

Frequency of Rheumatoid Arthritis in Young Females Presenting with Multiple Joints Pain Using Acr Diagnostic Criteria

SAIFULLAH KHAN KHALIL¹, AAMIR KAMRAN², SYED ZAHOO UL HASSAN ZAIDI³, SYED MUHAMMAD QASIM KHAN⁴, RASHIDA PARVEEN⁵, AFZAAL AKBAR⁶

¹Assistant Professor Pharmacology, Swat Medical College, Marghazar Road, Saidu Sharif, Swat

²Registrar Orthopaedic Unit Hayatabad Medical Complex, Peshawar

³Faculty of Eastern Medicine, Hamdard University, Karachi

⁴Pharm D (Pharmacy), Abasyn University Peshawar

⁵Department of Pharmaceutical Sciences, Faculty of Pharmacy, Superior University Lahore

⁶Pharm D (Pharmacy), Abasyn University Peshawar

Corresponding author: Dr Aamir Kamran, Email: Khattak_khan91@yahoo.com

ABSTRACT

Introduction: Rheumatoid arthritis (RA) is a chronic autoimmune disease causing morbidity and mortality in all population worldwide. It is present in 1 to 2 % in world's population

Objective: To determine the frequency of RA in young females presenting with multiple joints pain using ACR Diagnostic Criteria for RA.

Methodology: This study was Descriptive Cross-Sectional Study done at the Department of Medicine, Hayatabad Medical Complex, Peshawar for duration of six months from February 2021 to August 2021. In this study a total of 156 patients were observed to assess the frequency of rheumatoid arthritis in young females with multiple joints pain using ACR Diagnostic Criteria.

Results: Serology of RF and ACPA among 156 patients was positive in 93(59.6%) patients and Negative in 63(40.4). Acute phase reactants CRP and ESR among 156 patients were abnormal in 91(58.3%) and Normal in 65(41.7%) patients

Conclusion: Our study concludes that in young females with multiple joints pain, the frequency of rheumatoid arthritis was high by using ACR diagnostic criteria. Young females are prone to develop rheumatoid arthritis but remain undiagnosed and this may lead to adverse outcome and failure in prognosis and treatment. If it is diagnosed in early stage it may add to good health outcome and quality of life.

Key words: Rheumatoid arthritis; multiple joints pain, ACR Diagnostic Criteria

INTRODUCTION

Rheumatoid Arthritis (RA) is a chronic autoimmune disease-causing morbidity and mortality in all population worldwide ¹. It is present in 1 to 2 % in world's population. Women are affected more by the RA in young age almost 3 times more than men in young age ². In Pakistan the prevalence of RA is about 0.142% ³.

The increased incidence of RA in females is unknown but it is suggested that sex hormones play important role in the mechanism of RA ⁴. Literature shows that the signs and symptoms and incidence of RA and its flare decreases in pregnancy and increases post partum ⁵. RA is more common in females with null parity suggesting that hormonal variations play some role in the RA ^{6, 7}. The contraceptive use is considered to decrease the incidence of RA ⁸. The prevalence of RA in old age has no gender difference and affects both the gender equally. First degree relative have more risk of developing RA ⁹.

According to a study by Shamim R et al, out of 316 patients presenting with arthralgia, 85 had RA. Out of 85 patients with RA, 60 (70.5%) were females ¹⁰. The prevalence of RA varies in region and race. Tropical countries are less affected by the RA ¹¹.

The pathogenesis of the RA is unknown but an autoimmune process is involved in the pathogenesis of the RA. RA results due to the complex autoimmune reactions like CD4 T cells, osteoclasts, mono nuclear phagocytes, B cells and cytokines etc. synovial cell hyperplasia and the endothelial cell activation leads to inflammation and

destruction of the cartilages and bones ¹².

The clinical manifestations of the RA involve the persistent symmetric poly arthritis, constitutional symptoms and extrairticular involvement. RA usually affects joints of hands, feet, wrists, elbows and knees etc. It is a systemic disease affecting other parts of the body as well. It affects the cardiovascular and respiratory systems most commonly ¹³. The RA is diagnosed on the basis of criteria by the American college of rheumatology (ACR). This criteria addresses the treatment approaches in RA patients ¹⁴. Being a progressive disease, it can lead to joint destruction and disability if not properly treated. New standards for RA classification provide opportunities for early treatment. Initiating treatment with Disease modifying anti rheumatic drugs (DMARDs), especially in combination with short-term corticosteroids, can prevent progression and even alter the natural course of RA ¹⁵. In a recent study it is stated that involvement of multi-joint specially hand arthritis was the largest predictor of poor outcomes ¹⁶.

Young females are prone to develop RA but remain undiagnosed and this may lead to adverse outcome and failure in prognosis and treatment. If it is diagnosed in early stage it may add to good health outcome and of life quality. Our study was piloted to assess the frequency of RA in young females using the ACR diagnostic criteria in patients presented in tertiary care hospital. As the epidemiology of RA varies in gender, races and geography so this study will help us to determine the burden of disease in our population in young females.

MATERIALS AND METHODS

This descriptive cross-sectional study was carried out at the Department of Medicine, Hayatabad Medical Complex, Peshawar from February 2021 to August 2021. By using WHO calculator, sample size was 156. The inclusion criteria for our study include all female patients presenting with multiple joints pain with the duration of 6 months or more having age range 15-35 years whereas the exclusion criteria includes pregnant females, serious chronic disease like heart failure, patients with history of addiction, psychiatric illness, untreated malignancy or neurological disorder. Proper approval to this study was taken from institutional research and ethical committee. Informed consent was signed from all the subjects of our study.

Based on inclusion criteria all patients presenting with multiple joints pain (4-10 joints without large joint or > 10 joints with at least one large joint) with the duration of 6 months or more (as per operational definition above) were evaluated according to the ACR criteria for RA. 10 CC of blood were obtained in all the patients and were immediately sent to the hospital laboratory for detecting RF, ACPA, CRP and ESR as these are the components of the ACR criteria for RA. All the laboratory investigations were done from single hospital laboratory under supervision of single pathologist having minimum of five years of experience. RA was labeled on the basis of ACR criteria score having scores of 6 or more out of 10. All the information like age, weight, height, BMI, ACR score in patients with multiple joints pain, ACR score in RA patients, joint involvement, serology, acute phase reactants and duration of multiple joints pain was recorded in a pre-design proforma. All the data analysis was carried out by SPSS version 16. Mean and standard were computed for quantitative (continues) variables whereas frequencies and percentages were calculated for categorical and nominal data.

RESULTS

Totally 156 subjects were enrolled in our study. Among 156 subjects, 48(30.8%) were having age 15-20 years, 32(20.5%) have age 21-25 years, 40(25.6%) have 26-30 years while 36(23.1%) participants have 31-35 years of age. The mean age (SD) was 25 (3.87) years. (Figure 1) Serology of RF and ACPA among 156 patients was positive in 93(59.6%) subjects and negative in 63(40.4) subjects. (Figure 2) Acute phase reactants CRP and ESR among 156 patients were abnormal 91(58.3%) subjects while normal in 65(41.7%) subjects. (Figure 3)

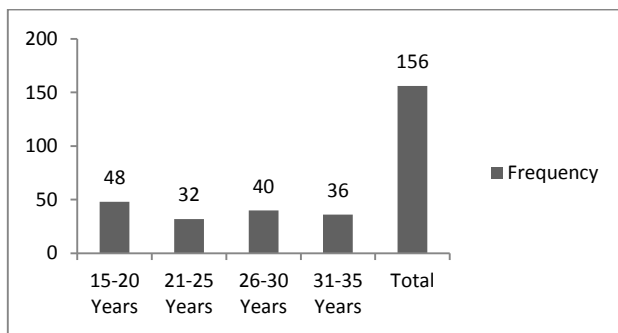


Figure 1: Subjects distribution based on age

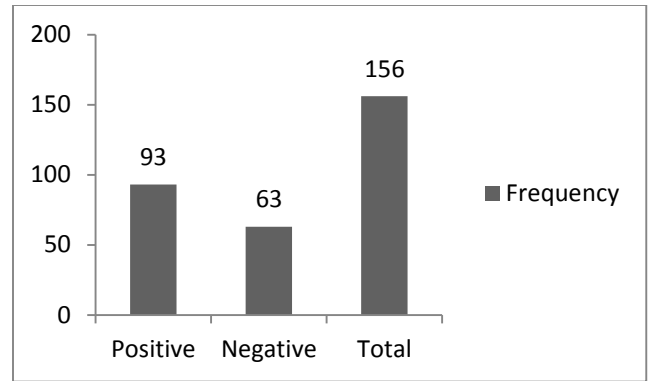


Figure 2: Distribution of subjects based on serology of RF and ACPA

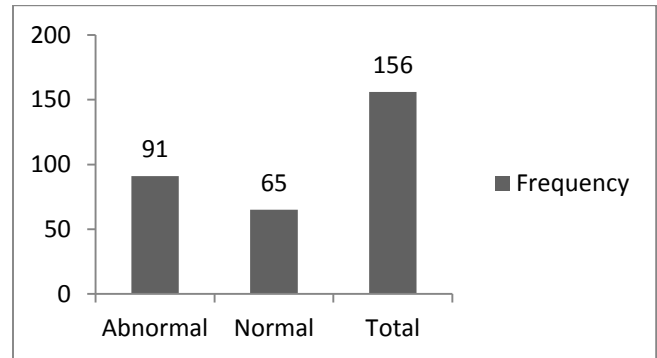


Figure 3: Distribution of patients based on acute phase reactants CRP and ESR

DISCUSSION

Pain, disability, and death are common outcomes of rheumatoid arthritis. Articular and extra-articular tissues both are affected by this inflammatory rheumatic illness¹⁷. In the vast majority of patients, chronic inflammation results in erosive joint deterioration and functional disability^{18, 19}. The beginning of illness differs from patient to patient, depending on the kind, number, and degree of joint involvement. The intensity of the inflammatory process, as well as genetic makeup, incidence of swollen joints, serum autoantibody and inflammatory process severity, may all influence the course of the illness^{20, 21}. In our study, Serology of RF and ACPA among 156 patients was positive in 93(59.6%) subjects and negative in 63(40.4) subjects. (Figure 2) Acute phase reactants CRP and ESR among 156 patients were abnormal 91(58.3%) subjects while normal in 65(41.7%) subjects. A previous study reported similar results to our study²². Another study done by Chunhua Xun et al. also reported comparable results to our study²³.

Early RA presents with symptoms that are similar to those seen in other inflammatory diseases. Patients with early RA are frequently characterized as undifferentiated arthritis, which is difficult to distinguish from those other inflammatory arthritis until a definitive diagnosis. In the past, patients with early RA were those who had symptoms for <2 years, with a predilection for <12 months. However, many rheumatologists are now willing to acknowledge individuals who had symptoms for no more than six weeks.. At this time, "early" RA is defined as those who have had

symptoms for less than three months²⁴. However, not all researchers have adopted this classification, since some rheumatologists feel individuals have either developed RA or undifferentiated inflammatory arthritis.

Early therapy of RA may have a favorable influence on RA's prognosis, avoiding joint erosions and reducing the progression of erosive disease²⁵. An early diagnosis and therapy might have an influence even if the disease is in remission²⁶. It is difficult to distinguish early RA from non-RA at the outset of illness, and the use of ACR criteria for early diagnosis has limitations. Because there are insufficient clinical or laboratory evidences at the outset of arthritis, this criterion is insufficiently sensitive to detect early RA²⁷.

CONCLUSION

Our study concludes that in young females with multiple joints pain, the frequency of rheumatoid arthritis was high by using ACR diagnostic criteria. Young females are prone to develop rheumatoid arthritis but remain undiagnosed and this may lead to adverse outcome and failure in prognosis and treatment. If it is diagnosed in early stage it may add to good health outcome and quality of life.

REFERENCES

- Butterworth RF. Neurosteroids in hepatic encephalopathy: novel insights and new therapeutic opportunities. *The Journal of steroid biochemistry and molecular biology*. 2016;160:94-7.
- Luo M, Guo J-Y, Cao W-K. Inflammation: A novel target of current therapies for hepatic encephalopathy in liver cirrhosis. *World J Gastroenterol*. 2015;21(41):11815.
- Patidar KR, Bajaj JS. Covert and overt hepatic encephalopathy: diagnosis and management. *Clin Gastroenterol Hepatol*. 2015;13(12):2048-61.
- Shawcross DL, Dunk AA, Jalan R, Kircheis G, De Knegt RJ, Laleman W, et al. How to diagnose and manage hepatic encephalopathy: a consensus statement on roles and responsibilities beyond the liver specialist. *Eur J Gastroenterol Hepatol*. 2016;28(2):146.
- Riggio O, Efrati C, Catalano C, Pediconi F, Mecarelli O, Accornero N, et al. High prevalence of spontaneous portal-systemic shunts in persistent hepatic encephalopathy: a case-control study. *Hepatology*. 2005;42(5):1158-65.
- Poordad FF. The burden of hepatic encephalopathy. *Aliment Pharmacol Ther*. 2007;25:3-9.
- Bustamante J, Rimola A, Ventura P-J, Navasa M, Cirera I, Reggiardo V, et al. Prognostic significance of hepatic encephalopathy in patients with cirrhosis. *J Hepatol*. 1999;30(5):890-5.
- Schaffner F. Abdominal enlargement and masses. *Gastroenterology*. Haubrich WS, Schaffner F, Berk JE, Eds. WB Saunders, Philadelphia; 1998.
- Volk ML, Tocco RS, Bazick J, Rakoski MO, Lok AS. Hospital re-admissions among patients with decompensated cirrhosis. *The American journal of gastroenterology*. 2012;107(2):247.
- Stepanova M, Mishra A, Venkatesan C, Younossi ZM. In-hospital mortality and economic burden associated with hepatic encephalopathy in the United States from 2005 to 2009. *Clin Gastroenterol Hepatol*. 2012;10(9):1034-41. e1.
- Bajaj JS, Wade JB, Gibson DP, Heuman DM, Thacker LR, Sterling RK, et al. The multi-dimensional burden of cirrhosis and hepatic encephalopathy on patients and caregivers. *The American journal of gastroenterology*. 2011;106(9):1646.
- Ferenci P, Lockwood A, Mullen K, Tarter R, Weissenborn K, Blei AT. Hepatic encephalopathy—definition, nomenclature, diagnosis, and quantification: final report of the working party at the 11th World Congresses of Gastroenterology, Vienna, 1998. *Hepatology*. 2002;35(3):716-21.
- Brusilow SW. Hyperammonemic encephalopathy. *Medicine (Baltimore)*. 2002;81(3):240-9. doi:10.1097/00005792-200205000-00007.
- Donovan JP, Schafer DF, Shaw Jr BW, Sorrell MF. Cerebral oedema and increased intracranial pressure in chronic liver disease. *The Lancet*. 1998;351(9104):719-21.
- CHATAURET N, BUTTERWORTH RF. Effects of liver failure on inter-organ trafficking of ammonia: implications for the treatment of hepatic encephalopathy. *J Gastroenterol Hepatol*. 2004;19:S219-S23.
- Schafer D, Jones EA. Hepatic encephalopathy and the γ -aminobutyric-acid neurotransmitter system. *The Lancet*. 1982;319(8262):18-20.
- Birch JT, Bhattacharya S. Emerging trends in diagnosis and treatment of rheumatoid arthritis. *Prim Care*. 2010;37(4):779-92.
- El Miedany Y, Youssef S, Mehanna AN, El Gaafary M. Development of a scoring system for assessment of outcome of early undifferentiated inflammatory synovitis. *Joint Bone Spine*. 2008;75(2):155-62.
- Rahmani M, Chegini H, Najafizadeh SR, Azimi M, Habibollahi P, Shakiba M. Detection of bone erosion in early rheumatoid arthritis: ultrasonography and conventional radiography versus non-contrast magnetic resonance imaging. *Clin Rheumatol*. 2010;29(8):883-91.
- Gossec L, Combescurie C, Rincival N, Saraux A, Combe B, Dougados M. Relative clinical influence of clinical, laboratory, and radiological investigations in early arthritis on the diagnosis of rheumatoid arthritis. Data from the French Early Arthritis Cohort ESPOIR. *The Journal of rheumatology*. 2010;37(12):2486-92.
- Finckh A, Liang MH, van Herckenrode CM, De Pablo P. Long-term impact of early treatment on radiographic progression in rheumatoid arthritis: a meta-analysis. *Arthritis Care & Research: Official Journal of the American College of Rheumatology*. 2006;55(6):864-72.
- Shen C, Sun XG, Liu N, Mu Y, Hong CC, Wei W, et al. Increased serum amyloid A and its association with autoantibodies, acute phase reactants and disease activity in patients with rheumatoid arthritis. *Mol Med Rep*. 2015;11(2):1528-34. doi:10.3892/mmr.2014.2804.
- Xun C, Zhao Y. Comparison of serological markers between ACPA⁺ and ACPA⁻ of RA patients. *Rheumatol Int*. 2012;32(5):1143-6. doi:10.1007/s00296-010-1757-y.
- Aletaha D, Eberl G, Nell V, Machold K, Smolen J. Practical progress in realisation of early diagnosis and treatment of patients with suspected rheumatoid arthritis: results from two matched questionnaires within three years. *Ann Rheum Dis*. 2002;61(7):630-4.
- Goekoop-Ruiterman YP, de Vries-Bouwstra JK, Allaart CF, van Zeben D, Kerstens PJ, Hazes JMW, et al. Comparison of treatment strategies in early rheumatoid arthritis: a randomized trial. *Ann Intern Med*. 2007;146(6):406-15.
- Finckh A. Early inflammatory arthritis versus rheumatoid arthritis. *Curr Opin Rheumatol*. 2009;21(2):118-23.
- Arnett FC, Edworthy SM, Bloch DA, Mcshane DJ, Fries JF, Cooper NS, et al. The American Rheumatism Association 1987 revised criteria for the classification of rheumatoid arthritis. *Arthritis & Rheumatism: Official Journal of the American College of Rheumatology*. 1988;31(3):315-24.