



شبكة المعلومات الجامعية

Ain Shams University

Information Network

بِسْمِ اللَّهِ الرَّحْمَنِ الرَّحِيمِ

جامعة عين شمس

شبكة المعلومات الجامعية

@ ASUNET

جامعة عين شمس

التوثيق الالكتروني والميكرو فيلم

قسم

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



يجب أن

تحفظ هذه الأفلام بعيدا عن الغبار

في درجة حرارة من ١٥-٢٥ مئوية ورطوبة نسبية من ٢٠-٤٠%

To be Kept away from Dust in Dry Cool place of
15-25- c and relative humidity 20-40%



شبكة المعلومات الجامعية التوثيق الالكتروني والميكرو فيلم

بعض الوثائق الأصلية تالفة

بالرسالة صفحات لم ترد بالاصل

**EVALUATION OF EARLY MEASUREMENT OF
GRANULOCYTE COLONY –STIMULATING FACTOR
AND INTERLEUKIN, AND ITS IMPACT ON THE
SEVERITY, PROGNOSIS AND OUTCOME OF
SYSTEMIC INFLAMMATORY RESPONSE
SYNDROME .**

616,028

THESIS

SUBMITTED TO

FACULTY OF MEDICINE

ALEXANDRIA UNIVERSITY

FOR PARTIAL FULFILLMENT OF THE REQUIREMENTS OF

DOCTORAL DEGREE IN CRITICAL CARE MEDICINE

BY

DR. GALAL HASSAN SHALABY ABDEL NABY

MBBCH. M S.

ASSISTANT LECTURER

CRITICAL CARE MEDICINE DEPARTMENT

FACULTY OF MEDICINE

ALEXANDRIA UNIVERSITY.

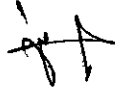
CKA Q
CP

2004

The Supervisors

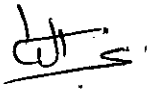
Prof Dr: Assem Abdel Razek Abd Rabbih

*Professor of Anaesthesia and Intensive Care
Faculty of Medicine
University of Alexandria*



Prof Dr: Nagwa Mahmoud Elkobbia

*Professor of Anaesthesia and Intensive Care
Faculty of Medicine
University of Alexandria*



Prof Dr: Salah Ahmed Marzouk

*Professor of Clinical Pathology
Faculty of Medicine
University of Alexandria*



CO-WORKER

Dr: Emad Eldin Abdel Monem Aly

*Assist. Professor of Anaesthesia and Intensive Care
Faculty of Medicine
University of Alexandria*



I would like to dedicate this work to

my beloved Wife, my Mother

the soul of my Father,

& my kids Ahmed

& Adham.

Acknowledgment

I would like to express my deepest gratitude and appreciation to *Professor Dr. Assem Abdel Razek* for his helpful comments, sincere guidance, and utmost support and I want to thank him for his effort, time and patience.

I am thankful to *Professor Dr. Nagwa Elkobia* for her thoughtful remarks and consideration.

Special thanks to *Professor Dr. Salah Marzouk* for his great help in doing the research work.

I would like to express my gratitude to *Assistant Professor Dr. Emad Abdel Moneim* for his great care, overwhelming kindness, and relevant direction.

Special thanks to all my colleagues for their active participation and help, and to the nursing staff for their valuable contribution and patients' care.

I am indebted to all patients with a great insight and a lot of knowledge.

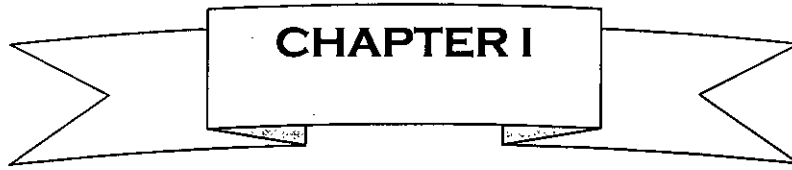
CONTENTS

CHAPTER	Page
I INTRODUCTION	1-66
II AIM OF THE WORK	67
III PATIENTS	68
IV METHODS	69-80
V RESULTS	81-131
VI DISCUSSION	132-157
VII SUMMARY	158-163
VIII CONCLUSIONS & RECOMMENDATIONS	164-165
IX REFERENCES	166-200
APPENDIX	
PROTOCOL	
ARABIC SUMMARY	

List of abbreviation

ACCP	American College of Chest Physician
APACHE	Acute physiology and chronic health evaluation
ARDS	Adult respiratory distress syndrome
AT III	Antithromin III
ATP	Adenosine Tri-Phosphate
BUN	Blood urea nitrogen
cfu/mL	Colony forming unit / mille liter
CO₂	Carbon dioxide
DO₂	Oxygen delivery
ELISA	Enzyme Linked Immunosorbant Assay
F_IO₂	Fraction of inspired oxygen
G -ve	Gram negative strain
G +ve	Gram positive strain
G-CSF	Granulocyte-colony stimulating factor
GM-CSF	Granulocyte-Macrophage-colony stimulating factor
GFR	Glomerular filtration rate
GIT	Gastro-Intestinal Tract
ICU	Intensive Care Unit
IFN	Interferon
Ig	Immunoglobulin
IL	Interleukin
KDa	Kilo Dalton
LODS	Logistic organ dysfunction score
LPs	Lipopoly- Saccharide
MOD	Multiple Organ Dysfunction Syndrome
MODS	Multiple Organ Dysfunction Score

MPM₀	Mortality Prediction Model on Admission
MPM₂₄	Mortality Prediction Model after 24 hours
NO	Nitric Oxide
PAF	Platelet Activating Factor
P_aO₂	Arterial Partial Pressure of oxygen
PCT	Procalcitonin
PDGF	Platelet Derived Growth Factor
pHi	Intra-mucosal pH
PMNLs	Polymorph Nuclear Leucocytes
P_sCO₂	Sublingual Capnometry
PUFA	Poly-unsaturated Fatty Acid
rhAPC	Recombinant human Activated Protein C
REE	Resting Energy Expenditure
SAPS	Simplified Acute Physiology Score
SIRS	Systemic Inflammatory Response Syndrome
SOFA	Sequential Organ Failure Assessment
TGF	Transforming Growth Factor
TEE	Total Energy Expenditure
TNF	Tumor Necrosis Factor
VO₂	Oxygen Consumption
WBCs	White Blood Cells



INTRODUCTION

SYSTEMIC INFLAMMATORY RESPONSE SYNDROME

In 1991, experts from a variety of disciplines met for a consensus conference sponsored by the American College of Chest Physicians and the Society of Critical Care Medicine and proposed new definitions for sepsis and the adverse sequelae of sepsis. ⁽¹⁾ The attendees recognized that sepsis was the systemic inflammatory response to a documented infection, but acknowledged that this response also may be observed in a number of other clinical conditions. ⁽²⁻⁵⁾ It was stated that sepsis was a continuum of injury response, ranging from sepsis to septic shock, to multi-system organ dysfunction syndrome (MODS). As patient progresses from sepsis to MODS, the associated mortality rate is expected to increase. ⁽²⁾

Definition of SIRS:

The American College of Chest Physicians/Society of Critical Care Medicine consensus panel defined the systemic inflammatory response syndrome (SIRS), sepsis, severe sepsis, and septic shock ⁽¹⁻⁵⁾ :

Infection: Infection is a microbial phenomenon characterized by an inflammatory response to the presence of microorganisms or the invasion of normally sterile host tissue by those organisms.

Systemic inflammatory response syndrome (SIRS): is a widespread inflammatory response to a variety of severe clinical insults. This syndrome is clinically recognized by the presence of two or more of the following:

- Temperature $>38^{\circ}\text{C}$ or $<36^{\circ}\text{C}$,
- Heart rate >90 beats/min ,
- Respiratory rate >20 breaths/min or $\text{PaCO}_2 <32$ mmHg,
- WBC $>12,000$ cells/mm³, <4000 cells/mm³, or presence of more than 10% immature (band) forms.

Sepsis: Sepsis is the systemic response to infection. Thus, in sepsis, the clinical signs describing SIRS are present together with definitive evidence of infection.

Severe sepsis: Sepsis is considered severe when it is associated with manifestation of organ dysfunction, hypoperfusion, or hypotension. These manifestations may include, lactic acidosis (lactate concentration of >2 mmol/L), oliguria (urine output of <0.5 ml/kg body weight for 2 hours), thrombocytopenia (platelet count of $<100,000$ /mm³), hypoxemia (P_{aO_2}/F_{iO_2} of <250) or an acute alteration in mental status without sedation.

Septic shock: Septic shock is sepsis with systolic hypotension (<90 mmHg) despite adequate fluid resuscitation combined with perfusion abnormalities that may include, but are not limited to, lactic acidosis, oliguria, or an acute alteration in mental status. Patients who are on inotropic or vasopressor agents may not be hypotensive at the time that perfusion abnormalities are measured.

Multiple organ failure: Multiple organ failure refers to the presence of altered organ function in an acutely ill patient such that homeostasis cannot be maintained without intervention. The multiple organ dysfunction syndrome is classified as either primary or secondary.

- Primary MODS is the result of a well-defined insult in which organ dysfunction occurs early and can be directly attributable to the insult itself (e.g., renal failure due to rhabdomyolysis).
- Secondary MODS is organ failure not in direct response to the insult itself, but as a consequence of a host response. In the context of the definitions of sepsis and SIRS, MODS represents the more severe end of the spectrum of severity of illness characterized by SIRS/sepsis.