## Study of the relationship between inflammation and depression in haemodialysis patients

#### **Thesis**

Submitted for Partial Fulfillment of Master Degree in Nephrology

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## Acknowledgment

First of all I cannot give a word to fulfill my deepest thanks to "Allah" the most gracious and the most merciful.

I would like to express my deep thanks, recognition, and everlasting gratitude to **Prof. Dr. Osama Mahmoud Mohamed,** Professor of Internal Medicine and Nephrology, Faculty of Medicine, Ain Shams University, for his kind support.

I would also like to thank to Dr. AbulRahman Nabil khedr A. Professor of Internal Medicine and Nephrology, Faculty of Medicine, Ain Shams University, for constant help and generous cooperation.

I am deeply thankful to **Dr. Nesreen Mohamed Mohsen**, A.Professor Neurology and psychatry ,Faculty
of Medicine – Ain Shams University for her great help,
outstanding support, active participation and guidance.

Also, I wish to express my gratitude and most sincere thanks to my mother and my wife.

Ahmed Mohamed Salem

2016

## **List of contents**

•	List of tables	II
•	List of figures	Ш
•	List of abbreviations	V
•	Introduction	1
•	Aim of the work	4
•	Review of literature:	
	> Chronic Kidney Disease	5
	> Depression	30
	> MIA Syndrome	.42
	> Relation between Depression and	
	Inflammation in CKD	60
•	Patients& methods	. 83
•	Results	86
•	Discussion	99
•	Summary	105
•	Conclusion	107
•	Recommendations	108
•	References	.109
•	Arabic summary	

## **List of Tables**

Number	Name	page
Table 1	Recommended equations for estimation of Glomerular Filtration Rate	2
Table 2	stages of CKD	7
Table 3	Criteria for CKD	8
Table 4	CKD nomenclature used by KDOQI	9
Table 5	Indication for renal replacement therapy	26
Table 6	Treatment options for depression in patients with ESRD	55
Table 7	causes for protein malnutrition in patients receiving maintenance hemodialysis	77
Table 8	The potential causes of inflammation in uremia:	66
Table 9	relation between demographic data of patients and presence of depressive symptoms	93
Table 10	relation between clinical data of patients and presence of depressive symptoms	95
Table 11	relation between laboratory investigations of patients and presence of depressive symptoms	97
Table 12	Regression analysis for factors affecting presence of depressive symptoms:	97
Table 13	results of Beck Depression Inventory	98

## List of figures

Number	Name	Page
Figure 1	results of Beck Depression Inventory	92
Figure 2	sex distribution of studied patients	93
Figure 3	relation between marital status and presence of depressive symptoms	94
Figure 4	causes of ESRD in our patients	94
Figure 5	relation between co-morbidity and presence of depressive symptoms (positive correlation)	96

## **Abbreviations**

CKD	Chronic kidney disease
ESRD	End stage renal disease
RRT	Renal replacement therapy
HD	hemodialysis
MIA	malnutrition inflammation atherosclerosis
	malnutrition inflammation-atherosclerosis-
MIAD	depression
TNF	tumor necrosis factor
IL1	interleukin 1
IL6	interleukin 6
BUN	Blood urea nitrogen
HS CRP	High sensitive c reactive protein
НВ	Hemoglobin level
GFR	Glomerular filtration rate
NKF	National Kidney Foundation
KDOQI	Kidney Disease Outcomes Quality Initiative
CRI	Chronic renal injury
P/Cr	Protein, creatinine ratio
HTN	Hypertension
MDRD	Modification of Diet in Renal Disease

### **List of Abbreviations**

KDIGO	Kidney Disease Improving Global Outcomes
AKI	acute kidney injury
ARVD	atherosclerotic Reno-vascular disease
ADPKD	autosomal dominant polycystic kidney disease
ACR	albumin-to-creatinine ratio
RAAS	renin-angiotensin-aldosterone system
PD	Peritoneal dialysis
CAPD	Continuous ambulatory peritoneal dialysis
CCPD	Continuous cycler-assisted peritoneal dialysis
USFTF	US Preventive Services Task Force
MDD	with major depressive disorder
BDI	Beck Depression Inventory
PSTF	PREVENTIVE SERVICES TASK FORCE
SLE	Systemic lupus erythematosis
GN	Glomerulonephritis
IHD	Ischemic heart disease
CVD	Cardiovascular disease
ECT	electroconvulsive therapy
SSRI	selective serotonin reuptake inhibitors
MAOI	monoamine oxidase inhibitors
(GH/IGF-1)	growth hormone/insulin like growth factor-1

### List of Abbreviations

VLDL	Very low-density lipoprotein
LDL	Low Density Lipoprotein

## Introduction

#### **INTRODUCTION**

Chronic kidney disease (CKD) is determined by the presence of kidney injury and by the level of renal function, assessment according to the glomerular filtration rate. Following the criteria proposed by the National Kidney Foundation (2002), the CKD is divided into five stages, classified according to the degree of the patient's renal function. Until the fourth stage of the disease, the so-called "conservative treatment" is recommended. In more advanced stages, called end-stage renal disease (ESRD), -i.e., when the kidneys can no longer maintain homeostasis of the body, the patient will depend on one of the modalities of renal replacement therapy (RRT): dialysis or kidney transplant (*Kimmel*, et al 2002)

The medical risk factors associated with increased mortality in hemodialysis (HD) patients are well known, but the psychosocial factors that may affect outcome have not been clearly defined. One key psychosocial factor, depression, has been considered a predictor of mortality. (*Kimmel PL*, *et al*,(2000),)

Depression is the most common psychiatric illness in patient with CKD affecting 20-30% patients on dialysis with 10 % having a major depressive disorders, however, there is a difficulty in recognizing its true extent in this population. This is due to methodological variations among studies (such as the diversity of instruments applied which does not allow a comparison of results) and the difficulty of diagnosis generated by similarity of somatic

#### Introduction

symptoms present in depression and uremic symptoms, which leads to an increased number of false-positive cases (*Kimmel, et al 2001*, 2002)

Depression issue in hemo-dialyzed patients has been addressed many times in the past nowadays, rapid and pronounce development in renal replacement therapy was observed which was followed by deep change in profile of hemodialyzed population (*Ricardo AC*, et al (2010))

Etiopathology of depression in patients with CKD is multifactorial, both medically and psychologically based. Additional factors also influence development of depression in ESRD patients. Chronic pain and sleep disturbances affect health-related quality of life (HRQOL) and depression in patients with all stages of CKD (*Odden ,et al ,2006*) and with ESRD with regard to not clear causality and order. Additionally, numerous authors underline role of patient's inflammatory status in depression development (*Ibrahim S, et al ,2008*).

Since there is little data about prevalence of depression in hemodialyzed population we designed a study haemodialyzed patients to address this issue and we will we will try to identify potential risk factors for depression development in this population with regard to inflammatory status. The question arises: is depression a new player in MIA syndrome? Confirming this hypothesis may result in changing MIA (malnutrition inflammation atherosclerosis)

to MIAD syndrome abbreviation (malnutrition inflammation—atherosclerosis—depression) in the future.

Many prior observations support the likelihood that pro inflammatory cytokines, interleukin 1 (IL-1), interleukin 6 (IL-6), and tumor necrosis factor (TNF) in particular, are involved in the pathogenesis of hemodialysis-related disease (Capuron, et al.,2011)

These cytokines were found to have a significant influence on neurotransmitter metabolism and neuroendocrine function and thus have been associated with the vegetative symptoms of depression (Miller DB, et al 2005)

There is a growing body of evidence that supports a relationship between cytokines and emotional symptoms However, the results of prior studies have not been consistent, and it remains unclear whether these results might be relevant to patients receiving hemodialysis. Moreover, malnutrition ,commonly observed in dialysis patients, has been associated with chronic inflammation(*Pertosa G,et al 2000*)

Malnutrition has also been associated with emotional symptoms in patients receiving hemodialysis, The complexity of the relationships among inflammation, nutritional status, and emotions in patients receiving hemodialysis, remains poorly understood. (*Czira ME*, *et al*, *2011*)

## Aim of the Work

### AIM OF THE WORK

The aim of this thesis is to study the prevalence of depressive symptoms in hemodialyzed patients and to identify the possible role of inflammation in depression development.

# Review of Literature