THE INTEGRITY OF GLUTATHIONE SYSTEM IN CHILDREN WITH CHRONIC RENAL FAILURE

THESIS

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بسم الله الرحمن الرحيم

« قالوا سبحانك لا علم لنا إلا ما علمتنا إنك أنت العليم الحكيم .»

صدق الله العظيم

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To ...

My Family

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LIST OF ABBREVIATIONS

2,3-DPG 2,3-diphosphoglyceric acid.

Al Aluminum

ATP Adenosine triphosphate

BFU-E Burst forming unit-erythroid

BUN Blood urea nitrogen

CAPD Continuous ambulatory peritoneal dialysis

CFU-E Colony forming unis-erythroid

CRF Chronic renal failure
DNA Deoxyribonucleic acid

Epo Erythropoietin

ESRD End stage renal disease

F Female

GFR Glomerular filtration rate

GSH Reduced glutathione
GSSG Oxidized glutathione
H₂O₂ Hydrogen peroxides

Hb Hemoglobin
HD Hemodialysis
LOOH Hydroperoxides

M Male

Mn Manganese

NADpH Reduced nicotinamide adenine dinucleotide phosphate

Ni Nickel

OF Osteitis fibrosa
Plase A₂ Phospholipase A₂

pLOOH Phospholipid hydroperoxide

RBCs Red blood corpuscles
RIA Radioimmunoassay

Se-GSH-PX Selenium glutathione peroxidase

Si Silicon

SLE Systemic lupus erythematosus

Zn Zinc

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Introduction

INTRODUCTION

Anemia occurs with relatively high incidence in patients affected by chronic renal failure (CRF) (*Erslew*, 1970).

It has been described as normocytic, normochromic anemia. Erythropoietin deficiency seems to be the main cause of anemia in these patients (*Giardini et al.*, 1984). The red blood cells (RBCs) of uremic patients, although reduced in number, show a shortened life span (*Jacob et al.*, 1975). The biochemical alterations involved in this event have yet to be completely clarified.

Different mechanisms have been implicated in aging and lysis of RBCs; lipid peroxidation of the RBC membrane induced by free radicals could be the most important (*Hirshman et al.*, 1981). In vitro studies demonstrate that plasma from CRF patients contains various oxidants (*Costagliola et al.*, 1987).

RBCs have several lines of defence against oxidative stresses. Reduced glutathione (GSH), together with its related enzymes, is one of the major scavengers of activated oxygen radicals in RBCs (Aggarwal et al., 1984).

Red cells release oxidized glutathione (GSSG) when exposed to oxidative stresses, most probably as a consequence of an increased intracellular content of GSSG due, in turn, to increase glutathione peroxidase activity (*Costagliola et al.*, 1986).

Aim of the Work

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The aim of this study is to investigate the RBC and plasma levels of reduced and oxidized glutathione in patients with CRF in order to test indirectly the presence in plasma of oxidants which affect the integrity of the glutathione system and the possible role of hemodialysis (HD).