


RESEARCH

Open Access



Pain catastrophizing negatively impacts drug retention rate in patients with Psoriatic Arthritis and axial Spondyloarthritis: results from a 2-years perspective multicenter GIRRCS (Gruppo Italiano di Ricerca in Reumatologia Clinica) study

Damiano Currado^{1,2}, Francesca Saracino^{1,2}, Piero Ruscitti³, Annalisa Marino¹, Ilenia Pantano⁴, Marta Vomero¹, Onorina Berardicurti^{1,2*} , Viktoriya Pavlych³, Claudio Di Vico⁴, Francesco Caso⁵, Luisa Costa⁵, Marco Tasso⁵, Federica Camarda⁶, Francesca Misceo⁷, Francesco De Vincenzo⁸, Addolorata Corrado⁹, Luisa Arcarese¹, Amelia Rigon¹, Marta Vadacca¹, Erika Corberi^{1,2}, Lyubomyra Kun^{1,2}, Francesca Trunfio^{1,2}, Andrea Pilato^{1,2}, Ludovica Lamberti^{1,2}, Francesco Paolo Cantatore⁹, Federico Perosa⁷, Giuliana Guggino⁶, Raffaele Scarpa⁵, Paola Cipriani³, Francesco Ciccia⁴, Roberto Giacomelli^{1,2} and Luca Navarini^{1,2}

Abstract

Background Chronic pain and inflammation are common features of rheumatic conditions such as Psoriatic Arthritis (PsA) and Axial Spondyloarthritis (axSpA), often needing prolonged medication treatment for effective management. Maintaining drug retention is essential for both achieving disease control and improving patients' quality of life. This study investigates the influence of pain catastrophizing, a psychological response to pain, on the drug retention rates of PsA and axSpA patients.

Methods A two-year prospective multicenter observational study involved 135 PsA and 71 axSpA patients. Pain Catastrophizing Scale (PCS) was employed to assess pain catastrophizing. Univariable and multivariable regression analyses were utilized to identify factors associated with drug retention.

Results In the PsA group, patients early discontinuing therapy showed higher baseline disease activity as well as higher incidence of comorbid fibromyalgia. Notably, pain catastrophizing, specifically the domains of Helplessness, Magnification, and Rumination, were significantly elevated in PsA patients who interrupted the treatment. Multivariable analysis confirmed pain catastrophizing as an independent predictor of drug suspension within two years.

In axSpA, drug discontinuation was associated with female gender, shorter disease duration, higher baseline disease activity as well as elevated levels of pain catastrophizing. Univariable analysis supported the role of pain

*Correspondence:

Onorina Berardicurti
o.berardicurti@policlinicocampus.it

Full list of author information is available at the end of the article



© The Author(s) 2024. **Open Access** This article is licensed under a Creative Commons Attribution-NonCommercial-NoDerivatives 4.0 International License, which permits any non-commercial use, sharing, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if you modified the licensed material. You do not have permission under this licence to share adapted material derived from this article or parts of it. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit <http://creativecommons.org/licenses/by-nc-nd/4.0/>.

catastrophizing, including its domains, as predictors of treatment interruption. However, limited events in axSpA patients precluded a multivariate analysis.

Conclusion This prospective study emphasizes the impact of pain catastrophizing on drug retention in patients with PsA and axSpA.

Keywords Psoriatic Arthritis, Axial Spondyloarthritis, Pain Catastrophizing, Retention Rate

Background

Spondyloarthritis (SpA), a group of interconnected chronic rheumatic diseases, displaying various clinical features, shares common clinical signs and distinctive genetic traits [1, 2]. The most prevalent forms of SpA are Psoriatic Arthritis (PsA) and Axial Spondyloarthritis (axSpA). Individuals with SpA often show a higher prevalence of obesity, type 2 diabetes, hypertension, metabolic syndrome, an increased risk of cardiovascular issues, and psychological comorbidities [3–7]. These conditions significantly impact the pain perception and the quality of life in SpA patients. In recent years, several studies were published focusing the extended-term usage of conventional synthetic disease-modifying anti-rheumatic drugs (csDMARDs) and biologic and targeted synthetic agents (bDMARDs and tsDMARDs) in spondyloarthritis, especially analysing the real-world data; despite of these works the factors associated to the retention rate of bDMARDs remain a pivotal question to be addressed [8].

Psychosocial factors are acknowledged as crucial elements influencing the pain experienced by patients with inflammatory arthritides, with particular attention given to pain catastrophizing (PC). The concept of catastrophizing was originally introduced by Albert Ellis and subsequently adapted by Aaron Beck to describe a maladaptive cognitive style observed in patients with anxiety and depressive disorders. Catastrophizing refers to the irrational anticipation of negative future events. In the context of pain, catastrophizing is broadly defined as an exaggeration of negative cognitive and emotional responses during actual or anticipated painful situations [9–11]. To assess catastrophization, the PC Scale (PCS) is employed, which evaluates three domains encompassing various aspects of pain catastrophizing. The first component, referred to as “rumination,” comprises four items related to ruminative thoughts, worry, and the inability to suppress pain-related thoughts. The second component, termed “magnification,” consists of three items reflecting the exaggeration of pain’s unpleasantness and the anticipation of negative outcomes. The third component, labeled “helplessness,” encompasses five items from the Coping Strategies Questionnaire (CSQ) and an additional item indicating the inability to cope with painful situations [12].

In a previous study we showed that PC negatively impacts, the achievement of remission and/or low disease activity in inflammatory arthritis [13]. A largely unexplored topic is the relationship between PC and drug retention rate. Patients with PC may be more likely to struggle with adhering to their prescribed treatment plans; this may include non-compliance with medications, missing appointments, or reluctance in changing their lifestyle and engaging physical therapy. Thus, we planned a multicentric prospective observational study to address the possible impact of PC and its related domains on two years drug retention rate in patients with PsA and axSpA.

Methods

A multicenter, prospective (two years) observational study was conducted on enrolled participants with Psoriatic Arthritis and Axial Spondyloarthritis in 7 Rheumatology Clinics of Italy, widely distributed throughout the Country. Consecutive outpatients were recruited between January 2021 and July 2021. At the initial assessment, PsA participants met the Classification Criteria for Psoriatic Arthritis (CASPAR), while axSpA participants met the 2009 Assessment of Spondyloarthritis International Society (ASAS) Criteria [14, 15]. The study received approval from the Ethics Committee of the University Campus Bio-Medico of Rome (approval no. 78.20 OSS) and was conducted in accordance with the Declaration of Helsinki and its subsequent revisions. Inclusion criteria encompassed individuals of both genders, aged over 18 years, who fulfilled the CASPAR or 2009 ASAS criteria. Exclusion criteria included a history of psychiatric disorders as per DSM-V prior to recruitment, a history of malignancy, pregnancy, age exceeding 75, or an inability to provide informed consent for participation in the study. Upon enrollment, the following PsA disease activity scores were recorded by the examining clinicians: Disease Activity for Psoriatic Arthritis (DAPSA), Minimal Disease Activity (MDA), Bath Ankylosing Spondylitis Disease Activity Index (BASDAI), Psoriasis Area Severity Index (PASI), and Leeds Enthesitis Index (LEI). Additionally, the following axSpA disease activity scores were collected: BASDAI, Bath Ankylosing Spondylitis Functional Index (BASFI), and Ankylosing Spondylitis Disease Activity Score-C-Reactive Protein (ASDAS-CRP)

[16]. To assess the drug discontinuation rate (considering csDMARDs and/or bDMARDs), any occurrence of withdrawal therapy and adverse events recognized or suspected as linked to therapies were recorded during the two-years follow-up. PC, including its domains of Helplessness, Rumination, and Magnification (as continuous variables), was assessed using the PCS.

Patients fulfilling the 2016 American College of Rheumatology revised criteria for fibromyalgia were further identified in our cohort [17, 18].

Continuous variables were presented as the median with the interquartile range (25th–75th percentile), while categorical variables were expressed as percentages (%). The normality of the data was assessed using the Shapiro–Wilk test. Contingency tables were analyzed using the Chi-squared test, whereas rank-based comparisons were conducted using the Mann–Whitney test and Kruskal–Wallis test with Holm’s pairwise comparison adjustments. To identify variables associated with drug retention rate, both univariable and multivariable regression analyses were employed. In the multivariable analysis, variables with a p -value of less than 0.1 in the univariate analysis, along with age and sex, were considered. Multivariable analysis of retention predictors was computed using the Cox proportional-hazards model.

All the statistical analyses were carried out by using Stata version 14, and statistical significance was defined as p -values less than 0.05.

Results

The study population included 135 patents affected by Psoriatic Arthritis (PsA) and 71 by Axial Spondyloarthritis (axSpA). The main demographic, anthropometric and clinical characteristics of the study populations are reported in Table 1 for PsA and in Table 2 for axSpA.

In PsA patient cohort, the one-year Retention Rate was 79.26%. [Drug suspension related to primary failure in 7.20% of patients, secondary failure in 60.71% of patients, side effects in 14.23% and other reasons in 17.86% of patients]. Moreover, the two-year Retention Rate was 69.63% ($n=94$), and most of the patients (73%) discontinued the drug for secondary failure. A large percentage of PsA patients discontinuing therapy were at the same time assuming steroid (drug suspenders 34.15% vs still treated 18.09%, $p=0.041$), had higher values of baseline DAPSA (drug suspenders 20.3 still treated 11.06, $p=0.0011$), and were more frequently affected by fibromyalgia (drug suspenders 47.37% vs still treated 21.84%, $p=0.004$). The PCS showed significantly increased levels of PC (drug suspenders 29 vs still treated 16, $p=0.0002$) and all its specific domains: Helplessness (drug suspenders 13 still treated 7, $p=0.0004$), magnification (drug suspenders 4 vs still treated 2, $p=0.0127$) and rumination

(drug suspenders 12.5 vs still treated 6, $p=0.0005$) in PsA patients discontinuing therapy.

To further evaluate the relationship between PC and the drug retention rate of these patients, univariable (Table 3) and multivariable linear regression (Table 4) analysis were also performed.

The univariable linear regression, as reported in Table 3, confirmed fibromyalgia comorbidity, corticosteroids assumption, higher disease activity at baseline DAPSA and incremented PC levels as best predictors for drug suspension, within two years of follow-up. Of note, the multivariable logistic regression (adjusted for age, sex, DAPSA and CCS use) showed significant relationships between drug discontinuation and: high PCS, helplessness domain, rumination domain, but not for magnification domain.

As far as axSpA patients are concerned, the one-year Retention Rate was 84.51%. [Drug suspension related to primary failure in 9.09% of patients, secondary failure in 54.55% of patients, side effects in 27.27% and other reasons in 9.09% of patients]. Moreover, the two-year Retention Rate was 80.28%, and the most frequent cause of suspension (45%) was secondary failure. In axSpA, the drug discontinuation was significantly more frequent in female patients (69.23%, $p=0.04$), in patient with a shorter disease duration (drug suspenders 59 months vs still in treatment 87 months, $p=0.035$), with higher baseline BASDAI disease activity (7 vs 3.75, $p=0.0002$) and higher level of PC (31 vs 17.5, $p=0.0001$). Of note, the univariable logistic regression showed that female gender, increased baseline BASDAI, and higher PCS and all its components: helplessness, magnification, rumination) may be significant predictors of treatment discontinuation in the first two years of follow-up (Table 5).

Due to the limited number of events (discontinuation of csDMARDs and/or bDMARDs at 2 years of follow up), a multivariate analysis was not performed for axSpA participants.

Discussion

Rheumatic conditions, including PsA and axSpA, are characterized by chronic pain and inflammation, which often require long-term pharmacological treatments. The drug retention, in these conditions, is crucial to achieve optimal disease control and a better quality of life (8, 19, 20). However, several factors may influence drug retention, and in this context, the role of PC is still largely unknown [9]. In this study we described, as primary aim, that PC, independent of treatments efficacy, is significantly associated with drug discontinuation both in PsA and axSpA patients. Furthermore, we observed that all the specific PC domains, such as Helplessness, Rumination and Magnification, were significantly

Table 1 Clinical characteristics of PsA participants

	PsA (135)	PsA 1 year Retention (107)	PsA 1 year suspension (28)	p	PsA 2 year Retention (94)	PsA 2 year Suspension (41)	p
Age	56 (47–64)	55 (46–63)	61 (51.5–65)	0.11	55.5 (46–62)	61 (48–65)	0.12
Female (%)	59.26	53.27	82.14	0.006	55.32	68.29	0.16
Disease duration (months)	84 (48–144)	80 (48–144)	111.5 (52–150)	0.41	80 (48–144)	97 (51–144)	0.69
BMI	26.6 (24.6–30.2)	26.4 (24.35–30.07)	27.14 (25–31.91)	0.60	26.2 (24.2–29.7)	27.2 (25–31.2)	0.40
Fibromyalgia (%)	29.6	22.22	57.69	< 0.0001	21.84	47.37	0.004
Smokers (%) Yes	23.88	24.53	21.43	0.62	26.88	17.07	0.005
ex	6.7	5.66	10.71		2.15	17.07	
Peripheral arthritis (%)	95.56	94.39	100	0.20	93.62	100	0.098
Axial involvement (%)	48.89	46.73	57.14	0.33	48.94	48.78	0.99
Enthesitis (%)	35.56	35.51	35.71	0.98	36.17	34.15	0.82
Dactylitis (%)	14.07	14.95	10.71	0.57	15.96	9.76	0.34
Psoriasis (%)	74.07	74.44	71.43	0.72	74.47	73.17	0.87
Nail psoriasis (%)	12.59	14.02	7.14	0.33	14.89	7.32	0.22
csDMARDs use (%)	49.63	47.66	57.14	0.37	44.68	60.98	0.08
bDMARDs use(%)	74.07	71.96	82.14	0.27	71.28	80.49	0.26
Ccs (%)	22.96	18.69	39.29	0.02	18.09	34.15	0.04
NSAIDs (%)	34.81	30.84	50.00	0.06	31.91	41.46	0.24
SNRI (%)	7.41	6.54	10.71	0.45	6.38	9.76	0.49
Anticonvulsivant drugs use (%)	8.15	7.48	10.71	0.58	7.45	9.76	0.65
TJ/68	2 (0–6)	1 (0–5)	3.5 (0–9)	0.08	1 (0–4)	4 (0–9)	0.01
SJ/66	0 (0–0)	0 (0–0)	0 (0–1)	0.03	0 (0–0)	0 (0–1)	0.051
PtGA	5 (2–7)	4.5 (2–7)	7 (5–8)	0.002	4 (2–7)	7 (5–8)	0.0003
PP	6 (3–8)	5 (1–7)	8 (6–9)	0.0005	5 (1–7)	8 (5–9)	< 0.0001
EGA	1 (0–2)	1 (0–2)	2 (1–3)	0.012	0.5 (0–2)	2 (1–3)	0.001
CRP	0.37 (0.14–0.8)	0.3 (0.12–0.64)	0.45 (0.31–0.92)	0.0351	0.315 (0.13–0.73)	0.4 (0.21–0.87)	0.33
DAPSA	13.34 (5.21–22.2)	11.5 (3.4–21.51)	20.31(11.78–23.25)	0.0065	11.06 (3.36–19.85)	20.3 (11.56–23.86)	0.0011
PCS	18 (6–31)	16 (5–29)	29.5 (17–38.5)	0.0001	16 (5–28)	29 (12–38)	0.0002
Helplessness	9 (2–14)	7 (2.12)	13(9–26)	0.0001	7 (2–11)	13 (5–16)	0.0004
Rumination	8 (2–13)	6 (2–12)	13(7–26)	0.0003	6 (2–11)	12.5 (5–16)	0.0005
Magnification	3 (1–5)	2 (1–5)	5 (2–6)	0.0059	2 (1–4)	4 (2–6)	0.0127
HADS anxiety	7 (3–11)	5 (3–10)	9.5 (7–14.5)	0.0002	5 (3–10)	9 (4.14)	0.003
HADS depression	4 (1–8)	3.5 (1–7)	7 (3–10)	0.0091	3 (1–7)	6 (2–10)	0.02
Acceptance and Action Questionnaire II (AAQ)	14 (9–24)	12 (9–22)	22 (9–30)	0.026	12 (9–22)	20 (9–28)	0.07
Trait Hope Scale (THS)	24 (21–26)	24 (21.5–26)	22 (16.5- 25)	0.0392	24 (22–26)	22 (17–25)	0.02
THS agency	12 (10–13)	12 (10–13)	11 (7.5- 12)	0.0361	12 (11–13)	12 (8–12)	0.1
THS pathway	12 (11–13)	12 (11–14)	11 (8.5–13)	0.0449	12 (11–14)	11 (8.5–13)	0.01

PsA Psoriatic Arthritis, BMI Body Mass Index, csDMARDs conventional synthetic disease-modifying anti-rheumatic drugs, bDMARDs biologic disease-modifying anti-rheumatic drugs, CCS corticosteroids, NSAIDs Non-steroidal anti-inflammatory drugs, SNRI Serotonin-norepinephrine reuptake inhibitor, TJ tender joints, SJ swollen joints, PtGA patient global assessment, PP patient pain, EGA Examiner global assessment, CRP C-reactive protein, DAPSA Disease Activity in Psoriatic Arthritis, PCS Pain Catastrophizing Scale, HADS Hospital Anxiety and Depression Scale

associated with therapy discontinuation after 2 years follow-up. We observed that PsA patients suspending therapy, before the end of the second year of follow

up, were more frequently affected by fibromyalgia and showing higher baseline disease activity, thus suggesting the correlations between fibromyalgia and higher

Table 2 Clinical characteristics of axSpA participants

	axSpA (71)	axSpa 1 year retention (60)	axSpa 1 year suspension (11)	p	axSpa 2 year retention (57)	axSpa 2 year suspension (14)	p
Age	49 (37–58)	51 (38–60)	44 (35–46)	0.10	49 (37–58)	46 (43–56)	0.78
Female (%)	43.66	40.32	66.67	0.13	37.93	69.23	0.040
Disease duration (months)	72 (48–120)	87 (50–132)	48 (18–60)	0.0083	87 (50–132)	59 (21–75)	0.035
BMI	26.04 (23.5–30.4)	26.22 (23.63–30.45)	24.7 (22.6–30.1)	0.61	26.4 (23.9–30.5)	24.14 (20.35–29.6)	0.20
Fibromyalgia (%)	12.5	12.73	11.11	0.89	13.46	8.33	0.63
Smokers (%) yes	28.17	25.81	44.44	0.36	25.86	38.46	0.34
ex	9.86	11.29	0.00		12.07	0.00	
Peripheral arthritis (%)	54.53	0	100	<0.0001	0	100	<0.0001
HLA B27 (%)	43.08	45.61	25	0.27	47.17	25.00	0.16
Positive MRI (%)	74.65	74.19	77.78	0.82	75.86	69.23	0.62
Positive Rx (%)	49.28	50	44.44	0.76	50	46.15	0.80
Active Uveitis (%)	1.41	1.61	0	0.91	1.72	0	0.80
Past uveitis (%)	19.72	19.35	22.22	0.91	20.69	15.38	0.80
csDMARDs use (%)	22.12	12.96	11.76	0.90	25.86	15.38	0.42
bDMARDs use (%)	91.23	42.86	9.38	0.011	93.10	76.92	0.08
Ccs (%)	5.63	6.45	0	0.43	6.9	0	0.33
NSAIDs (%)	47.89	45.16	66.67	0.23	44.83	61.54	0.28
SNRI (%)	2.82	1.61	11.11	0.11	0	15.38	0.002
Tricyclic antidepressant (%)	4.23	4.84	0	0.5	3.45	7.69	0.49
Anticonvulsivant drugs use (%)	8.45	8.06	11.11	0.76	6.9	15.38	0.32
Myorelaxant agent (%)	9.86	8.06	22.22	0.18	8.62	15.38	0.46
Opioid drugs (%)	4.23	3.23	11.11	0.27	3.45	7.69	0.49
LEI	0 (0–0)	0 (0–0)	0 (0–1.5)		0 (0–0)	0 (0–1.5)	
CRP	0.2 (0.08–0.5)	0.14 (0.07–0.44)	0.55 (0.42–2.23)	0.0213	0.13 (0.07–0.43)	0.5 (0.32–1.4)	0.017
ESR	9 (4–19)	8 (4–19)	12(9–14)	0.20	8 (4–18)	14 (9–25)	0.07
BASDAI	4.17 (2.1–6.3)	3.88 (2–5.8)	7 (6–9)	0.0023	3.75 (2–5.7)	7 (6–8.8)	0.0002
PtGA	5 (3–8)	4.5 (2.5–7)	8 (7–10)	0.0081	4 (2–75)	8 (7–10)	0.0014
PhyGA	1 (0–3)	1(0–3)	3 (1–6.5)	0.23	1 (0–3)	1 (25–6.5)	0.14
ASDAS-PCR	2.32 (1.29–3.16)	2.25 (1.25–2.99)	3.37 (2.48–4.12)	0.0192	2.19 (1.24–2.84)	3.35 (2.45–3.91)	0.0042
BASFI	5.7 (0.3–16)	4.9 (0.3–16)	7.9 (6.8–9.5)	0.18	4.9 (0.3–16)	6.8 (4.6–9.5)	0.32
PCS	16 (6–28)	13.5 (6–25.5)	40 (30–44)	0.0002	12.5 (6–25.5)	31 (25–44)	0.0001
Helplessness	7 (2–12)	5 (2–10)	18.5 (13.5–21)	0.0002	5 (2–10)	16 (11–20)	0.0004
Rumination	7 (4–12)	6(3–11)	16 (9.5–18.5)	0.0018	6 (2–11)	13 (7–18)	0.0018
Magnification	2 (0–4)	2 (0–3)	5.5 (2.5–6.5)	0.007	2 (0–3)	4 (2–6)	0.0712
HADS anxiety	7 (4–10)	6 (3–10)	9 (8–11)	0.023	7 (3–10)	9 (5–11)	0.14
HADS depression	5 (2–6)	4 (1–6)	6 (5–10)	0.044	4 (2–6)	5 (3–9)	0.27
Acceptance and Action Questionnaire II (AAQ)	14 (9–22)	13 (9–22)	17(15–21)	0.22	13 (9–22)	17 (13–21)	0.41
Trait Hope Scale (THS)	24 (22–28)	25 (22–28)	22 (19–24)	0.0371	25 (22–28)	22 (20–25)	0.14
THS agency	12 (10–14)	12 (11–14)	10 (9–13)	0.1	12 (11–14)	11 (10–13)	0.33
THS pathway	12 (11–14)	12 (11–15)	11 (10–12)	0.0192	12 (11–15)	12 (10–12)	0.081
Compassionate Engagement and Action Scales – self compassion	74.5 (64–88)	75 (67–90)	68 (63–75)	0.17	75 (67–90)	68 (63–75)	0.09

Table 2 (continued)

	<i>axSpA</i> (71)	<i>axSpa</i> 1 year retention (60)	<i>axSpa</i> 1 year suspension (11)	<i>p</i>	<i>axSpa</i> 2 year retention (57)	<i>axSpa</i> 2 year suspension (14)	<i>p</i>
Compassionate Engagement and Action Scales- compassion from others	66 (52–83)	68.5 (55–83)	61 (52–69)	0.32	69 (55–83)	62 (52–69)	0.22

axSpA axial Spondyloarthritis, *BMI* Body Mass Index, *MRI* Magnetic Resonance Imaging, *csDMARDs* conventional synthetic disease-modifying anti-rheumatic drugs, *bDMARDs* biologic disease-modifying anti-rheumatic drugs, *CCS* corticosteroids, *NSAIDs* Non-steroidal anti-inflammatory drugs, *SNRI* Serotonin–norepinephrine reuptake inhibitor, *LEI* Leeds enthesitis Index, *CRP* C-reactive protein, *ESR* erythrocyte sedimentation rate, *BASDAI* Bath Ankylosing Spondylitis Disease Activity Index, *PtGA* patient global assessment, *PhyGA* Physician global assessment, *ASDAS* Ankylosing Spondylitis Disease Activity Score, *BASFI* Bath Ankylosing Spondylitis Function Index, *PCS* Pain Catastrophizing Scale, *HADS* Hospital Anxiety and Depression Scale

Table 3 Univariable linear regression (PsA participants)

Two year discontinuation (PsA participants) Independent variables	Univariable		
	OR	95%CI	<i>p</i>
Fibromyalgia	3.22	1.42 7.28	0.005
CCS	2.35	1.02 5.40	0.04
TJ	1.10	1.02 1.18	0.01
PP	1.09	1.00 1.18	0.04
PtGa	1.3	1.12 1.49	<0.0001
DAPSA	1.06	1.02 1.10	0.001
PCS	1.05	1.02 1.08	<0.0001
Helplessness	1.12	1.05 1.19	0.001
Rumination	1.13	1.05 1.21	<0.001
Magnification	1.22	1.05 1.42	0.009

PsA Psoriatic Arthritis, *CCS* corticosteroids, *TJ* tender joints, *SJ* swollen joints, *PP* patient pain, *PtGA* patient global assessment, *DAPSA* Disease Activity in Psoriatic Arthritis, *PCS* Pain Catastrophizing Scale

levels of disease activity with drug discontinuation in PsA patients [21, 22]. Furthermore, in a previous paper, we published that PC specifically correlated with the number of tender joints, patient-reported pain, and patient global assessment, thus, confirming its predominantly

impact on the subjective dimension of the disease activity scores, independent of inflammation [13, 23, 24]. In this setting we recently published how various aspects of the PCS may influence the achievement of low disease activity and remission, as measured by DAPSA and BASDAI. Of note, the significant correlation between PCS and the disease activity of other rheumatic diseases was already described in scientific literature [11, 25–27]. In detail, multivariable linear regression analyses showed that PC and its components helplessness and rumination were independent predictors of drug suspension within two years. In contrast, a predictive relationship was not observed for magnification. This discrepancy may be due to our decision to deliberately exclude individuals with a documented history of psychiatric disorders, as outlined in DSM-V. In fact, it has been reported that magnification shows a significant association with both physical and mental health-related quality of life (QOL) and depressed mood [28]. These results partially mirror what we already published in which females with longer disease durations significantly correlate with drug discontinuation [8]. Nonetheless, another study, despite demonstrating the mentioned relation between female and low retention

Table 4 Multivariable regression (PsA participants)

	OR	95%CI	<i>P</i>		OR	95% CI	<i>P</i>
Age	1.02	0.98–1.05	0.32	Age	1.02	0.99–1.06	0.23
Sex	0.99	0.41–2.40	0.99	Sex	1.02	0.39–2.63	0.97
DAPSA	1.04	0.99–1.08	0.10	DAPSA	1.03	0.99–1.08	0.15
PCS	1.04	1.00–1.07	0.03	Rumination	1.10	1.02–1.20	0.02
CCS	1.98	0.78–4.99	0.15	CCS	1.99	0.76–5.20	0.16
Age	1.02	0.99–1.06	0.20	Age	1.02	0.98–1.06	0.26
Sex	1.17	0.47–2.94	0.72	Sex	1.29	0.53–3.14	0.58
DAPSA	1.03	0.98–1.08	0.20	DAPSA	1.04	1.00–1.09	0.04
Helplessness	1.09	1.01–1.18	0.03	Magnification	1.15	0.97–1.36	0.10
CCS	1.91	0.78–4.95	0.18	CCS	1.87	0.72–4.81	0.20

DAPSA Disease Activity in Psoriatic Arthritis, *PCS* Pain Catastrophizing Scale, *CCS* corticosteroids

Table 5 Univariable linear regression (axSpA participants)

Two year discontinuation (AxSpA participants) Independent variables	Univariable		
	OR	95%CI	p
Sex	3.68	1.01 13.40	0.04
BASDAI	1.81	1.28 2.57	0.001
Asdas CRP	2.49	1.30 4.72	0.006
PCS	1.13	1.05 1.20	< 0.0001
Helplessness	1.24	1.09 1.40	0.001
Rumination	1.25	1.07 1.45	0.004
Magnification	1.35	1.02 1.78	0.03

axSpA axial Spondyloarthritis, BASDAI Bath Ankylosing Spondylitis Disease Activity Index, ASDAS Ankylosing Spondylitis Disease Activity Score, PCS Pain Catastrophizing Scale

rate for anti-TNFi assumption, did not found differences in both genders for methotrexate [29].

For axSpA, drug suspension was significantly associated with higher levels of PC, female gender, shorter disease duration, as well as higher baseline BASDAI. Univariable logistic regression analysis strongly confirmed that female gender, higher baseline BASDAI, and higher PC levels, along with its components, were predictors of treatment suspension within two years. Available literature already reported a higher retention rate in axSpA male patients, as well as the relationship between drug withdrawal and higher BASDAI levels [30–32].

Conclusions

Taking together all these data, we may assume that PC is not just a psychological response to pain but may have important consequences on the patient’s ability to adhere to treatments and a better knowledge of this problem may help healthcare providers in managing patients with rheumatic conditions.

We are aware that our study has some limitations including the relatively low number of participants, which does not allow us to derive robust conclusion; on the other hand, despite the limitations, the screening for any previous psychological intervention in our participants allow us to select a well-defined cohort to assess the role of PC. Moreover, our Italian multicenter approach had the advantage to grant a sample representative of the whole national population; these results could therefore be generalized to Caucasian populations.

In conclusion, PC has emerged as a significant factor affecting drug retention in PsA and axSpA patients.

These findings highlight the critical need for clinicians to assess and address PC to improve treatment outcomes. Given the potential impact on patient management and quality of life, further research is essential to explore the mechanisms underlying this relationship and to develop

targeted interventions. Future studies should also examine the generalizability of these results to diverse populations.

Abbreviations

SPA	Spondyloarthritis
PsA	Psoriatic Arthritis
axSpA	Axial Spondyloarthritis
csDMARDs	Conventional synthetic disease-modifying anti-rheumatic drugs
bDMARDs	Biologic disease-modifying anti-rheumatic drugs
PC	Pain catastrophizing
PCS	Pain Catastrophizing Scale
CSQ	Coping Strategies Questionnaire
CASPAR	Classification Criteria for Psoriatic Arthritis
ASAS	Assessment of Spondyloarthritis International Society)
DAPSA	Disease Activity for Psoriatic Arthritis
MDA	Minimal Disease Activity
BASDAI	Bath Ankylosing Spondylitis Disease Activity Index
PASI	Psoriasis Area Severity Index
LEI	Leeds Enthesitis Index
BASFI	Bath Ankylosing Spondylitis Functional Index
ASDAS-CRP	Ankylosing Spondylitis Disease Activity Score-C-Reactive Protein
BMI	Body Mass Index
CCS	Corticosteroids
NSAIDs	Non-steroidal anti-inflammatory drugs
SNRI	Serotonin–norepinephrine reuptake inhibitor
TJ	Tender joints
SJ	Swollen joints
PtGA	Patient global assessment
PP	Patient pain
EGA	Examiner global assessment
CRP	C-reactive protein
HADS	Hospital Anxiety and Depression Scale

Acknowledgements

None.

Authors’ contributions

All authors made substantial contributions to the conception or design of the work, the acquisition, and interpretation of data. All authors contributed to the critical review and revision of the manuscript and approved the final version. All the authors agreed to be accountable for all aspects of the work. DC study design, data acquisition, statistical analysis, interpretation of data, writing of the first draft of the paper; FS study design, data acquisition, statistical analysis, interpretation of data, writing of the first draft of the paper; PR study design, data acquisition, interpretation of data; AM study design, data acquisition, interpretation of data; IP study design, data acquisition, interpretation of data; MV study design, interpretation of data; OB study design, interpretation of data, statistical analysis, interpretation of data, writing of the first draft of the paper; VP study design, data acquisition, interpretation of data; CDV study design, data acquisition, interpretation of data; FC study design, data acquisition, interpretation of data; LC study design, data acquisition, interpretation of data; MT study design, data acquisition, interpretation of data; FC study design, data acquisition, interpretation of data; FM study design, data acquisition, interpretation of data; FDV study design, data acquisition, interpretation of data; AC study design, data acquisition, interpretation of data; LS study design, data acquisition, interpretation of data; AR study design, data acquisition, interpretation of data; MV study design, data acquisition, interpretation of data; EC study design, data acquisition, interpretation of data; LK study design, data acquisition, interpretation of data; FT study design, data acquisition, interpretation of data; AP study design, data acquisition, interpretation of data; LL study design, data acquisition, interpretation of data; FPC study design, data acquisition, interpretation of data; FP study design, data acquisition, interpretation of data; GG study design, data acquisition, interpretation of data; RS study design, data acquisition, interpretation of data; PC study design, data acquisition, interpretation of data; FC study design, data acquisition, interpretation of data; RG study design, data acquisition, interpretation of data, interpretation of data, writing of the first draft of the paper; LN study design,

data acquisition, interpretation of data, interpretation of data, writing of the first draft of the paper.

Funding

The authors received no specific funding for this work.

Availability of data and materials

The datasets used and analysed during the current study are available from the corresponding author on reasonable request.

Data availability

No datasets were generated or analysed during the current study.

Declarations

Ethics approval and consent to participate

The study received approval from the Ethics Committee of the University Campus Bio-Medico of Rome (approval no. 78.20 OSS) and was conducted in accordance with the Declaration of Helsinki and its subsequent revisions.

Consent for publication

Not applicable.

Competing interests

The authors declare no competing interests.

Author details

¹Clinical and Research Section of Rheumatology and Clinical Immunology, Fondazione Policlinico Universitario Campus Bio-Medico, Rome, Italy. ²Department of Medicine, School of Medicine, Rheumatology and Clinical Immunology, University of Rome "Campus Bio-Medico", Rome, Italy. ³Department of Biotechnological and Applied Clinical Sciences, Rheumatology Unit, University of L'Aquila, L'Aquila, Italy. ⁴Department of Precision Medicine, Rheumatology Unit, University of Campania L. Vanvitelli, Caserta, Italy. ⁵Department of Clinical Medicine and Surgery, Rheumatology Research Unit, University of Naples Federico II, Naples, Italy. ⁶Department of Health Promotion, Mother and Child Care, Rheumatology Section, Internal Medicine and Medical Specialties, University Hospital "P. Giaccone", Palermo, Italy. ⁷Department of Biomedical Science and Human Oncology (DIMO), Rheumatic and Systemic Autoimmune Diseases Unit, University of Bari Medical School, Bari, Italy. ⁸Department of Human Sciences, European University of Rome, Rome, Italy. ⁹Department of Medical and Surgical Sciences, Rheumatology Clinic, University of Foggia, Rione Biccarei, Foggia, FG 71122, Italy.

Received: 18 March 2024 Accepted: 3 September 2024

Published online: 18 September 2024

References

- Dougados M, Baeten D. Spondyloarthritis. *Lancet Lond Engl*. 2011;377(9783):2127–37.
- FitzGerald O, Ogdie A, Chandran V, Coates LC, Kavanaugh A, Tillett W, et al. Psoriatic arthritis. *Nat Rev Dis Primer*. 2021;7(1):59.
- Haque N, Lories RJ, de Vlam K. Comorbidities associated with psoriatic arthritis compared with non-psoriatic spondyloarthritis: a cross-sectional study. *J Rheumatol*. 2016;43(2):376–82.
- Scriffignano S, Perrotta FM, De Socio A, Lubrano E. Role of comorbidities in spondyloarthritis including psoriatic arthritis. *Clin Rheumatol*. 2019;38(1):3–10.
- Navarini L, Caso F, Costa L, Currado D, Stola L, Perrotta F, et al. Cardiovascular risk prediction in ankylosing spondylitis: from traditional scores to machine learning assessment. *Rheumatol Ther*. 2020;7(4):867–82.
- Navarini L, Margiotta DPE, Caso F, Currado D, Tasso M, Angeletti S, et al. Performances of five risk algorithms in predicting cardiovascular events in patients with Psoriatic Arthritis: An Italian bicentric study. *PLoS ONE*. 2018;13(10):e0205506.
- Caso F, Navarini L, Carubbi F, Picchianti-Diamanti A, Chimenti MS, Tasso M, et al. Mediterranean diet and Psoriatic Arthritis activity: a multicenter cross-sectional study. *Rheumatol Int*. 2020;40(6):951–8.
- Navarini L, Costa L, Tasso M, Chimenti MS, Currado D, Fonti GL, et al. Retention rates and identification of factors associated with anti-TNF α , anti-IL17, and anti-IL12/23R agents discontinuation in psoriatic arthritis patients: results from a real-world clinical setting. *Clin Rheumatol*. 2020;39(9):2663–70.
- Linton SJ, Nicholas MK, MacDonald S, Boersma K, Bergbom S, Maher C, et al. The role of depression and catastrophizing in musculoskeletal pain. *Eur J Pain Lond Engl*. 2011;15(4):416–22.
- Quartana PJ, Campbell CM, Edwards RR. Pain catastrophizing: a critical review. *Expert Rev Neurother*. 2009;9(5):745–58.
- Wilk M, Łosińska K, Pripp AH, Korkosz M, Haugeberg G. Pain catastrophizing in rheumatoid arthritis, psoriatic arthritis and axial spondyloarthritis: biopsychosocial perspective and impact on health-related quality of life. *Rheumatol Int*. 2022;42(4):669–82.
- Osman A, Barrios FX, Kopper BA, Hauptmann W, Jones J, O'Neill E. Factor structure, reliability, and validity of the Pain Catastrophizing Scale. *J Behav Med*. 1997;20(6):589–605.
- Currado D, Biaggi A, Pilato A, Marino A, Ruscitti P, Pantano I, et al. The negative impact of pain catastrophizing on disease activity: analyses of data derived from patient-reported outcomes in psoriatic arthritis and axial spondyloarthritis. *Clin Exp Rheumatol*. 2023;41(9):1856–61.
- Taylor W, Gladman D, Helliwell P, Marchesoni A, Mease P, Mielants H, et al. Classification criteria for psoriatic arthritis: development of new criteria from a large international study. *Arthritis Rheum*. 2006;54(8):2665–73.
- Sieper J, Rudwaleit M, Baraliakos X, Brandt J, Braun J, Burgos-Vargas R, et al. The Assessment of SpondyloArthritis international Society (ASAS) handbook: a guide to assess spondyloarthritis. *Ann Rheum Dis*. 2009;68 Suppl 2:ii1–44.
- Mease PJ. Measures of psoriatic arthritis: Tender and Swollen Joint Assessment, Psoriasis Area and Severity Index (PASI), Nail Psoriasis Severity Index (NAPSI), Modified Nail Psoriasis Severity Index (mNAPSI), Mander/Newcastle Enthesitis Index (MEI), Leeds Enthesitis Index (LEI), Spondyloarthritis Research Consortium of Canada (SPARCC), Maastricht Ankylosing Spondylitis Enthesis Score (MASES), Leeds Dactylitis Index (LDI), Patient Global for Psoriatic Arthritis, Dermatology Life Quality Index (DLQI), Psoriatic Arthritis Quality of Life (PsAQOL), Functional Assessment of Chronic Illness Therapy-Fatigue (FACIT-F), Psoriatic Arthritis Response Criteria (PsARC), Psoriatic Arthritis Joint Activity Index (PsAJAI), Disease Activity in Psoriatic Arthritis (DAPSA), and Composite Psoriatic Disease Activity Index (CPDAI). *Arthritis Care Res*. 2011;63(Suppl 11):S64–85.
- Bazzichi L, Giacomelli C, Consensi A, Giorgi V, Batticciotto A, Di Franco M, et al. One year in review 2020: fibromyalgia. *Clin Exp Rheumatol*. 2020;38 Suppl 123(1):3–8.
- Wolfe F, Clauw DJ, Fitzcharles MA, Goldenberg DL, Häuser W, Katz RL, et al. 2016 Revisions to the 2010/2011 fibromyalgia diagnostic criteria. *Semin Arthritis Rheum*. 2016;46(3):319–29.
- Armağan B, Kılıç L, Farisoğulları B, Yardımcı GK, Bilgin E, Bölek EÇ, et al. Drug retention rate and predictive factors of drug survival for secukinumab in radiographic axial spondyloarthritis. *Rheumatol Int*. 2023;43(1):147–56.
- Athanassiou P, Kotrotsios A, Kallitsakis I, Bounas A, Dimitroulas T, Garyfallos A, et al. The effects of golimumab on work productivity and quality of life among work-active axial spondyloarthritis and psoriatic arthritis patients treated in the routine care in Greece: the 'GO-UP' study. *Qual Life Res Int J Qual Life Asp Treat Care Rehabil*. 2022;31(5):1385–99.
- Ferrito M, Cincinelli G, Manara M, Di Taranto R, Favalli EG, Caporali R. Retention rate of tumor necrosis factor inhibitors, anti-interleukin 17, and anti-interleukin 12/23 drugs in a single-center cohort of psoriatic arthritis patients. *Reumatismo*. 2023;75(2).
- Mease PJ, Collier DH, Saunders KC, Li G, Kremer JM, Greenberg JD. Comparative effectiveness of biologic monotherapy versus combination therapy for patients with psoriatic arthritis: results from the Corrona registry. *RMD Open*. 2015;1(1):e000181.
- Navarini L, Currado D, Pilato A, Marino A, Biaggi A, Di Donato S, et al. Association between Patient Acceptable Symptom State and disease activity in psoriatic arthritis is disrupted by confounders, including comorbid fibromyalgia. *Clin Exp Rheumatol*. 2024;42(6):1150–5.

24. Navarini L, Currado D, Caso F, Costa L, Chimenti MS, D'Antonio A, et al. Duration of clinical remission and low disease activity impacts on quality of life and its domains in psoriatic arthritis patients: results from an Italian multicentre study. *Clin Exp Rheumatol*. 2022;40(7):1285–92.
25. Penhoat M, Saroux A, Le Goff B, Augereau P, Maugars Y, Berthelot JM. High pain catastrophizing scores in one-fourth of patients on biotherapy for spondylarthritis or rheumatoid arthritis. *Joint Bone Spine*. 2014;81(3):235–9.
26. Hayashi K, Morishima T, Ikemoto T, Miyagawa H, Okamoto T, Ushida T, et al. Pain Catastrophizing is independently associated with quality of life in patients with severe hip osteoarthritis. *Pain Med Malden Mass*. 2019;20(11):2220–7.
27. Kieskamp SC, Paap D, Carbo MJG, Wink F, Bos R, Bootsma H, et al. Central sensitization, illness perception and obesity should be considered when interpreting disease activity in axial spondyloarthritis. *Rheumatol Oxf Engl*. 2021;60(10):4476–85.
28. Craner JR, Gilliam WP, Sperry JA. Rumination, magnification, and helplessness: How do Different aspects of pain catastrophizing relate to pain severity and functioning? *Clin J Pain*. 2016;32(12):1028–35.
29. Generali E, Sciré CA, Cantarini L, Selmi C. Sex differences in the treatment of psoriatic arthritis: a systematic literature review. *Isr Med Assoc J IMAJ*. 2016;18(3–4):203–8.
30. Chimenti MS, Fonti GL, Conigliaro P, Sunzini F, Scrivo R, Navarini L, et al. One-year effectiveness, retention rate, and safety of secukinumab in ankylosing spondylitis and psoriatic arthritis: a real-life multicenter study. *Expert Opin Biol Ther*. 2020;20(7):813–21.
31. Fabbroni M, Cantarini L, Caso F, Costa L, Pagano VA, Frediani B, et al. Drug retention rates and treatment discontinuation among anti-TNF- α agents in psoriatic arthritis and ankylosing spondylitis in clinical practice. *Mediators Inflamm*. 2014;2014:862969.
32. Favalli EG, Selmi C, Becciolini A, Biggioggero M, Ariani A, Santilli D, et al. Eight-year retention rate of first-line tumor necrosis factor inhibitors in spondyloarthritis: a multicenter retrospective analysis. *Arthritis Care Res*. 2017;69(6):867–74.

Publisher's Note

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.