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شبكة المعلومات الحامعية

# بسم الله الرحمن الرحيم



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شبكة العلومات الحامعية



شبكة المعلومات الجامعية التوثيق الالكتروني والميكروفيلم





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شبكة المعلومات الجامعية

# جامعة عين شمس

التوثيق الإلكتروني والميكروفيلم

### قسو

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سامية محمد مصطفى

شبكة المعلومات الحامعية



بالرسالة صفحات لم ترد بالأصل





## Clinical utility of anti-CCP assay in patients with rheumatoid arthritis

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Finally, I ask ALLAH to accept this work from me.

#### List of abbreviations

ACR American College of Rheumatology

AFA antifilaggrin autoantibodies

AKA Antikeratin antibody

Anti-CCP Anticyclic citrullinated peptide

APF Antiperinuclear factor

ATTRACT . Anti- tumor necrosis factor trial in rheumatoid arthritis with

concomitant therapy

BAL Bronchoalveolar lavage

C:MC Carpo:metacarpal ratio

CAM Complementary and alternative medicine

CD Cluster of differentiation

CMC Carpometacarpal

Cox Cyclooxygenase enzyme

CRP C- reative parotein

CTS Carpal tunnel syndrome

DAS28 The 28 joint disease activity score

DIP Distal interphalangeal joint

DM Dermatomyositis

DMARDs Disease-modifying antirheumatic drugs

EBV Epstein-Barr virus

ELISA Enzyme linked immunosorbent assay

ESR Erythrocyte sedimentation rate

Fc Fragment crystallizable

FM Fibromyalgia

HAO Health Assessment Questionnaire

HAQ-DI Health Assessment Questionnaire – disability index

HCQ Hydroxychloroquine

HCV Hepatitis C virus

7.

HEVs High endothelial venules

HLA Human leucocytic antigen

HRCT High resolution CT scan

Ig Immunoglobulin

IL Interleukin

ILD Interstitial lung disease

IP Interphalangeal

JSN Joint space narrowing

MCB Metacarpal base

MCP Metacarpophalangeal joint

MCTD Mixed connective tissue disease

MHC Major histocompatibility complex

mRNA messenger ribonucleic acid

MTP Metatarsophalangeal joint

MTX Methotrexate

NSAIDs Non-steroidal anti-inflammatory drugs

OA Osteoarthritis

PAD Peptidylarginine deiminase

PIP Proximal interphalangeal joint

PM Polymyositis

PMR Polymyalgia rheumatica

PsA Psoriatic arthritis

PTB Patellar tendon-bearing orthosis

RA Rheumatoid arthritis

RF Rheumatoid factor

ROM Range of motion

RT-PCR Reverse transcriptase-polymerase chain reaction

SENS Simple Erosion Narrowing Score

SES Short Erosion Scale

SLE Systemic lupus erythematosus

SSc Systemic sclerosis

SSZ Sulfasalazine

Tc<sup>99m</sup> –DTPA Technetium<sup>99m</sup> diethylene triamine pentacetate radionuclide

TNF Tumor necrosis factor

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## INTRODUCTION AND AIM OF THE WORK

#### Introduction

Rheumatoid arthritis is a systemic autoimmune disease of unknown origin. The diagnosis of RA remains empirical and imprecise, particularly early in the course of diease. Because early initiation of disease modifying treatments can significantly improve long-term outcomes for patients with RA, there is considerable motivation to accurately diagnose RA in patients with inflammatory arthritis early in the course of disease (O'Dell, 2002; Mottonen et al., 2002).

The presence of rheumatoid factor (RF) was identified in patients with RA over 50 years ago (Rose et al., 1949), assays for RF remain one of the American College of Rheumatology (ACR) classification criteria for RA (Arnett et al., 1988). The RF assay, in its current manifestation, remains suboptimal as a diagnostic test, as it lacks sensitivity (54-88%) and specificity (48-92%) (Weinblatt et al., 1980; Schellekens et al., 2000).

Anti-cyclic citrullinated peptide (anti-CCP) antibody belongs to the family of antifilaggrin autoantibodies (AFA), accompanied by the antikeratin anti body (AKA) and antiperinuclear factor (APF) (Sebbag et al., 1995). The epitopes recognized by AFA, AKA, and APF were found to be generated by a post-translational modification, namely deimination of the natural amino acid arginine to the amino acid citrulline by activity of peptidylarginine deiminase. Based on that knowledge, Schellekens et al. (1998) developed an enzyme linked immunosorbent assay (ELISA) using a cyclic citrulinated peptide (CCP) derived from the sequence of human filaggrin as substrate (van Venrooji et al., 2002; Zendman et al., 2004). The second version of the anti-CCP test (anti-CCP2) demonstrated high specificity (89-98%) and good sensitivity (41-88%) for RA (Herold et al., 2005).

Initial studies characterising the frequency of antibodies to CCP in mixed cohorts containing patients with rheumatic diseases, infectious diseases, and healthy subjects, have shown it to be moderately sensitive (68%) but highly specific (98%) for RA (Schellekens et al., 2000). Anti-CCP gives us a tool, which is important to identify patients with early RA (Gao et al., 2005), and helps to diagnose other inflammatory and non-inflammatory diseases (especially connective tissue diseases) by reducing the rate of false-positive results in comparison with RF (Sauerland et al., 2005). Anti-CCP is also significantly associated with some parameters of both disease activity and severity (Samanci et al., 2005).

#### Aim of the work:

This work aims to evaluate the clinical utility of the newer version of anti-CCP (anti-CCP2) in patients with RA relative to patients with a diversity of rheumatic diseases. We intend to study the relationship between anti-CCP and disease activity parameters, functional disability, and radiological destruction in rheumatoid arthritis patients, as well as to examine the sensitivity and specificity of anti-CCP in those patients.