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بنيه النوالة مزالت ب

مَ على اللاز، « نلان اللان اللاز»

THE PROGNOSTIC VALUE OF MORPHOLOGICAL AND IMMUNOLOGICAL LYMPHOCYTE SUBTYPES IN CHRONIC LYMPHOCYTIC LEUKAEMIA

B. IYVAAI

Thesis

مُعْ مُعْمِعِلُهُ مِيكُودُ بِلْمِياً مُعِمَالًا لِلْمُعِمَالًا لِلْمُعِمَالًا لِلْمُعِمَالًا لِلْمُعِمَالًا للمُعْمِعَالًا للمُعْمِعِيلًا للمُعْمِعِيلًا للمُعْمِعِلًا للمُعْمِعِلِيلًا للمُعْمِعِلِيلًا للمُعْمِعِلًا للمُعْمِعِلِيلُوعِيلًا للمُعْمِعِلًا للمُعْمِعِلِيلًا للمُعْمِعِلِيلًا للمُعْمِعِلِيلًا للمُعْمِعِلِيلًا للمُعْمِعِلًا للمُعْمِعِلًا للمُعْمِعِلًا للمُعْمِعِلِيلًا للمُعْمِعِلِيلًا للمُعْمِعِلِيلُوعِلِيلًا للمُعْمِعِلِيلًا للمُعْمِعِلِيلًا للمُعْمِعِلًا لللمُعْمِعِلًا للمُعْمِعِلًا لللمُعْمِعِلِيلًا للمُعْمِعِلًا للمُعْمِعِلًا للمُعْمِعِلًا للمُعْمِعِلَّا للمُعْمِعِلَا للمُعْمِعِلْ للمُعْمِعِلْ للمُعْمِعِلِيلِيلِيلِيلًا للمُعْمِعِلًا للمُعْمِعِلًا للمُعْمِعِلًا للمُعْمِعِلًا للمُعْمِعِلًا للمُعْمِعِلًا للمُعْمِعِلًا للمُعْمِعِلَا للمُعْمِعِلَا للمُعْمِعِلَا للمُعْمِعِلِيلًا للمُعْمِعِلَا للمُعْمِعِلِيلِيلًا للمُعْمِعِلَا للمُعْمِعِلَا للمُعْمِعِلَا للمُعْمِعِلَا لللْمُعِمِعِلَّا لللْمُعِمِعِلًا للمُعْمِعِلًا للمُعْمِعِلِيلًا للمُعْمِعِلَا للمُعْمِعِلَا للمُعْمِعِلَا للمُعْمِعِلَا للمُعْمِعِلَّا للمُعْمِعِلَا للمُعْمِعِلَّا للمُعْمِعِلَّا للمُعْمِعِلْمِعِلَّا للمُعْمِعِلًا للمُعْمِعِلِيلُوعِ للمُعْمِعِلَّا لِمُعْمِعِلَّا لِمُعْمِعِلِيلِيلُوعِ لللْمُعِمِعِلِيلًا للمُعْمِعِلِيلِيلِيلِيلِعِلْمِعِلَّا لِعِلْمِعِلَّا لِمِعِلَّا لِمُعْمِعِلًا لم

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By

Wael Yassin M. El-Sadek

M.B.Ch.B.

Master Clinical Haematology Haematology Department Medical Research Institute University of Alexandria

SUPERVISORS

Prof. Dr. Laila El-Said Zyada

Professor and Head of Haematology Dept. M.R.T. Alexandria University

Prof. Dr. Mohamed Ragab Hussin

Professor of Kaematology
M.R.T.
Alexandria University

Prof. Dr. Hossam Kamel Mahmoud

Professor of Medical Oncology N.G.T Gairo University

Dr. Hanaa Abd El-Megid El-Agan

Assistant Drofessor of Kaematology M.R.I. Alexandria University

Dr. Homam Mohamed Sharshira

Lecturer of Haematology M.R.T. Alexandria University

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LIST OF ABBREVIATIONS

Ags Antigen

AIHA Autoimmunhaemolytic anaemia

ANAE α-naphthylacetate esterase

ATLL Adult T-cell leukaemia/lymphoma

 β_2 -M β_2 -microglobulin

BCGF B-cell growth factor

CLL Chronic lymphocytic leukaemia

CD Cluster of differentiation

CALLA Common acute lymphoblastic leukaemia antigen

CNS Central nervous system

CTcLs Cutaneous T-cell lymphoma

2-cdA 2-Chlorodeoxyadenosine

dcF Deoxycoformycin
EBV Epstein-Barr virus

FDC Follicular dendritic cell

GM-CSF Granulocyte, macrophages-colni stimulating factor

HCL Hairy cell leukaemia

HTLV Human T-lymphotrophic virus

INF-γ Interferon-γ

IWCLL International working group on CLL

LPL Lymphoplasmacytic lymphoma

L.Ns Lymph nodes

LGL Large granular lymphocytes

MHC Major histocompatibility complex

MoAbs Monoclonal antibodies

MR BC-R Mice red blood cell-rosette
Non-hodgkin's lymphoma

NHL Non-hodgkin's lymph NK Natural killer

PLL Prolymphocytic leukaemia

PRCA Pure red cell aplasia

PHA Phytohaemoagglutinum

SCID Severe combined immunodeficiency disease

sIg Surface immunoglobulin

SLVL Splenic lymphoma with villous lymphocytes

TCR T-cell receptor

TNF- α Tumour necrosis factor- α

WBC White blood cells

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INTRODUCTION

INTRODUCTION

LYMPHOCYTES

Lymphocytes are the milestone of immunity. They are derived from the multipotential haemopoietic precursor cells. Lymphocytes are engaged in two broad immunologic tasks:

- I. Humoral immunity; which involves synthesis and secretion of specific antibodies by B. lymphocytes. They are designated as B cells referring to their birthplace, which is Bursa of Fabricius in birds and the bone marrow in mammals.
- II. Cell mediated immunity which involves activation of T lymphocytes to express a variety of effectors functions. T cells are the subpopulation of lymphocytes that regulate and participate in cell mediated immune response. T cells can recognize and react to foreign or tumor antigens (Ags) on host cells and kill host cells infected by antigenic viruses. They are termed T cells because they are derived from the thymus.⁽¹⁾

The thymus and bone marrow comprise the primary lymphoid organs while the secondary lymphoid organs are the spleen, lymph nodes, and mucosa associated lymphoid tissue of the gut and pharynx.⁽²⁾

Natural killer cells (NK) are a subset of lymphocytes that arise from a precursor cell in the bone marrow. Studies in rodents indicate that precursors of NK cells originate in the bone marrow and after maturation,

migrate to the spleen via the blood stream.⁽³⁾ Most human NK functions are mediated by large granular lymphocytes (LGL).⁽⁴⁾ The terms LGL and NK cells are used interchangeably. The NK designation is a functional one. However, NK activity can be mediated by other cells, whereas LGL is a morphological term, therefore not all LGL are NK cells.⁽⁴⁾

Thus we have three distinct lineage of lymphocytes: B cells, T cells, and NK cells. They differ greatly from each other in terms of origin, life span, and functions. The most important methods used for the classification of each lymphocyte lineage are based on the identification of certain glycoproteins displayed on the membrane of lymphocytes and collectively referred to as markers. Monoclonal antibodies are used to detect these markers. These monoclonal typing reagents led to the discovery of previously unrecognized lymphoid subsets, each expressing a unique surface differentiation antigen. Many international workshops have led to exchange monoclonal antibodies and to test for cross reactivity. Monoclonal antibodies showing similar reactivity have been given a common [CD] designation [CD] cluster of differentiation]. (5)