (12)	2 (2.2)	610 (12.1)
IL-23/IL-12+23 inhibitors	344 (8.2)	58 (7.9)	5 (5.6)	407 (8.1)
tsDMARDs	103 (2.4)	28 (3.8)	10 (11.2)	141 (2.8)
JAK inhibitors	45 (1.1)	10 (1.4)	7 (7.9)	62 (1.2)
Apremilast	58 (1.4)	18 (2.4)	3 (3.4)	79 (1.6)
No DMARD treatment	573 (13.6)	128 (17.4)	21 (23.6)	722 (14.3)
Other therapies				
Glucocorticoids (#)	251 (6.1) (N=4116) (Missing=104)	89 (12.3) (N=722) (Missing=14)	17 (20) (N=85) (Missing=4)	357 (7.3) (N=4923) (Missing=122)
0 mg/d < Glucocorticoids <= 7.5 mg/d	165 (4) (N=4078) (Missing=142)	48 (6.9) (N=697) (Missing=39)	11 (13.3) (N=83) (Missing=6)	224 (4.6) (N=4858) (Missing=187)
Glucocorticoids > 7.5 mg/d	46 (1.1) (N=4078) (Missing=142)	16 (2.3) (N=697) (Missing=39)	4 (4.8) (N=83) (Missing=6)	66 (1.4) (N=4858) (Missing=187)
NSAIDs	948 (24.8) (N=3824) (Missing=396)	137 (20.3) (N=674) (Missing=62)	12 (15.8) (N=76) (Missing=13)	1097 (24) (N=4574) (Missing=471)
Acitretin	22 (0.5)	5 (0.7)	2 (2.2)	29 (0.6)

Data are N (column %) for categorical variables or mean (SD) for continuous variables. Data refers to patients with non-missing values for the respective variable; total N for patients with non-missing values is given in parentheses for variables with missing values; the total number of missing values is also given in parenthesis, for the applicable variables. (#) Includes patients with a missing glucocorticoid dosage.

bDMARD, biological disease-modifying anti-rheumatic drugs; BMI, body mass index; csDMARD, conventional synthetic disease-modifying anti-rheumatic drugs; DMARD, disease-modifying anti-rheumatic drugs; IL, interleukin; JAK, Janus kinase; N, number; NSAID, non-steroidal anti-inflammatory drugs; TNF; tumour necrosis factor; tsDMARD, targeted synthetic disease-modifying anti-rheumatic drugs.

Supplementary Table 3. Multivariable ordinal logistic regression analysis of factors associated with COVID-19 severity, stratified by IMID (PsA, axSpA and PsO)												
	Psoriatic arthritis				Axial spondyloarthritis				Psoriasis (without arthritis)			
N total	2293				1818				912			
N deaths / hospitalizations without death / other	51 / 392 / 1850				26 / 229 / 1563				12 / 115 / 785			
	N deaths / hospitalizations without death / neither	OR	95% CI		N deaths / hospitalizations without death / neither	OR	95% CI		N deaths / hospitalizations without death / neither	OR	95% CI	
Male sex (vs. female)	26 / 205 / 822	1.50	1.18	1.91	21 / 132 / 837	1.41	1.04	1.93	10 / 82 / 458	2.12	1.32	3.42
Pandemic time period												
Until June 15 th , 2020	24 / 207 / 513	1	[Reference]		13 / 123 / 423	1	[Reference]		8 / 65 / 338	1	[Reference]	
June 16, 2020 – December 31, 2020	16 / 117 / 910	0.41	0.31	0.55	11 / 71 / 792	0.36	0.26	0.52	1 / 29 / 308	0.52	0.31	0.88
January 1, 2021, and later	11 / 68 / 427	0.53	0.37	0.75	2 / 35 / 348	0.35	0.22	0.56	3 / 21 / 139	0.89	0.49	1.63
Comorbidities												
Hypertension alone (vs. no hypertension, no CVD)	18 / 141 / 466	1.06	0.79	1.41	8 / 68 / 255	1.49	1.01	2.19	2 / 34 / 120	1.69	0.97	2.95
CVD alone (vs. no hypertension, no CVD)	2 / 18 / 31	1.46	0.76	2.83	3 / 11 / 22	2.77	1.25	6.13	2 / 9 / 22	1.71	0.67	4.36
CVD and hypertension (vs. no hypertension, no CVD)	11 / 37 / 81	1.07	0.66	1.74	5 / 15 / 33	1.72	0.82	3.62	5 / 11 / 22	2.00	0.83	4.83
COPD or asthma	14 / 49 / 124	1.95	1.34	2.82	5 / 23 / 86	1.40	0.81	2.40	2 / 14 / 46	2.02	0.94	4.35
Other chronic lung disease	3 / 21 / 23	2.41	1.28	4.55	5 / 7 / 18	2.75	1.18	6.40	3 / 6 / 13	2.94	1.11	7.78
Chronic kidney disease	9 / 26 / 26	2.56	1.46	4.48	3 / 11 / 10	3.04	1.22	7.54	2 / 5 / 10	1.10	0.34	3.54
Obesity alone (vs. no obesity, no diabetes)	9 / 75 / 306	1.40	1.02	1.94	1 / 33 / 189	1.37	0.87	2.15	1 / 30 / 175	1.31	0.77	2.26
Diabetes mellitus alone (vs. no obesity, no diabetes)	6 / 60 / 115	1.85	1.25	2.74	5 / 23 / 60	1.54	0.88	2.70	4 / 19 / 36	3.33	1.68	6.61

Obesity and diabetes mellitus (vs. no obesity, no diabetes)	10 / 35 / 93	1.85	1.18	2.92	3 / 9 / 27	1.82	0.79	4.21	1 / 17 / 42	2.00	0.93	4.30
Cancer and known smoking habits (vs. no cancer, never smoked)	2 / 19 / 30	1.20	0.62	2.33	0 / 4 / 11	1.01	0.26	3.89	2 / 6 / 16	1.28	0.45	3.60
Cancer and unknown smoking habits (vs. no cancer, never smoked)	2 / 4 / 8	2.74	0.86	8.74	3 / 3 / 5	2.42	0.59	9.92		n.a.		
No cancer and ever smoked or unknown smoking habits (vs. no cancer, never smoked)	24 / 155 / 934	0.83	0.63	1.09	11 / 83 / 716	0.88	0.59	1.32	3 / 43 / 272	0.98		
Medication												
No DMARD therapy	8 / 49 / 177	1	[Reference]		8 / 47 / 254	1	[Reference]		5 / 32 / 141	1	[Reference]	
Antimalarials	0 / 3 / 12	0.78	0.17	3.45	0 / 1 / 2	1.96	0.13	30.73		n.a.		
Methotrexate	11 / 101 / 342	0.97	0.64	1.47	2 / 14 / 35	1.70	0.80	3.61	0 / 18 / 72	0.77	0.37	1.58
Leflunomide	2 / 16 / 36	1.24	0.60	2.59		n.a.				n.a.		
Sulfasalazine	7 / 15 / 54	1.86	0.98	3.54	5 / 17 / 82	1.25	0.66	2.37				
Cyclosporine	0 / 1 / 10	0.61	0.07	5.13		n.a.						
TNF inhibitors	11 / 104 / 780	0.47	0.32	0.71	10 / 128 / 1038	0.75	0.49	1.13	3 / 16 / 207	0.38	0.19	0.75
IL-17 inhibitors	1 / 49 / 251	0.56	0.34	0.90	1 / 21 / 142	0.71	0.38	1.33	0 / 18 / 127	0.53	0.25	1.12
IL-23/IL-12+23 inhibitors	3 / 31 / 111	0.76	0.43	1.34		n.a.			2 / 27 / 225	0.43	0.23	0.82
JAK inhibitors	7 / 9 / 35	1.65	0.80	3.42	0 / 1 / 10	0.52	0.06	4.72		n.a.		
Apremilast	1 / 14 / 45	0.91	0.43	1.91		n.a.			2 / 4 / 13	1.14	0.34	3.76
Disease activity (DA) and glucocorticoids (GCs)												
Remission/low DA, no GCs	30 / 273 / 1431	1	[Reference]		18 / 159 / 1274	1	[Reference]		9 / 83 / 64	1	[Reference]	
Remission/low DA, GCs	9 / 35 / 121	1.63	1.02	2.59	4 / 21 / 53	2.54	1.41	4.57	0 / 2 / 3	2.18	0.24	20.25
Moderate/high DA, no GCs	8 / 63 / 241	1.44	1.01	2.04	3 / 37 / 213	1.60	0.99	2.59	3 / 30 / 141	1.11	0.65	1.90

Moderate/high DA, GCs	4 / 21 / 57	2.02	1.12	3.64	2 / 11 / 23	2.85	1.25	6.52		n.a.		
NSAIDs	8 / 75 / 479	0.75	0.55	1.02	6 / 69 / 565	0.72	0.48	1.09	2 / 7 / 23	1.69	0.65	4.40

Results for ordinal mixed effects logistic regression analysis in patients with specific disease entities. Shown are fixed effects, random effects for country are not shown. Missing values are imputed via multiple imputation, patient numbers may thus be rounded. The model was additionally adjusted for age employing five-knot restricted cubic splines. Significant associations highlighted in bold.

axSpA, axial spondyloarthritis; bDMARD, biological disease-modifying anti-rheumatic drugs; BMI, body mass index; csDMARD, conventional synthetic disease-modifying anti-rheumatic drugs; DMARD, disease-modifying anti-rheumatic drugs; IL, interleukin; JAK, Janus kinase; N, number; NSAID, non-steroidal anti-inflammatory drugs; PsA, psoriatic arthritis; PsO, psoriasis without arthritis; TNF; tumour necrosis factor; tsDMARD, targeted synthetic disease-modifying anti-rheumatic drugs.

Supplementary Table 4. Sensitivity analysis – Multivariable ordinal logistic regression analysis of factors associated with COVID-19 severity (confirmed cases)				
N deaths/hospitalisations/neither	75 / 680 / 3421			
	N deaths/ hospitalizations/neither	OR	95% CI	
Male sex (vs. female)	47 / 387 / 1733	1.48	1.23	1.78
Pandemic time period				
Until June 15 th , 2020	36 / 361 / 739	1	[Reference]	
June 16, 2020 – December 31, 2020	24 / 203 / 1858	0.28	0.23	0.35
January 1, 2021, and later	15 / 116 / 824	0.36	0.28	0.47
Comorbidities				
Hypertension alone (vs. no hypertension, no CVD)	21 / 228 / 678	1.32	1.05	1.66
CVD alone (vs. no hypertension, no CVD)	6 / 35 / 62	1.84	1.14	2.97
CVD and hypertension (vs. no hypertension, no CVD)	18 / 55 / 117	1.28	0.86	1.90
COPD or asthma	19 / 78 / 202	1.83	1.35	2.47
Other chronic lung disease	10 / 34 / 47	2.35	1.48	3.74
Chronic kidney disease	11 / 38 / 36	2.39	1.48	3.87
Obesity alone (vs. no obesity, no diabetes)	9 / 130 / 554	1.36	1.06	1.74
Diabetes mellitus alone (vs. no obesity, no diabetes)	14 / 97 / 171	1.91	1.40	2.60
Obesity and diabetes mellitus (vs. no obesity, no diabetes)	10 / 53 / 135	1.65	1.13	2.42
Cancer and known smoking habits (vs. no cancer, never smoked)	3 / 28 / 47	1.09	0.63	1.88
Cancer and unknown smoking habits (vs. no cancer, never smoked)	4 / 6 / 11	2.80	1.04	7.51
No cancer and ever smoked or unknown smoking habits (vs. no cancer, never smoked)	32 / 256 / 1662	0.84	0.67	1.04
Rheumatic disease				
Psoriatic arthritis	43 / 354 / 1535	1	[Reference]	
Axial spondyloarthritis	21 / 215 / 1308	1.08	0.86	1.36
Psoriasis (without arthritis)	11 / 111 / 578	0.55	0.41	0.74
Medication				
No DMARD therapy	20 / 124 / 492	1	[Reference]	
Antimalarials	0 / 4 / 13	1.09	0.29	4.05
Methotrexate	12 / 122 / 371	1.00	0.73	1.39
Leflunomide	1 / 13 / 36	0.79	0.37	1.72
Sulfasalazine	10 / 32 / 113	1.54	0.97	2.44
Cyclosporine	0 / 1 / 12	0.41	0.05	3.46
TNF inhibitors	18 / 226 / 1628	0.60	0.46	0.79
IL-17 inhibitors	1 / 80 / 412	0.67	0.47	0.96

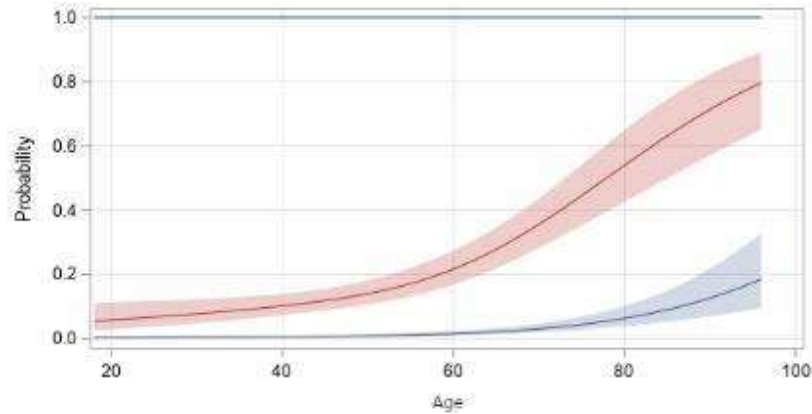
IL-23/IL-12+23 inhibitors	3 / 53 / 264	0.66	0.43	0.9999
JAK inhibitors	7 / 8 / 41	1.43	0.71	2.87
Apremilast	3 / 17 / 42	1.34	0.69	2.59
Disease activity (DA) and glucocorticoids (GCs)				
Remission/low DA, no GCs	49 / 477 / 2748	1	[Reference]	
Remission/low DA, GCs	10 / 57 / 156	1.78	1.22	2.60
Moderate/high DA, no GCs	12 / 119 / 449	1.46	1.12	1.90
Moderate/high DA, GCs	4 / 27 / 68	1.81	1.06	3.08
NSAIDs	13 / 138 / 878	0.76	0.59	0.97
<p>Results for ordinal mixed effects logistic regression analysis in patients with a confirmed COVID-19 diagnosis. Shown are fixed effects, random effects for country are not shown. Missing values are imputed via multiple imputation, patient numbers may thus be rounded. The model was additionally adjusted for age employing four-knot restricted cubic splines. Significant associations highlighted in bold.</p> <p>bDMARD, biological disease-modifying anti-rheumatic drugs; BMI, body mass index; csDMARD, conventional synthetic disease-modifying anti-rheumatic drugs; DMARD, disease-modifying anti-rheumatic drugs; IL, interleukin; JAK, Janus kinase; N, number; NSAID, non-steroidal anti-inflammatory drugs; TNF, tumour necrosis factor; tsDMARD, targeted synthetic disease-modifying anti-rheumatic drugs.</p>				

Supplementary Table 5. Sensitivity analysis – Multivariable binary logistic regression analysis of factors associated with COVID-19 severity (outcome hospitalisation)				
N hospitalizations /other	837 / 4220			
	N hospitalizations/other	OR	95% CI	
Male sex (vs. female)	479 / 2130	1.50	1.26	1.79
Pandemic time period				
Until June 15 th , 2020	442 / 1285	1	[Reference]	
June 16, 2020 – December 31, 2020	250 / 2018	0.40	0.33	0.49
January 1, 2021, and later	145 / 917	0.51	0.40	0.66
Comorbidities				
Hypertension alone (vs. no hypertension, no CVD)	272 / 847	1.31	1.05	1.62
CVD alone (vs. no hypertension, no CVD)	45 / 76	1.92	1.22	3.02
CVD and hypertension (vs. no hypertension, no CVD)	87 / 136	1.36	0.93	1.98
COPD or asthma	111 / 257	1.69	1.27	2.25
Other chronic lung disease	45 / 55	2.39	1.49	3.82
Chronic kidney disease	58 / 46	2.29	1.44	3.66
Obesity alone (vs. no obesity, no diabetes)	153 / 676	1.38	1.09	1.74
Diabetes mellitus alone (vs. no obesity, no diabetes)	117 / 212	1.92	1.43	2.58
Obesity and diabetes mellitus (vs. no obesity, no diabetes)	75 / 162	1.75	1.22	2.51
Cancer and known smoking habits (vs. no cancer, never smoked)	33 / 57	1.19	0.70	2.02
Cancer and unknown smoking habits (vs. no cancer, never smoked)	13 / 13	2.06	0.80	5.33
No cancer and ever smoked or unknown smoking habits (vs. no cancer, never smoked)	330 / 1933	0.87	0.71	1.06
Rheumatic disease				
Psoriatic arthritis	450 / 1850	1	[Reference]	
Axial spondyloarthritis	260 / 1576	1.08	0.87	1.35
Psoriasis (without arthritis)	127 / 794	0.47	0.36	0.63
Medication				
No DMARD therapy	151 / 573	1	[Reference]	
Antimalarials	4 / 14	1.20	0.33	4.39
Methotrexate	49 / 449	1.12	0.82	1.53
Leflunomide	18 / 42	1.13	0.56	2.27
Sulfasalazine	44 / 136	1.25	0.79	1.97
Cyclosporine	1 / 18	0.31	0.04	2.46
TNF inhibitors	278 / 2026	0.59	0.46	0.77
IL-17 inhibitors	91 / 520	0.69	0.49	0.96
IL-23/IL-12+23 inhibitors	63 / 344	0.74	0.50	1.10

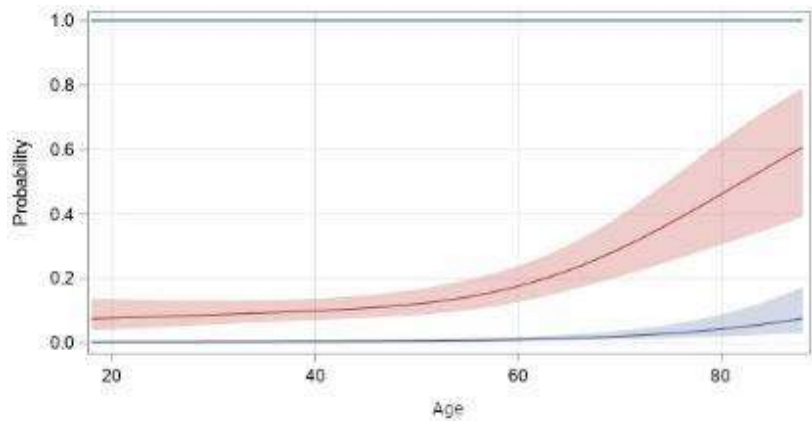
JAK inhibitors	17 / 45	1.34	0.68	2.65
Apremilast	21 / 58	1.07	0.57	2.00
Disease activity (DA) and glucocorticoids (GCs)				
Remission/low DA, no GCs	580 / 3358	1	[Reference]	
Remission/low DA, GCs	74 / 177	1.97	1.38	2.82
Moderate/high DA, no GCs	145 / 504	1.36	1.07	1.74
Moderate/high DA, GCs	38 / 82	2.34	1.43	3.81
NSAIDs	171 / 1072	0.78	0.61	0.99
<p>Results for binary mixed effects logistic regression analysis in all patients, for the outcome of hospitalization. Shown are fixed effects, random effects for country are not shown. Missing values are imputed via multiple imputation, patient numbers may thus be rounded. The model was additionally adjusted for age employing four-knot restricted cubic splines. Significant associations highlighted in bold.</p> <p>bDMARD, biological disease-modifying anti-rheumatic drugs; BMI, body mass index; csDMARD, conventional synthetic disease-modifying anti-rheumatic drugs; DMARD, disease-modifying anti-rheumatic drugs; IL, interleukin; JAK, Janus kinase; N, number; NSAID, non-steroidal anti-inflammatory drugs; TNF; tumour necrosis factor; tsDMARD, targeted synthetic disease-modifying anti-rheumatic drugs.</p>				

Supplementary Table 6. Sensitivity analysis – Multivariable binary logistic regression analysis of factors associated with COVID-19 severity (outcome death)				
N deaths / survivors	89 / 5070			
	N deaths/ survivors	OR	95% CI	
Male sex (vs. female)	57 / 2605	2.00	1.22	3.26
Pandemic time period				
Until June 15 th , 2020	45 / 1723	1	[Reference]	
June 16, 2020, and later	44 / 3347	0.67	0.41	1.10
Comorbidities				
Three or more comorbidities (vs. 0 – 2 comorbidities)	36 / 435	3.34	1.98	5.63
Smoking habits				
Ever smoked (vs. never smoked)	20/ 1209	0.76	0.41	1.39
Unknown smoking habits (vs. never smoked)	24 / 1141	1.99	0.95	4.16
Medication				
No DMARD therapy	21 / 714	1	[Reference]	
Methotrexate	13 / 596	0.57	0.26	1.24
Leflunomide	2 / 61	0.58	0.11	3.03
Sulfasalazine	12 / 171	2.64	1.13	6.17
TNF inhibitors	24 / 2360	0.50	0.26	0.98
IL-17 inhibitors	2 / 625	0.11	0.02	0.51
IL-23/IL-12+23 inhibitors	5 / 403	0.39	0.13	1.14
JAK inhibitors	7 / 58	7.49	2.61	21.47
Apremilast	3 / 82	1.05	0.25	4.40
Disease activity (DA) and glucocorticoids (GCs)				
Moderate/high DA (vs. remission/low DA)	20 / 857	1.12	0.60	2.06
GC therapy (vs. no GC therapy)	18 / 352	1.91	1.002	3.64
NSAIDs	16 / 1252	0.81	0.42	1.54
<p>Results for binary mixed effects logistic regression analysis in all patients, for the outcome death. Shown are fixed effects, random effects for country are not shown. Missing values are imputed via multiple imputation, patient numbers may thus be rounded. The model was additionally adjusted for age employing three-knot restricted cubic splines. Significant associations highlighted in bold.</p> <p>bDMARD, biological disease-modifying anti-rheumatic drugs; BMI, body mass index; csDMARD, conventional synthetic disease-modifying anti-rheumatic drugs; DMARD, disease-modifying anti-rheumatic drugs; IL, interleukin; JAK, Janus kinase; N, number; NSAID, non-steroidal anti-inflammatory drugs; TNF; tumour necrosis factor; tsDMARD, targeted synthetic disease-modifying anti-rheumatic drugs.</p>				

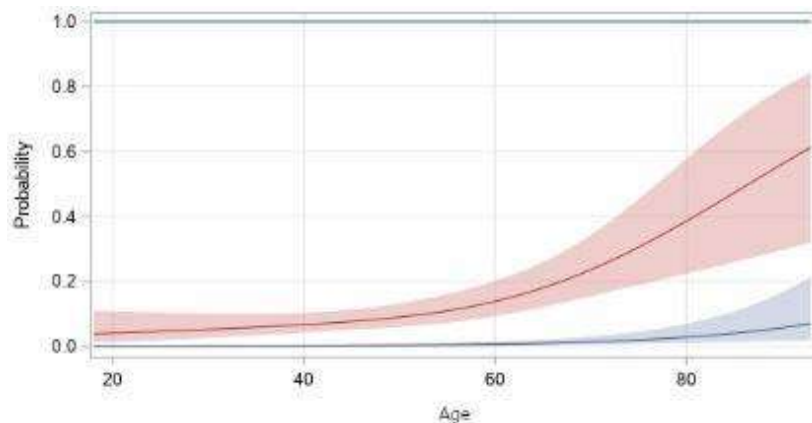
Supplementary figure 1. Relationship between age and probability of hospitalisation (red) and death (blue) in patients with psoriatic arthritis (disease-specific model), estimated by three-knot restricted cubic splines, with 95% confidence intervals.



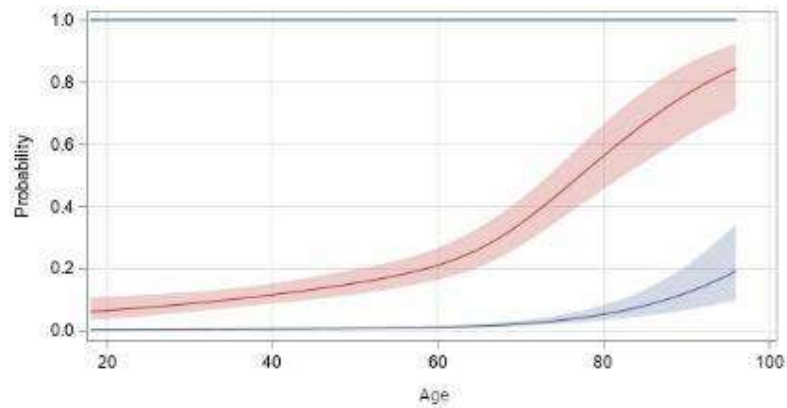
Supplementary figure 2. Relationship between age and probability of hospitalisation (red) and death (blue) in patients with axial spondyloarthritis (disease-specific model), estimated by three-knot restricted cubic splines, with 95% confidence intervals.



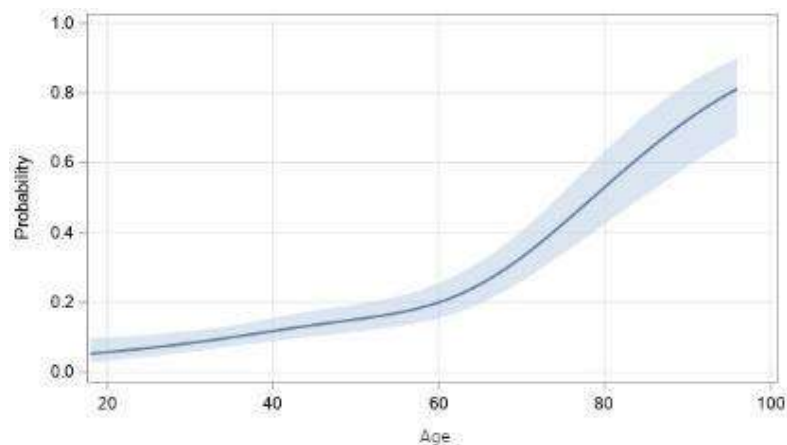
Supplementary figure 3. Relationship between age and probability of hospitalisation (red) and death (blue) in patients with psoriasis without arthritis (disease-specific model), estimated by three-knot restricted cubic splines, with 95% confidence intervals.



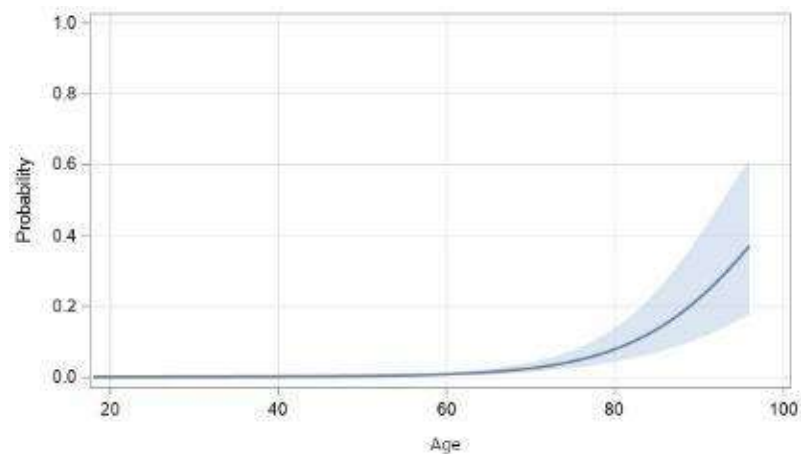
Supplementary figure 4. Relationship between age and probability of hospitalisation (red) and death (blue) estimated by four-knot restricted cubic splines, with 95% confidence intervals, for one imputation stratum (model with confirmed cases only).



Supplementary figure 5. Relationship between age and probability of hospitalisation estimated by four-knot restricted cubic splines, with 95% confidence interval (binary outcome model: hospitalisation).



Supplementary figure 6. Relationship between age and probability of death estimated by three-knot restricted cubic splines, with 95% confidence interval, for one imputation stratum (binary outcome model: death).



SUPPLEMENTARY METHODS

Choice of reference category for medication exposure

Regarding the choice of the reference category for disease-modifying antirheumatic drugs (DMARDs), the most populated categories and therefore possible choices were TNF inhibitors (TNFi), methotrexate, and absence of DMARD therapy. While the TNFi group was the largest group overall, patients receiving TNFi were younger, had less comorbidities, smoked less, took steroids less often, and had better disease control than the overall average, and therefore this group was felt to be inappropriate as a reference group. The proportion of patients receiving methotrexate, on the other hand, was much higher among psoriatic arthritis (PsA) patients compared to patients with axial spondyloarthritis (axSpA) or psoriasis without arthritis (PsO). Moreover, in spondyloarthritis patients with purely axial disease, conventional synthetic DMARDs like methotrexate are ineffective and rarely prescribed. Thus, absence of DMARD therapy was adopted as the medication reference group for DMARDs.

Assumptions of the proportional odds model

In the proportional odds model, the odds ratio for the events “hospitalized or deceased” vs. “neither hospitalized nor deceased” is assumed to be equal to the odds ratio for the events “deceased” vs. “not deceased (hospitalized or not hospitalized)”. This is the odds ratio reported as results for the model. Potential deviations from this assumption were assessed graphically by plotting the stratified means for the levels of the ordinal outcome together with the expected values given that the proportional odds assumption holds,[1] without detecting deviations of concern overall (data not shown).

Statistical interactions

Four two-way interactions were modeled in an additive sense[2]:

1. Between hypertension and cardiovascular disease (CVD): hypertension alone, CVD alone, hypertension combined with CVD; vs. no hypertension and no CVD.
2. Between obesity and diabetes: obesity alone, diabetes alone, obesity combined with diabetes; vs. no obesity and no diabetes.
3. Between smoking status and cancer: cancer and known smoking habits, cancer unknown smoking habits, no cancer and ever smoked or unknown smoking habits; vs. no cancer and never smoked. The interaction between smoking status and cancer was modelled in a non-standard way due to the group of patients

with missing information on smoking habits presenting an OR exceeding the ORs for patients that had never or ever smoked in preliminary analyses, suggesting it was more appropriate to consider it as a separate subgroup within the interaction.

4. Between disease activity and prednisolone-equivalent glucocorticoid (GC) use: remission/low disease activity and GC use, moderate/high disease activity and no GC use, moderate/high disease activity and GC use; vs. remission/low disease activity and no GC use.

Patients excluded and handling of missing data

Patients under the age of 18, patients with missing primary outcome as well as missing values for age, sex, pandemic time period, DMARD exposure, patients treated with more than one biological DMARD, patients treated with DMARDs not typically used or licensed for PsO, PsA or axSpA, PsA (abatacept, B-cell inhibitors, IL-1 and IL-6 inhibitors), and patients diagnosed with multiple inflammatory rheumatic diseases (except for Sjögren's syndrome, and if not receiving B-cell therapies, azathioprine, mycophenolate, ciclosporin, cyclophosphamide or tacrolimus) were excluded from the analysis (529 patients excluded in total).

Further, for some analyses, medications without patients at all or medications with all patients falling into the same outcome category had to be excluded for statistical reasons in some of the secondary and sensitivity analyses (leflunomide, cyclosporine, IL-23 inhibitors, and apremilast, for the analyses focusing on patients with axSpA; antimalarials, leflunomide, sulfasalazine, cyclosporine, JAK inhibitors, and glucocorticoids in the presence of moderate/high disease activity, for the analysis focusing on patients with PsO; antimalarials and cyclosporine, for the analysis using the binary outcome mortality). Missing values for comorbidities, glucocorticoid therapy, disease activity and NSAIDs were derived by multiple imputation using full conditional specification.[3]

References

1. Harrell F. Regression Modeling Strategies: With Applications to Linear Models, Logistic Regression, and Survival Analysis. Chapter 5: Resampling, Validating, and Simplifying the Model. 2001 01/01; 3:88-103.
2. Rothman KJ. Modern epidemiology: Little, Brown & Co, Boston, 1986.
3. van Buuren S. Multiple imputation of discrete and continuous data by fully conditional specification. Stat Methods Med Res 2007;16:219–42.