

Emotional repression in patients with chronic inflammatory rheumatism

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Abstract

Background: This preliminary study was undertaken to assess and compare the psychological profiles of patients with chronic inflammatory rheumatism (CIRs), rheumatoid arthritis (RA) or spondyloarthritis (SpA), to try to identify a particular emotional profile compared to patients without CIRs. Emotional repression, i.e., a tendency to inhibit the expression of negative feelings and/or unpleasant thoughts, has been the most studied.

Methods: This monocenter observational pilot study included patients followed in a university rheumatology department who underwent systematic assessment of different psychological parameters by an experienced psychiatrist. The clinical and biological characteristics of their rheumatism were collected. Comparisons were performed using Chi 2 or Fisher's exact tests.

Results: Fifty-nine patients were assessed: 47 with CIRs (27 RA and 20 SpA) and 12 non-CIR controls (nine osteoarthritis, and one each viral disease, osteoporosis or osteomalacia). The rates of severe emotional repression, and the rates of severe early life events, were higher in the group of CIR patients than in the control group (respectively $P = 0.02$, $P = 0.02$). In contrast, the rates of severe psychological and somatic complaints were significantly higher than in the control group (respectively, $P < 0.01$ and $P = 0.01$).

Conclusion: Our results confirm the importance of emotional repression in coping with various traumatic life events which contribute to the etiology and/or evolution of CIRs. They suggest that these patients would probably benefit from psychotherapy in addition to medical management of their CIRs.

Declarations

Ethics Approval and Consent to Participate: A declaration of conformity to a reference methodology with the CNIL (National Commission for Computing and Liberties) was carried out under the number 2214762v0.

Consent for publication: All patients were informed and accepted the use of their computerized medical data

Availability of data and supporting materials section: data are available at the university hospital of Amiens.

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Authors' Contributions:

Psychological assessment: Cécile Lalanne, Alain Dervaux

Clinical assessment: Thibault Rabin, Sarah Salomon, Patrice Fardellone and Vincent Goeb

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Study design: Cécile Lalanne, Alain Dervaux, Vincent Goeb

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Background

Several factors, notably genetic, environmental, hormonal, infectious and/or psychological, can be involved in the development of chronic inflammatory rheumatisms (CIRs)(1)(2). However, the degree of implication of these different factors in CIR onset has not been determined.

Non-medication treatments are being used more-and-more frequently, especially regular physical activity, educational therapeutic workshops with support groups, mindfulness meditation... Over the past few years, self-reported evaluation criteria and patient-reported outcomes have been accorded increasing importance in the assessment of their disease, suggesting that the rheumatologist's perception of his/her patients' psychological well-being is essential in routine practice to correctly determine the efficacy of disease management(3).

Numerous studies on traumatic life events and stressful situations suggested that these factors could be implicated in triggering disease onset and its symptomatic progression of some diseases such as lupus, diabetes, multiple sclerosis or rheumatoid arthritis (RA)(4)(5). The hypothesis of the present study was that a particular psychological profile, namely emotional repression, was associated with autoimmune deregulation, and investigated a possible link between autoimmunity and emotional regulation.

The objective of this preliminary and exploratory study was to examine CIR patients' emotional function by assessing their psychological profiles to update and put into perspective their global management. Our secondary objectives were to determine whether RA and spondyloarthritis (SpA) patients have different emotional profiles, and whether a particular profile was associated with different rheumatological characteristics, in particular a biological inflammatory syndrome, the presence of sacroiliitis, a visualized structural involvement, a rheumatoid factor (RF)- or anti-citrullinated peptide antibody (ACPA)-positivity, or human leukocyte antigen (*HLA*)-*B27* allele carriage.

Patients And Methods

Population and study design

This monocenter cross-sectional observational, pilot study was conducted, between December 2012 and June 2014, in a university hospital rheumatology department during routine follow-up in a day hospital consultation by an experienced psychiatrist who assessed patients' psychological parameters.

Patients were included for a work-up in the framework of global CIR management, which includes dermatological, dental and gynecological consultations; rheumatologist-requested assessment of psychiatric manifestations (anxiety disorders, depressive disorders, sleep disorders, conjugal violence, childhood traumas); or requested by the patient. Among the 59 patients addressed to this consultation, none refused to participate. Exclusion criteria were: age <18 or >65 years old, cognitive disorders, substance-dependence, or acute somatic or any psychiatric disorders requiring emergency hospitalization.

The participating patients were followed in the rheumatology department by seven different rheumatologists. CIR patients fulfilled the RA, meeting the American College of Rheumatology/European League Against Rheumatism classification 2010 criteria; SpA, satisfying the Assessment of Spondyloarthritis international Society (ASAS) 2009 criteria; or psoriatic arthritis fulfilling CIASsification criteria for Psoriatic ARthritis (CASPAR) classification criteria. The 12 patients with ankylosing spondyloarthritis and 8 with psoriatic arthritis were combined under spondyloarthritides, due to their weak number, to facilitate statistical analyses. Even though subtle differences exist between their clinical characteristics, these pathologies share common pathophysiological mechanisms and genetic backgrounds, such as HLA-B27-positivity.

The non-CIR controls were patients followed in the department for other rheumatological diseases.

Psychological assessment tools

An experienced psychiatrist collected information during a semi-structured, semi-directed interview lasting 1.5 to 2 hours. The following psychological parameters were assessed, **depressive symptoms and their severity**, based on the Montgomery-Asberg Depression Rating Scale (MADRS)(6); **history of major depression episodes; anxiety symptoms**, using the Beck Anxiety Inventory, a 21-question multiple-choice self-report inventory, which assess severity (minimal, mild, moderate, severe anxiety)(7); **alexithymia**, a behavioral trait of persons unable to identify and describe their interior feelings, with limited imaginative capacity, and tending to focus their thoughts externally rather than resorting to introspection(8)(9), and its severity, using the 20-item Toronto Alexithymia Scale (TAS-20)(10); **social desirability intensity**, i.e., the tendency to seek approval of others and preserve one's self-image, using the Marlowe-Crowne Social Desirability scale (MCSD), a 33-item self-report questionnaire (11); **the severity of emotional repression**, a general term that describes a tendency to inhibit the expression of negative feelings or disagreeable thoughts, in accordance with the Weinberger classification; combination of the State-*Trait* Anxiety Inventory (*STAI*) (12) and the MCSD enables classification as repressors individuals with a low *STAI* score and high MCSD.

The psychiatrist used a Likert-type scale to assess the severity of the psychological parameters in a specific questionnaire which assessed : the **severity of somatic and psychological complaints**, i.e., the place the patient accords disease symptoms during the interview, with complaints rated severe when the responses to the majority of questions focus on them;**emotional-expressivity intensity** was graded as mild, when the patient had several emotional passages without breaking down or flights of fancy;

moderate, when facial expressions accompanying break downs or exuberant joy agreed with the verbal response and changed as the discussion progressed; or severe, when the patient had a tendency for hyper-expressivity throughout the interview; **life events and their intensities**, with type 1 being those occurring before the age of 15 years and type 2 those appearing during the 3 years preceding rheumatological disease onset; intensity was classified as: mild, when stressful events were non-existent or only mild; moderate, when stressful events did not incur traumatic stress (e.g., emotional deprivation concerning socio-economic difficulties, severe or disabling diseases of close friends/relatives), or severe, a traumatic event (sudden death of a person caring for a child, physical or sexual assault, patient witnessed a death or thought he/she would die); **somatic escalade**, which corresponds to the occurrence of a series of several somatic disorders or another chronic pathology before disease onset; **actual stress level**; importance the patient accorded to his/her rheumatological disease's **impact on professional activities**, corresponding to the occupational repercussions experienced, was classified as mild for minor suffering, moderate for notable suffering or severe for a painful experience (difficult loss of professional environment and activity, feeling of injustice or elevated fear of losing one's job); **manual labor**, whether **physical activity was enjoyable before and after disease onset**.

For the different psychological parameters, we combined mild and moderate intensities for comparison against severe intensities. Indeed, mild-to-moderate psychological disorders were easily identified in our patients exhibiting severe anxiety or depressive comorbidities and emotional experiences, often affected by chronic pain, but retaining severe intensity allows a more significant clinical difference. That approach represents an attempt to adjust for the population recruitment bias.

During the same day-hospitalization, various patient characteristics, notably rheumatological, were collected from the electronic medical files using the DxCare (medical information and prescription software) program: biological inflammatory syndrome, defined as an erythrocyte sedimentation rate ≥ 30 mm/1st h and/or C-reactive protein ≥ 5 mg/L; ACPA-positivity, defined as ≥ 7 U/mL by enzyme-linked immunosorbent assay (ELISA); RF >15 IU/mL by ELISA; magnetic resonance imaging (MRI) detection of grade-2 bilateral or grade-3 unilateral sacroiliitis, with active, inflammatory (edema) or chronic (erosion, bone condensation, bone bridges, fat conversion) lesions.

Statistical analyses

Continuous variables were expressed as median [Q1-Q3]. Categorical variables were expressed as number (percentages), (percentages calculated were calculated excluding missing data) and were compared by Chi² test or Fisher's exact test, as appropriate. Missing data were not replaced. For all analyses, $P < 0.05$ was considered statistically significant. Because of the exploratory character of the comparisons, the type I error was not adjusted for multiplicity. All statistical analyses were performed with SAS release 9.4 (SAS Institute Inc, Cary, NC).

Results

Patient characteristics

Our study population was comprised of 59 patients, 47 (79.7 %) with CIRs and 12 (20.3%) non-CIR patients followed in the department for pathologies other than inflammatory rheumatisms. Participants' characteristics according to their disease are summarized in Table 1.

The non-CIR control group consisted of 12 women (median age 65.0 years [55.0-72.0]): nine (75%) with arthroses and three (25%) with viral disease, osteoporosis or osteomalacia; 75% with structural involvement.

Among the 47 CIR patients (72.3% women and 27.7% men; median age 55.0 years [39.0-60.0]), 27 (57.4%) had RA and 20 (42.6%) SpA. Among the 27 RA patients (81.5% women median age 57 years [52-63]), most were RF-positive, ACPA-positive or had structural involvement and a solid third a biological inflammatory syndrome. Among the 20 SpA patients (60.0% women; median age 41.5 years [34.5-53.0]), a third had a biological inflammatory syndrome, and almost half were *HLA-B27*-positive and 55.6% had MRI-detected sacroiliitis.

Psychological assessments

Too few patients were *HLA-B27*-positive or had sacroiliitis and thus their associations with psychological parameters could not be analyzed.

Assessments of patients with CIRs vs. other diseases

Comparisons of these two patient groups for the parameters chosen found that they differed significantly for emotional repression, somatic complaints, psychological complaints, type-1 life-event severity, manual labor or enjoyment of physical activity before disease diagnosis (Table 2).

Assessments of RA vs. SpA patients

Comparisons of these two patient groups for the parameters chosen found that they differed significantly for previous depressive episode, professional impact and manual labor (Table 3).

Assessments of patients with vs. without biological inflammatory syndromes

Comparisons of these patients' psychological data showed their significant differences for emotional repression (Table 4).

Assessments of patient with vs. without structural involvement

Significant differences for patients according to structural involvement status were observed for somatic complaints, emotional repression severity, professional impact and enjoyment of physical activity prior to disease onset (Table 5).

Assessments of RF- or ACPA-positive patients

No psychological finding was significantly associated with these antibodies (data not show).

Discussion

CIR patients, compared to those with other rheumatological pathologies, presented significantly more frequent severe emotional repression, whereas those with other pathologies had significantly more frequent severe psychological and somatic complaints, potentially attesting to their strong emotional repression.

Temoshok (13) previously described the personality profile of RA patients as type C, which regroups submission, conciliatory approach, repression of hostility, self-effacement of personal needs and depressive vulnerability. Grossarth-Maticek and al. (14)(15) made similar observations in patients with cancer. Bayle and al.(16), devised a psychological vulnerability score and obtained converging results very close to the type C personality, which was significantly higher for patients with secondary Raynaud's syndrome than a control group with idiopathic or primary Raynaud's syndrome.

Nagano and al. (17) showed that, for RA patients, rational/anti-emotional behaviors, characterized by an extreme tendency to squelch emotional behaviors and rationalize negative experiences, was associated with poorer prognoses. Ishii and al. (18) found that RA patients, who were easily brought to tears in response to stress, had better responses to treatment and better overall prognoses.

Our results and those reported in the literature highlight the importance of emotional dysregulation, especially emotional repression, on CIR etiology and prognosis. However, the present study and its findings cannot be generalized with respect to identifying it as a factor favoring CIR or a secondary coping strategy of CIR. The repressive behavior has sometimes been analyzed as a consequence of the disease diagnosis, rather than its cause (19)(20).

The intensity of key early life events was significantly more severe for CIR patients than those with other diseases. Cutolo and Straub (21)(22) showed that stressful events preceded RA onset for 86% of their patients.

Reported findings also underscore the role of trying life events in favoring CIR onset or flares, whereas others accorded greater emphasis to the role of minor life events and daily stress. In contrast, we found no significant difference for the actual stress level presented by CIR and other-pathology patients, probably because of a lack of statistical power. Rimón and Laasko (23) described that higher stress at RA onset predicted a poorer prognosis for the disease. O'Donovan and al. (24) also found that veterans

experiencing trauma and developing PTSD could enhance the risk of developing autoimmune diseases, including RA. Based on their study of Vietnam veterans, Boscarino and al. (25) reported that those with RA had more PTSD symptoms, compared to their control counterparts.

The results of those studies are in agreement with a biopsychosocial model linking psychological stress and stressful life events with various immune-function changes in the etiology of autoimmune diseases. It could be thought that the impact of stress, resulting from an intense life event, modulated by adjustment strategies, like emotional repression, would increase vulnerability to autoimmune diseases because of dysregulated immune-system function (26)(27)(28)(29)(30) and enhanced inflammatory activity.

Herein, no significant difference was found between CIR and non-CIR patients regarding frequency of depression disorders, depression severity, history of prior depressive episode(s) or anxiety symptoms (100% of each group).

However, depressive and anxiety disorders are described as the most frequent comorbidities psychiatric of CIR patients. Reynier-Legarçon and al. (31) found that patients with autoimmune diseases (systemic lupus erythematosus, systemic sclerosis or primary Sjögren syndrome) presented more severe depressive and anxiety symptoms controls from general population. Baerwald and al. (32) found that the rates of depressive disorders for RA patients were significantly higher in for the general population. We were not able to replicate those findings, probably because of a lack of statistical power. In contrast, the homogeneity of our patients in terms of depressive and anxiety symptoms attenuates any potential bias linked to their emotional dysregulation associated with these psychological comorbidities.

Also, no significant difference were found between RA and SpA groups for the frequency of depression and depression severity. On the other hand, RA presented significantly more frequent prior depressive episodes than those with SpA, which has never been reported previously. Nonetheless, some findings indicated that anxiety symptoms, depression and perception of the disease impacted the physical quality of life differently for SpA and RA patients. Some authors noted that RA patients had more physical quality-of-life difficulties, while those with psoriatic arthritis and ankylosing spondylitis had more mental quality-of-life issues (33)(34)(35). Hyphantis and al. (36) showed that SpA patients' quality of life was associated with anxiety linked to the disease but not because of depressive symptoms, unlike those with RA. Our results, in line with those reported previously, suggest a link between depression and RA patients, compared to those with SpA.

Together, these results suggest that CIR (RA or SpA) patients would probably benefit from aid with emotional management (depression, disease representation and living with it) from psychotherapists and antidepressants for symptomatic depressive episode. Moreover, these symptoms can contribute to a poorer prognosis of their rheumatological disease (37). Bijsterbosch and al. (38) showed that arthritis patients' perceptions of their disease were predictive of the functional disability and that cognitive-behavioral therapy could modify the representations of the disease and obtain a better functional result.

A biological inflammatory syndrome was significantly associated with low disorder of emotional regulation (mild/moderate emotional repression). However, examination of the literature find more association between emotional disorders and a biological inflammatory syndrome.

Smoak and al. (39) found increased nuclear factor- κ B activity in patients with PTSD resulting from childhood violence, compared to healthy controls. Howren et al. (40) reported that patients with depression had significantly higher interleukin-1 and -6, tumor necrosis factor- α and C-reactive protein levels. Goldsmith and al. (41) also found in a meta-analysis of 68 studies, that patients with schizophrenia, bipolar disorder or major depressive episodes had increased inflammatory cytokines levels. In addition, depressive symptoms were more frequent in patients with autoimmune diseases and respond to anti-inflammatory drugs. Gobin and al. (42) found that antidepressants lowered production of inflammatory cytokines, e.g. interleukins-1 β and -6 and tumor necrosis factor- α .

Several monoclonal antibodies targeting relevant inflammatory pathways for the treatment of CIRs were associated with neuropsychiatric adverse events, notably depression, or suicidal ideation or behavior (43). Indeed, development of brodalumab, a molecule targeting interleukin-17, was stopped after suicidal behaviors were observed during clinical trials(44).

Our results, in line with the literature suggest a bidirectional relationship between a depressive syndrome and a biological inflammatory syndrome but need to be confirmed by a longitudinal study to establish conclusions. It remains to be determined whether systemic inflammation itself induces emotional symptoms by acting on certain neuronal cells or whether these emotional disorders are at the origin of a biological inflammatory syndrome that subsequently triggers the onset of CIRs.

One of the main limitations of this preliminary study is its lack of statistical power due to of the small sample. However, this pilot study investigated potential associations between CIRs and biopsychosocial factors, and its findings are merely indicative of what items should be examined in greater depth during future studies. In addition, because this was an observational study, no conclusions can be drawn about any association with causality. Because this was a cross-sectional investigation, it was not possible to know whether the primary emotional regulation disorders, possibly contributing to the rheumatological disease onset, or secondary disorders, should be interpreted as adaptive or coping modalities to handle a functional handicap or at limiting pain caused by the disease. Our study suffers from two selection biases: it was monocentric, recruiting patients consulting at a university hospital, who could represent a particular sociodemographic status; and most patients were followed in a day hospital focusing on more severe pathologies and comorbidities. Furthermore, the clinical psychological parameters chosen for assessment, although not subjected to consensual agreement, do correspond to clinical entities. Nonetheless, participants in this study benefited from combined psychiatric and rheumatological work-up during a day of hospitalization, which is novel, enabling evaluation of a large number of psychological factors and to compare them to concomitant rheumatological findings.

Conclusion

To summarize, the results of our pilot study showed an association between emotional dysregulation, more specifically, a tendency towards emotional repression and intense life events, and the CIR etiological or evolutionary process. These findings join those already published in other fields, like cancerology, or other autoimmune diseases, e.g., lupus or Gougerot-Sjögren syndrome. Moreover, among the highly frequent psychiatric disorders identified, RA patients had significantly more previous depressive episodes than those with SpA. Our observations could suggest that CIR patients would probably benefit from psychotherapy in conjunction with medical management.

Our observations also show how informative a multimodal assessment of emotional regulation in future studies could contribute to improving management of CIR patients.

Abbreviations

ACPA: Anti-citrullinated peptide antibodies; CIR: Chronic inflammatory rheumatism; HLA: Human leukocyte antigen; MRI, Magnetic resonance imaging; PTSD; Post-traumatic stress disorder; RA: Rheumatoid arthritis; RF: rheumatoid factor; SpA: Spondyloarthritis

References

1. Pretorius E, Akeredolu O-O, Soma P, Kell DB. Major involvement of bacterial components in rheumatoid arthritis and its accompanying oxidative stress, systemic inflammation and hypercoagulability. *Exp Biol Med.*2017;242:355-73.
2. Dougados M, Baeten D. Spondyloarthritis. *Lancet.*2011;377:2127-37.
3. Söderlin MK, Bergsten U, Svensson B, BARFOT Study Group. Patient-reported events preceding the onset of rheumatoid arthritis: possible clues to aetiology. *Musculoskeletal Care.* 2011;9:25-31.
4. Halliday JL. Psychological Aspects of Rheumatoid Arthritis. *Proc R Soc Med.* 1942;35:455-7.
5. Moos RH. Personality factors associated with rheumatoid arthritis: a review. *J Chronic Dis. J Chronic Dis.* 1964;17:41-55.
6. Montgomery SA, Asberg M. A new depression scale designed to be sensitive to change. *Br J Psychiatry J Ment Sci.*1979;134:382-9.
7. Beck AT, Ward CH, Mendelson M, Mock J, Erbaugh J. An inventory for measuring depression. *Arch Gen Psychiatry.*1961;4:561-71.
8. Sifneos PE. Alexithymia: past and present. *Am J Psychiatry.* 1996;153:137-42.
9. Sifneos PE. The prevalence of « alexithymic » characteristics in psychosomatic patients. *Psychother Psychosom.* 1973;22:255-62.
10. Bagby RM, Taylor GJ, Parker JD. Construct validity of the Toronto Alexithymia Scale. *Psychother Psychosom.* 1988;50:29-34.
11. Crowne DP, Marlowe D. A new scale of social desirability independent of psychopathology. *J Consult Psychol.*1960;349–354.

12. Gaudry E, Vagg P, Spielberger CD. Validation of the State-Trait Distinction in Anxiety Research. *Multivar Behav Res.*1975;10:331-41.
13. Temoshok L. Personality, coping style, emotion and cancer: towards an integrative model. *Cancer Surv.* 1987;6:545-67.
14. Grossarth-Maticek R, Vetter H, Frentzel-Beyme R, Heller WD. Precursor lesions of the GI tract and psychosocial risk factors for prediction and prevention of gastric cancer. *Cancer Detect Prev.* 1988;13:23-9.
15. Grossarth-Maticek R, Eysenck HJ, Boyle GJ, Heep J, Costa SD, Diel IJ, et al. Interaction of psychosocial and physical risk factors in the causation of mammary cancer, and its prevention through psychological methods of treatment. *J Clin Psychol.*2000;56:33-50.
16. Bayle O, Consoli SM, Baudin M, Vayssairat M, Fiessinger JN, Housset E. Idiopathic and secondary Raynaud's phenomenon. A comparative psychosomatic approach. *Presse Med.* 1990;19:741-5.
17. Nagano J, Morita T, Taneichi K, Nagaoka S, Katsube S, Asai T, et al. Rational/antiemotional behaviors in interpersonal relationships and the functional prognosis of patients with rheumatoid arthritis: a Japanese multicenter, longitudinal study. *Biopsychosoc Med.*2014;8:8.
18. Ishii H, Nagashima M, Tanno M, Nakajima A, Yoshino S. Does being easily moved to tears as a response to psychological stress reflect response to treatment and the general prognosis in patients with rheumatoid arthritis? *Clin Exp Rheumatol.*2003;21:611-6.
19. Butow PN, Hiller JE, Price MA, Thackway SV, Kricker A, Tennant CC. Epidemiological evidence for a relationship between life events, coping style, and personality factors in the development of breast cancer. *J Psychosom Res.*2000;49:169-81.
20. Zachariae R, Jensen AB, Pedersen C, Jørgensen MM, Christensen S, Lassesen B, et al. Repressive coping before and after diagnosis of breast cancer. *Psychooncology.*2004;13:547-61.
21. Cutolo M, Straub RH. Stress as a risk factor in the pathogenesis of rheumatoid arthritis. *Neuroimmunomodulation.* 2006;13:277-82.
22. Straub RH, Cutolo M. Does stress influence the course of rheumatic diseases? *Clin Exp Rheumatol.*2006;24:225-8.
23. Rimón R, Laakso RL. Life stress and rheumatoid arthritis. A 15-year follow-up study. *Psychother Psychosom.* 1985;43:38-43.
24. O'Donovan A, Cohen BE, Seal KH, Bertenthal D, Margaretten M, Nishimi K, et al. Elevated risk for autoimmune disorders in Iraq and Afghanistan veterans with posttraumatic stress disorder. *Biol Psychiatry.*2015;77:365-74.
25. Boscarino JA, Forsberg CW, Goldberg J. A twin study of the association between PTSD symptoms and rheumatoid arthritis. *Psychosom Med.*2010;72:481-6.
26. Webster Marketon JI, Glaser R. Stress hormones and immune function. *Cell Immunol.* 2008;252:16-26.

27. Borovikova LV, Ivanova S, Zhang M, Yang H, Botchkina GI, Watkins LR, et al. Vagus nerve stimulation attenuates the systemic inflammatory response to endotoxin. *Nature*.2000;405:458-62.
28. Levite M. Neuropeptides, by direct interaction with T cells, induce cytokine secretion and break the commitment to a distinct T helper phenotype. *Proc Natl Acad Sci U S A*.1998;95:12544-9.
29. Carpenter LL, Gawuga CE, Tyrka AR, Lee JK, Anderson GM, Price LH. Association between plasma IL-6 response to acute stress and early-life adversity in healthy adults. *Neuropsychopharmacol Off Publ Am Coll Neuropsychopharmacol*.2010;35:2617-23.
30. Janusek LW, Tell D, Gaylord-Harden N, Mathews HL. Relationship of childhood adversity and neighborhood violence to a proinflammatory phenotype in emerging adult African American men: An epigenetic link. *Brain Behav Immun*.2017;60:126-35.
31. Reynier-Legarçon E, Clément JP, Calvet B, Bézananary H, Liozon E, Gondran G, et al. Personnalité et maladies auto-immunes systémiques. *Rev Médecine Interne*.2016;37:38.
32. Baerwald C, Manger B, Hueber A. Depression as comorbidity of rheumatoid arthritis. *Z Rheumatol*.2019;78:243-8.
33. 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;54:2665-73.
34. Salaffi F, Carotti M, Gasparini S, Intorcchia M, Grassi W. 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*.2009;7:25.
35. Dagfinrud H, Mengshoel AM, Hagen KB, Loge JH, Kvien TK. Health status of patients with ankylosing spondylitis: a comparison with the general population. *Ann Rheum Dis*. 2004;63:1605-10.
36. Hyphantis T, Kotsis K, Tsifetaki N, Creed F, Drosos AA, Carvalho AF, et al. The relationship between depressive symptoms, illness perceptions and quality of life in ankylosing spondylitis in comparison to rheumatoid arthritis. *Clin Rheumatol*.2013;32:635-44.
37. Michelsen B, Kristianslund EK, Sexton J, Hammer HB, Fagerli KM, Lie E, et al. Do depression and anxiety reduce the likelihood of remission in rheumatoid arthritis and psoriatic arthritis? Data from the prospective multicentre NOR-DMARD study. *Ann Rheum Dis*. 2017;76:1906-10.
38. Bijsterbosch J, Scharloo M, Visser AW, Watt I, Meulenbelt I, Huizinga TWJ, et al. Illness perceptions in patients with osteoarthritis: change over time and association with disability. *Arthritis Rheum*.2009;61:1054-61.
39. Smoak KA, Cidlowski JA. Mechanisms of glucocorticoid receptor signaling during inflammation. *Mech Ageing Dev*.2004;125:697-706.
40. Howren MB, Lamkin DM, Suls J. Associations of depression with C-reactive protein, IL-1, and IL-6: a meta-analysis. *Psychosom Med*.2009;71:171-86.
41. Goldsmith DR, Rapaport MH, Miller BJ. A meta-analysis of blood cytokine network alterations in psychiatric patients: comparisons between schizophrenia, bipolar disorder and depression. *Mol Psychiatry*. 2016;21:1696-709.

42. Gobin V, Van Steendam K, Denys D, Deforce D. Selective serotonin reuptake inhibitors as a novel class of immunosuppressants. *Int Immunopharmacol.*2014;20:148-56.
43. Minnema LA, Giezen TJ, Souverein PC, Egberts TCG, Leufkens HGM, Gardarsdottir H. Exploring the Association between Monoclonal Antibodies and Depression and Suicidal Ideation and Behavior: A VigiBase Study. *Drug Saf.* 2019;42:887-895.
44. Lebwohl MG, Papp KA, Marangell LB, Koo J, Blauvelt A, Gooderham M, et al. Psychiatric adverse events during treatment with brodalumab: Analysis of psoriasis clinical trials. *J Am Acad Dermatol.*2018;78:81-89.5.

Tables

Table 1. Clinical and laboratory characteristics of the 59 patients

Characteristic	RA <i>n</i> = 27	SpA <i>n</i> = 20	All CIRs <i>n</i> = 47	Other diseases <i>n</i> = 12
Age, yr median [Q1-Q3]	57.0 [52.0-63.0]	41.5 [34.5-53.0]	55.0 [39.0-60.0]	65.0 [55.0-72.0]
Female/male sex ratio	22/5	12/8	34/13	12/0
Rheumatoid factor+	18/27 (66.7%)	1/8 (12.5%)	19/35 (54.3%)	0
ACPA+	17/27 (63%)	1/8 (12.5%)	18/35 (51.4%)	0
Elevated ESR and/or CRP	10/27 (37.0%)	6/19 (30%)	16/47 (34.0%)	0
<i>HLA-B27</i> +	0	9/19 (47.4%)	9/21 (42.9%)	0
Sacroiliitis	0	10/18 (55.6%)	10/20 (50%)	0
Structural involvement	14/27 (51.9%)	0	14/40 (35%)	9/11 (81.8%)

RA, rheumatoid arthritis; SpA, spondyloarthropathy; CIRs, chronic inflammatory rheumatism; ACPA, anti-citrullinated peptide antibodies; ESR, erythrocyte sedimentation rate first hour; CRP, C-reactive

Table 2 Psychological evaluations of patients with chronic inflammatory rheumatism (CIRs) vs. those with other diseases

Psychological parameter	All CIRs n = 47	Other diseases n = 12	<i>P</i>
Depression	38/47 (80.9%)	11/12 (91.7%)	0.67
Intensity			0.40
Mild/moderate	32/38 (84.2%)	8/11 (72.7%)	
Severe	6/38 (15.8%)	3/11 (27.3%)	
Previous depression	37 (78.7%)	11 (91.7%)	0.43
Anxiety	47/47 (100%)	12/12 (100%)	-
Intensity			0.17
Mild/moderate	21/47 (44.7%)	8/12 (66.7%)	
Severe	26/47 (55.3%)	4/12 (33.3%)	
Alexithymia	46/47 (97.9%)	11/12 (91.7%)	0.37
Intensity			1.00
Mild/moderate	20/46 (43.5%)	5/11 (45.5%)	
Severe	26/46 (56.5%)	6/11 (54.5%)	
Social desirability			0.43
Mild/moderate	37/47 (78.7%)	11/12 (91.7%)	
Severe	10/47 (21.3%)	1/12 (8.3%)	
Emotional repression			0.02
Mild/moderate	25/47 (53.2%)	11/12 (91.7%)	
Severe	22/47 (46.8%)	1/12 (8.3%)	
Somatic complaints			0.01
Mild/moderate	32/47 (68.1%)	3/12 (25.0%)	
Severe	15/12 (31.9%)	9/12 (75.0%)	
Psychological complaints			<0.01
Mild/moderate	33/47 (70.2%)	2/12 (16.7%)	
Severe	14/47 (29.8%)	10/12 (83.3%)	
Emotional expressivity intensity			1.00
Mild/moderate	32/47 (68.1%)	8/12 (66.7%)	
Severe	15/47 (31.9%)	4/12 (33.3%)	
Life event 1	29/44 (65.9%)	8/11 (72.7%)	1.00
Intensity			0.02
Mild/moderate	15/29 (51.7%)	8/8 (100%)	
Severe	14/29 (48.3%)	0	
Life event 2	44/47 (93.6%)	11/12 (91.7%)	1.00
Intensity			1.00
Mild/moderate	30/44 (68.2%)	8/11 (72.7%)	
Severe	14/44 (31.8%)	3/11 (27.3%)	
Somatic escalate	14/47 (29.8%)	7/12 (58.3%)	0.09
Actual stress level			1.00
Mild/moderate	44/47 (93.6%)	12/12 (100%)	
Severe	3/47 (6.4%)	0	
Professional impact			0.06
Mild/moderate	22/44 (50%)	7/8 (87.5%)	
Severe	22/44 (50%)	1/8 (12.5%)	
Manual labor	28/47 (59.6%)	2/12 (16.7%)	<0.01
Physical activity enjoyable prediagnosis	23/47 (48.9%)	2/12 (16.7%)	0.04
Physical activity enjoyable postdiagnosis	8/47 (17%)	1/12 (8.3%)	0.67

Table 3 Psychological evaluations of patients with rheumatoid arthritis (RA) vs. those with spondyloarthropathy (SpA)

Psychological parameter	RA <i>n</i> = 27	SpA <i>n</i> = 20	<i>P</i>
Depression	21/27 (77.8%)	17/20 (85%)	0.71
Intensity			0.20
Mild/moderate	16/21 (76.2%)	16/17 (94.1%)	
Severe	5/21 (23.8%)	1/17 (5.9%)	
Previous depression	26/27 (96.3%)	11/20 (55.0%)	<0.01
Anxiety	27/27 (100%)	20/20 (100%)	-
Intensity			0.07
Mild/moderate	9/27 (33.3%)	12/20 (60.0%)	
Severe	18/27 (66.7%)	8/20 (40.0%)	
Alexithymia	27/27 (100%)	19/20 (95.0%)	0.43
Intensity			0.17
Mild/moderate	14/27 (51.9%)	6/19 (31.6%)	
Severe	13/27 (48.1%)	13/19 (68.4%)	
Social desirability			0.48
Mild/moderate	20/27 (74.1%)	17/20 (85.0%)	
Severe	7/27 (25.9%)	3/20 (15.0%)	
Emotional repression			0.33
Mild/moderate	16/27 (59.3%)	9/20 (45.0%)	
Severe	11/27 (40.7%)	11/20 (55.0%)	
Somatic complaints			0.38
Mild/moderate	17/27 (63.0%)	15/20 (75.0%)	
Severe	10/27 (37.0%)	5/20 (25.0%)	
Psychological complaints			0.21
Mild/moderate	17/27 (63.0%)	16/20 (80.0%)	
Severe	10/27 (37.0%)	4/20 (20.0%)	
Emotional expressivity intensity			0.13
Mild/moderate	16/27 (59.3%)	16/20 (80.0%)	
Severe	11/27 (40.7%)	4/20 (20.0%)	
Life event 1	16/24 (66.7%)	13/20 (65.0%)	0.91
Intensity			0.84
Mild/moderate	8/16 (50.0%)	7/13 (53.8%)	
Severe	8/16 (50.0%)	6/13 (46.2%)	
Life event 2	24/27 (88.9%)	20/20 (100%)	0.25
Intensity			0.38
Mild/moderate	15/24 (62.5%)	15/20 (75.0%)	
Severe	9/24 (37.5%)	5/20 (25.0%)	
Somatic escalate	6/27 (22.2%)	8/20 (40.0%)	0.19
Actual stress level			0.25
Mild/moderate	24/27 (88.9%)	20/20 (100%)	
Severe	3/27 (11.1%)	0/20	
Professional impact			<0.01
Mild/moderate	17/25 (68%)	5/19 (26.3%)	
Severe	8/25 (32%)	14/19 (73.7%)	
Manual labor	12/27 (44.4%)	16/20 (80.0%)	0.01
Physical activity enjoyable prediagnosis	10/27 (37.0%)	13/20 (65.0%)	0.06
Physical activity enjoyable postdiagnosis	5/27 (18.5%)	3/20 (15.0%)	1.00

Table 4 Psychological evaluations of CIRs patients with biological inflammatory syndromes vs. those without

Psychological parameter	Biological inflammatory syndrome		<i>P</i>
	Yes, <i>n</i> = 16	No, <i>n</i> = 31	
Depression	13/16 (81.3%)	25/31 (80.6%)	1.00
Intensity			
Mild/moderate	11/13 (84.6%)	21/25 (84.0%)	1.00
Severe	2/13 (15.4%)	4/25 (16.0%)	
Previous depression	14/16 (87.5%)	23/31 (74.2%)	0.46
Anxiety	16/16 (100%)	31/31 (100%)	1.00
Intensity			0.25
Mild/moderate	9/16 (56.3%)	12/31 (38.7%)	
Severe	7/16 (43.8%)	19/31 (61.3%)	
Alexithymia	16/16 (100%)	30/31 (96.8%)	1.00
Intensity			0.22
Mild/moderate	5/16 (31.3%)	15/30 (50.0%)	
Severe	11/16 (68.8%)	15/30 (50.0%)	
Social desirability			0.7
Mild/moderate	12/16 (75.0%)	25/31 (80.6%)	
Severe	4/16 (25.0%)	6/31 (19.4%)	
Emotional repression			0.03
Mild/moderate	12/16 (75%)	13/31 (41.9%)	
Severe	4/16 (25%)	18/31 (58.1%)	
Somatic complaints			0.56
Mild/moderate	10/16 (62.5%)	22/31 (71.0%)	
Severe	6/16 (37.5%)	9/31 (29.0%)	
Psychological complaints			1.00
Mild/moderate	11/16 (68.8%)	22/31 (71.0%)	
Severe	5/16 (31.3%)	9/31 (29.0%)	
Emotional expressivity intensity			0.21
Mild/moderate	9/16 (56.3%)	23/31 (74.2%)	
Severe	7/16 (43.8%)	8/31 (25.8%)	
Life event 1	11/14 (78.6%)	18/30 (60.0%)	0.31
Intensity			0.81
Mild/moderate	6/11 (54.5%)	9/18 (50.0%)	
Severe	5/11 (45.5%)	9/18 (50.0%)	
Life event 2	15/16 (93.8%)	29/31 (93.5%)	1.00
Intensity			1.00
Mild/moderate	10/15 (66.7%)	20/29 (69.0%)	
Severe	5/15 (33.3%)	9/29 (31.0%)	
Somatic escalate	7/16 (43.8%)	7/31 (22.6%)	0.18
Actual stress level			0.54
Mild/moderate	16/16 (100%)	28/31 (90.3%)	
Severe	0/16	3/31 (9.7%)	
Professional impact			0.75
Mild/moderate	8/15 (53.3%)	14/29 (48.3%)	
Severe	7/15 (46.7%)	15/29 (51.7%)	
Manual labor	8/16 (50.0%)	20/31 (64.5%)	0.34
Physical activity enjoyable prediagnosis	7/16 (43.8%)	16/31 (51.6%)	0.61
Physical activity enjoyable postdiagnosis	3/16 (18.8%)	5/31 (16.1%)	1

Table 5 Psychological evaluations of CIRs patients with structural involvement vs. those without

Psychological parameter	Structural involvement		<i>P</i>
	Yes, <i>n</i> = 14	No, <i>n</i> = 26	
Depression	11/14 (78.6%)	20/26 (76.9%)	1.00
Intensity			1.00
Mild/moderate	9/11 (81.8%)	17/20 (85%)	
Severe	2/11 (18.2%)	3/20 (15%)	
Previous depression	14/14 (100%)	19/26 (73.1%)	0.08
Anxiety	14/14 (100%)	26/26 (100%)	1.00
Intensity			0.08
Mild/moderate	4/14 (28.6%)	15/26 (57.7%)	
Severe	10/14 (71.4%)	11/26 (42.3%)	
Alexithymia	14/14 (100%)	25/26 (96.2%)	1.00
Intensity			0.58
Mild/moderate	6/14 (42.9%)	13/25 (52.0%)	
Severe	8/14 (57.1%)	12/25 (48.0%)	
Social desirability			0.22
Mild/moderate	13/14 (92.9%)	19/26 (73.1%)	
Severe	1/14 (7.1%)	7/26 (26.9%)	
Emotional repression			0.19
Mild/moderate	10/14 (71.4%)	13/26 (50.0%)	
Severe	4/14 (28.6%)	13/26 (50.0%)	
Somatic complaints			0.03
Mild/moderate	6/14 (42.9%)	21/26 (80.8%)	
Severe	8/14 (57.1%)	5/26 (19.2%)	
Psychological complaints			0.48
Mild/moderate	8/14 (57.1%)	19/26 (73.1%)	
Severe	6/14 (42.9%)	7/26 (26.9%)	
Emotional expressivity intensity			<0.01
Mild/moderate	5/14 (35.7%)	21/26 (80.8%)	
Severe	9/14 (64.3%)	5/26 (19.2%)	
Life event 1	9/13 (69.2%)	14/24 (58.3%)	0.72
Intensity			1.00
Mild/moderate	5/9 (55.6%)	7/14 (50%)	
Severe	4/9 (44.4%)	7/14 (50%)	
Life event 2	12/14 (85.7%)	25/26 (96.2%)	0.28
Intensity			1.00
Mild/moderate	8/12 (66.7%)	16/25 (64.0%)	
Severe	4/12 (33.3%)	9/25 (36.0%)	
Somatic escalate	6/14 (42.9%)	4/26 (15.4%)	0.12
Actual stress level			0.54
Mild/moderate	14/14 (100%)	23/26 (88.5%)	
Severe	0/14	3/26 (11.5%)	
Professional impact			<0.01
Mild/moderate	11/13 (84.6%)	10/25 (40.0%)	
Severe	2/13 (15.4%)	15/25 (60.0%)	
Manual labor	7/14 (50%)	15/26 (57.7%)	0.64
Physical activity enjoyable prediagnosis	3/14 (21.4%)	15/26 (57.7%)	0.03
Physical activity enjoyable postdiagnosis	2/14 (14.3%)	4/26 (15.4%)	1.00