ORIGINAL ARTICLE

The relationship between depressive symptoms, illness perceptions and quality of life in ankylosing spondylitis in comparison to rheumatoid arthritis

Thomas Hyphantis • Konstantinos Kotsis • Niki Tsifetaki • Francis Creed • Alexandros A. Drosos • André F. Carvalho • Paraskevi V. Voulgari

Received: 28 September 2012 / Revised: 27 December 2012 / Accepted: 28 December 2012 / Published online: 18 January 2013 © Clinical Rheumatology 2013

Abstract Anxiety and depressive symptoms as well as cognitive variables are important in determining outcome in rheumatic diseases. We aimed to compare psychological distress symptoms and illness perceptions in ankylosing spondylitis (AS) and rheumatoid arthritis (RA) and to test whether their associations with health-related quality of life (HROoL) were similar in these rheumatologic disorders. In 55 AS and 199 RA patients, we administered the Patient Health Questionnaire (PHQ-9), the Symptom Check-List and the Brief-Illness Perception Questionnaire to assess psychological variables and the World Health Organization Quality of Life Instrument, Short Form to assess HRQoL. We used hierarchical regression analyses to determine the associations between psychological variables and HRQoL after adjusting for demographic variables and disease parameters. The prevalence of clinically significant depressive symptoms (PHO-9 \geq 10) was 14.8 % in AS and 25.1 % in RA patients, but adjustment for demographics rendered these differences in depressive symptoms' severity nonsignificant. Psychological distress levels and HRQoL were

T. Hyphantis (⊠) · K. Kotsis Department of Psychiatry, Medical School, University of Ioannina, Ioannina 45110, Greece e-mail: tyfantis@cc.uoi.gr

N. Tsifetaki · A. A. Drosos · P. V. Voulgari Rheumatology Clinic, Department of Internal Medicine, Medical School, University of Ioannina, Ioannina, Greece

F. Creed Psychiatry Research Group, Medical School, University of Manchester, Manchester, UK

A. F. Carvalho Psychiatry Research Group, Faculty of Medicine,

Federal University of Ceará, Fortaleza, CE, Brazil

similar in both disorders. Illness concern (b=-0.37) was the only significant independent correlate of physical HRQoL in AS. In RA, depression (b=-0.25), illness concern (b=-0.14) and worries about the consequences of the disease (b=-0.31) were the independent correlates of physical HRQoL. These findings suggest that cognitive variables are important correlates of HRQoL in AS, whereas in RA depressive symptoms and illness perceptions equally contribute to HRQoL. Our data encourage the design of psychotherapeutic trials targeting disease-related cognitions in AS in an attempt to improve patient's physical HRQoL.

Keywords Ankylosing spondylitis · Anxiety · Depression · Illness representations · Quality of life

Introduction

Ankylosing spondylitis (AS) is a chronic inflammatory disease of the sacroiliac joints and spine that begins usually in young adulthood and is more common in men. Prevalence among white populations varies from 0.5 to 1.0 % [1]. In our catchment area, the prevalence is 29.5/100,000 adults [2]. Apart from the axial and articular manifestations, AS may present with extra-articular manifestations such as enthesitis and acute anterior uveitis; involvement of other systems may also occur, such as pulmonary and cardiovascular involvement, neurologic lesions, inflammatory bowel disease and psoriasis. AS causes severe pain and disability; a significant number of patients have spinal osteoporosis, resulting in vertebral fractures and thoracic kyphosis [1], while a remarkable proportion require assistance at work or even withdraw from the workforce [3]. The high health care utilization [4] and the impaired health-related quality of life (HRQoL) compared to healthy controls [5, 6] highlight the disease's burden. Therefore, the identification of treatable psychosocial factors associated with HRQoL is a research priority, which could ultimately lead to the development of beneficial psychological interventions.

Evidence suggests that patients with rheumatologic disorders have high levels of psychological distress, especially depressive symptoms, which significantly impair their HRQoL [4, 7]. The prevalence of current major depressive disorder (MDD) in rheumatologic disorders ranges between 17 and 22.5 % [8, 9], but the prevalence of lifetime MDD is even higher, up to 47 % [10]. Although depressive symptoms make unique contributions to health outcomes in rheumatologic patients, few studies have investigated the prevalence and severity of depressive and other symptoms of psychological distress in AS. A recent study [11], using the Zung Self-Rating Depression Scale (ZDS) [12], reported that 27.4 % of AS patients presented clinically significant depressive symptoms, while in other studies the prevalence of depression ranged from 20 to 31 % [13-15]. However, no study has investigated the prevalence and severity of depressive symptoms using validated instruments based on the Diagnostic and Statistical Manual of Mental Disorders (DSM-IV) diagnostic criteria [16], such as the Patient Health Questionnaire (PHQ-9) [17], which we have shown to be a valid MDD screening inventory in rheumatologic populations [18].

Cognitive variables are also important in determining outcome in rheumatologic patients [8, 9]. Patients develop representations of their illness to make sense of and adapt to the difficulties their illness causes. These representations are organized in various components and may influence patients' ability to cope [19]. Each patient creates his/her own ideas about the identity, cause, timeline or the consequences of the illness and beliefs about its cure and controllability. Several illness perceptions are associated with HRQoL in rheumatoid arthritis (RA) [8, 9], systemic sclerosis [20] or psoriatic arthritis [21]. To the best of our knowledge, however, no study has investigated illness perceptions in AS and their associations with HRQoL. Furthermore, we have shown that the effect of anxiety, depression and illness perceptions on physical HRQoL is rather complex and differs between psoriatic arthritis and RA [21]. We examine here whether the same applies to AS patients. This study aimed to test the following hypotheses:

- That AS patients present remarkable levels of clinically significant psychological distress symptoms, at least similar to RA patients;
- That both psychological distress (i.e. anxiety and depressive symptoms) and illness perceptions are independent correlates of impaired HRQoL in AS patients, after adjusting for age, gender, education and disease's parameters;

 That the effect of anxiety, depression and illness perceptions on physical HRQoL differs between AS and RA, given the different clinical presentations and physical consequences of these diseases.

Methods

Participants and procedures

Data were collected during the baseline assessment of the cohort study "Psychological factors and HROoL in rheumatologic diseases" in consecutive patients treated at the Rheumatology Department, University Hospital of Ioannina, Greece, which provides secondary and tertiary care for 350,000 inhabitants. The main objective of this ongoing prospective cohort is to evaluate treatment response and to determine the independent contribution of psychosocial variables in the formation of HRQoL in patients with rheumatologic disorders 3 years later. In the present cross-sectional study, we recruited a sample of AS attending a follow-up clinic during the 6-month period of 1 April 2010 to 1 October 2010. Diagnosis of AS was based on the modified New York criteria [22]. Exclusion criteria were inability to read and write Greek, history of psychotic illness, alcohol and/or drug abuse or dementia. Patients were examined and assessed for eligibility by experienced rheumatologists (NT and PVV). After complete explanation of the study to 65 eligible patients, 55 agreed to participate (response rate, 84.6 %). No statistically significant differences were found in age between participants and non-participants.

To test whether AS patients' psychological distress levels, illness perceptions and HRQoL differed from that of patients with another rheumatologic disease, 199 consecutive RA patients attending the same department during the same time period served as disease controls. RA was chosen because it shares similarities with AS and was used for comparison in previous studies evaluating HRQoL in AS [5, 23, 24]. RA diagnosis was confirmed using the American College of Rheumatology criteria [25].

The procedures complied with the ethical standards on human experimentation (World Medical Association, Helsinki Declaration). The study protocol was approved by the hospital's ethics committee. Signed informed consent was obtained from all participants.

Measures

Socio-demographic and clinical data were collected, and medical records were reviewed for coexisting medical conditions. The current use of anti-inflammatory, antirheumatic or antidepressant agents was also recorded. The severity of AS was defined using the Bath Ankylosing

Spondylitis Disease Activity Index (BASDAI), a composite score ranging from 0 to 10 based on the grading of fatigue, spinal pain, joint pain/swelling, areas of localized tenderness and morning stiffness; lower score represents less severe disease activity [26]. RA disease activity was estimated by disease activity for 28-joint indices score (DAS-28) [27]. The same rheumatologist (NT) assessed DAS-28 and BASDAI. Patients recorded the severity of pain on a visual analogue scale (VAS) from 0 to 100; higher scores indicate more severe pain. Psychological data were collected via a semi-structured interview by the same mental health professional (KK). All patients with PHO-9 scores indicating clinically significant depressive symptoms (i.e. PHQ-9≥10) were referred to the outpatient department of psychiatry for further assessment and where clinically indicated, treatment. The following selfreported questionnaires were administered:

- 1. Patient Health Questionnaire (PHQ-9): Depressive symptoms' severity was assessed using the PHQ-9, a brief screen for depression whose specific items establish DSM-IV criteria-based diagnosis of MDD. It rates the frequency of symptoms over the past 2 weeks on a 0-3 Likert-type scale; summed scores range from 0 to 27 [17], yielding an index of depressive symptom severity ranging from no depression to severe depression (Table 1). We have shown that, at a threshold of 10 or more, PHQ-9 has a sensitivity of 81.2 % and specificity of 86.8 % in diagnosing MDD among Greek rheumatologic patients [18]. A definitive MDD diagnosis can only be established with a structured interview or a full clinical examination; therefore, we used here the cutoff of ≥ 10 to define patients with "clinically significant depressive symptoms".
- Symptom Distress Checklist (SCL-90-R): To assess specific psychopathological profiles, the SCL-90-R was used. This 90-item multidimensional self-report symptom inventory measures a broad range of psychological distress symptoms in psychiatric and medical patients, namely: anxiety, depression, somatization, hostility, obsessive-compulsiveness, interpersonal sensitivity, phobic anxiety, paranoid ideation and psychoticism [28]. This instrument is also useful for the assessment of treatment response [29]. Higher scores indicate more severe symptoms. The SCL-90-R was chosen because it has been extensively validated in rheumatology samples [30] and is standardized for use in the Greek population [31].
- Brief Illness Perception Questionnaire (B-IPQ): Illness perceptions were measured with the Greek version of the B-IPQ [32]. The B-IPQ uses a single-item approach to assess perceptions on a scale from 0 to 10. Five items assess cognitive illness representations: consequences (how much does your illness affect your life?); timeline (how long do you think your illness will continue?);

personal control (how much control do you feel you have over your illness?); treatment control (how much do you think your treatment can help your illness?) and identity (how much do you experience symptoms from your illness?). Two items assess emotional representations: concern (how concerned are you about your illness?) and emotions (how much does your illness affect you emotionally?). One item assesses illness comprehensibility (how well do you feel you understand your illness?). The last question is an open-ended response item in which patients list the three most important causal factors for their illness. Responses can be grouped into categories, such as: stress, lifestyle, hereditary, etc. [32, 33].

Health-related quality of life

World Health Organization (WHO) defines quality of life (QoL) as the individual's perceptions of their position in life in the context of the culture and value systems in which they live and in relation to their goals, expectations, standards and concerns [34]. This definition is surrounded by conceptual limitations since it reflects the view that OoL refers to a subjective evaluation that is embedded in a cultural, social and environmental context. In an effort to overcome these limitations, WHO introduced the WHOQOL-100, a crossculturally valid assessment which is available in several culture-specific and language-specific versions [35]. Its short form, the World Health Organization Quality of Life Instrument, Short-Form (WHOQOL-BREF) was developed for use when time is restricted and when respondent's burden must be minimised. The WHOQOL-BREF has similar psychometric properties to the WHOQOL-100 [34]. WHOQOL-BREF assesses four domains: physical, mental, social relations and environment HRQoL. It also includes facets assessing overall OoL and satisfaction with general health. Each item is rated on a five-point Likert scale, and scores are transformed to a 0-100 scale. Higher scores indicate better HRQoL. The WHOQOL-BREF was chosen because data from 23 countries, including Greece, showed that it is a valid instrument for the assessment of HRQoL across different cultures [34] and has adequate test-retest reliability, internal consistency and factor structure in rheumatologic populations [36]. Furthermore, the WHOQOL-BREF has been used to assess interventions aimed at improving HRQoL in patients with chronic medical conditions [37]. Finally, this instrument has been standardized for use within the Greek population [38].

A major concern in assessing HRQoL is that psychological distress is highly correlated with certain aspects of HRQoL. Thus, it has been suggested that psychological distress symptoms should always be considered thoroughly when assessing HRQoL to take the measurement overlap into account [39]. Since we showed that psychological distress mediates the relationship of psychosocial variables Table 1Demographic and clinical characteristics, causal representations and univariatecomparisons between ankylosing spondylitis (AS) and rheumatoid arthritis (RA)

	AS (N=55) values	RA (N=199) values	d	р
Demographics				
Age (years) (mean±SD)	42.9±10.9	55.2±13.6	-0.941	< 0.001 ^a
Years of education (mean±SD)	11.0 ± 3.5	8.0±4.2	0.739	<0.001 ^a
Female gender, $N(\%)$	8 (14.5 %)	164 (82.4 %)	-1.493	<0.001 ^b
Divorced/widowed/separated, N (%)	1 (1.8 %)	19 (9.7 %)	-0.357	0.08 ^b
Clinical parameters				
Disease duration (years) (mean±SD)	15.3±11.5	13.7 ± 10.5	0.149	$0.479^{\rm a}$
Pain (VAS-pain) (mean±SD)	$0.88 {\pm} 0.77$	$0.94 {\pm} 0.88$	-0.070	$0.752^{\rm a}$
RA DAS-28 (mean±SD)	_	2.72 ± 0.96	_	—
AS BASDAI (mean±SD)	1.97 ± 1.38	_	_	—
Comorbidity (N, %)				
Coronary artery disease	1 (1.8 %)	3 (1.5 %)	_	_
Thyroid disease	_	1 (0.5 %)	_	_
Diabetes	_	1 (0.5 %)	_	_
Myasthenia	_	1 (0.5 %)	_	_
Drug treatment (N, %)				
Disease-modifying anti-rheumatic drugs (methotrexate, leflunomide, hydoxyclorocine, cyclosporine A, sulfasalazine)	0 (0.0 %)	170 (85.4 %)	-0.779	0.001 ^b
Corticosteroids	0 (0.0 %)	16 (8.0 %)	-0.304	0.003 ^b
Biological agents				
(Etanercept, infliximab, adalimumab)	49 (89.1 %)	45 (22.6 %)	1.477	0.001 ^b
Antidepressants	0 (0.0 %)	14 (7.2 %)	-0.266	0.045 ^b
Patients' perceptions regarding the causal factors in their illness (<i>N</i> , %) (Causal representation item of B-IPQ)			0.496	0.03 ^b
Psychological	14 (25.4 %)	77 (59.7 %)	0.190	0.05
Risk factors (e.g. diet, lack of exercise)	35 (63.6 %)	36 (27.9 %)		
Immunity	4 (7.3 %)	14 (10.9 %)		
Depression (PHQ-9)	+ (7.3 70)	14 (10.9 70)	-0.449	0.001 ^b
Score 0 (no depression)	13 (24.1 %)	10 (5.0 %)	0.115	0.001
Score 1–4 (minimal depression)	19 (35.2 %)	81 (40.7 %)		
Score 5–9 (mild depression)	19 (33.2 %)	58 (29.1 %)		
Score 10–14 (moderate depression)	7 (13.0 %)	36 (18.1 %)		
Score 15–19 (moderate depression)	0 (0.0 %)	10 (5.0 %)		
Score 20–27 (severe depression)	0 (0.0 %) 1 (1.9 %)	4 (2.0 %)		
PHQ-9≥10	1 (1.9 %) 8 (14.8 %)	4 (2.0 %) 50 (25.1 %)	-0.279	0.09 ^b

d Cohen's d as the effect size; BASDAI AS severity index, DAS-28 RA disease activity for 28-joint indices score, B-IPQ Brief Illness Perception Questionnaire, HAQ-DI Health Assessment Questionnaire– Disability Index, PHQ-9 Patient Health Questionnaire 9 ^aTwo tailed t tests

^bChi-square tests

with physical HRQoL in RA [40], we used physical HRQoL as an outcome measure. This domain also showed greater standardized response mean than other scales measuring disability in RA and may have increased statistical power in detecting change in rheumatologic patients [36].

Statistical analysis

The Statistical Package for the Social Sciences (SPSS) 15.0 (SPSS Inc., Chicago, IL, USA) for Windows was used.

Summary statistics for all variables were calculated. Kolmogorov–Smirnov test for normality showed that parametric statistics were appropriate for the main independent variables and the outcome variable. Chi-square analyses for categorical data and two-tailed *t* tests for continuous data were carried out to assess the differences in major demographic and clinical variables between AS and RA. To assess differences in psychological distress, illness perceptions and HRQoL between AS and RA patients, one-way analyses of variance (ANOVA) were performed followed by one-way analyses of covariance (ANCOVA) adjusted for age, sex, education and medications.

To test whether the associations between psychological distress and illness perceptions with physical HRQoL differed between AS and RA, we used hierarchical multiple regression analyses. First, we conducted univariate correlations between the psychological variables and physical HRQoL. Selection of independent variables for subsequent multivariate hierarchical regression analyses was based on the results of these analyses. As there were many significant illness perception components, preliminary separate stepwise multiple regression analyses were performed with dependent variable the physical HRQoL and independent variables the eight components of B-IPQ to determine the most important illness perceptions components associated with physical HRQOL in each disease.

A series of hierarchical models of regression analysis was next performed separately for AS and RA, with the dependent variable being the physical HRQoL. Demographic variables were entered in step 1, disease parameters in step 2 and the most significant illness perceptions in step 3. SCL-90-R anxiety and depression symptoms were entered in final step 4. The magnitude of the R^2 change at each step was used to determine the variance explained by each set of variables. Collinearity between independent variables was tested based on variance inflation factors (VIF) and tolerances for individual variables. Although significant, the emotional response component of B-IPQ was not included in hierarchical analyses as it showed high collinearity with the depression subscale of SCL-90-R. After this, all tolerance values were greater than 0.2 and all VIFs less than 2, indicating that multicollinearity was not biasing the regression models [41].

Results

Patient's characteristics

The median disease duration of AS patients was 12 years. The proportion of physical comorbidities was low in both diseases (2.3–2.5 %; Table 1), and disease activity was low. Most patients were on medications (AS=89.1 %; RA= 85.4 %). The use of medications was only weakly associated with physical HRQoL in both AS (r=0.096, p=0.494) and RA (r=-0.093, p=0.202). More AS than RA patients had received biological agents, while the majority of RA patients had received disease-modifying anti-rheumatic drugs (DMARD) or a combination of DMRADs; no AS patient was on antidepressants compared to 17 (8.5 %) RA patients. AS patients were younger and had received more education; the proportion of males was higher within the AS sample. AS and RA patients had similar disease duration

and pain levels (Table 1). Given these differences, all analyses performed to compare AS and RA were further adjusted for age, gender, education and medication.

Prevalence of clinically significant depressive symptoms and dimensions of psychological distress

As shown in Table 1, eight (14.8 %) AS patients presented PHQ-9 scores indicative of clinically significant depressive symptoms (PHQ-9≥10), compared to 50 (25.1 %) RA patients, but this difference failed to reach statistical significance (p=0.09). In general, however, AS patients presented less severe depressive symptoms than RA patients, as indicated by the SCL-90-R depression subscale (p=0.012) and the PHQ-9 total score (p=0.022; Table 2). However, adjustment for age, gender and education rendered these differences non-significant (Table 2). Furthermore, there was no significant difference between AS and RA patients in the other SCL-90-R psychological distress symptom dimensions (Table 2).

Illness perceptions and HRQoL

The AS patients attributed fewer somatic symptoms to their disease (identity), and they believed their disease had fewer consequences compared to RA patients, but adjustment for age, gender, education and medication also rendered these differences non-significant (Table 2). A significant proportion of AS patients (64.0 %) attributed their illness to various risk factors (e.g. diet, lack of exercise). On the other hand, RA patients considered their disease mainly a result of psychological factors (59.7 %). There were no other significant differences between AS and RA patients in illness perceptions (Table 2).

Participants with AS had better physical HRQoL (including experience of physical pain, incapacity to work and get around, and problems with energy and sleep) and better social relations HRQoL (i.e. less dissatisfaction with personal relationships, social support and sexual life) than their RA counterparts, but adjustment for confounders rendered these differences non-significant (Table 2). Mental (i.e. negative emotions like despair, believing that life is meaningless and feeling psychologically incapacitated) and environment HRQoL (i.e. feeling that one's environment is unsafe or unhealthy, lack of financial resources and lack of leisure opportunities), as well as overall HRQoL and general health were also similar in both diseases (Table 2).

Psychological distress, illness perceptions and physical HRQoL

Univariate analyses indicated that among illness perceptions, consequences, timeline, identity, illness concern and

	One-way ANOVA			ANCOVA adjusted for age, sex, years of education and medication				
	AS (N=55)	RA (N=199)	F	р	AS (N=55)	RA (N=199)	F	р
Psychological distress								
Anxiety	$0.49 {\pm} 0.07$	$0.54 {\pm} 0.04$	0.311	0.578	$0.57 {\pm} 0.10$	$0.52 {\pm} 0.04$	0.247	0.620
Depression	$0.66 {\pm} 0.07$	$0.91 {\pm} 0.04$	6.452	0.012 *	$0.85 {\pm} 0.10$	$0.87 {\pm} 0.04$	0.022	0.883
Somatization	$0.84 {\pm} 0.10$	$0.90{\pm}0.05$	0.267	0.606	1.09 ± 0.12	$0.84 {\pm} 0.05$	2.851	0.093
Interpersonal sensitivity	$0.63 {\pm} 0.07$	$0.70 {\pm} 0.04$	0.631	0.428	$0.70 {\pm} 0.10$	$0.69 {\pm} 0.04$	0.009	0.925
Hostility	$0.62 {\pm} 0.09$	$0.49 {\pm} 0.04$	2.324	0.129	$0.55 {\pm} 0.10$	$0.51 {\pm} 0.04$	0.130	0.718
Phobic anxiety	$0.27 {\pm} 0.05$	$0.28 {\pm} 0.03$	0.041	0.841	$0.35 {\pm} 0.08$	$0.27 {\pm} 0.03$	0.874	0.351
Obsessive-compulsive	0.73 ± 0.10	$0.69 {\pm} 0.05$	0.140	0.708	0.81 ± 0.12	$0.68 {\pm} 0.05$	0.920	0.338
Paranoid ideation	$0.80{\pm}0.09$	$0.66 {\pm} 0.05$	1.564	0.212	$0.86 {\pm} 0.11$	$0.66 {\pm} 0.05$	2.112	0.147
Psychoticism	$0.38 {\pm} 0.06$	$0.27 {\pm} 0.03$	3.416	0.066	$0.36{\pm}0.06$	$0.28 {\pm} 0.03$	0.960	0.328
Illness perceptions								
Consequences	$4.52 {\pm} 0.45$	$5.94 {\pm} 0.21$	9.262	0.003 **	$5.73 {\pm} 0.48$	5.63 ± 0.21	0.030	0.863
Timeline	$8.96 {\pm} 0.27$	$8.88 {\pm} 0.17$	0.049	0.824	9.41 ± 0.38	$8.74 {\pm} 0.17$	2.169	0.142
Personal control	7.15 ± 0.40	$6.40 {\pm} 0.21$	2.758	0.098	$6.51 {\pm} 0.50$	6.55 ± 0.22	0.006	0.939
Treatment control	8.43 ± 0.30	$7.83 {\pm} 0.17$	2.605	0.108	8.22 ± 0.41	$7.84 {\pm} 0.18$	0.711	0.400
Identity	$4.74 {\pm} 0.47$	$5.63 {\pm} 0.19$	4.010	0.046 *	$5.91 {\pm} 0.47$	$5.31 {\pm} 0.21$	1.148	0.285
Illness concern	$5.74 {\pm} 0.50$	$5.65 {\pm} 0.25$	0.028	0.868	$5.98 {\pm} 0.60$	$5.57 {\pm} 0.27$	0.334	0.564
Comprehensibility	$7.80 {\pm} 0.37$	$7.50 {\pm} 0.18$	0.566	0.453	$7.36 {\pm} 0.42$	7.61 ± 0.18	0.252	0.616
Emotions	$5.28 {\pm} 0.45$	$5.97 {\pm} 0.24$	1.782	0.183	$5.64 {\pm} 0.58$	$5.88 {\pm} 0.26$	0.130	0.718
Quality of life								
Physical	$63.83 {\pm} 2.65$	58.12 ± 1.40	3.620	0.050 *	56.37±3.17	60.01 ± 1.43	0.988	0.321
Mental	$69.07 {\pm} 2.48$	64.93±1.26	2.240	0.136	64.90 ± 3.09	65.88±1.36	0.073	0.787
Social relations	67.32 ± 2.68	60.14 ± 1.55	4.696	0.031*	62.81±3.55	61.18±1.55	0.154	0.695
Environment	$60.39 {\pm} 1.78$	63.03 ± 1.12	1.257	0.263	$61.08 {\pm} 2.66$	62.83 ± 1.16	0.313	0.576
Overall QoL	62.27 ± 30.38	62.37±24.2	0.001	0.980	61.02±4.33	62.90±1.95	0.135	0.714
General health	53.64 ± 32.06	$54.85 {\pm} 26.71$	0.081	0.777	52.70±4.75	55.42±2.14	0.235	0.628
PHQ-9 total score	4.61 ± 0.64	6.39±0.36	5.303	0.022 *	$5.58 {\pm} 0.85$	6.17±0.38	0.337	0.562

Table 2 Psychological distress symptoms, illness perceptions and quality of life in ankylosing spondylitis (AS) and rheumatoid arthritis (RA)

*p<0.05; **p<0.01

emotional response were significantly associated with physical HRQoL in both samples (Table 3). Preliminary stepwise multiple regression analyses with physical HRQoL as the dependent variable and all B-IPQ components as independent variables showed that the most significant correlates of physical HRQoL in both diseases were emotional response, consequences and illness concern (Table 4). Based on these results, emotional response, consequences and illness concern were selected for inclusion in the multivariate hierarchical analysis. However, emotional response scores showed high collinearity with SCL-90-R depression score, and it could not be included in the regression equations.

Table 5 shows that, in AS, demographic variables (model 1) and disease parameters (pain and BASDAI score—model 2) explained 38.2 % of the variance (p=0.001). Addition of illness perceptions (model 3) increased the variance by

12.9 % (p=0.001); B-IPQ illness concern was significantly associated with physical HRQoL; BASDAI scores remained significant, but pain severity was no longer significant. Addition of anxiety and depressive symptoms showed that they were not significantly associated with physical HRQoL (model 4). In the final model, illness severity and illness concern were the variables independently associated with AS patients' physical HRQoL.

A similar set of analyses in RA showed that demographic variables (model 1) and disease parameters (longer disease duration, more active disease and pain-model 2) explained 26.6 % of the variance (p<0.001) (Table 6). Addition of illness perceptions in model 3 increased the variance by 25.8 % (p=0.001); both B-IPQ consequences and illness concern were significantly associated with physical HRQoL, but disease duration was no longer significant. Addition of anxiety and depression added an additional

 Table 3
 Univariate comparisons between ankylosing spondylitis and rheumatoid arthritis patients' physical HRQoL and demographic, clinical and psychological variables studied

	Univariate comparisons with physical HRQoL (WHOQOL-BREF)				
	Ankylosing spondylitis (<i>N</i> =55)	Rheumatoid arthritis (<i>N</i> =199)			
Age	-0.427 ***	-0.211 **			
Gender	0.217	0.125			
Education	0.409 **	0.272 ***			
Disease duration	-0.302	-0.247 **			
BASDAI	-0.608 ***	_			
DAS-28	_	-0.167 **			
Pain	-0.474 **	-0.598 ***			
Psychological distress (SCL-90-R)					
Anxiety symptoms	-0.456 ***	-0.470 ***			
Depressive symptoms	-0.478 ***	-0.596 ***			
Illness perceptions (B-IPQ)					
Consequences	-0.640 ***	-0.679 ***			
Timeline	-0.325 *	-0.367 ***			
Personal control	0.249	0.239 ***			
Treatment control	0.006	0.233 ***			
Identity	-0.692 ***	-0.600 ***			
Illness concern	-0.650***	-0.439 ***			
Comprehensibility	0.040	0.058			
Emotions	-0.726***	-0.489 ***			

Values shown are beta regression coefficients based on univariate regression analyses with dependent variable the physical HRQoL *p<0.05; **p<0.01; ***p<0.001

6.6 % (p<0.001). At this step, depressive symptoms were significantly associated with physical HRQoL, but disease activity was no longer significant (model 4). In the final model, pain, B-IPQ consequences and illness concern and

Table 4 Preliminary stepwise multiple regression analyses with dependent variable the physical HRQoL and independent variables the 8 B-IPQ components separately in ankylosing spondylitis (N=55) and rheumatoid arthritis (N=199) patient samples SCL-90-R depressive symptoms were the significant independent correlates of RA patients' physical HRQoL.

Discussion

Our results showed a prevalence of clinically significant depressive symptoms of 14.8 % in AS patients, while an additional 25.9 % experienced mild depressive symptoms as measured by the PHO-9. These rates were lower than those observed in RA (25.1 and 29.1 %, respectively), but adjustment for gender, age and educational level rendered the differences non-significant, confirming our first hypothesis and indicating that these differences could be attributed to socio-demographic factors. The RA sample, for example, comprised mostly women (82.4 %), whereas AS sample included mainly men (85.5 %) as anticipated [1], and a greater female prevalence of depression is a widely replicated epidemiological finding. Like others [8] we found that, in RA, both depressive symptoms and illness perceptions were significant independent correlates of physical HRQoL. Conversely, in the AS sample, the most prominent psychological construct independently associated with HRQoL was illness concern, while symptoms of anxiety and depression had a minor contribution to physical HRQoL. Thus, our second hypothesis was partially rejected; in AS, only illness perceptions were independently associated with HRQOL. Our findings also indicate that the effects of depression and illness perceptions on physical HRQoL are rather complex and differ between AS and RA, thereby confirming our third hypothesis.

The prevalence of clinically significant depressive symptoms in our AS sample (14.8 %) is lower than previously reported figures [13–15]. In a study with 62 AS patients [11], the prevalence of depression estimated by the ZDS was 27.4 %. Others found prevalence rates

	Ankylosing sp	Ankylosing spondylitis		Rheumatoid arthritis		
	beta	p value	beta	p value		
Illness perceptions (B-IPQ))					
Emotions	-0.447	<.001	-0.043	.578		
Illness concern	-0.421	<.001	-0.185	.009		
Consequences	-0.333	.019	-0.447	<.001		
Identity	-0.096	.161	-0.142	.012		
Personal control	0.139	.123	0.082	.115		
Treatment control	-0.108	.267	0.016	.774		
Timeline	-0.099	.388	-0.127	.018		
Comprehensibility	0.018	.924	0.071	.130		
Regression statistics	$R^{2}_{Adj} = 0.507, p$	$R^2_{\rm Adj}$ =0.507, p=.019		<i>p</i> <.001		

Table 5 Hierarchical models ofthe factors associated with thephysical HRQoL in ankylosingspondylitis patients ($N=55$)		Model 1	Model 2	Model 3	Model 4
	Demographics				
	Age, years	-0.321 *	-0.115	-0.021	0.001
	Education	0.293 *	0.264 *	0.152	0.151
	Disease parameters				
	BASDAI score	_	-0.334 **	-0.306 **	-0.264 *
	Pain (VAS-Pain)	_	-0.264 *	-0.142	-0.128
	Illness perceptions				
Independent variables were se- quentially entered in linear re- gression models in which the dependent variable was the physical HRQoL. Values shown are standardized (beta) regres- sion coefficients <i>PASI</i> Psoriasis Area and Severity Index * $p < 0.05$; ** $p < 0.01$; *** $p < 0.001$	Consequences	_	_	-0.112	-0.057
	Illness concern	_		-0.371 ***	-0.370 ***
	Psychological distress				
	Depressive symptoms (SCL-90-R)	_	_	_	-0.066
	Anxiety symptoms (SCL-90-R)	_	_	_	-0.071
	Adjusted R^2 of model	0.228	0.382	0.512	0.520
	Incremental adjusted R^2	0.228	0.154	0.129	0.008
	Significance of F change	< 0.01	0.001	0.001	0.553

of 39.8 % [42] and up to 55.5 % [43], measured with the Hospital Anxiety and Depression Scale [44] in Turkish and Moroccan AS patients, respectively. The different instruments and cutoffs used for assessing 'depression' and the demographic diversities among studies may explain these differences, which underscore the need for studies using structured interviews for assessing MDD in AS. It is worth noting that the prevalence of depressive symptoms in our AS sample rises up to 40.7 % if patients with 'mild' depressive symptoms are considered. Additionally, no AS patient had received treatment for a depressive illness. These findings highlight the under-recognition and undertreatment of depression among rheumatologic patients, and should encourage rheumatologists to assess psychological distress symptoms as an important step in the care of AS patients.

We also found that AS patients had better physical and social relations HRQoL than RA patients, but adjustment for demographic variables rendered these differences non-significant. Overall HRQoL, general health and mental and environment HRQoL were similar in

Table 6 Hierarchical models of the factors associated with the physical HRQoL in rheumatoid arthritis patients (<i>N</i> =199)		Model 1	Model 2	Model 3	Model 4
	Demographics				
	Age, years	-0.058	-0.048	-0.103	-0.077
	Education	0.231 **	0.142	0.075	0.081
	Disease parameters				
	DAS-28	_	-0.189 ***	-0.111 *	-0.073
	Disease duration	_	-0.134 *	-0.048	-0.064
	Pain (VAS Pain)	_	-0.425 ***	-0.199 ***	-0.182 ***
Independent variables were se-	Illness perceptions				
quentially entered in linear regres- sion models in which the dependent variable was the physi- cal HRQoL Values shown are standardized (beta) regression coefficients	Consequences	_	_	-0.441 ***	-0.314 ***
	Illness concern	_	_	-0.226 **	-0.146 **
	Psychological distress				
	Anxiety symptoms (SCL-90-R)	_	_	_	-0.087
	Depressive symptoms (SCL-90-R)	_	_	_	-0.250 ***
DAS-28 Disease activity for 28-joint indices score *p<0.05; **p<0.01; ***p<0.001	Adjusted R^2 of model	0.065	0.266	0.524	0.590
	Incremental adjusted R^2	0.065	0.201	0.258	0.066
	Significance of F change	0.001	0.001	0.001	0.001

both disorders. Other studies found that RA patients presented worse physical but better mental HRQoL than AS patients [5, 6]. Demographic diversities among studies may explain also these differences since direct comparison of the consequences of RA and AS is difficult due to inherently different age and sex distributions [23]. Cultural differences may also exist: a previous study found meaningful differences in AS patients' HRQoL across three European countries [4]. Nevertheless, our findings are consistent with the conclusion of Zink et al. [24] that RA and AS have similar detrimental impacts on well-being.

Our main new finding that AS patients' physical HRQoL was associated with illness concern (i.e. the degree that patients are concerned about their illness) but not with depressive symptoms, in contrast with RA where depressive symptoms and illness consequences (i.e. the degree that patients believe that the illness affect their life) were equally major correlates, indicates that, in AS, cognitive illness representations may play a major role, greater than symptoms of psychological distress. It is possible that AS patients worry about the more severe forms of the disease leading to limited neck movement, increased stiffness of the entire spine, loss of normal posture and development of thoracic kyphosis, disabilities that do not occur to the same extent in RA. The importance of illness concerns is underlined by evidence strongly associating this component with slower return to work following a sick leave [32] given also that the impact of AS on employment and work disability is considerable [3]. To our knowledge, this is the first study to assess illness perceptions and psychological distress in relation to HRQoL in AS. These novel findings may contribute to a better understanding of the factors associated with impaired physical HRQoL in AS and may lead to a better appreciation and management of these treatable aspects of HROoL.

This study's main limitation is its cross-sectional design which prevents solid cause-effect inferences. The generalizability of our findings is also limited by the relatively small AS sample. However, we recruited a remarkable proportion of AS patients with a high response rate (84.6 %), given that between 1983 and 2002, 113 AS cases were diagnosed in our catchment area [2]. Nevertheless, our findings need to be replicated in prospective studies with larger samples. Another drawback is the use of self-report measures only; since depression can only be diagnosed by a structured interview, studies using such interviews are needed to confirm our findings. It is also possible that factors not included here, such as the patients' defensive profile, spirituality or social support, may have interfered with the results. Finally, since in B-IPQ only one item is used per 'subscale', B-IPQ might be inherently less reliable than the full-length instrument IPQ or IPQ-R. Thus, studies using the full-length inventory are awaited to confirm our findings.

The main clinical implication of our study is that, apart from recognition and treatment of depression clinicians should also consider AS patients' illness perceptions since we found that they could be important predictors of outcome in AS. Since depressive symptoms are frequent in rheumatologic disorders, most studies examining psychological correlates of health outcomes focus on depression, but illness perceptions should not be overlooked. Therefore, clinicians should also assess AS patients' illness representations and refer them for psychotherapeutic interventions. B-IPQ may be a useful and time-efficient tool for rheumatologists to assess illness representations in order to prevent further deterioration of AS patients' HRQoL.

Our findings generate relevant hypothesis. For instance, the use of antidepressants and/or more insight-oriented psychotherapies is justified for RA patients given their psychologically oriented causal attributions and the role that depression plays in outcome. Conversely, cognitive behavioural therapy (CBT) targeting illness cognitions might be more effective for AS patients. Evidence suggests that, in patients with arthritis, CBT can change illness representations resulting in better outcomes [45]. It should be noted, however, that any psychological intervention should only be provided after a thorough psychiatric evaluation. Moreover, when HRQoL is assessed, specific psychological distress measurements should be used too, in order to detect the influence of psychopathology on HRQOL. Our findings call also for additional investigation into the specific paths that form AS patients' HRQoL and for clinical psychotherapeutic trials targeting illness representations in AS in order to help rheumatologists and consultationliaison psychiatrists to schedule proper therapeutic interventions targeting AS patients' HRQoL.

Disclosures None.

References

- 1. Klippel JH, Stone JH, Crofford LJ, White PH (eds) (2008) Primer on the rheumatic diseases, 13th edn. Springer, New York
- Alamanos Y, Papadopoulos NG, Voulgari PV et al (2004) Epidemiology of ankylosing spondylitis in Northwest Greece, 1983–2002. Rheumatology (Oxford) 43(5):615–618
- Boonen A, Chorus A, Miedema H et al (2001) Employment, work disability, and work days lost in patients with ankylosing spondylitis: a cross sectional study of Dutch patients. Ann Rheum Dis 60:353–358
- 4. Boonen A, van der Heijde D, Landewé R et al (2003) Direct costs of ankylosing spondylitis and its determinants: an analysis among three European countries. Ann Rheum Dis 62:732–740
- 5. Salaffi F, Carotti M, Gasparini S, Intorcia M, Grassi W (2009) The health-related quality of life in rheumatoid arthritis, ankylosing spondylitis, and psoriatic arthritis: a comparison with a selected sample of healthy people. Health Qual Life Outcomes 7:25
- Dagfinrud H, Mengshoel AM, Hagen KB, Loge JH, Kvien TK (2004) Health status of patients with ankylosing spondylitis: a

comparison with the general population. Ann Rheum Dis 63 (12):1605–1610

- Hudson M, Thombs BD, Steele R, Panopalis P, Newton E, Baron M, Canadian Scleroderma Research Group (2009) Health-related quality of life in systemic sclerosis: a systematic review. Arthritis Rheum 61:1112–1120
- Murphy H, Dickens C, Creed F, Bernstein R (1999) Depression, illness perception and coping in rheumatoid arthritis. J Psychosom Res 46:155–164
- 9. Baubet T, Ranque B, Taieb O et al (2011) Mood and anxiety disorders in systemic sclerosis patients. Presse Med 40:e111–119
- Bachen EA, Chesney MA, Criswell LA (2009) Prevalence of mood and anxiety disorders in women with systemic lupus erythematosus. Arthritis Rheum 61(6):822–829
- Günaydin R, Göksel Karatepe A, Ceşmeli N, Kaya T (2009) Fatigue in patients with ankylosing spondylitis: relationships with disease-specific variables, depression, and sleep disturbance. Clin Rheumatol 28(9):1045–1051
- Zung WWK (1965) A self-rating depression scale. Arch Gen Psychiatry 12:63–70
- Barlow JH, Macey SJ, Struthers GR (1993) Gender, depression, and ankylosing spondylitis. Arthritis Care Res 6:45–51
- Ward MM (1999) Health-related quality of life in ankylosing spondylitis: a survey of 175 patients. Arthritis Care Res 12:247–255
- Bakker C, van der Linden S, van Santen-Hoeufft M, Bolwijn P, Hidding A (1995) Problem elicitation to assess patient priorities in ankylosing spondylitis and fibromyalgia. J Rheumatol 22:1304–1310
- American Psychiatric Association (1994) Diagnostic and statistical manual of mental disorders, 4th edn. American Psychiatric Press, Washington, DC
- Kroenke KS, Robert L (2002) The PHQ-9: a new depression diagnostic and severity measure. Psychiatr Ann 32:509–515
- Hyphantis T, Kotsis K, Voulgari PV, Tsifetaki N, Creed F, Drosos AA (2011) Diagnostic accuracy, internal consistency and convergent validity of the Greek version of PHQ-9 in diagnosing depression in rheumatological disorders. Arthritis Care Res (Hoboken) 63:1313–1321
- Leventhal H, Nerenz DR, Steele DJ (1984) Illness representation and coping with health threats. In: Baum A, Taylor SE, Singer JE (eds) The handbook of psychology and health. Vol IV: social psychological aspects of health. Erlbaum, Hillsdale, pp 219–252
- Richards HL, Herrick AL, Griffin K, Gwilliam PD, Loukes J, Fortune DG (2003) Systemic sclerosis: patients' perceptions of their condition. Arthritis Rheum 49:689–696
- 21. Kotsis K, Voulgari PV, Tsifetaki N et al (2012) Anxiety and depressive symptoms and illness perceptions in psoriatic arthritis and associations with physical health-related quality of life. Arthritis Care Res (Hoboken) 64(10):1593–601
- 22. Van der Linden SM, Valkenburg HA, Cats A (1984) Evaluation of diagnostic criteria for ankylosing spondylitis: a proposal for modification of the New York criteria. Arthritis Rheum 27:361–368
- 23. Chorus AM, Miedema HS, Boonen A, Van Der Linden S (2003) Quality of life and work in patients with rheumatoid arthritis and ankylosing spondylitis of working age. Ann Rheum Dis 62 (12):1178–1184
- 24. Zink A, Braun J, Listing J, Wollenhaupt J (2000) Disability and handicap in rheumatoid arthritis and ankylosing spondylitis results from the German rheumatological database. German Collaborative Arthritis Centers. J Rheumatol 27:613–622
- 25. Arnett FC, Edworthy SM, Bloch DA et al (1988) The American Rheumatism Association 1987 revised criteria for the classification of rheumatoid arthritis. Arthritis Rheum 31:315–24
- 26. Garrett S, Jenkinson T, Kennedy LG et al (1994) A new approach to defining disease status in ankylosing spondylitis: the Bath Ankylosing Spondylitis Disease Activity Index (BASDAI). J Rheumatol 21:2286–2291

- 27. Prevoo ML, van't Hof MA, Kuper HH, van Leeuwen MA, van de Putte LB, van Riel PL (1995) Modified disease activity scores that include twenty-eight-joint counts. Development and validation in a prospective longitudinal study of patients with rheumatoid arthritis. Arthritis Rheum 38:44–48
- Derogatis LR (1994) SCL-90-R: administration, scoring, and procedure manual, 3rd edn. National Computer Systems, Minneapolis
- 29. Demyttenaere K, Desaiah D, Petit C, Croenlein J, Brecht S (2012) Time course of improvement of different symptom clusters in patients with major depression and pain treated with duloxetine or placebo. Curr Med Res Opin 28(1):41–48
- Parker JC, Buckelew SP, Smarr KL, Buescher KL, Beck NC, Frank RG et al (1990) Psychological screening in rheumatoid arthritis. J Rheumatol 17:1016–1021
- Donias S, Karastergiou A, Manos N (1991) Standardization of the symptom checklist-90-R rating scale in a Greek population. Psychiatrike 2:42–48 (in Greek with English abstract)
- Broadbent E, Petrie KJ, Main J, Weinman J (2006) The Brief Illness Perception Questionnaire. J Psychosom Res 60:631–637
- Sluiter JK, Frings-Dresen MH (2008) Quality of life and illness perception in working and sick-listed chronic RSI patients. Int Arch Occup Environ Health 81(4):495–501
- 34. Skevington SM, Lotfy M, O'Connell KA, WHOQOL Group (2004) The World Health Organization's WHOQOL-BREF quality of life assessment: psychometric properties and results of the international field trial. A report from the WHOQOL group. Qual Life Res 13(2):299–310
- Power M, Harper A, Bullinger M (1999) The World Health Organization WHOQOL-100: tests of the universality of quality of life in 15 different cultural groups worldwide. Health Psychol 18:495–505
- 36. Taylor WJ, Myers J, Simpson RT, McPherson KM, Weatherall M (2004) Quality of life of people with rheumatoid arthritis as measured by the World Health Organization Quality of Life Instrument, short form (WHOQOL-BREF): score distributions and psychometric properties. Arthritis Rheum 51:350–357
- Moullec G, Ninot G (2010) An integrated programme after pulmonary rehabilitation in patients with chronic obstructive pulmonary disease: effect on emotional and functional dimensions of quality of life. Clin Rehabil 24(2):122–136
- Ginieri-Coccossis M, Triantafillou E, Tomaras V, Soldatos C, Mavreas V, Christodoulou G (2012) Psychometric properties of WHOQOL-BREF in clinical and health Greek populations: incorporating new culture-relevant items. Psychiatrike 23:130–142
- 39. Aigner M, Forster-Streffleur S, Prause W, Freidl M, Weiss M, Bach M (2006) What does the WHOQOL-Bref measure? Measurement overlap between quality of life and depressive symptomatology in chronic somatoform pain disorder. Soc Psychiatry Psychiatr Epidemiol 41(1):81–86
- Bai M, Tomenson B, Creed F et al (2009) The role of psychological distress and personality variables in the disablement process in rheumatoid arthritis. Scand J Rheumatol 38:419–430
- Miles J, Shevlin M (2003) Applying regression and correlation. Sage, London, pp 126–132
- 42. Baysal O, Durmuş B, Ersoy Y et al (2011) Relationship between psychological status and disease activity and quality of life in ankylosing spondylitis. Rheumatol Int 31(6):795–800
- 43. Hakkou J, Rostom S, Aissaoui N et al (2011) Psychological status in Moroccan patients with ankylosing spondylitis and its relationships with disease parameters and quality of life. J Clin Rheumatol 17(8):424–8
- Zigmond AS, Snaith PR (1983) The hospital anxiety and depression scale. Acta Psychiatr Scand 67:361–370
- 45. Bijsterbosch J, Scharloo M, Visser AW et al (2009) Illness perceptions in patients with osteoarthritis: change over time and association with disability. Arthritis Rheum 61(8):1054– 1061