

# Prevalence of Inflammatory Back Pain and Axial Spondyloarthritis Among University Employees in Izmir, Turkey

Fatos Onen, Dilek Solmaz, Pinar Cetin, Ismail Sari, Ali Balci, Merih Birlik, Servet Akar, and Nurullah Akkoc

**ABSTRACT. Objective.** To estimate the prevalence of inflammatory back pain (IBP) and axial spondyloarthritis (axSpA) using the Assessment of SpondyloArthritis International Society (ASAS) classification criteria among employees in a university.

**Methods.** In the first stage of the study, a face-to-face interview was done using a standard questionnaire to investigate IBP in 381 subjects randomly selected from 2894 employees at Dokuz Eylul University in Izmir, Turkey. In the second stage, subjects with back pain for  $\geq 3$  months and age at onset  $< 45$  years were evaluated for axSpA using the ASAS criteria. Both the European Spondyloarthropathy Study Group (ESSG) criteria and Amor criteria were used for the classification of the whole group of spondyloarthritis (SpA).

**Results.** There were 131 male and 250 female subjects (mean age: 38.0 yrs). Twenty-five subjects (6.6%) were classified as having IBP according to the ASAS criteria. The prevalence of IBP according to the Berlin and Calin criteria was 7.1% and 21.5%, respectively. The prevalence of axSpA was estimated at 1.3% according to the ASAS classification criteria (0.5% for radiographic axSpA and 0.8% for nonradiographic axSpA). A total of 7 patients (1.8%) fulfilled both the Amor and ESSG criteria for the whole group of SpA.

**Conclusion.** This is the first prevalence study of IBP and axSpA using ASAS classification criteria in the Turkish population. The prevalence estimates of IBP and axSpA reported here are within the upper range of other studies in European countries and the United States. (First Release July 15 2015; J Rheumatol 2015;42:1647–51; doi:10.3899/jrheum.141600)

## Key Indexing Terms:

PREVALENCE      INFLAMMATORY BACK PAIN      AXIAL SPONDYLOARTHRITIS

Spondyloarthritis are among the most prevalent inflammatory rheumatic diseases<sup>1</sup>. There is a considerable diagnostic delay (8.9 yrs) in ankylosing spondylitis (AS), the prototype of this group, mainly because of the requirement of radiographic sacroiliitis for its diagnosis<sup>2</sup>. Low awareness of inflammatory back pain (IBP), the first and most common symptom of spondyloarthritis (SpA), in daily practice is also a major reason for the diagnostic delay<sup>3</sup>. New classification criteria developed by the Assessment of SpondyloArthritis International Society (ASAS) provide that patients with SpA can be classified as either patients with axial SpA (axSpA)

or those with peripheral SpA. The ASAS axSpA criteria cover the entire spectrum of axial disease including AS and non-radiographic axSpA (nr-axSpA)<sup>4</sup>. Early diagnosis of axSpA can create an opportunity for better treatment strategies.

The number of epidemiological studies regarding IBP<sup>5,6,7,8</sup> and axSpA<sup>9</sup> is limited. In a review of the literature, we found no population-based study that investigated their prevalence using the new ASAS classification criteria. We sought to determine the prevalence of IBP and axSpA using these criteria among university employees in Izmir, Turkey.

## MATERIALS AND METHODS

We conducted our study at the Health Sciences Campus at Dokuz Eylul University in Izmir, which has 2894 medical and nonmedical staff aged between 18 and 67 years. A sample of 395 subjects was selected randomly by a computer from the list of all employees, based on the IBP prevalence of 5% in the general population<sup>5</sup>, using OpenEpi (version 2.3) and CI  $\pm 2\%$ . A total of 381 of these 395 subjects agreed to participate, an acceptance rate of 96.5%. In the first stage of the study, 6 trained medical students, using a standard questionnaire, interviewed participants face to face. Questionnaire responses were used to determine whether participants met the ASAS criteria for IBP<sup>10</sup>. Subjects were also evaluated for IBP based on the Berlin<sup>11</sup> and Calin criteria<sup>12</sup> (Table 1).

In the second stage, the subjects with back pain for more than 3 months

From the Dokuz Eylul University School of Medicine, Division of Rheumatology, and Department of Radiodiagnostic, Izmir, Turkey.

F. Onen, MD, Professor; D. Solmaz, MD, Assistant Professor; P. Cetin, MD, Specialist; I. Sari, MD, Associate Professor, Dokuz Eylul University School of Medicine, Division of Rheumatology; A. Balci, MD, Associate Professor, Dokuz Eylul University School of Medicine, Department of Radiodiagnostic; M. Birlik, MD, Associate Professor; S. Akar, MD, Professor; N. Akkoc, MD, Professor, Dokuz Eylul University School of Medicine, Division of Rheumatology.

Address correspondence to Dr. F. Onen, Dokuz Eylul University School of Medicine, Department of Internal Medicine, Division of Rheumatology, 35340 Inciralti/Izmir-Turkey. E-mail: fatos.onen@deu.edu.tr

Accepted for publication May 27, 2015.

Table 1. Inflammatory back pain criteria sets.

Calin Criteria <sup>12</sup>	Berlin Criteria <sup>11</sup>	ASAS Criteria <sup>10</sup>
Age at onset < 40 yrs Duration of back pain > 3 mos	Morning stiffness > 30 min Improvement with exercise, not with rest	Age at onset < 40 yrs Insidious onset
Insidious onset	Awakening at second half of the night because of pain	Improvement with exercise
Morning stiffness Improvement with exercise	Alternating buttock pain	No improvement with rest Pain at night (with improvement upon getting up)
IBP if 4/5 are present	IBP if 2/4 are present	IBP if 4/5 are present

ASAS: Assessment of SpondyloArthritis International Society; IBP: inflammatory back pain.

and symptom onset before age 45 years were invited to the rheumatology outpatient clinic to participate in the study. Subjects who accepted the invitation were evaluated for axSpA by 2 rheumatologists using the ASAS classification criteria<sup>4</sup>. Pelvic radiographs and magnetic resonance imaging (MRI) of the sacroiliac joint (SIJ) were obtained. They were read by an experienced rheumatologist and a radiologist blinded to the patients' identities. The patients with at least 2 features of SpA and negative imaging or who had no SIJ imaging were tested for HLA-B27.

The subjects were also evaluated to determine whether they met the modified New York (mNY) criteria for AS<sup>13</sup> and both the European Spondyloarthritis Study Group (ESSG) criteria<sup>14</sup> and Amor criteria<sup>15</sup> for SpA.

Ethics approval for our study was obtained from the Local Research Ethics Committee (22 March 2012/11-15).

*Statistical analysis.* Stats Direct Statistical Software (version 3.0.23, Stats Direct Ltd.) was used for calculating the prevalence and 95% CI.

## RESULTS

In the first stage of our study, among the 395 subjects who had been contacted, 381 (131 men, 250 women; mean age: 38.1 ± 9.5 yrs) had agreed to take part; an acceptance rate of 96.5%. Among them, 299 (78.5%; 95% CI 69.8–87.9) had experienced back pain at least once in their lifetime. A total of 172 subjects (45.1%; 95% CI 38.7–52.4) had back pain lasting more than a month and 127 (33.3%; 95% CI 27.8–39.7) more than 3 months.

Among 381 subjects, 25 (6.6%; 95% CI 4.3–9.7) were classified as having IBP according to ASAS criteria<sup>10</sup>. The prevalence of IBP according to Berlin<sup>11</sup> and Calin criteria<sup>12</sup> was found to be 7.1% (95% CI 4.7–10.3) and 21.5% (95% CI 17.1–26.7), respectively (Table 2). The prevalence of IBP according to the ASAS criteria was 8.4% (95% CI 5.4–12.3) among all patients with back pain. It was estimated to be 14.5% (95% CI 9.4–21.5) and 19.7% (95% CI 12.7–29.0) among those with chronic back pain lasting more than 1 month and 3 months, respectively.

The prevalence of IBP in women (8.0%; 95% CI 4.9–12.4) was found to be higher than that in men (3.8%; 95% CI 1.2–8.9), according to the ASAS criteria. IBP was also more prevalent among women than men according to other IBP criteria sets (Table 2).

There were 115 patients (30%) with back pain for more

than 3 months and symptom onset before the age of 45 years. They were invited to undergo further clinical evaluation at the rheumatology outpatient clinic, and 95 (82.6%) agreed to do so. Twenty-five of them had at least 1 SpA feature. All these patients had IBP based on the ASAS criteria and 7 of them had other SpA features (6 gluteal pain, 3 enthesitis, 1 psoriasis, 1 peripheral arthritis, and 1 dactylitis; Figure 1).

Pelvic radiographs were obtained in 21/25 patients with at least 1 SpA feature. Two of them demonstrated bilateral grade 3 sacroiliitis. In 14 out of 19 patients with normal radiographs, MRI of SIJ could be performed. Two of these patients demonstrated signs of acute sacroiliitis (Figure 1).

There were 7 patients with chronic back pain and at least 2 SpA features. One of them had MRI sacroiliitis. Among the remaining 6 patients, SIJ imaging was not available for 3 and the other 3 had negative MRI of SIJ. Five of the 7 patients with chronic back pain and at least 2 SpA features could be tested for HLA-B27 and 1 of them was found to be positive. This patient was classified as nr-axSpA according to the clinical arm of the ASAS criteria (Figure 1).

In total, 5 subjects (1.3%) with at least 1 SpA feature (all had IBP) were classified as having axSpA according to the ASAS classification criteria<sup>4</sup>. Two of them (0.5%) had radiographic sacroiliitis and met the mNY criteria for AS<sup>13</sup>. The prevalence of nr-axSpA was estimated to be 0.8%. There were 7 patients (1.8%) who fulfilled both the Amor<sup>15</sup> and ESSG criteria<sup>14</sup> for the whole group of SpA (Table 2). One of them had psoriasis and met the Classification for Psoriatic Arthritis (CASPAR) criteria<sup>16</sup> for psoriatic arthritis (Figure 1).

We found the prevalence of axSpA to be 1.5% in men and 1.2% in women. The prevalence of r-axSpA was higher in men than in women (0.8% vs 0.4%), but nr-axSpA was found to be slightly more prevalent in women than in men (0.80% vs 0.76%). The prevalence of SpA based on the Amor and ESSG criteria among women and men was 2.0% and 1.5%, respectively (Table 2).

Two patients were newly diagnosed as having nr-axSpA and 1 as having r-axSpA (AS) during the study period. Previously, they had been diagnosed by a nonrheumatologist with lumbar disc hernia (LDH).

Table 2. Prevalence of inflammatory back pain and axial spondyloarthritis/spondyloarthritis according to various classification criteria among university employees.

	Total % (95% CI)	Female % (95% CI)	Male % (95% CI)
<b>IBP Criteria</b>			
ASAS	6.6 (4.3–9.7)	8.0 (4.9–12.4)	3.8 (1.2–8.9)
Berlin	7.1 (4.7–10.3)	8.8 (5.5–13.3)	3.8 (1.2–8.9)
Calin	21.5 (17.1–26.7)	24.2 (18.3–30.9)	16.8 (10.5–25.4)
<b>axSpA (ASAS)</b>			
axSpA (ASAS)	1.3 (0.4–3.0)	1.2 (0.2–3.5)	1.5 (0.2–5.5)
nr-axSpA (ASAS)	0.8 (0.16–2.3)	0.8 (0.1–2.9)	0.76 (0.02–4.3)
r-axSpA (ASAS)/AS (mNY)	0.5 (0.06–1.9)	0.4 (0.01–2.2)	0.8 (0.02–4.3)
SpA (Amor)	1.8 (0.7–3.8)	2.0 (0.7–4.7)	1.5 (0.2–5.5)
SpA (ESSG)	1.8 (0.7–3.8)	2.0 (0.7–4.7)	1.5 (0.2–5.5)

IBP: inflammatory back pain; ASAS: Assessment of SpondyloArthritis international Society; axSpA: axial spondyloarthritis; nr-axSpA: nonradiographic axSpA; r-axSpA: radiographic axSpA; AS: ankylosing spondylitis; mNY: modified New York criteria for AS; SpA: spondyloarthritis; ESSG: European Spondyloarthropathy Study Group.

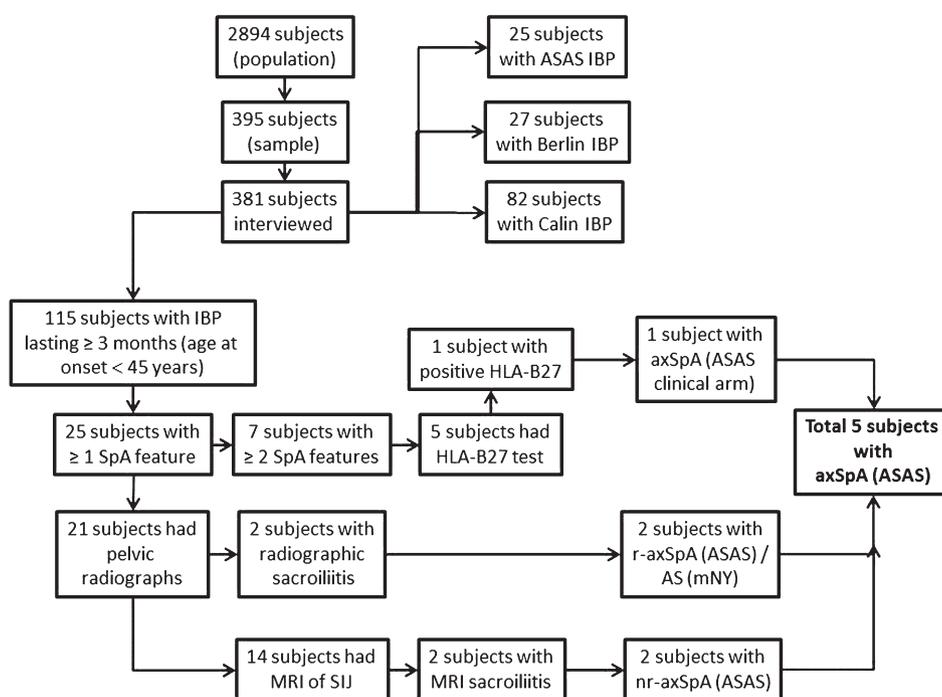


Figure 1. Study design and results in each stage. IBP: inflammatory back pain; ASAS: Assessment of SpondyloArthritis International Society; SpA: spondyloarthritis; axSpA: axial SpA; r-axSpA: radiographic axSpA; SIJ: sacroiliac joint; MRI: magnetic resonance imaging; AS: ankylosing spondylitis; mNY: modified New York criteria; nr-axSpA: nonradiographic axSpA.

## DISCUSSION

Our epidemiological study estimated a prevalence of 6.6% for IBP based on the ASAS criteria among the health campus employees of a university in Izmir, a major city in Turkey. The only previous study using the ASAS criteria for IBP<sup>6</sup> estimated a slightly higher prevalence of IBP (7.7%) among a primary care patient population in the United Kingdom. In that study, the prevalence was 15.4% by the Berlin criteria, which was also higher than the prevalence of 7.1% using the same criteria in our study<sup>6</sup>. The higher results in the UK

survey may be related to the study population including subjects with previous back pain history. On the other hand, the higher prevalence for IBP by the Calin criteria in the 2 studies could be due to a higher sensitivity and lower specificity of these criteria compared to the others<sup>17</sup>. The reported IBP prevalence in different studies may also vary according to geographic and genetic factors as well as criteria used and other methodological differences. A study based on the COPCORD (Community Oriented Program for the Control of Rheumatic Diseases) in the Mexican population<sup>7</sup> and a

retrospective analysis of data from the National Health and Nutrition Examination Survey (NHANES) II (1971–1975) in the United States<sup>8</sup> estimated significantly lower IBP prevalences of 3% and 0.8%, respectively. No standardized criteria sets to define IBP were used in these studies but the 2009–2010 NHANES survey used a data collection instrument specifically designed for IBP. In the latter survey, the age-adjusted prevalence of IBP was found to be 5.0% by Calin criteria among US adults. The IBP prevalence was 5.6% based on the ESSG criteria, and 5.8% and 6.0% based on the Berlin Criteria 8a and 7b, respectively<sup>5</sup>.

IBP was found to be about twice as prevalent among women than men in our study. The NHANES (2009–2010)<sup>5</sup> and UK studies<sup>6</sup> also reported somewhat higher frequencies in women than in men although the differences were not significant.

We estimated a prevalence of 1.3% for axSpA in the second stage of the study. More than half the patients with axSpA were found to have nr-axSpA with the prevalence of 0.8%. The only previous study<sup>9</sup> using the ASAS classification criteria estimated the prevalence of axSpA in the United States at 0.7%, which was lower than what we found. However, that was a retrospective cohort study and the diagnosis of axSpA was established on identification of elements of the ASAS criteria in medical records of at-risk patients treated by a representative sample of US rheumatologists. Thus, this may have led to underestimation of the prevalence of axSpA. In that study, the prevalence estimates of AS and nr-axSpA were 0.35% and 0.35%, respectively<sup>9</sup>. Our previous study among adults in an urban area of Izmir demonstrated a prevalence of 0.5% for AS based on the mNY criteria<sup>18</sup>. In our current study, we estimated the same prevalence for AS (mNY)/r-axSpA (ASAS).

The major disadvantage of the mNY criteria, the most commonly used criteria in epidemiological studies, is the requirement of radiographic sacroiliitis of at least grade 2 bilaterally or grade 3 unilaterally for the classification of AS<sup>13</sup>. Therefore, a diagnostic delay up to 10 years is commonly seen in AS, due to slow progression of radiographic damage<sup>19</sup>. Further, the mNY criteria focus on axial involvement, and other important SpA features such as extraspinal and extraarticular findings, family history, and response to nonsteroidal antiinflammatory drugs are not included<sup>13</sup>. The new ASAS axSpA criteria include MRI as a sensitive method for detection of inflammation in the sacroiliac joints, which is not well demonstrated by plain radiographs. They also include HLA-B27 testing when imaging was not performed. Therefore, these criteria allow for the identification of nr-axSpA. ASAS criteria have been demonstrated to be sensitive (82.9%) and specific (84.4%) for the classification of axSpA<sup>4</sup>.

Our current study confirmed that SpA was one of the most common forms of inflammatory rheumatic diseases in the general white population<sup>1</sup>. The prevalence of SpA by both

the ESSG and Amor criteria was estimated to be 1.8%, which was higher than that in the previous general population study in Izmir (a prevalence of 1.05% by the ESSG criteria)<sup>18</sup>.

The previous survey in the Turkish general population showed that AS affected men more frequently than women, but the rate of SpA, as a whole group, was slightly higher in women<sup>18</sup>. In our current study, these findings were confirmed. The prevalence of r-axSpA (AS) in men was established to be about twice as high as in women and the whole group of SpA was slightly more prevalent in women than men, with a ratio of 1.33. This study also showed that the prevalence of nr-axSpA was similar among women and men. There were no other population-based studies that compared the prevalence of nr-axSpA between men and women. Results from the German Spondyloarthritis Inception Cohort (GESPIC)<sup>19</sup>, which included patients with axSpA, demonstrated that AS was more prevalent in men; however, nr-axSpA was characterized by a higher prevalence among women. More recently, data from GESPIC were confirmed in the Herne<sup>20</sup> and Swiss Clinical Quality Management<sup>21</sup> cohorts, which included patients with r-axSpA and nr-axSpA. These results are in line with the well-known finding that male patients with AS have more severe radiographic changes<sup>22</sup>.

There were 3 patients who were newly diagnosed as having axSpA during this study period. One of them had r-axSpA. Although all these patients had IBP, previously they had been diagnosed by a nonrheumatologist with LDH. This result is consistent with that found in our previous study, which showed that an initial diagnosis of LDH is associated with a later diagnosis of AS<sup>3</sup>.

Our study suggests that axSpA is a more prevalent rheumatic disease than previously thought. Awareness of IBP and other symptoms of SpA in clinical practice may provide a chance for early diagnosis and treatment. The new ASAS criteria for axSpA that include MRI of SIJ and HLA-B27 testing may help to better classify patients with nr-axSpA.

## ACKNOWLEDGMENT

We thank the third-year medical students of the Special Study Module Group of Dokuz Eylul University School of Medicine for their assistance with this study.

## REFERENCES

1. Braun J, Bollow M, Remlinger G, Eggens U, Rudwaleit M, Distler A, et al. Prevalence of spondylarthropathies in HLA-B27 positive and negative blood donors. *Arthritis Rheum* 1998;41:58-67.
2. Feldtkeller E, Khan MA, van der Heijde D, van der Linden S, Braun J. Age at disease onset and diagnosis delay in HLA-B27 negative vs. positive patients with ankylosing spondylitis. *Rheumatol Int* 2003;23:61-6.
3. Gerdan V, Akar S, Solmaz D, Pehlivan Y, Onat AM, Kisacik B, et al. Initial diagnosis of lumbar disc herniation is associated with a delay in diagnosis of ankylosing spondylitis. *J Rheumatol* 2012; 39:1996-9.
4. Rudwaleit M, van der Heijde D, Landewe R, Listing J, Akkoc N,

- Brandt J, et al. The development of Assessment of SpondyloArthritis international Society classification criteria for axial spondyloarthritis (part II): validation and final selection. *Ann Rheum Dis* 2009;68:777-83.
5. Weisman MH, Witter JP, Reveille JD. The prevalence of inflammatory back pain: population-based estimates from the US National Health and Nutrition Examination Survey, 2009-10. *Ann Rheum Dis* 2013;72:369-73.
  6. Hamilton L, Macgregor A, Warmington V, Pinch E, Gaffney K. The prevalence of inflammatory back pain in a UK primary care population. *Rheumatology* 2014;53:161-4.
  7. Pelaez-Ballestas I, Flores-Camacho R, Rodriguez-Amado J, Sanin LH, Valerio JE, Navarro-Zarza E, et al. Prevalence of back pain in the community. A COPCORD-based study in the Mexican population. *J Rheumatol Suppl* 2011;86:26-30.
  8. Dillon CF, Hirsch R. The United States National Health and Nutrition Examination Survey and the epidemiology of ankylosing spondylitis. *Am J Med Sci* 2011;341:281-3.
  9. Strand V, Rao SA, Shillington AC, Cifaldi MA, McGuire M, Ruderman EM. Prevalence of axial spondyloarthritis in United States rheumatology practices: Assessment of SpondyloArthritis International Society criteria versus rheumatology expert clinical diagnosis. *Arthritis Care Res* 2013;65:1299-306.
  10. Sieper J, van der Heijde D, Landewe R, Brandt J, Burgos-Vagas R, Collantes-Estevez E, et al. New criteria for inflammatory back pain in patients with chronic back pain: a real patient exercise by experts from the Assessment of SpondyloArthritis international Society (ASAS). *Ann Rheum Dis* 2009;68:784-8.
  11. Rudwaleit M, Metter A, Listing J, Sieper J, Braun J. Inflammatory back pain in ankylosing spondylitis: a reassessment of the clinical history for application as classification and diagnostic criteria. *Arthritis Rheum* 2006;54:569-78.
  12. Calin A, Porta J, Fries JF, Schurman DJ. Clinical history as a screening test for ankylosing spondylitis. *JAMA* 1977;237:2613-4.
  13. van der Linden S, Valkenburg HA, Cats A. Evaluation of diagnostic criteria for ankylosing spondylitis. A proposal for modification of the New York criteria. *Arthritis Rheum* 1984;27:361-8.
  14. Dougados M, van der Linden S, Juhlin R, Huitfeldt B, Amor B, Calin A, et al. The European Spondylarthropathy Study Group preliminary criteria for the classification of spondylarthropathy. *Arthritis Rheum* 1991;34:1218-27.
  15. Amor B, Dougados M, Mijiyawa M. [Criteria of the classification of spondylarthropathies]. [Article in French] *Rev Rhum Mal Osteoartic* 1990;57:85-9.
  16. Taylor W, Gladman D, Helliwell P, Marchesoni A, Mease P, Mielants H. Classification criteria for psoriatic arthritis: development of new criteria from a large international study. *Arthritis Rheum* 2006;54:2665-73.
  17. Solmaz D, Akar S, Soysal O, Akkoc Y, Can G, Gerdan V, et al. Performance of different criteria sets for inflammatory back pain in patients with axial spondyloarthritis with and without radiographic sacroiliitis. *Clin Rheumatol* 2014;33:1475-9.
  18. Onen F, Akar S, Birlik M, Sari I, Khan MA, Gurler O, et al. Prevalence of ankylosing spondylitis and related spondyloarthritides in an urban area of Izmir, Turkey. *J Rheumatol* 2008;35:305-9.
  19. Rudwaleit M, Haibel H, Baraliakos X, Listing J, Marker-Hermann E, Zeidler H, et al. The early disease stage in axial spondylarthrititis: results from the German Spondyloarthritis Inception Cohort. *Arthritis Rheum* 2009;60:717-27.
  20. Kiltz U, Baraliakos X, Karakostas P, Igelmann M, Kalthoff L, Klink C, et al. Do patients with non-radiographic axial spondylarthrititis differ from patients with ankylosing spondylitis? *Arthritis Care Res* 2012;64:1415-22.
  21. Ciurea A, Scherer A, Exer P, Bernhard J, Dudler J, Beyeler B, et al. Tumor necrosis factor alpha inhibition in radiographic and nonradiographic axial spondyloarthritis: results from a large observational cohort. *Arthritis Rheum* 2013;65:3096-106.
  22. Lee W, Reveille JD, Davis JC Jr., Learch TJ, Ward MM, Weisman MH. Are there gender differences in severity of ankylosing spondylitis? Results from the PSOAS cohort. *Ann Rheum Dis* 2007;66:633-8.