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PATHOGENESIS OF ALOPECIA AREATA

THESIS

SUBMITTED IN PARTIAL FULFILLMENT FOR MASTER DEGREE IN DERMATOLOGY, VENEREOLOGY AND ANDROLOGY

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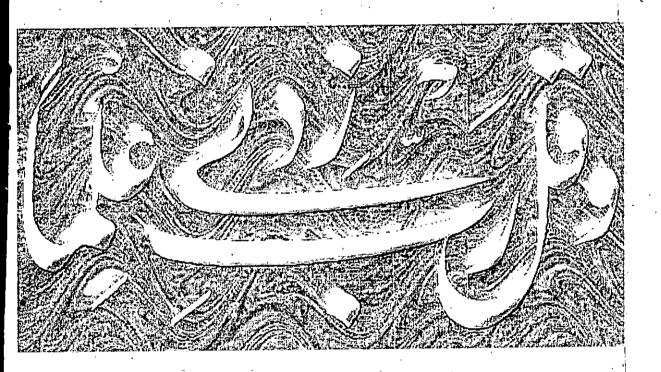
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صدق الله العظيم

DEDICATION

To My Dear And Great Family, My Father, My Mother, Hanan, Ahmed, Amr.

Without Their Encouragement, Inspiration And Patience . This Work Would Never Been Brought To Fruition.

To All Who Taught Me A Letter From First Day

Of School Up Till Now.

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LIST OF ABBREVIATIONS

AA : Alopecia areata

AMA : Antimitochondrial antibody
ANA : Antinuclear antibodies

ASM : Antismooth muscle antibodies

AT : Alopecia totalis

ATP : Adenosine triphosphate AU : Alopecia universalis

bFGF : basic fibroblast growth factor

BL: Basal lamina

CBCL : Child Behavior Checklist

CD16+ and CD56+ : Natural killer cells

CD19+ : B cells

CD1a+ and CD36+ : Dendritic cells (thymocytes and Langerhans

cells

CD3+ : Matrure T lymphocytes CD4+ : T helper lymphocytes

CD8+ : T suppressor lymphocytes

CDI : Childern's Depression Inventory
CGRP : Calcitonin gene-related peptide
CMAS : Childern Manifest Anxiety Scale

CMV : Cytomegalovirus

CRSD : Carroll Rating Scale for Depression

CTS : Connective tissue sheath

DC : Dark cells

DCP : Diphenylcyclopropenone

DP : Dormal papilla
DPCP : Diphencyprone

EGF : Epidermal growth factor

ELAM-1 : Endothelial leukocyte adhesion molecule-1.

EMH : Exclamation mark hairs

G-CSF : Granulocyte-colony stimulating factor

GM-CSF: Granulocyte macrophage-colony stimulating

factor

HF : Hair follicle

HIV : Human immuno-deficiency virus

HLA : Human leukocyte antigen

HSR : High stress reactor

ICAM-1 : Intercellular adhesion molecule-1

IFN-γ : Interferon gamma

IgA: Immunoglobulin AIgG: Immunoflobulin GIgM: Immunoflobulin M

IL-1 : Interleukin-1

IL-1α : Interleukin-1 alphaIL-1β : Interleukin-1 beta

IL-1ra : Interleukin-1 receptor antagonist

IL-2R : Interleukin-2 receptor

KD : Kilodalton

LAA : Localized alopecia areata LFA-1 : Lymphocyte surface integrin

LG : Langerhans granules
MAA : Multiple alopecia areata
MX : Epithelial hair matrix

NK : Natural killer ORS : Outer root sheath

PCA : Antiparietal cell antibody
PCR : Polymerase chain reaction

 PGE_2 : Prostaglandin E_2

PUVA : Psoralen photochemotherapy SADBE : Squaric acid dibutylester

SCID : Severe combined immunodeficiency

SDS-PAGE: Sodium dodecyl sulphate-8% polyacrylamide

gel electrophoresis.

SP : Substance P

 $TGF-\alpha$: Transforming growth factor-alpha

TMA : Thyroid microsome antibody
 TNF-α : Tumour necrosis factor-alpha
 TNF-β : Tumour necrosis factor-beta

VB-repertoire : Variable region of T cell receptor chain

Patrodicion

INTRODUCTION

Hair has no vital function in humans, yet its psychological functions are extremely important, as any clinical dermatologist or cosmetician can readily attest from routine daily practice. If the inevitability of scalp baldness makes it reluctantly tolerable to genetically disposed men, in women, loss of hair from the scalp is even more distressing than the growth of body or facial hair in excess of the culturally acceptable norm (**Dawber et al., 1998**).

There are several types of alopecia, i.e.: androgenetic alopecia (Olsen, 1994; Simpson, 1997); congenital alopecia and hypotrichosis including total or partial absence of hair of developmental origin (Sinclair and De Berker, 1997); total alopecia (Dawber, 1997); diffuse alopecia of endocrine origin (Comaish, 1985 and Holt and Marks, 1977); alopecia of chemical origin (Simpson, 1997); alopecia of nutritional and metabolic origin (Weismann, 1980; and Gummer, 1985); chronic diffuse alopecia (Fiedler and Hafeez, 1994), alopecia in central nervous system disorders (Mikula and Stiedl, 1961; and Tarnow, 1971), and alopecia areata (Gupta et al., 1990; Hordinsky, 1994; Skinner et al., 1995a; and Tosti et al., 1996b and Messenger and Simpson, 1997).

Alopecia areata (AA) is a relatively common, reversible hair follicle disorder (Bystryn and Tamesis, 1991 and Tobin et al., 1994a), characterized by the sudden appearance of sharply defined round or oval patches of hair loss. Although the condition occurs at all ages, the first attack usually appears in patients under 25 years of age (Schachner and Hansen, 1996). It is characterized by the sudden and premature involution of early anagen hair follicle (HF) at the site of disease activity

(Messenger et al., 1986). Hair shafts may be defective or totally lacking in affected HF. The location, extent, and time course of AA lesions are unpredictable, with lesions ranging from small patches to total loss of scalp hair (alopecia totalis) and loss of all body hair (alopecia universalis) (Tobin, 1997).

It is not at present possible to attribute all or indeed any case of AA to a single cause. Among the many factors that appear to be implicated in at least a proportion of cases are the patient's genetic constitution, the atopic state, non-specific immune and organ-specific autoimmune reactions, and possibly emotional stress (Messenger and Simpson, 1997).