

Association of Nail Psoriasis With Disease Activity Measures and Impact in Psoriatic Arthritis: Data From the Corrona Psoriatic Arthritis/Spondyloarthritis Registry

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ABSTRACT. Objective. To examine the association of nail psoriasis with disease activity, quality of life, and work productivity in patients with psoriatic arthritis (PsA).

> Methods. All patients with PsA who enrolled in the Corrona PsA/Spondyloarthritis Registry between March 2013 and October 2018 and had data on physician-reported nail psoriasis were included and stratified by presence vs absence of nail psoriasis at enrollment. Patient demographics, disease activity, quality of life (QOL), and work productivity at enrollment were compared between patients with vs without nail psoriasis using t-tests or Wilcoxon rank-sum tests for continuous variables and chi-square or Fisher exact tests for categorical variables.

> Results. Of the 2841 patients with PsA included, 1152 (40.5%) had nail psoriasis and 1689 (59.5%) did not. Higher proportions of patients with nail psoriasis were male (51.9% vs 44.1%) and disabled from working (12.3% vs 7.8%) compared with patients without nail psoriasis (all P < 0.05). Patients with nail psoriasis had higher disease activity than those without nail psoriasis, including higher tender and swollen joint counts, worse Disease Activity Index for Psoriatic Arthritis and Psoriatic Arthritis Disease Activity Score values, and increased likelihood of having enthesitis and dactylitis (all P < 0.05). Patients with nail psoriasis had worse pain, fatigue, and work and activity impairment than those without nail psoriasis (all P < 0.05).

> Conclusion. Patients with PsA who have nail psoriasis had worse disease activity, QOL, and work productivity than those without nail involvement, emphasizing the importance of identification and management of nail disease in patients with PsA.

Key Indexing Terms: nail disease, psoriasis, psoriatic arthritis, registry

Psoriatic arthritis (PsA) is a chronic, immune-mediated inflammatory disease with a diverse array of symptoms that affect the musculoskeletal system, skin, and/or nails1. The Group for Research and Assessment of Psoriasis and Psoriatic Arthritis (GRAPPA) recognizes 6 clinical domains of PsA: peripheral disease, axial disease, enthesitis, dactylitis, skin disease, and nail disease². Symptoms in these domains may occur alone or

in combination and range from mild to severe2. In addition to clinical signs and symptoms, patients with PsA can experience reduced health-related QOL (HRQOL) from the pain, stiffness, fatigue, and reduced physical function that result from the disease³.

Nail disease is an important feature of PsA, and prevalence estimates of nail disease among patients with PsA range from

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41% to 93%⁴. Manifestations of nail psoriasis differ based on the involved structure within the nail unit. Psoriasis in the nail matrix can present as pitting, crumbling, leukonychia, red spots of the lunula, and transverse grooves, whereas psoriasis in the nail bed may present as oil-drop discoloration, splinter hemorrhages, subungual hyperkeratosis, or onycholysis^{5,6}. Additionally, imaging studies have indicated a close link between nail psoriasis and enthesitis (inflammation of the tendons, ligaments, or joint capsule fiber insertion into the bone)^{7,8}, a hallmark manifestation of early PsA that can lead to structural changes, pain, and disability⁹.

Patients with psoriatic disease who have nail involvement have worse HRQOL than those without nail involvement 10,11. Nail psoriasis can result in pain and substantial functional impairment, which may inhibit daily activities and reduce work productivity^{11,12,13,14}. Nail psoriasis is also associated with an increased prevalence of anxiety and depression, as well as psychological distress due to embarrassment, self-consciousness, and feeling stigmatized by what is perceived as a disfiguring disease^{6,10,15}.

Although nail disease is an important feature of PsA, little is known about how patients with PsA and nail disease differ from those without nail disease with respect to patient and clinical characteristics and disease burden. A better understanding of the burden of nail disease in patients with PsA may help raise awareness of the importance of assessing nail involvement in patients with PsA and affect treatment choices. In patients with psoriasis (without PsA), nail involvement is associated with higher psoriasis disease severity, including a higher percentage of affected body surface area (BSA) and worse Psoriasis Area and Severity Index (PASI) scores^{11,16,17,18}, more pain and psychological distress, reduced HRQOL, and substantial functional impairment and disability^{11,13,15,17,18}. However, limited data are available regarding the burden of nail disease in PsA. Few real-world studies have examined characteristics of patients with PsA and nail psoriasis, particularly in the United States. The objective of this study was to examine the association of nail disease with patient demographics, disease activity, QOL, and work productivity of patients with PsA. We addressed this question using real-world data from the US-based Corrona PsA/Spondyloarthritis (SpA) Registry.

MATERIALS AND METHODS

Study design. This was a cross-sectional study conducted using data collected at the time of patient enrollment in the Corrona PsA/SpA registry. Data source. The Corrona PsA/SpA Registry is a large, independent, prospective, observational cohort initiated in March 2013 that comprises patients diagnosed with PsA or SpA by a rheumatologist. The registry includes patients recruited by 49 participating rheumatologists from 45 private and academic practice sites across 27 states in the USA. As of October 1, 2018, data on approximately 3572 patients with PsA/SpA had been collected.

All participating investigators were required to obtain full board approval for conducting noninterventional research with a limited data set involving human participants. The Corrona PsA/SpA Registry and its investigators have been reviewed and approved by a central institutional review board (IRB; New England Independent Review Board No. 120160070). For academic investigative sites that did not receive a waiver

to use the central IRB, full board approval was obtained from the respective governing IRB. All research was conducted in compliance with the current (2013) version of the Declaration of Helsinki. All registry participants were required to provide written informed consent and authorization prior to participating.

Study population. This study included all patients aged ≥ 18 years enrolled in the Corrona PsA/SpA Registry between March 2013 and October 2018 with a diagnosis of PsA and nonmissing data on physician-reported nail psoriasis at the enrollment visit.

Exposure. Patients were stratified by presence vs absence of physician-reported nail psoriasis at the time of enrollment. Presence of nail psoriasis was defined as a nonzero response on the nail psoriasis visual analog scale (VAS) of 0−100. The physician nail psoriasis VAS has been used in previous studies of patients with PsA and shown to correlate well with both the modified Nail Psoriasis Severity Index (mNAPSI) and patient-reported nail VAS scores 19. No disease severity cutoffs have been established for the nail psoriasis VAS and we observed substantial variability in the distribution of nail psoriasis VAS scores in our study population. The lack of clear cutoffs precluded meaningful analysis of the correlation between the severity of nail involvement and disease activity measures. Thus, for the purposes of these analyses, patients were classified as those with nail psoriasis (nail psoriasis VAS score ≥ 1) and those without nail psoriasis (nail psoriasis VAS score = 0).

Study assessments. Data were collected at registry enrollment using questionnaires completed during office visits by patients and their treating rheumatologists. Data collected included patient demographics (age, sex, race, insurance type, and work status), clinical characteristics (BMI, symptom duration, disease duration, history of physician-reported comorbidities, treatment history, and current treatment), laboratory measurements (C-reactive protein and erythrocyte sedimentation rate), disease activity measures, and patient-reported outcome measures (PROM).

Disease activity measures evaluated in this study included nail psoriasis VAS, 68 tender joint count (TJC), 66 swollen joint count (SJC), Disease Activity Index for Psoriatic Arthritis (DAPSA) score, Psoriatic Arthritis Disease Activity Score (PASDAS), enthesitis, dactylitis, percentage of affected BSA, physician global assessment (PGA) of arthritis (VAS, 0–100), PGA of arthritis and psoriasis (VAS, 0-100), and minimal disease activity (MDA). MDA was defined as "yes" if a patient met ≥ 5 of the 7 following categories: TJC \leq 1, SJC \leq 1, BSA \leq 3%, patient pain VAS \leq 15, patient global activity VAS ≤ 20, Health Assessment Questionnaire-Disability Index (HAQ-DI) \leq 0.5, and tender entheseal points \leq 1²⁰. PROM included patient-reported pain (VAS, 0-100), patient-reported fatigue (VAS, 0-100), patient global assessment (PtGA) of arthritis (VAS, 0-100), PtGA of arthritis and psoriasis (VAS, 0-100), morning stiffness, HAQ-DI (0-3), EuroQol 5-dimension questionnaire (EQ-5D; 0-1), and EQ VAS (0-100). Work productivity was assessed using the Work Productivity and Activity Impairment Questionnaire.

Statistical analysis. Descriptive analyses of patient demographics, disease activity, QOL, and work productivity were assessed at enrollment. Categorical variables were summarized using frequency counts and percentages. Continuous variables were summarized using means and SD. Patients with and without nail psoriasis were compared using *t*-tests or Wilcoxon rank-sum tests for continuous variables and chi-square or Fisher exact tests for categorical variables. Statistical analyses were performed using Stata 15.1 (StataCorp).

RESULTS

Demographics, clinical characteristics, and treatment profile. Of the 2925 patients with PsA enrolled in the Corrona PsA/SpA Registry between March 2013 and October 2018, information was available for 2841 about the presence of nail psoriasis at registry enrollment and were included in the analysis. Of the included patients, 1152 had nail psoriasis (prevalence, 40.5%).

Overall, the mean age was 53.9 years, 52.7% of patients were female, most patients were White (94.0%), were overweight or

obese (83.7%), and had a history of psoriasis (86.5%; Table 1). The mean symptom and disease durations were 11.3 and 8.0

Table 1. Demographics, clinical characteristics, and treatment profile of patients with PsA at enrollment, stratified by presence of nail psoriasis.

Characteristic ^a	Overall b , $N = 2841$	With Nail Psoriasis, $n = 1152$	Without Nail Psoriasis, n = 1689	P^*
Age, yrs	53.9 (13.2) [2804]	53.1 (12.9) [1133]	54.4 (13.3) [1671]	< 0.01
Female, n (%) [n]	1486 (52.7) [2818]	549 (48.1) [1142]	937 (55.9) [1676]	< 0.01
Race, n (%)	n = 2772	n = 1120	n = 1652	0.11
White	2607 (94.0)	1046 (93.4)	1561 (94.5)	
Black	19 (0.7)	5 (0.4)	14 (0.8)	
Other	146 (5.3)	69 (6.2)	77 (4.7)	
Work status, n (%)	n = 2811	n = 1135	n = 1676	< 0.01
Full time	1514 (53.9)	614 (54.1)	900 (53.7)	
Part time	230 (8.2)	93 (8.2)	137 (8.2)	
Disabled from working	270 (9.6)	140 (12.3)	130 (7.8)	
Retired	602 (21.4)	213 (18.8)	389 (23.2)	
Other	195 (6.9)	75 (6.6)	120 (7.2)	
BMI, kg/m ²	31.8 (7.5) [2783]	31.8 (7.1) [1132]	31.7 (7.7) [1651]	0.39
Normal/underweight (< 25.0), n (%)	452 (16.2)	160 (14.1)	292 (17.7)	0.04
Overweight (≥ 25.0–29.9), n (%)	836 (30.0)	354 (31.3)	482 (29.2)	
Obese (≥ 30.0), n (%)	1495 (53.7)	618 (54.6)	877 (53.1)	
Symptom duration, yrs	11.3 (10.3) [2759]	11.4 (10.6) [1120]	11.1 (10.2) [1639]	0.79
Disease duration, yrs	8.0 (8.6) [2779]	8.0 (8.8) [1126]	8.0 (8.5) [1653]	0.50
History of comorbid conditions, n (%)				
CVD^{c}	345 (12.1)	135 (11.7)	210 (12.4)	0.57
Depression	429 (15.1)	205 (17.8)	224 (13.3)	< 0.01
Diabetes mellitus	417 (14.7)	181 (15.7)	236 (14.0)	0.20
Any cancer (excluding NMSC)	221 (7.8)	89 (7.7)	132 (7.8)	0.93
Hypertension	1101 (38.8)	442 (38.4)	659 (39.0)	0.73
Hyperlipidemia	679 (23.9)	291 (25.3)	388 (23.0)	0.16
Metabolic syndrome	441 (15.5)	187 (16.2)	254 (15.0)	0.39
Crohn disease	27 (1.0)	13 (1.1)	14 (0.8)	0.42
Ulcerative colitis	26 (0.9)	8 (0.7)	18 (1.1)	0.31
Anxiety	118 (4.2)	49 (4.3)	69 (4.1)	0.83
Uveitis	50 (1.8)	23 (2.0)	27 (1.6)	0.43
Psoriasis	2458 (86.5)	1012 (87.8)	1446 (85.6)	0.09
Nail psoriasis ^d	1304 (45.9)	1152 (100.0)	152 (9.0)	< 0.01
Fibromyalgia	157 (5.5)	63 (5.5)	94 (5.6)	0.91
Serious infections ^e	181 (6.4)	62 (5.4)	119 (7.0)	0.08
History of medication use, n (%)				
Biologic	890 (31.3)	359 (31.2)	531 (31.4)	0.88
tsDMARD	144 (5.1)	52 (4.5)	92 (5.4)	0.27
csDMARD	814 (28.7)	319 (27.7)	495 (29.3)	0.35
Prednisone	394 (13.9)	147 (12.8)	247 (14.6)	0.16
Current medication use, n (%)				
Biologic	1698 (59.8)	700 (60.8)	998 (59.1)	0.37
Biologic monotherapy	972 (34.2)	393 (34.1)	579 (34.3)	0.93
tsDMARD	176 (6.2)	60 (5.2)	116 (6.9)	0.07
tsDMARD monotherapy	114 (4.0)	34 (3.0)	80 (4.7)	0.02
csDMARD	1440 (50.7)	602 (52.3)	838 (49.6)	0.17
csDMARD monotherapy	699 (24.6)	286 (24.8)	413 (24.5)	0.82
Prednisone	190 (6.7)	65 (5.6)	125 (7.4)	0.07

^a All values were calculated based on available data and are presented as mean (SD) [n] unless otherwise stated. ^b Patients with nonmissing nail VAS. ^c CVD includes combined histories of myocardial infarction, acute coronary syndrome, coronary artery disease, congestive heart failure, peripheral arterial thromboembolic event, peripheral artery disease, cardiac revascularization procedure, ventricular arrhythmia, cardiac arrest, unstable angina, stroke, transient ischemic attack, peripheral ischemia or gangrene (necrosis), pulmonary embolism, carotid artery disease, or other cardiovascular event. ^d Determined by physician-reported nail psoriasis VAS > 0 or clinical changes associated with nail changes or typical psoriatic nail dystrophy (may include the history of nail changes or typical psoriatic nail dystrophy prior to enrollment). ^c Includes infections that led to hospitalization or intravenous antibiotics: joint/bursa, cellulitis, sinusitis, diverticulitis, sepsis, pneumonia, bronchitis, gastroenteritis, meningitis, urinary tract infection, upper respiratory tract infection, mycobacterium tuberculosis, or infection of other specified site. ^c Wilcoxon rank-sum test or *t*-test for continuous variables and chi-square or Fisher exact test for categorical variables. csDMARD: conventional synthetic disease-modifying antirheumatic drug; CVD: cardiovascular disease; NMSC: nonmelanoma skin cancer; PsA: psoriatic arthritis: tsDMARD: targeted synthetic antirheumatic drug; VAS: visual analog scale.

years, respectively. Prior to registry enrollment, the majority of patients (68.7%) were biologic-naïve. At the time of registry enrollment, 59.8% of patients were receiving a biologic.

Patients with nail psoriasis were slightly younger (mean age, 53.1 vs 54.4 yrs) and more likely to be male (51.9% vs 44.1%) compared with patients without nail psoriasis (both P < 0.01; Table 1). Higher proportions of patients with nail psoriasis were disabled from working (12.3% vs 7.8%; P < 0.01 for distribution across work status category) and had a history of depression (17.8% vs 13.3%; P < 0.01) compared with patients without nail psoriasis. Symptom duration, disease duration, and prior and current medication use were comparable in patients with vs without nail psoriasis.

Disease activity and PROM. Overall, patients with nail psoriasis had higher disease activity at enrollment than those without nail psoriasis (Table 2). Patients with nail psoriasis had a significantly higher mean percentage of affected BSA (7.9% vs 3.5%), TJC

(6.0 vs 3.5), and SJC (2.4 vs 1.7) and worse mean DAPSA (18.1 vs 13.4) and PASDAS (3.8 vs 3.3) than patients without nail psoriasis (all P < 0.01). Patients with nail disease were less likely to have MDA (35.0% vs 47.6%) and had an increased prevalence of enthesitis (27.7% vs 17.0%) and dactylitis (12.2% vs 7.4%) compared with those without nail psoriasis (all P < 0.01). Additionally, patients with nail psoriasis had higher mean PGA of arthritis (22.7 vs 17.0) and PGA of arthritis and psoriasis (27.2 vs 18.8) scores than those without nail psoriasis (both P < 0.01).

Patients with nail psoriasis also had overall worse physical function and HRQOL than patients without nail psoriasis (Table 3). Patients with nail psoriasis reported higher mean pain (41.7 vs 36.1) and fatigue (43.9 vs 39.0) and had higher scores on their PtGA of arthritis (41.8 vs 38.0), and PtGA of arthritis and psoriasis (41.6 vs 37.4) than those without nail psoriasis (all P < 0.01). In addition, patients with nail psoriasis had worse mean HAQ-DI (0.7 vs 0.6) and EQ VAS (68.6 vs 72.1) scores

Table 2. Disease activity among patients with PsA at enrollment, stratified by presence of nail psoriasis.

Characteristic ^a	Overall, N = 2841	With Nail Psoriasis, n = 1152	Without Nail Psoriasis, n = 1689	P^*
Nail psoriasis (VAS 0–100)	7.7 (17.0)	18.9 (22.5)	0	-
TJC (0-68)	4.6 (8.5) [2766]	6.0 (9.9) [1127]	3.5 (7.1) [1639]	< 0.01
SJC (0-66)	2.0 (3.9) [2764]	2.4 (4.4) [1125]	1.7 (3.5) [1639]	< 0.01
MDA, n/m (%) ^b	1075/2530 (42.5)	359/1027 (35.0)	716/1503 (47.6)	< 0.01
DAPSA	15.2 (13.9) [1637]	18.1 (15.7) [643]	13.4 (12.4) [994]	< 0.01
DAPSA group, n (%)	n = 1637	n = 643	n = 994	< 0.01
Remission	322 (19.7)	90 (14.0)	232 (23.3)	
Low	633 (38.7)	231 (35.9)	402 (40.4)	
Moderate	442 (27.0)	195 (30.3)	247 (24.8)	
High	240 (14.7)	127 (19.8)	113 (11.4)	
PASDAS	3.5 (1.6) [1591]	3.8 (1.6) [619]	3.3 (1.5) [972]	< 0.01
PASDAS group, n (%)	n = 1591	n = 619	n = 972	< 0.01
Inactive	680 (42.7)	215 (34.7)	465 (47.8)	
Moderate	359 (22.6)	132 (21.3)	227 (23.4)	
Active	467 (29.4)	228 (36.8)	239 (24.6)	
Very active	85 (5.3)	44 (7.1)	41 (4.2)	
PGA of arthritis	19.3 (20.8) [2823]	22.7 (22.0) [1139]	17.0 (19.6) [1684]	< 0.01
PGA of arthritis and psoriasis	22.2 (22.0) [2784]	27.2 (23.6) [1127]	18.8 (20.2) [1657]	< 0.01
Enthesitis, n/m (%)	606/2841 (21.3)	319/1152 (27.7)	287/1689 (17.0)	< 0.01
SPARCC Enthesitis Index (1-16)	3.7 (2.9) [606]	3.8 (2.9) [319]	3.5 (2.8) [287]	0.14
Dactylitis, n/m (%)	265/2841 (9.3)	140/1152 (12.2)	125/1689 (7.4)	< 0.01
Dactylitis count (1–20)	2.3 (2.1) [265]	2.4 (2.3) [140]	2.2 (1.8) [125]	0.50
CRP, mg/L	8.7 (19.5) [1769]	9.6 (24.6) [702]	8.1 (15.2) [1067]	0.27
ESR, mm/h	17.2 (16.8) [1777]	17.6 (17.3) [687]	17.0 (16.4) [1090]	0.62
BSA, % affected	5.3 (11.3) [2761]	7.9 (14.8) [1125]	3.5 (7.6) [1636]	< 0.01
BSA (categorical), n (%)	n = 2761	n = 1125	n = 1636	< 0.01
No disease (0%)	784 (28.4)	201 (17.9)	583 (35.6)	
Mild disease (> 0–3%)	1149 (41.6)	469 (41.7)	680 (41.6)	
Moderate disease (> 3% to ≤ 10%)	512 (18.5)	258 (22.9)	254 (15.5)	
Severe disease (> 10%)	316 (11.4)	197 (17.5)	119 (7.3)	

^a All values were calculated based on available data and are presented as mean (SD) [n] unless otherwise stated. ^b MDA is defined as "yes" if a patient met ≥ 5 of the 7 following categories: TJC ≤ 1, SJC ≤ 1, BSA ≤ 3%, patient pain VAS ≤ 15, patient global activity VAS ≤ 20, HAQ-DI ≤ 0.5, and tender entheseal points ≤ 1. * t-Test or Wilcoxon rank-sum test for continuous variables and chi-square or Fisher exact test for categorical variables. BSA: body surface area; CRP: C-reactive protein; DAPSA: Disease Activity Index for Psoriatic Arthritis; ESR: erythrocyte sedimentation rate; HAQ-DI: Health Assessment Questionnaire—Disability Index; MDA: minimal disease activity; n/m: number of responders/number of patients with sufficient data for evaluation; PASDAS: Psoriatic Arthritis Disease Activity Score; PGA: physician global assessment; PsA: psoriatic arthritis; SJC: swollen joint count; SPARCC: Spondyloarthritis Research Consortium of Canada; TJC: tender joint count; VAS: visual analog scale.

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Table 3. Patient-reported outcome measures among patients with PsA at enrollment, stratified by presence of nail psoriasis.

Characteristic ^a	Overall, $N = 2841$	With Nail Psoriasis, $n = 1152$	Without Nail Psoriasis, n = 1689	P^*
Patient-reported pain (VAS 0-100)	38.4 (29.6) [2684]	41.7 (30.1) [1081]	36.1 (29.1) [1603]	< 0.01
Patient-reported fatigue (VAS 0-100)	41.0 (29.6) [2796]	43.9 (29.4) [1135]	39.0 (29.6) [1661]	< 0.01
PtGA of arthritis (VAS 0-100)	39.6 (29.9) [2795]	41.8 (29.5) [1132]	38.0 (30.1) [1663]	< 0.01
PtGA of arthritis and psoriasis (VAS 0-100)	39.1 (29.7) [2789]	41.6 (29.0) [1130]	37.4 (30.0) [1659]	< 0.01
Morning stiffness	n = 2762	n = 1118	n = 1644	0.1
< 30 min	980 (35.5)	380 (34.0)	600 (36.5)	
≥ 30 min	1782 (64.5)	738 (66.0)	1044 (63.5)	
HAQ-DI (0-3)	0.7 (0.6) [2700]	0.7 (0.7) [1087]	0.6 (0.6) [1613]	< 0.01
EQ-5D (0-1)	0.8 (0.2) [2680]	0.7 (0.2) [1080]	0.8 (0.2) [1600]	< 0.01
EQ VAS (0-100)	70.7 (21.3) [2790]	68.6 (21.4) [1127]	72.1 (21.1) [1663]	< 0.01
WPAI domains				
Current employment, n/m (%)	1681/2745 (61.2)	678/1109 (61.1)	1003/1636 (61.3)	0.93
% work time missed	5.1 (15.7) [1443]	5.9 (16.9) [575]	4.6 (14.9) [868]	0.26
% impairment while working	19.0 (23.4) [1534]	20.3 (23.8) [605]	18.1 (23.1) [929]	0.02
% overall work impairment	21.6 (25.6) [1379]	23.2 (26.1) [540]	20.5 (25.3) [839]	0.02
% activity impairment	28.9 (29.0) [2569]	32.5 (30.2) [1026]	26.6 (28.0) [1543]	< 0.01

^a All values were calculated based on available data and are presented as mean (SD) [n] unless otherwise indicated. * Wilcoxon rank-sum test *t*-test or for continuous variables and chi-square or Fisher exact test for categorical variables. EQ-5D: EuroQol 5-dimension questionnaire; HAQ-DI: Health Assessment Questionnaire—Disability Index; MDA: minimal disease activity; n/m: number of responders/number of patients with sufficient data for evaluation; PsA: psoriatic arthritis; PtGA: patient global assessment; VAS: visual analog scale; WPAI: Work Productivity and Activity Impairment Questionnaire.

than patients without nail psoriasis (both P < 0.01). Patients with nail psoriasis also reported greater mean percentages of impairment while working (20.3% vs 18.1%; P = 0.02), overall work impairment (23.2% vs 20.5%; P = 0.02), and activity impairment (32.5% vs 26.6%; P < 0.01) than those without nail psoriasis.

DISCUSSION

Although nail psoriasis is recognized as a common feature of PsA, few studies have examined the relevance of nail disease for treatment and outcomes in PsA. Whereas a handful of studies have quantified the prevalence of nail psoriasis in PsA, to our knowledge, our study is among the first to evaluate the burden of nail disease in a real-world population of patients with PsA. We found that patients with PsA in our study who had nail psoriasis had a higher percentage of affected BSA and higher PsA disease activity, including higher TJC, SJC, PASDAS, and DAPSA scores, than those without nail psoriasis. Additionally, the prevalence of work disability and depression was higher among patients with PsA with nail psoriasis, and patients with nail psoriasis had worse pain, fatigue, and EQ VAS scores, and more work and activity impairment than those without nail psoriasis. Our results show that nail psoriasis was associated with an overall higher burden of disease in all domains of PsA.

Nail psoriasis is closely linked with enthesitis, an inflammatory change that occurs early in the development of PsA^{8,21}. This relationship may be due in part to the close anatomical association of the nail matrix with the extensor tendon of the distal interphalangeal joint, an entheseal site frequently affected in early PsA^{7,8,9,22}. In our study population, a higher proportion of patients with nail psoriasis had enthesitis compared with those without nail involvement, which may be reflective of the

relationship between nail psoriasis and enthesitis. In patients with PsA, enthesitis is associated with more peripheral and axial joint damage, higher disease activity, worse HRQOL, and more functional impairment^{23,24}. Thus, when selecting therapies for a patient with PsA with nail involvement, it may be beneficial to consider therapies that can effectively treat nail disease. Available treatments for nail psoriasis include topical therapies, corticosteroid injections, conventional synthetic disease-modifying antirheumatic drugs (csDMARD), and biologics^{5,6,25,26,27}. Current guidelines recommend that choice of therapy should take into account the presence of symptoms in other PsA domains, such as severe skin involvement, presence of enthesitis or dactylitis, and presence of inflammatory bowel disease, as well as patient factors such as medication history and preferences for mode of administration and frequency of dosing^{2,28,29,30,31,32}.

In our real-world population of US patients with PsA, 40.5% of patients had nail psoriasis at the time of registry enrollment. The prevalence of nail involvement in our study population is lower than that observed in previous real-world studies of patients with varied prior biologic experience, which have reported rates of nail involvement ranging from 50% to 87% among patients with PsA^{33–39}. Notably, approximately 31% of patients in our study population had received biologic therapy and 29% had received csDMARD prior to enrollment, which may have ameliorated nail disease in those patients before their participation in the study^{5,26}. Additionally, the majority of previous studies specifically enrolled patients with PsA and psoriasis, whereas the presence of psoriasis was not a criterion for enrollment in our study.

This study is subject to the general limitations of real-world observational studies. Although patients enrolled in the Corrona PsA/SpA Registry are assumed to be generally representative of

patients with PsA in the USA, there may be differences in generalizability overall, given that most patients are enrolled in the registry at the time of therapy initiation 40. All comparisons were descriptive; no adjustments were made to account for differences in patient characteristics, such as age and sex, which may have influenced the other differences observed in patients with vs without nail psoriasis. In this cross-sectional study, adjusting for some of the disease activity elements at baseline was colinear with the outcome of interest and was therefore considered inappropriate for this analysis. Future cohort studies that examine the implications of nail disease on disease activity outcomes should address the effect of higher baseline disease activity. The presence of nail psoriasis was based on physician reporting and may have been underreported. Information on duration and subtypes (e.g., pitting, onycholysis) of nail psoriasis was not captured. The overall severity of nail involvement was evaluated on a VAS, but the link between VAS severity and disease burden was not specifically assessed, as cutoffs have not been established for this measure.

Overall, the results of this study highlight the burden of nail psoriasis in patients with PsA. Nail psoriasis was associated with greater severity of psoriasis and PsA symptoms, more disability and functional impairment, and worse HRQOL. These findings emphasize the importance of identification and management of nail disease in patients with PsA. Further research is needed to assess whether nail disease affects treatment response in patients with PsA.

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