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# جامعة عين شمس

التوثيق الالكتروني والميكروفيلم



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Plasma Endothelin-1 Level in Patients with Schistosomal and Post Viral Liver Cirrhosis with Portal Hypertension: Its Relation to Portal Haemodynamics and Risk of Bleeding

#### THESIS

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The Soul of my Father

My Beloved Mother

My sincere husband

E to all my family



#### List of Abbreviations

ALT : Alanine Transaminase

AST : Aspartate Transaminase

AVH : Acute Viral Hepatitis

CI : Congestion index

CIPV : Congestion index of portal vein

ECE : Endothelin converting enzyme

EDCFs : Endothelium derived contractile factors

ELISA : Enzyme linked Immunosorbent Assay

Et-1 : Endothelin-1

EtA : Endothelin A receptor

EtB : Endothelin B receptor

GEV : Gastrooesophageal varices

HBV : Hepatitis B virus

HCV : Hepatitis C virus

HCC: Hepatocellular carcinoma

HSCs : Hepatic stellate cells

HSS : Hepatosplenic schistosomiasis

IGV : Isolated gastric varices

IL10 : Interleukin 10

IL2 : Interleukin 2

IHA : Indirect haemagglutination test

MVPV : Mean velocity of portal vein

NIEC: North Italian Endoscopic club

NO : Nitric Oxide

NOS : Nitric Oxide Synthase

OV : Oesophageal varices

PBV : Portal blood flow velocity

PCR : Polymerase chain reaction

PHG : Portal hypertensive gastropathy

PPF : Periportal fibrosis

Proth. activity: Prothrombin activity

PV : Portal vein

Red WM : Red wale marking

ROS : Reactive oxygen species

S. Bil : Serum bilirubin

S. alb : Serum albumin

S. alk ph. : Serum alkaline phosphatase

SHF : Schistosomal hepatic fibrosis

SMV : Superior Mesenteric Vein

SV : Splenic vein

TGF $\beta$  : Transforming growth factor  $\beta$ 

TFA : Trifluroacetic acid

Th1 : T helper 1

Th2 : T helper 2

US : Ultrasound

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# INTRODUCTION

### Chapter (I)

## HEPATIC SCHISTOSOMIASIS

Schistosomiasis is a chronic debilitating disease caused by helminthic infection. Three closely related species of trematodes, belonging to the family schistosomatidae *Schistosoma mansoni*, schistosoma haematobium and schistosoma japonicum mainly affect humans. Other species which are essentially animal parasites but may occasionally infect man include: *S.bovis*, *S.matteei*, *S.intercalatum* and *S.mekongi*. The two species present in Egypt are *S.haematobium* which is highly endemic in rural areas and *S.mansoni* which is almost limited to the Nile Delta and rarely found in south of Cairo. (2)

#### Incidence

Hepatic schistosomiasis is propably the world's most prevalent chronic liver disease. (3) In Egypt, it cause 70% of liver cirrhosis. Half of these cases are pure schistosomal, while the other half is complicated mostly by nutritional defects or sequelae of viral hepatitis. (4)

Hashem,<sup>(5)</sup> by study of autopsy material found that hepatic schistosomiasis occurs in 15% of cases infected with schistosoma haematobium; in 72% of *Schistosoma mansoni* and in 36% of mixed infection. Elwi and Attai,<sup>(6)</sup> reported an incidence of 10,37 and 32.0% respectively.

#### Pathogenesis and pathology

The pathogenesis of liver affection in schistosomiasis is produced by prolonged and repeated embolization of the liver by ova that are shifted back from the terminal radicals of the mesenteric and haemorrhoidal veins where they fail to emerge through their walls. (6) Deposition of ova may occur either in the large portal radicals, producing the coarse type of SHF (Symmer's clay pipe stem) or in the small portal tracts producing the fine diffuse type. (7) Eggs deposited which are the important initiating factor of hepatic schistosomiasis are surrounded by intense eosinophilic and histiocytic cellular reaction forming granulomata in and around the adjacent vessels with swelling of its endothelium and sometimes cellular reaction of their walls. The reaction may be so intense as to block the vessel. This reaction is gradually replaced by fibroblasts associated with new capillary formation. The liver parenchyma between the fibrotic areas is typically well preserved correlating with the maintenance of nearly normal hepatic function which is characteristic of hepatic schistosomiasis. (8,9)

Diffusible toxin of dead or living worms, the presence of dead worms and dietary deficiency are other factors which may contribute to the pathogenesis.<sup>(10)</sup>

Hepatic schistosomiasis has the following special pathological pattern which differentiate it from other types of cirrhosis: (11)

1- Diffuse cellular granulomatous eosinophilic infiltration ending in coarse or fine diffuse fibrosis of the portal tracts.