

شبكة المعلومات الحامعية

# بسم الله الرحمن الرحيم



-Caro-



شبكة المعلومات الحامعية



شبكة المعلومات الجامعية التوثيق الالكتروني والميكروفيلم





ببكة المعلم مات المامعية

## hossam maghraby

## جامعة عين شمس

التوثيق الإلكتروني والميكروفيلم

## قسو

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



يجب أن

تحفظ هذه الأقراص المدمجة يعبدا عن الغيار





شبكة المعلومات الجامعية



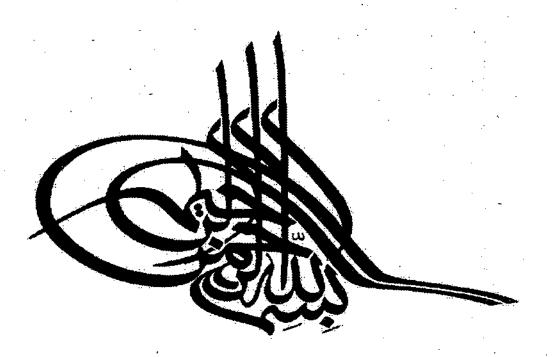


شبكة المعلومات الحامعية



بالرسالة صفحات لم ترد بالأصل





STUDY OF HEAT SHOK PROTEIN 32 (HSP – 32) IN POST RENAL TRANSPLANT PATIENTS: CORRELATION TO RENAL HAEMODYNAMICS, GRAFT FUNCTION AND SURVIVAL.

**Thesis** 

B17719

Submitted to the Faculty of Medicine
University of Alexandria
In partial fulfillment of the requirements for
Master Degree of Internal Medicine

By

Mohamed Mohamed Mohamed Sakr

(MBBCh, Alex)

Faculty of Medicine
University of Alexandria
2005

#### **SUPERVISORS**

## Prof. Dr. Hayam Abdel meguid El Aggan

Professor of Internal Medicine and Nephrology

Faculty of Medicine

University of Alexandria

¥

Gle To

#### Prof. Dr. Sherif El – Sayed Hegab

Professor of Diagnostic Radiology

Faculty of Medicine

University of Alexandria

Prof. Dr. Mohamed Moustafa Mohamed Rizk

Professor of Clinical Pathology

Faculty of Medicine

University of Alexandria

Dr. Hesham Kamal El Sayegh

Lecturer of Medicine and Nephrology

Faculty of Medicine

University of Alexandria

To my parents Who gave me a lot of support

## 

	Chapter	Page
	Introduction	1-68
H.	Aim of the work	69.
III.	Subjects	70
IV.	Methods	71-81
$\mathbb{V}$ .	Results	82-130
VI.	Discussion	131-146
VII.	Summary	147-150
VIII.	Conclusion	151-152
IX.	Recommendations	153
Х.	References	154-189
XI.	Protocol	
XII	Arabic summary	1-4

#### List of Abbreviation

AAs Amino Acids.

APC Antigen Presenting Cells.

ARE Antioxidant Responsive Element.

ATN Acute Tubular Necrosis.

CAN Chronic Allograft Nephropathy.

cGMP cyclic Guanosine Monophosphate.

CMV Cytomegalovirus.

CO Carbon monoxide.

COHb Carboxyhemoglobin.

CSA Cyclosporine A.

EMIT Enzyme multiplied Immunoassay Technique.

ESRD End Stage Renal Disease.

Fe<sup>++</sup>. Free Iron.

FPIA Fluorescence Polarization Immunoassay.

GRE - Glucocorticoid Response Elements.

**HLA** Human Leukocyte Antigen.

HO Heme Oxygenase.

HPLC High Performance Liquid Chromatography.

HSB-1 Heat Shock Binding Protein-1.

HSF Heat Shock Factors.

**HSP** Heat Shock Protein.

H<sub>2</sub>O<sub>2</sub> Hydrogen Peroxide.

ICAM Intercellular Adhesion Molecule.

IFN-γ Interferon Gamma.

IL Interleukin.

iNOS inducible Nitric Oxide Synthase.

IR/I Ischemia Reperfusion Injury.

MAbs Monoclonal Antibodies.

MARE Maf Recognition Element protein.

MHC Major Histocompatibility Complex.

MMF Mycophynolate Mofetil

NF-AT Nuclear Factor of Activated T-cell.

NO Nitric Oxide.

NOS Nitric Oxide Synthase.

PCR Polymerase Chain Reaction.

PG-E2 Prostaglandin E2.

PI Pulsatility Index.

PKC Protein Kinase-C.

RI Resistive Index.

ROS Reactive Oxygen Species.

RRT Renal Replacement Therapy.

SMC Smooth Muscle Cell.

SRL Sirolmus.

TCR T-Cell Receptor.

TGF-B Transforming Growth Factor Beta.

TNF-α Tumor Necrosis Factor Alpha.

TOR Target Of Rapamycin.



#### INTRODUCTION

#### RENAL TRANSPLANTATION

Chronic renal failure is a progressive persistent decline in renal function that the kidney can no more maintain the consistence of the internal medium this accompanied by the retention of end products of proteins (eg: urèa, creatinine, uric acid), disturbance of electrolytes and acid base balance, anemia and hypertension. These patients need renal replacement therapy in the form of hemodialysis, peritoneal dialysis or renal transplantation. (1)

Despite tremendous increase in knowledge and skill in the management of end stage renal disease (ESRD) patients, such individuals, particularly those treated by dialysis, remain unwell. Impaired quality of life, dependence on others, poor rehabilitation, and depressed sexual function all contribute to the physical and emotional disabilities that may persist even in well dialysed ESRD patients. (2)

Renal replacement therapy (RRT) describe various substitution treatment available for severe acute renal failure and ESRD patients, it includes dialysis (hemodialysis and peritoneal dialysis) and renal transplantation. (3)

The improvement in immunosuppression along with better treatment and prophylaxis for infectious complications, have made, renal transplantation became the preferred mode of RRT for all patients with ESRD, unless they have systemic malignancy, chronic infection, severe cardiovascular diseases or neuropsychiatric disorders. (3)

The advantages of renal transplantation are clearly established as it prolongs the survival of the recipients, improve quality of life, so, they can enjoy unrestricted activities and can return to their previous full term employment. (4)

Efforts were mainly directed toward reducing fatality and morbidity after renal transplantation, patients mortality tend to be highest during the first year after operation, infection was the cause of death of 50 % of patients. The second major cause of death during the first year post transplant is cardiovascular complications especially in elderly and diabetic patients. Also in spite of increasing success of renal transplantation, rejection of the transplanted kidney by the recipient still remains the major problem to be overcome. (5)

#### Transplantation Immunobiology

#### Major Histocompatibility Complexes:

Genes encode histocompatibility antigens, (cell surface proteins), are referred to as histocompatibility genes. There are more than 30 of such genes. The histocompatibility antigens are responsible for the graft's being recognized as similar to one's own tissues or as foreign. They are classified according to their relative potencies as either major or minor. The group of histocompatibility genes that have a central role in antigen recognition and transplantation immunobiology, this group of genes had been defined as the major histocompatibility complex (MHC). (6)

The MHC is a complex of genes found in humans and they are located on chromosome number 6. They encode polymorphic cell surface molecules, alloantigens known as human leukocyte antigen (HLA). The term of HLA is used as a synonym for the human MHC proteins. The HLA molecules are involved in the presentation of antigen to T-Cells and are