

# بسم الله الرحمن الرحيم



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شبكة المعلومات الجامعية التوثيق الالكتروني والميكرونيلم





# جامعة عين شمس

التوثيق الإلكتروني والميكروفيلم

# قسم

نقسم بالله العظيم أن المادة التي تم توثيقها وتسجيلها علي هذه الأقراص المدمجة قد أعدت دون أية تغيرات



يجب أن

تحفظ هذه الأقراص المدمجة يعيدا عن الغيار





# Neutrophil to lymphocyte ratio and platelet to lymphocyte ratio in evaluation of inflammation and nutritional Status in pre dialysis chronic kidney disease patients

#### Thesis

Submitted for partial fulfillment of Master Degree in Internal Medicine

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CKD Chronic kidney disease
ESRD End stage renal disease
GFR Glomerular filtration rate
GIT Gastrointestinal tract

**CVD** Cardiovascular disease

**GERD** Gastroesophageal reflux disease

**DM** Diabetes mellitus

**ESAs** Erythropoiesis- stimulating agents

HTN Hypertension NK Natural killer

MIA Syndrome malnutrition – inflammation -

atherosclerosis

CRP C-reactive protein
IL1β Iterleukin-1 beta
IL-6 Interleukin-6
IL-1 Interlukin-1

**hs-CRP** High-sensitivity C-reactive protein

PTX3 Pentraxin

ROS Reactive oxygen species
PMN Polymorph nuclear cells
TLR-4 Toll-like receptors 4
TLR-2 Toll-like receptors-2

CLRs C-type lectin receptorsTNF-α Tumor necrosis alpha

**G-CSF** Growing factor

**DAMPs** Damage-associated molecular patterns

**PRRs** Pattern recognizing receptors

**RIGs** Retinoic acid-inducible gene

**NF-kB** Nuclear factor-kappa B

NLRs Intracellular Node-like receptors HIN-200 Hematopoietic interferon - 200

AP-1 Activator protein-1

RCC Renal cell carcinoma

MPs Membrane microparticles
TLC Total leukocyte count

**DLC** Differential leukocyte count

**PAMPs** Pathogen-associated molecular patterns

NLR Neutrophil – lymphocyte ratio
PLR Platelet – lymphocyte ratio

HD HemodialysisMHD Maintenance HD

**PEW** Protein energy wasting

**ISRNM** International society of renal nutrition and

metabolism

**RRT** Renal replacement therapy

**RAAS** Renin angiotensin aldosterone system

**IGF-1** Insulin -like growth factor- 1

**IGF-1BP** Insulin -like growth factor binding proteins

BMI Body mass index

**IHME** Institute for health metrics and evaluation

**ORG** Obesity-related glomerulopathy

%UBW %usual body weightTSF Triceps- skin fold

FFM Fat free mass

**FM** Fat Mass

MAC Mid arm circumference

MAMC Arm muscle circumference

ECW Extracellular water BCM Body cell mass

**ECM** Extra-cellular mass

**APPR** Acute phase protein response

**Hcy** Homocysteine

PD Peritoneal dialysis
TBN Total body nitrogen
TBK Total body potassium

DXA Dual x-ray absorptiometry
BIA Bio-impedance analysis

**SGA** Subjective global assessment

**m-SGA** Modified subjective global assessment

**DMS** Dialysis Malnutrition Score

MIS Malnutrition Inflammation Score

**CAPD** Continuous ambulatory peritoneal dialysis

**PG-SGA** Patient-Generate Subjective Global

Assessment

**PhilSPEN** Philippine Society of Parenteral and Enteral

Nutrition

RDN Registered dietitian nutritionist
 nPCR Normalized protein catabolic rate
 PNA Protein equivalent of total nitrogen

appearance

**QOL** Quality of life

MNT Medical Nutrition Therapy

**NEAP** Net Acid production

LC n-3 PUFA Long Chain Omega-3 Polyunsaturated Fatty

Acids

IDPN Total and Intradialytic Parenteral NutritionDASH Dietary approaches to stop hypertension

**DGA** Dietary guidelines for American

HDF HemodiafiltrationHDx Expanded HD

**HRO** High retention onset

MCO Medium cut-off

**MDRD** Modification of Diet in Renal Disease

**ACR** Albumin: creatinine ratio

**HGS** Hand grip strength

WHO World health organization

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# Introduction

Chronic kidney disease (CKD) is defined as decline in glomerular filtration rate (GFR) to less than 60 ml/min/1.73 m² with or without structural kidney damage for more than 3 months (*Levey et al.,2011*), it considered as a significant emerging worldwide health problem (*Levey et al., 2007*) that leading to an economic burden on health care system (*Freeman et al., 2018*), yet over the past 27 years CKD burden has not decline as a burden of other important noncommunicable diseases, in 2017 there is 1.2 million deaths as a result of CKD, by 2040 the number has been projected to rise to 2.2 million in a best-case scenario and up to 4 million in a worst-case scenario (*Cockwell & Fisher, 2020*).

In CKD patients the mortality mainly been attributed to cardiovascular disease (CVD) for many years, the evidence of non-cardiovascular mortality in CKD patient also increased during last years like a mortality from infection or malignancies, the relation between some traditional risk factors for CVD and non-cardiovascular mortality in uremic environment also described (*De jager et al.,2014*).

Inflammation and malnutrition considered as a components of CKD that can leading to poor outcome

(Maraj et al., 2018), Sustained low grade inflammatory status in CKD have been proved in many studies (Kutsal et al., 2016), (Ahbap et al, 2016), (Abd ElHafeez et al., 2018) and considered as an essential part of CKD since 1990 (Akchurin & Kaskel ,2015), with prevalence of 30%-60% for inflammation and 40% for malnutrition in uremic patients which described as independent risk factors for mortality in CKD (Kutsal et al., 2016).

Inflammation is considered as important link between increased risk of both cardiovascular and non-cardiovascular mortality in CKD patients ( *De jager et al.,2014*), and the role of inflammatory mediators is proven in many studies after detection of high levels of these mediators in patients with CKD (*Maraj et al., 2018*), (*Pecoits- Filho et al., 2002*).

The relation of inflammation and malnutrition to CVD is mentioned in previous studies known as malnutrition, inflammation, and atherosclerosis referred as MIA syndrome, considered as a silent factor for increased cardiovascular mortality rates in CKD patients, this increasing in cardiovascular mortality is not enough to be explained by the effect of traditional risk factors like diabetes mellitus (DM), hypertension (HTN), hyperlipidemia (*Turkmen et al., 2012*).

mediators play important Inflammatory an development of atherosclerotic heart disease and considered as a strong indicator for its progression, acute phase reactant like c-reactive protein (CRP) and proinflammatory cytokines like interleukin-6 (IL-6) & tumor necrosis factor alpha (TNFα) are will known conventional inflammatory mediator as will as a total leukocyte count (TLC) the classical inflammatory markers in many cardiovascular studies, the differential leukocyte count ( DLC ) is introduced in evaluation of inflammatory response that related to CVD, neutrophilia and relative lymphocytopenia are founded to be independent predictor of mortality in patient with heart failure (Okyay et al., 2013).

Neutrophil to lymphocyte ratio (NLR) introduced as inflammatory marker in many cardiac and non-cardiac diseases (*Sibarani et al.*, 2018), it is easily determined and cost effective predictor of mortality in patient with heart failure and myocardial infarction (*Okyay et al.*, 2013)

In some researches NLR is found to be associated with CKD and its progression *(Sibarani et al., 2018)* and reported to be closely related to inflammation in both hemodialysis (HD) and peritoneal dialysis (PD) with limited data regarding this association in pre-dialysis CKD patients.