Ain Shams University Faculty of Medicine Internal Medicine Department



Significance of Leukocyte Esterase Reagent Strips in Spontaneous Bacterial Peritonitis (SBP) and associated Pleural Effusion in Cirrhotic Patient

Thesis Submitted to the Faculty of Medicine, Ain Shams University In Partial Fulfillment for the Requirements of the Master Degree In '' Internal Medicine ''

Ayman Mohamed Sallam M.B.,B.CH

Supervisors Prof. Dr. Sayed Abd-Elnaby Shalaby

Professor of Internal Medicine Faculty of Medicine Ain Shams University

Dr. Amany Talaat Kamal

Assistant Professor of Internal Medicine Faculty of Medicine Ain Shams University

Dr. Moataz Mohamed Sayed

Lecturer of internal Medicine Faculty of Medicine Ain Shams University

Faculty of Medicine Ain Shams University 2010

أهمية كاشف اليكوصيت استريز في حالات الالتهاب البكتيري التلقائي في السائل البيريتوني المصحوب بارتشاح بللورى لمرضى تليف الكبد

رسالة مقدمة توطئة للحصول على درجة الماجستير في أمراض الباطنة العامة

مقدمة من الطبيب أيمن محمد سلام بكالوربوس الطب والجراحة

تحت إشراف أد/ سيد عبد النبي شلبي أدر سيد أستاذ أمراض الباطنة

كلية الطب - جامعة عين شمس

أد/ أمانى طلعت كمال

أستاذ مساعد أمراض الباطنة

كلية الطب . جامعة عين شمس

د/ معتز محمد سید

مدرس أمراض الباطنة

كلية الطب . جامعة عين شمس

كلية الطب

جامعة عين شمس

Acknowledgement

First of all and above all, great thanks to **ALLAH**, whose blessings on me can not be counted.

The sincerest thanks, deepest appreciation and greatest admiration to **Prof. Dr. Sayed Abd-Elnaby Shalaby**, Professor of Internal Medicine, Faculty of Medicine, Ain Shams University, for his constructive supervision, and encouragement, He continuously advised me and spared no time or effort to offer his help, I owe special feelings of gratitude and thanks to him.

I would like to acknowledge the help of Ass. Prof.Dr. Amany Talaat Kamal, Assistant Professor of Internal Medicine, Faculty of Medicine, Ain Shams University and thank her for continuous expert guidance and continuous supervision throughout this work.

I'm specially grateful and specially indebted to **Dr. Moataz Mohamed Sayed**, Lecturer Professor of Internal Medicine, Faculty of Medicine, Ain Shams Universit, for his sincere and experienced guidance, kindness, continuous supervision and creative suggestions.

Last, I want to thank my family, my friends and my patients without their help, this work could not have been completed.



List of Abbreviations

ADH : Anti Diuretic H ormone.

ANP : Atrial Natriuretic Peptide.

APC : Antigen Presenting Cells.

BT : Bacterial Translocation.

CGRP : Calcitonin Gene-Related Peptide.

CNNA : Culture-Negative Neutrocytic Ascites.

CSF : CerebroSpinal Fluid.

CT : Computed Tomography.

CTP : Child-Turcotte-Pugh.

GALT : Gut-Associated Lymphoid Tissue.

GI : Gastro-Intestinal.

HE : Hepatic Encephalopathy.

HRS : Hepatorenal Syndrome.

Ig : Immunoglobulins.

IL-6 : Interleukin-6.

LDH : Lactate Dehydrogenase.

LE : Leukocyte Esterase.

LVP : Large Volume Paracentesis.

MELD : Model For End-Stage Liver Disease.

List of Abbreviations



MLN : Mesenteric Lymph Nodes.

MNNB : Monomicrobial Non-Neutrocytic Bacterascites.

MRI : Magnetic Resonance Imaging.

NO : Nitric oxide.

PMN : Polymorphonuclear leukocyte count.

PRR : Pattern Recognition Receptor.

PVS's : PeritoneoVenous Shunt.

RAAS : Rennin-Angiotensin-Aldosterone System.

SAAG : Serum – Ascites Albumin Gradient.

SBEM : Spontaneous Bacterial Empyema.

SBP : Spontaneous Bacterial Peritonitis.

SNS : Sympathetic Nervous System.

TGF-B : Transforming Growth Factor-B.

TIPS : Transjugular Intra Hepatic PortoSystemic

Shunt.

TLR : Toll Like Receptor.

US : Ultra-Sonography.



List of Figures

Item	Page
Fig. (1): Age of patients in the studied groups	93
Fig. (2): Sex of patients in the studied groups	94
Fig. (3): Clinical findings of patients in the studied groups	95
Fig. (4): Haemoglobin level of patients in the studied groups	97
Fig. (5): Total leukocytic and Platelet count of patients in the studied groups	97
Fig. (6): Creatinine level in the studied groups	98
Fig. (7): Urea level of patients in the studied groups	98
Fig. (8): Liver transaminases level in the studied groups .	99
Fig. (9): Serum albumin level in the studied groups	99
Fig. (10): Bilirubin level (total and direct) in the studied groups	100
Fig. (11): Prothrombin time in the studied groups	100

List of Figures



Item	Page
Fig. (12): Ascitic analysis in the studied groups	101
Fig. (13): Ascitic fluid analysis before and 48 hours after	
treatment in group (I) (SBP)	102
Fig. (14): Ascitic fluid reagent strip testing in the studied	
groups	103
Fig. (15): Ascitic fluid reagent strip testing before and	
48hours after treatment in group (I) (SBP)	104
Fig. (16): Show comparison between group (I) and group	
(II) as regard results of pleural fluid reagent	107
strip testing	106
Fig. (17): Show comparison between pleural fluid reagent strip testing before and 48hours after treatment in group (I)	107
Fig. (18): Ascitic and pleural fluid cultures in the studied	
grouns	108



List of Tables

Item	Page
Table (1): Classification for the causes of cirrhosis according to prevalence	6
Table (2): Child-Pugh Classification	13
Table (3): Predictive factors for development of hepatorenal syndrome in patients with cirrhosis and ascites	64
Table (4): Diagnostic criteria of hepatorenal syndromea	65
Table (5): Vasoactive factors potentially involved in the regulation of renal perfusion in cirrhosis and in the pathogenesis of hepatorenal syndrome	66
Table (6): Risk factors for development of spontaneous bacterial empyema	72
Table (7): Diagnostic criteria of spontaneous bacterial empyema	72
Table (8): Show comparison between group (I) and group (II) as regard age	93
Table (9): Show comparison between group (I) and group (II) as regard sex	94
Table (10): Show comparison between group (I) and group (II) as regard clinical findings	95
Table (11): Show comparison between group (I) and group (II) as regard laboratory tests	96
Table (12): Show comparison between group (I) and group (II) as regard ascitic fluid (A. F.) analysis	101

List of Tables



Item	Page
Table (13): Show comparison between ascitic fluid analysis before and 48 hours after treatment in group (I) (SBP)	102
Table (14): Show comparison between group (I) and group (II) as regard results of ascitic fluid reagent strip testing	103
Table (15): Show comparison between ascitic fluid reagent strip testing before and 48hours after treatment in group (I) (SBP)	104
Table (16): Show correlation between ascitic fluid strip testing result and the ascitic fluid cell count	105
Table (17): Sensitivity, specificity, positive and negative predictive value of the ascitic fluid reagent strip test	105
Table (18): Show comparison between group (I) and group (II) as regard results of pleural fluid reagent strip testing	106
Table (19): Show comparison between pleural fluid reagent strip testing before and 48hours after treatment in group (I)	107
Table (20): Show correlation between Spontanous Bacterial Peritonitis (SBP)and Spontanous Bacterial Empayema (SBEM) in the studied groups unction	108
Table (21): Show correlation between Spontanous Bacterial Peritonitis (SBP)and Spontanous Bacterial Empayema (SBE) as regad ascitic and pleural fluid cultures in the studied groups	108
Table (22): Show organisms detected in ascitic fluid cultures of patients in group (I) (SBP)	109



Contents

Item	Page
Introduction	1
Aim of the Work	3
Review of Literature	4
- Liver Cirrhosis	4
- Ascites	18
- Spontaneous Bacterial Peritonitis	33
Patients and Methods	74
Results	85
Discussion	110
Summary	119
Conclusion and Recommendation	122
References	123
A rabic summary	



Introduction

Ascites is a collection of extracellular fluid in the peritoneal cavity resulting from imbalance between inflow and outflow through peritoneal membrane (*Bataller et al.*, 1997).

Patients with cirrhosis and ascites have a greatly increased total body sodium and water. The main reason for this is a marked retention of sodium and water by the kidney (*Ring-larson and Herinken*, 1986).

Besides variceal bleeding, spontaneous bacterial peritonitis (SBP) is another serious complication that can develop in cirrhotic patients (*Jepsen et al.*, 2003).

Symptoms of SBP include: fevers, chills, nausea, vomiting, abdominal tenderness and general malaise. Patients may complain of abdominal pain and worsening ascites (*Filik and Unal*, 2004).

Unfortunately symptoms of SBP are not present in all cirrhotic patients who develop SBP (*Boixeda et al.*, 1996). In addition many hospitalized cirrhotic patients develop SBP during their non-SBP related admissions especially those with gastrointestinal bleeding (*Deschenes and Villeneuve*, 1999). Therefore routine diagnostic paracentesis is the recommended practice for patients with ascites who develop signs or risk factors for SBP (*Deschenes and Villeneuve*, 1999). The standard criteria for diagnosis of SBP are ascitic fluid PMN (polymorphoneuclear) cell



count of $> 250 \text{ mm}^3$ and/or a positive ascitic fluid bacterial culture (*Rimola et al.*, 2000).

Cefotaxime or other third-generation cephlosporins have been considered the first-choice empirical antibiotics in the treatment of cirrhotic patients with SBP and is efficacius in approximately 90% of cases (*Strauss and Caly, 2006*). Broad-spectrum quinolones which almost completely absorbed after oral administration and diffuse rabidly through the ascitic fluid are currently used for oral treatment of uncomplicated SBP (*Strauss and Caly, 2006*).

Prophylactic oral norfloxacin is extremely useful in preventing SBP in patients that are at high risk for developing SBP such as hospitalized cirrhotic patients with gastrointestinal hemorrhage or low ascitic fluid protein (*Guarner and Soriano*, 1997).

Recently, leukocyte esterase activity testing by dipstick has been used for a rapid diagnosis of infection in many body fluids such as urine, pleural fluid and cerebrospinal fluid (*Rungsun et al.*, 2006).

The aim of this study is to evaluate the usefulness of dipstick in rapid diagnosis of SBP in cirrhotic patients with pleural effusion.



Aim of the Work

Evaluation of leukocyte esterase reagent strips as a rapid bedside diagnosis of SBP and associated pleural fluid effusion in cirrhotic patient and evaluation of responding to medical treatment .



Liver Cirrhosis

History:

Cirrhosis was first described in the fourth century B.C. hippocratic aphorism: "In case of jaundice, it is a bad sign when the liver becomes hard (*Chen and Chen, 1984*). The word "cirrhosis" is a neologism that derives from Greek kirrhos, meaning "tawny" (the orange-yellow colour of the diseased liver). While the clinical entity was known before, it was Rene Leannec who gave it the name "cirrhosis (*Rogun, 2006*).

Definition:

Cirrhosis is defined anatomically as a diffuse hepatic process characterized by fibrosis and nodule formation. Cirrhosis represents the final common histologic pathway for a wide variety of chronic disease. The progression of liver injury to cirrhosis may occur over weeks to years (Sherlock and Dooley, 2002).

Pathogenesis of Cirrhosis:

The response of the liver to necrosis are limited; the most important are collapse of hepatic lobules, formation of diffuse fibrous septa and nodular regrowth of liver cells. Thus, irrespective of the aetiology, the ultimate histological pattern of the liver is the same, or nearly the same (*Iredule*, 2003).