THE PROGNOSTIC VALUE OF ANTI CITRULLINATED PROTEIN ANTIBODY TEST IN RHEUMATOID ARTHRITIS

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ABSTRACT

Background: Rheumatoid arthritis (RA) is a chronic, systemic inflammatory disorder that may affect many tissues and organs, but principally attacks synovial joints. About 1% of the world's population is afflicted by rheumatoid arthritis, women three times more often than men. Onset is most frequent between the ages of 40 and 50, but any age group can be affected.

Aim of Work : To study the value of Anti cyclic citrullinated protein (Anti-CCP) test as a predictor of disease course, disability, prognosis and response to treatment in Rheumatoid arthritis patients of a specific age group with regard to history, clinical and radiological findings. The results for Anti-CCP will be compared to Rheumatoid factor test. **Patients:** 80 female cases were collected from rheumatology outpatient clinics, with disease duration of 3- 5 years and age range from 35 to 40 years **Methods**: All cases underwent thorough history taking, clinical examination, radiological and laboratory assessment as well as disability and health assessment scoring. Each patient will undergo a quantitative test of Anti cyclic citrullinated protein (Anti-CCP) and Rheumatoid factor. **Results :** Our study showed that there was high significant relation between positive and negative RF and Anti-CCP titres. (p=0.0000) for the patients in the studied group. There was a high significant correlation between Anti-CCP and RF titres (r=0.81, p<0.05)

Also there was a significant correlation between Anti-CCP and ACR disability class. In addition, there was a positive but not significant correlation between Anti-CCP and STLW score. (Clinical findings score- Swelling, Tenderness, limitation of movement, Warmth) and with VAS (Visual analogue scale) **Conclusion:** Our study found that Anti-CCP has a reasonably good prognostic value, especially as regards disability and functional status prediction. It also showed that Anti-CCP is very comparable to Rheumatoid factor when comparing prognostic values. It is only slightly more indicative in terms of clinical, radiological and health assessment.

Keywords: Rheumatoid Arthritis, Anti-CCP, ACPA, Rheumatoid factor, prognosis, radiographic damage, disability, functional status.

INTRODUCTION

Rheumatoid arthritis (RA) is a chronic, systemic inflammatory disorder that may affect many tissues and organs, but principally attacks synovial joints. The process produces an inflammatory response of the synovium (synovitis) secondary to hyperplasia of synovial cells, excess synovial fluid, and the development of pannus in the synovium (**Majithia and Geraci 2007**). A high percentage of patients with rheumatoid arthritis have substantial titres of antibody to peptides containing citrulline, and citrullinated peptides are found in rheumatoid synovium (**Yelin** *et al.*, **1999**). The Anti citrullinated protein antibody (ACPA) tests, also known as the anti-cyclic citrullinated peptide (anti-CCP) antibody test, is an enzyme-linked immunosorbent assay that tests for the presence of antibodies that recognizes specific antigens containing citrulline (**Schellekens** *et al.*, **1998**). The specificity of anti-CCP is around 95-98% as regards undifferentiated forms of arthritis that do not develop into RA. IgM RF is often found in the same patients, but with much lower specificity for RA. One study using a CCP1 assay showed a sensitivity of 55% and a specificity of 97% for RA, when both anti-CCP and IgM RF were positive in the early stage of arthritis (**Jansen** *et al.*, **2002**). Several observations have indicated that anti-CCP positive early RA patients may develop a more erosive disease than those without anti-CCP. Other investigators have confirmed this, and suggested the superiority of anti-CCP over IgM RF in predicting an erosive disease course. The use of anti-CCP results in the decision whether a patient should be treated aggressively at an early stage or not is an important area for research (Kroot *et al.*, **2000**).

Methods

This study was conducted on 80 Rhematiod patients were collected from Rheumatology outpatient clinics of Alhussein University Hospital, Sayed Galal University hospital and the National Research Center rheumatology clinic. The cases were diagnosed according to the 1987 classification criteria of the American College of Rheumatology (Arnett *et al.*, 1988).

Inclusion criteria: 1- All cases were females.(there age ranged from 35-40).

2- All cases had disease duration of 3 to 5 years since the onset of diagnostic symptoms.

3- All cases were receiving steroid doses of 5 to 15 mg/day in combination with Non steroidal anti inflammatory drugs and either Methotrexate (doses ranged from 7.5 to 17.5mg/week) or Leflonamide.

Exclusion Criteria: Pregnancy, DM, ischemic heart disease, hypertension, chronic liver disease, chronic kidney disease, other autoimmune diseases or any other chronic diseases.

All participants accept to participate in this study.

Thorough history taking and clinical examination were done for each case, clinical findings of local examination were made using the Swelling tenderness limitation of movement warmth "STLW" score system. The DAS28 score was calculated for all patients (Aletaha *et al.*, 2004).

DAS 28 Score:

Disease Activity Score (DAS) (Van Der Heijde *et al.*, 1993) provides a composite measure of disease activity based on joint counts, ESR, and patient global assessment of general health rather than measuring response. This is not dependent on baseline parameters to show a difference and therefore provides a more absolute measure of disease severity, which can be followed over time. The DAS28 based on joint counts of 28 joints. (Prevoo *et al.*, 1995).

The collected scores were calculated using the following formula:

 $[0.56 \times \sqrt{(TEN \ 28)]} + [0.28 \times \sqrt{(SW \ 28)]} + [0.70 \times (ESR \ after \ 1 \ hour \ in \ mm^{x})] + [0.014 \times (patient's \ assessment \ in \ mm)] = DAS \ 28 \ score.$ TEN28: Number of tender joints. SW28: Number of swollen joints. ESR: Erythrocyte sedimentation rate.

The net score indicates the disease activity according to the following:

 \leq 3.2 score indicates inactivity.

 $> 3.2 \le 5.1$ score indicates moderate activity.

>5.1 score indicates very high activity. (Van Der Heijde et al., 1993),

Radiological assessment was performed using the Sharp score:(Van der Heijde modification) (Sharp et al, 1985., Van Der Heijde D 1999.

The patient's disability status and abilities at the time of examination were determined based upon a health assessment questionnaire and the American College of Rheumatology revised criteria for functional status in RA, classification from class 1 to 4 (**Hochberg** *et al.*. **1992**). All patients were given a score percentage. The Health Assessment Questionnaire (HAQ) was translated into Arabic to be applicable for use with Egyptian patients.

The following laboratory investigations were done:

Blood sampling:

Venous blood samples (random) were obtained from each subject in the study one mL was delivered into EDTA tube for CBC; 2mL were delivered into tube containing trisodium citrate for ESR. The rest was left to clot, centrifuged, and the separated serum was divided in 2 aliquots one used for routine biochemical laboratory investigations (liver, kidney, CRP and random blood glucose) while the second aliquot was stored freezed at -20 C until used for determination of Anti CCP and RF.

The following laboratory investigations were performed to all patients participating in this study :

- 1- Erythrocyte sedimentation rate by Westergren method.
- 2- Liver, kidney function tests and Random blood sugar. (all were analzeyd by Hitachi Cobas C311 automated analyzer).
- 3- Complete Blood Picture by fully autmoated cell counter Sysmex Kx-21 N.
- 4- Quantitative determination of Rhumatiod factor by turbidimetry method using Kit of Spinreact.
- 5- Anti CCP antibodies (by enzyme linked immunosorbent assay for quantitative determination of anti-cyclic citrullinated peptide.

Kits: ORGENTEC Diagnostika GmbH.

Analyzer: Stat fax -2600 (ChroMate)

The principle of assay of anti-CCP antibody:

The anti-CCP antibody Kit is based on an ELISA method. The test utilizes microtitre plate wells coated with citrulinated synthetic peptides (antigen).

Diluted patients sample serum is applied to the wells and incubated. If specific antibodies are present, they will bind to the antigen in the wells. Unbound materials are washed away and any bound antibodies are detected by adding horse radish peroxidase (HRP) labeled anti-human IgG, followed by a second washing step and incubation with substrate. The presence of reacting antibodies will result in the development of colour, which is proportional to the quality of bound antibody, and this is determined photometrically.

Statistical analysis of data:

Statistics were done by computer using Epi - info. Software, version 6.04. A word processing, data base and statistics program (WHO, 2001). The tests used were: X mean,

SD standard deviation: to measure the central tendency of data and the distribution of data around their mean. Student's t test: for testing statistical significant difference between means of two samples. Median: is a measure of central tendency when extremes of values are found in data. X2 tests (Chi square test) to test statistical significant relation between different variable or grades (qualitative data) or percentages. Z test to test statistical significant difference between two percentages. F "analysis of variance" (or ANOVA test) to test statistical significant difference between Between more than 2 means i.e. differences between 3 or more groups at the same time. Pearson's correlation test (correlation coefficient r) to test a positive or negative linear relationship between two variables (one dependent and the other is independent variable).

RESULTS ANALYSIS:

Descriptive study.

The present study was carried out on 80 Rheumatoid arthritis female patients diagnosed according to the 1987 American College of Rheumatology criteria. All cases had disease duration of 3 to 5 years since the onset of diagnostic symptoms. Their age ranged from 35 to 40 years old), (Mean age was 38.01 ± 3.4) The BMI (Body mass index) of the study group: (Mean 31.98 ± 5.49), (Range: 21.7 to 53.9) and Waist to hip ratio: (Mean=1.15 ± 0.2), (Range: 0.82 to 1.75). Extra-articular manifestations were present in 55 cases (68%) and subcutaneous nodules were present in 10 cases (12.5%).

-DAS 28 score: (Disease Activity score): It had a mean value of 5.35, range from 2.02 to 7.35 in the study group.

-STLW score: (Swelling, Tenderness, And Limitation of movement, Warmth): It had a mean value of 29.5, range from 0 to 97 in the study group.

-Deformities: were present in 23 cases (29%).

-Radiographic progression: Modified Sharp score: It had a mean value of 24.8, range from zero to 128 in the study group. (Table 1).

Table (1): Demographic data and medical history of the studied group (80 cases):

Variables	Number of cases	Percentage
Onset		
Acute	35	45
Gradual	45	55
Course		
Intermittent	9	11.3
Progressive	68	85
Steady	3	3.7
	Mean	Range
Age /year	38.01 ±3.4	35 to 40 years
Duration /year	3.29 ±0.86	2 to 5 years
Weight/kg	81.63±14.37	42 to 145
Height /m	1.59 ± 0.07	1.37 to 1.74
Waist circumference/cm	98.2±15.42	65 to 134
Hip circumference /cm	87.74±17.68	52 to 133
BMI(kg/ m^2)	31.98 ±5.49	21.7 to 53.9
Waist / hip ratio	1.15 ±0.2	0.82 to 1.75

BMI: Body mass index

Table 1 shows the studied group with respect to age, duration, weight, height, waist circumference, hip circumference, Body Mass Index, Waist /Hip ratio. The present study

was conducted on 80 females of Rheumatoid arthritis. 45% were acute onset, 53.8% were gradual onset, 1.2% were insidious onset 11.3% were intermittent, 85% were progressive, and 3.7% were steady.

Anti-CCP and RF results:

- The mean value of RF titre in the studied group was 82.59 ± 113.19 IU/ml, the range was < 8 to 512 IU/ml and the median was 32 IU/ml.

- The mean value of Anti-CCP titre in the studied group was 303.19 ± 336.94 IU/ml, the range was 3.7 to 1213 IU/ml and the median was 194 IU/ml. Table (2).

Table (2): RF and Anti-CCP titre results of the studied group (80 cases) :

Variables	Mean	Range	Median
RF(I.U/ml)	82.59±113.19	<8-512	32
Anti CCP(I.U/ml)	303.19 ±336.94	3.7 - 1213	194

RF: Rheumatoid factor

Anti CCP: Anti- cyclic citrullinated protein.

Comparison study:

By comparing the results of Anti-CCP and RF, this study demonstrated that there was a high significant difference between RF and Anti-CCP positive and negative results. Table (3), Figure (1).

Table (3): Relation	ı between RF	and Anti- CCP:
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	Anti- CCP< 2	5 Anti- CCP>25	Total
RF<8	19	9	28 (35%)
RF>8	5	47	52 (65%)
Total	24(30%)	56(70%)	80

X2 29.4 p=0.0000



Fig (1): Shows distribution of cases according to RF and Anti-CCP positive and negative groups.

Variables	RF<8	RF>8	Test of	Р
	No=28	No.=52	significance	value
Anti-CCP	110.9±236.8	406.74±339.02	t=4.1	0.0000
Median	11.65	347		
Range	3.7-988	5-1213		
Sharp score	26.67±23.24	23.69±27.91	t=0.46	0.6461
Median	22	16		
Range	0-92	0-128		
HAQ	36.88±22.96	40.81±25.24	t=0.68	0.4922
Median	38.75	35		
Range	0-83.75	2.5-85		
ACR class	1.61±0.92	2.22±1.14	t=2.43	0.0174
Range	1-4	1-4		
VAS	5±2.75	6.45±2.51	t=2.38	0.0199
Range	0-10	0-10		
DAS28	5.29±1.38	5.39±1.46	T=0.26	0.7955
Range	2.02-7.35	2.22-7.57		
STLW	26±12.99	31.37±22.19	t=1.17	0.2440
Range	0-54	0-97		

Table (4): Difference between laboratory scores, pain and severity of disease as regard to positive or negative RF results:

RF: Rheumatoid factor

Anti-CCP: Anti cyclic citrullinated protein antibody

HAQ: Health assessment Questionnaire

ACR Class: American college of Rheumatology disability classification

DAS: Disease activity score

VAS: Visual analogue scale

STLW score: Swelling tenderness limitation of movement warmth

There was high statistical significant difference between positive or negative RF results in Anti-CCP readings.

There was high significant difference between positive or negative RF results in ACR class & VAS.

Variables	Anti-CCP	Anti-CCP >	Test of	P value
	<25- No =	25- No = 56	significance	
	24			
Sharp	18.96±20.13	27.72±28.28	t=1.33	0.1883
score				
Median	16	20		
Range	0-92	0-128		
HAQ	37.24±23.84	40.36±24.77	t=0.52	0.6036
Median	38.12	37.5		
Range	0-83.75	2.5-85		
ACR	$1.54{\pm}0.88$	2.2±1.13	T=2.53	0.0132
class				
Median	1	2		
Range	1-4	1-4		
VAS	5.04 ± 2.49	6.33±2.67	t=2.01	0.0485
Median	5	7		
Range	0-9	1-10		
DAS28	5.47±1.23	5.3±1.51	t=0.46	0.6482
Median	5.63	5.4		
Range	2.02-7.35	2.22-7.57		
STLW	24.96±12.08	31.43±21.79	t=1.36	0.1764
Median	23.5	29.5		
Range	0-54	0-97		

Table (5): Difference between clinical, radiological and disability scores as regard to positive or negative Anti-CCP:

Anti-CCP: Anti cyclic citrullinated protein antibody

HAQ: Health assessment Questionnaire

ACR Class: American college of Rheumatology disability classification

DAS: Disease activity score

VAS: Visual analogue scale

STLW score: Swelling tenderness limitation of movement warmth

There was high significant difference between positive or negative Anti-CCP in American College of Rheumatology disability class and in VAS (Visual analogue scale).

Correlation studies:

A correlation study was performed between the results of RF and Anti-CCP showing significant positive correlation Fig (3)

A correlation between Anti-CCP and clinical, radiological and disability scorers showing the following:

1- There was significant positive correlation between Anti-CCP and ACR disability class.

2- There was significant positive correlation between Anti-CCP and RF results.

3 - There was positive but not significant correlation between Anti-CCP and STLW scores

4 -In addition, there was positive but not significant correlation between Anti-CCP and VAS scores.

5 -The rest of items showed no statistical correlation. Table (6).

A correlation between Anti-CCP and other Lab results showed

No statistical significant correlation between Anti-CCP and the studied variables table (7).



Figure 3: Correlation between Anti-CCP and RF titres of the study group:

Table (6): A correlation between Anti-CCP and clinical, radiological and disability scorers.

Correlation- Anti-CCP	r	р	Significance
Sharp score	0.07	>0.05	NS
HAQ	0.04	>0.05	NS
ACR class	0.23	< 0.05	S
VAS	0.19	>0.05	NS
DAS28	0.14	>0.05	NS
STLW	0.2	>0.05	NS
RF	0.81	< 0.01	S
	a <u>a</u> , , , , ,		

NS: Not significant

S: Significant

Table (7): A correlation between Anti-CCP and Lab results.

Correlation- Anti-CCP	r	р	Significance
ESR	0.01	>0.05	NS
Haemoglobin	0.04	>0.05	NS
RBCs count	0.03	>0.05	NS
WBCs count	0.12	>0.05	NS
Platelets	0.01	>0.05	NS
Ceartinine	0.16	>0.05	NS
AST	0.13	>0.05	NS
ALT	0.02	>0.05	NS

RBCs: Red Blood Cells

WBCs: White Blood Cells

AST: Aspartate Transaminase

ALT: Alanine Transaminase

In the present study we aimed to investigate the vealue of Anti-CCP as a prognostic factor in Rheumatoid Arthritis and compare the results with Rheumatoid factor test.

Eighty female cases of Rheumatoid arthritis within the age group (Range 35 to 40 years old) were enrolled this study. All cases underwent detailed history taking, thorough clinical evaluation including functional impairment, radiological and laboratory assessment. (including Liver function tests, kidney function tests, random blood sugar, CBC, ESR, RF, and Anti–CCP).

Our results showed that there was a statistical significant higher RF titre in Anti-CCP positive patients compared with Anti-CCP negative patients (p < 0.01) (mean value for RF titre was 111.27± 124.25 IU/ml with positive Anti-CCP and 16.88±27.09 IU/ml with negative Anti-CCP), and significant positive correlation between Anti-CCP titre and RF titres (r=0.81, p<0.05). **Benjamin** *et al.* (2010) made a similar study, Most of his cases were anti-CCP (75%) positive and RF (80%) positive, with a high statistical significant positive correlation between Anti –CCP and RF (r=0.44, p<0.001). Kastbom *et al.* (2004) also studied Anti-CCP and R.F tests titres as predictors of disease course and prognosis and used the same disease duration as our study (3 years) but he used about 3 times the number of patients (242). Their results showed there was a significant propensity for Anti-CCP antibody positive patients to be positive also for RF (p=0.001) with 78% of the RF positive and 40% of the RF negative patients were Anti-CCP antibody positive. They also concluded that Anti-CCP was better as a predictor of the disease course over 3 years than rheumatoid factor. Nehir *et al.* (2006) also found a significant correlation between the levels of Anti-CCP and RF (r=0.57, p<0.001).

Our results showed a statistically significant difference between positive and negative Anti-CCP and ACR disability classification (ACR class was higher with positive than with negative Anti-CCP) and a significant positive correlation between Anti-CCP titre and ACR disability classification. In their study, **Farragher** *et al.* (2010) made a similar study over a period of five years, patients who were anti-CCP positive had a mean change on their HAQ scores of 0.20 points (95% CI 0.09 to 0.31), which reflected a worsening of functional disability. Those who were negative for the antibodies had no worsening in HAQ score, with a mean change of 0.005 (95% CI -0.06 to 0.07). Although they used the HAQ itself as a measure for functional status/disability over a slightly longer disease duration than in our study (5 years), their results came in accordance with ours as it showed strong positive correlation between positive Anti-CCP test and disability. Nehir *et al.* (2006) found by logistic regression analysis that only the HAQ score (functional disability) and the duration of morning stiffness are the variables significantly associated with anti-CCP positivity in patients with advanced RA, anti-CCP positive RA patients had more functional disability than anti-CCP negative ones according to HAQ.

Our results showed a statistically significant difference between positive and negative Anti-CCP with VAS (Visual Analogue Scale), with a mean VAS score of 5.04 ± 2.49 with negative APCA and 6.33 ± 2.67 with positive Anti-CCP (test of significance = 2.01, p = <0.05). Landmann *et al.* (2010) who correlated the level of anti-CCP and disease activity using the DAS-28, as in our study and used, a patient-reported visual analog scale (VAS) score, and the ESR. They found a significant positive correlation between Anti-CCP and VAS. On the other hand, Nehir *et al.* (2006) found no significant correlation between VAS and anti CCP titres, but detected a higher VAS score in Anti-CCP positive patients compared to Anti-CCP negative patients with a difference in mean of 30.1 ± 29.4 vs 19.7 ± 20.6 respectively.

Our study revealed that Anti-CCP has almost the exact same predictive value as that given by R.F. Lindqvist *et al.* (2005) found in a study of patients with early arthritis, that baseline values of anti-CCP (67%) and RF (69%) showed similar sensitivity for prediction of radiographic progression at 5 years, but they did not test disability and functional status. **Yasser** *et al.* (2010) found a positive correlation between Anti-CCP titre with clinical severity and functional disability, but did not express difference between findings of positive Anti-CCP and positive R.F.

We found that anti-CCP is a good prognostic marker for Rheumatoid arthritis patients especially with disability prediction, and gave more or less similar predictive values with Rheumatoid factor. The strongly positive correlation between the titres of both tests shows that they are likely to be similar with patients of different grades of severity. This agrees with the conclusion found by **Yasser** *et al.* (2010). Who found that the currently available data suggested that anti-CCP is of good prognostic value besides RF. This is in contrast to **Zendman** *et al.* (2006). Who found that Anti-CCP positivity has prognostic potential, but the absolute level of anti-CCP has not been demonstrated to be a useful measure of future severity of disease. Our results also disagree with Lindqvist *et al.* (2005). who stated that combining RF with anti-CCP results appeared to help in prediction: RF+/anti-CCP+ patients had greater progression than RF+/anti-CCP- and RF-/CCP- patients at 5 years, while in our study, the subgroup analysis of RF and Anti-CCP positive versus Anti-CCP negative alone.

Limitations: The number of patients in the study group was small when compared to similar studies, and so did not allow better analysis of the findings. Also, the disease duration of the study group was short when compared to similar studies.

The main limitation of the Health Assessment Questionnaire used in our study was that it was not tailored for Egyptian lifestyle especially the low socioeconomic levels and the illiteracy of many of the patients in our study group prevented them from reading and filling the Questionnaire themselves (although it was translated into Arabic), which hampered the accuracy of the results. Radiographic and clinical assessment of the cases including DAS score calculation was not done periodically (once at baseline then once again every year as in most similar studies) to enable more accurate detection of progression and severity. In addition, specialized radiologists were not involved in the study for a better estimation of the radiographic score.

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القيمة التكهنية لاختبار الاجسام المضادة للبروتين الستروليني في الالتهاب المفصلي الروماتويدي

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*قسم الروماتيزم بطب الاز هر **قسم الباطنة بالمركز القومي للبحوث***قسم الباثولوجيا الاكلينيكية بطب الاز هر

مرض التهاب المفاصلل الروماتويدى هو التهاب مزمن يؤثر على العديد من الانسجة و الاعضاء ولكنه يهاجم اساسا المفاصل الزلالية. يصاب حوالى ١ % من سكان العالم بمرض التهاب المفاصل الروماويدى، ونسبة حدوثه فى النساء ثلاث مرات اكثر من الرجال. وهو اكثر شيوعا مابين ٤٠ و ٥٠ سنة ولكن من الممكن ان يؤثر المرض على اى مرحلة عمرية.

وقد اشارت العديد من الدراسات الى ان الاجسام المضادة للبروتين الستروليني تكون ايجابية في وقت مبكر في نسبة كبيرة من مرضى التهاب المفاصل الروماتويدي. ولكن كان يجب التأكد من مدى امكانية استخدام هذا الاختبار للتنبؤ بمجرى المرض وبمدى تاثيره على حياة المرضى.

ولقد اجرى هذا البحث لدراسة قيمة اختبار الاجسام المضادة للبروتين الستروليني في مرضى الالتهاب المفصلي الروماتويدي كمتوقع لمجرى المرض ودرجة العجز والتكهن والاستجابة للعلاج تدبير مسلمات منان المالات ساندن من الالترام المناسف المسلمات المسانية المستروكيني المنتقل

وقد تم جمع ٨٠ حالة من النساء الاتى يعانين من مرض الالتهاب المفصلى الروماتويدى داخل الفئة العمرية (٣٥ اى ٤٠ سنة) من العيادات الخارجية لمستشفى الحسين الجامعى ومستشفى سيد جلال الجامعى والمركز القومى للبحوث وتم تشخيص الحالات بمعرفة اطباء الروماتيزم والطب الطبيعى والتاهيل.

وقد تم اختيار حالات البحث وفقا للأتي:

١ ـ جميع الحالات اناث في المرحلة العمرية من ٣٥-٤٠ سنة

٢- لايعانين من امراض مزمنة اخرى غير الالتهاب المفصلي الروماتويدي

۳۔ غیر حوامل

٤- مدة المعاناة من الالتهاب المفصلى الروماتويدى تتراوح مابين ٣ الى ٥ سنوات منذ ظهور الاعراض التشخيصية للمرض.

٥- جميع المرضى تحت العلاج.

وقد تم اخذ التاريخ المرضى (بعد موافقة جميع السيدات على الاشراك فى الدراسة) واجراء الفحص الاكلنيكى الكامل للحالات ثم اجراءتقييم لدرجة الاعاقة ودرجة العجز المرضى واخيرا اجراء الاختبارات المعملية الاتية: سرعة ترسيب، صورة دم كاملة، سكر عشوائى، وظائف كبد، وظائف كلى، معامل الروماتويد،

و قياس الاجسام المضادة للبروتين الستروليني بطريقة الاليزا.

واظهرت النتائج ان هناك علاقة احصائية ايجابية بين مستوى الاجسام المضادة للبروتين الستروليني ومستوى عامل الروماتويد في النتائج الايجابية والسلبية للمرضى و ايضا توجد علاقة احصائية ايجابية بين الاجسام المضادة للبروتين الستروليني وعامل الروماتويد مع المقياس المرئي المشابه.

ومن ناحية اخرى اظهرت النتائج وجود علاقة احصائيةايجابية بين مستوى الاجسام المضادة للبروتين الستروليني ومستوى عامل الروماتويد ودرجة الاعاقة وفقا لتقسيم الكلية الامريكية للامراض الروماتيزمية.

وقد خلصنا الى ان اختبار الاجسام المضادة للبروتين الستروليني هو اختبار تكهنى جيد ويمكن استخدامه لهذا الغرض مع مرضى الالتهاب المفصلي الروماتويدي وانه يتنبأ بمساروشدة المرض خاصة فيما يتعلق بدرجة الاعاقة.