

THE UNIVERSITY OF CHICAGO PRESS

D

CELL ADHESION IN THE HEMOPOIETIC SYSTEM

Essay

*Submitted in Partial Fulfillment for the
Master Degree
in
Clinical & Chemical Pathology*

By

Iman Mahmoud Al-Zoghby

616.0756

E

M

Under the Supervision of

5345

Prof. Dr. Zeinab Mohamed Tawfik

Professor of Clinical & Chemical Pathology
Faculty of Medicine - Ain Shams University

Handwritten signature

Prof. Dr. Hebat-Alla Adel Sedky

Professor of Clinical & Chemical Pathology
Faculty of Medicine - Ain Shams University

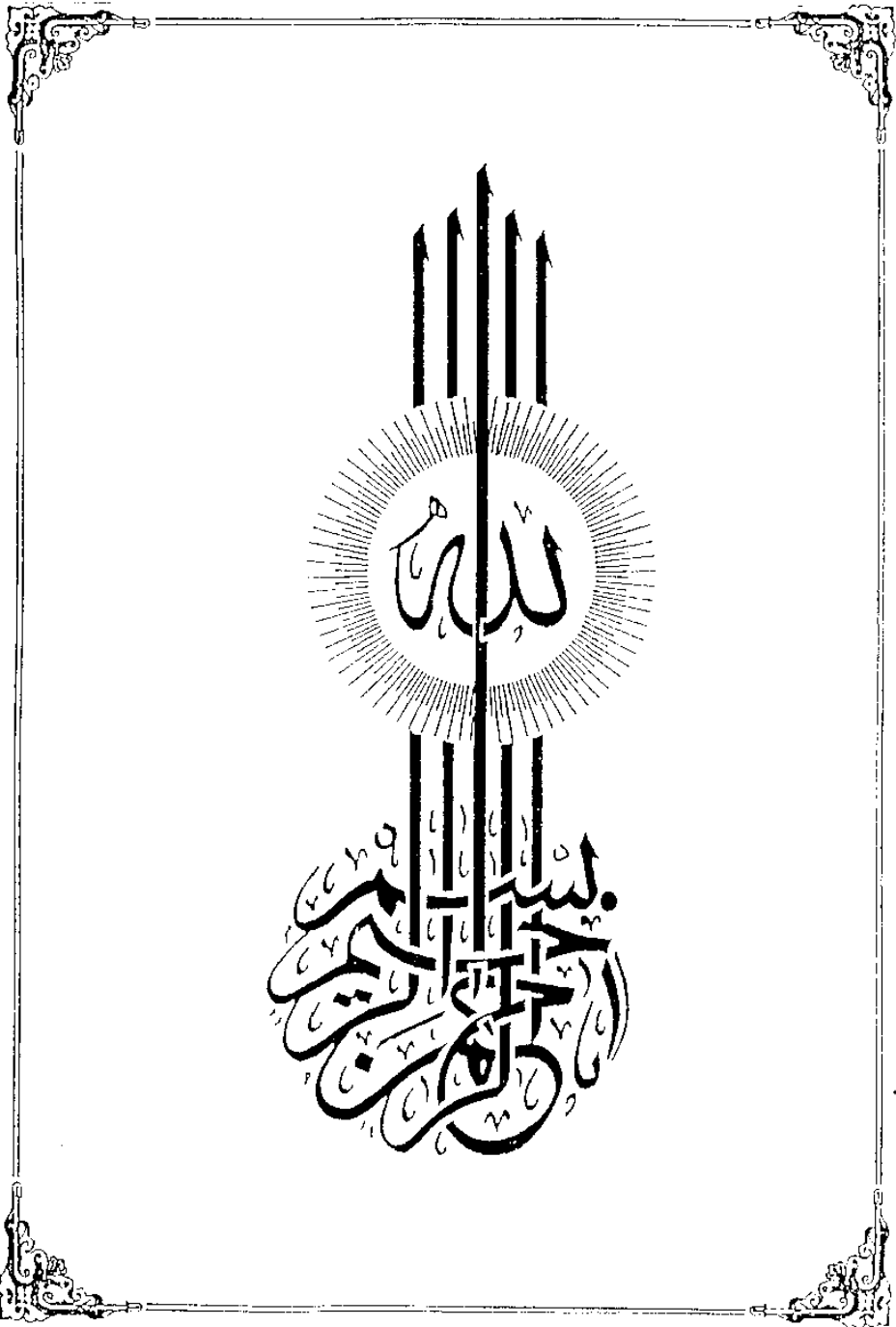
Handwritten signature

Dr. Manal Hashem Fayek

Lecturer of Clinical & Chemical Pathology
Faculty of Medicine - Ain Shams University

**Faculty of Medicine
Ain Shams University
1996**





بِسْمِ اللَّهِ الرَّحْمَنِ الرَّحِيمِ

«قالوا سبحانك لا علم لنا
إلا ما علمتنا انك انت
العليم الحكيم»

صدق الله العظيم

سورة البقرة - آية ٢٢



To My Father

ACKNOWLEDGEMENT

First of all thanks, to God.

I wish to express my thanks and gratitude to Prof. Dr. ZEINAB MOHAMED TAWFIK, Professor of Clinical Pathology, Ain Shams University, for her constructive criticism, valuable comments and kind advice.

My deepest thanks and special regards are due to Professor Dr. HEBAT-ALLA ADEL SEDKY, Professor of Clinical Pathology, Ain Shams University, for her generous help and continuous encouragement.

I am profoundly grateful to Dr. MANAL HASHEM FAYEK, Lecturer of Clinical Pathology, Ain Shams University, for her devotion, all the time and effort she sacrificed and the valuable friendship she offered me to start and complete this work.

Last no words could even express my deepest gratitude to my ever giving family, my husband and my son. Their favour is unforgettable.

LIST OF ABBREVIATIONS

AC:	Accessory cell.
ALL:	Acute lymphocytic leukemia.
AML:	Acute myeloid leukemia.
BFUe:	Burst forming unit erythroid.
BM:	Bone marrow.
BSS:	Bernard soulier syndrome.
CAMP:	Cyclic adenosine monophosphate.
CD:	Cluster of differentiation.
CFC:	Colony forming cells.
CFU-GM:	Colony forming unit granulocyte, macrophage.
CML:	Chronic myeloid leukemia.
CNS:	Central nervous system.
CSF:	Cerebrospinal fluid.
DIC:	Disseminated intravascular coagulopathy.
ECM:	Extracellular matrix.
EGF:	Epithelial growth factor.
ELAM:	Endothelial leucocyte adhesion molecule.
FAB:	French-American British.
Fab:	Fragment antigen binding.
FCM:	Flow cytometry.
Fg:	Fibrinogen.
FIB:	Fibrinogen.
Fn:	Fibronectin.
GMP-140:	Granule membrane protein 140.
GP:	Glycoprotein.
GT:	Glanzmann's thrombasthenia.
Hb:	Hemoglobin.
HC:	Hairy cell.
HCL:	Hairy cell leukemia.
HEV:	High endothelial venule.
ICAM:	Intracellular adhesion molecule.
IFN:	Interferon.
Ig:	Immunoglobulin.
IL:	Interleukin.
KD:	Kilo dalton.
LAD:	Leucocyte adhesion deficiency.
LAM:	Leucocyte adhesion molecule.

LECAM:	Leucocyte endothelial cell adhesion molecule.
LFA:	Lymphocyte function associated antigen.
LHR:	Lymphocyte homing receptor.
LN:	Laminin.
LPAM:	Lymphocyte Peyer's patch adhesion molecule.
LPS:	Lipopolysaccharide.
LTBMC:	Long term bone marrow culture.
mAb:	Monoclonal antibody.
MAdcAM-1:	Mucosal adressin cell adhesion molecule-1.
MHC:	Major histocompatibility complex.
NK:	Natural killer.
NCAM:	Neural cell adhesion molecule.
P150/95:	Protein 150/95.
PADGEM:	Platelet activation dependent granule co external membrane.
PECAM:	Platelet and endothelial cell adhesion molecule.
PMA:	Phytohaemagglutinin.
PPME:	A yeast polysaccharide of mannose and mannose-6-phosphate.
RGD:	Arginine-glycine-aspartate.
SCHO:	Stable chinese hamster ovary.
T-PBL:	T-peripheral blood lymphocyte.
TCR:	T-cell receptor.
TGF-B:	Transforming growth factor B.
TNF:	Tumour necrosis factor.
TSP:	Thrombospondin.
VCAM:	Vascular cell adhesion molecule.
VLA:	Very late activation molecule.
VnR:	Vitronectin receptor.
vWF:	von Willibrand factor.

TABLE OF CONTENTS

Page

- INTRODUCTION AND AIM OF THE WORK	1
- REVIEW OF LITERATURE:	
* Intracellular Adhesion Molecules:	
- Adhesion Pathways	3
- CD2-mediated adhesion pathway	4
- Selectin mediated adhesion pathway	12
- Integrins mediated adhesion pathway	21
- CD44 mediated adhesion pathway	33
- Vascular adressins	35
- Other adhesion receptors	36
- Regulation of adhesion receptors	39
- Signalling through adhesion receptors	42
* Cell Adhesion in Normal hemopoietic System:	
- Cell adhesion in hemopoiesis	43
- Cell adhesion and platelet function	51
- Cell adhesion and leucocyte function	63
* Cell Adhesion in Diseases:	
- Altered expression of adhesion molecules	68
- Diseases due to genetic deficiency of adhesion molecules:	
i- Bernard-Soulier Syndrome	68
ii- Glanzmann's thrombasthenia	71
iii- Leucocyte adhesion deficiency-1	72
- Diseases due to dysregulated expression of adhesion molecules:	
i- Thrombotic and inflammatory disorders ...	74
ii- Tumours and tumour metastasis	75
iii- Hematological malignancies	77
- Adhesion molecules and infections	85
* Key points For Clinical Practice	85
- SUMMARY	87
- REFERENCES	
- ARABIC SUMMARY.	



INTRODUCTION

INTRODUCTION

The term adhesion molecules refers to those cell surface structures that directly play a decisive mechanical role, the binding of a cell to its environment, either to the extracellular matrix or to another cell. This cellular adhesion plays a central role in many biological processes such as morphogenesis, cell migration, and direct cell-cell cooperation. In addition the phenomenon has important clinical consequences involving for instance, vascular thrombosis, the inflammatory process, tumour metastasis, bacterial and parasite infections (Edelman, 1985).

At present there are many clusters of differentiation (CD) molecules on the surface of cells of the haemopoietic system, defined by monoclonal antibodies (mAbs). Structurally, these molecules often traverse the cell membrane and are linked to the cytoskeleton, so that the cell can use them to gain traction to another cell or to the extracellular matrix as it moves (Roitt et al., 1993).

Moreover the events of cellular adhesion are closely linked to the events of cellular signalling. The inert coupling of adhesion to signalling is due to both cell stabilization by the adhesion molecules permitting other surface molecules to bind their ligands and transduction of powerful signals controlling cell activation/proliferation via adhesion molecules (Denning et al., 1987).

Cellular adhesion in man is mediated via multiple molecular pathways which involve specific intermolecular events. These pathways are made of multimolecular complexes, and each pathway may strongly influence the affinity of another (Roitt et al., 1993).

AIM OF THE WORK:

The aim of this work to prepare a review for the cellular adhesion pathways and their importance in hematological functions and diseases.