

بسم الله الرحمن الرحيم

 $\infty\infty\infty$

تم رفع هذه الرسالة بواسطة / سامية زكى يوسف

بقسم التوثيق الإلكتروني بمركز الشبكات وتكنولوجيا المعلومات دون أدنى مسئولية عن محتوى هذه الرسالة.

ملاحظات: لا يوجد

AIN SHAMS UNIVERSITY

Since 1992

Propries 1992



Serum Ferritin and cellular reactive protein (CRP) in non alcoholic Fatty liver disease(NAFLD) and non alcoholic steatohepatitis(NASH) patients

Thesis

Submitted for Partial Fulfillment of Master Degree In Gastroenterology and Hepatology

By

Mai Aboelkheir Awad Mohammed Dabiesh (M.B.,B.Ch.) Faculty of Medicine, Ain Shams University

Supervised by

Prof. Dr. Amira Ahmed Salem

Professor of Internal Medicine Faculty of Medicine - Ain Shams University

Prof. Dr. Khaled Hamdy Abdelmageed

Professor of Internal Medicine
Faculty of Medicine - Ain shams University

Dr. Mohamed Nabil Badawy

Lecturer of Internal Medicine Faculty of Medicine - Ain shams University

> Faculty of Medicine Ain Shams University

> > 2022

Acknowledgement

First of all, all gratitude is due to Allah almighty for blessing this work, until it has reached its end, as a part of his generous help, throughout my life.

Really I can hardly find the words to express my gratitude to **Prof. Dr. Amera Ahmed Salem**, Professor of Internal Medicine, Gastroenterology, Faculty of Medicine – Ain Shams University, for her supervision, continuous help, encouragement throughout this work and tremendous effort she has done in the meticulous revision of the whole work. It is a great honor to work under her guidance and supervision.

I would like also to express my sincere appreciation and gratitude to **Prof. Dr. Khaled Hamdy Abdelmageed**, Professor of Internal Medicine, Gastroenterology, Faculty of Medicine – Ain Shams University, for his continuous directions and support throughout the whole work.

I cannot forget the great help of **Dr. Mohammed Nabil Badawy**, Lecturer of Internal Medicine, Gastroenterology, Faculty
of Medicine – Ain Shams University, for his invaluable efforts,
tireless guidance and for his patience and support to get this work
into light.

Last but not least, I dedicate this work to my family, whom without their sincere emotional support, pushing me forward this work would not have ever been completed.

Mai Abou Elkheir Dabiesh

List of Contents

Ti	itle :	Page
•	List of Abbreviations	I
•	List of Tables	II
•	List of Figures	V
•	Introduction	1
•	Aim of the Study	3
•	Review of Literature	
	- NAFL	4
	- Ferritin	9
	- CRP	13
	- NASH	15
•	Patients and Methods	36
•	Results	47
•	Discussion	76
•	Summary	85
•	Conclusion	89
•	Recommendations	90
•	References	91
•	Arabic Summary	

List of Abbreviations

Abb.	Full-term		
AST	Aminotransferase, Aspartate		
BMI	• •		
CBC	Complete Blood Count		
CCL-2	CC-Chemokine Ligand 2		
CRP	Cellular Reactive Protein		
ELF	Enhanced Liver Fibrosis		
H.S	Highly Significant		
на	Hyaluronic Acid		
HOMA-IR	Homeostasis Model Assessment of Insulin Resistance		
IL-6	Interleukin-6		
LDL	Low Density Lipoprotein		
N.S	Non-Significant		
NAFLD	Nonalcoholic Fatty Liver Disease		
NASH	Non Alcoholic Steatohepatitis		
SF	Serum Ferritin		
Sig	Significant		
SPSS	Statistical Package for Social Sciences		
TNF-a	Tumor Necrosis Factor Alpha		
u.s	United States		
U/S	Ultrasound		
VLDL	Very Low Density Lipoprotein		

List of Tables

Table No.	Title Page
Table (1):	Comparison between the studied groups
Table (2):	Comparison between both groups as regard diabetes mellitus (DM)49
Table (3):	Comparison between both groups as regard abdominal ultrasound 50
Table (4):	Comparison between both groups as regard age
Table (5):	Comparison between both groups as regard body mass index (BMI)
Table (6):	Comparison between both groups as regard haemoglobin (Hb)
Table (7):	Comparison between both groups as regard platelets
Table (8):	Comparison between both groups as regard ALT level
Table (9):	Comparison between both groups as regard AST level
Table (10):	Comparison between both groups as regard albumin
Table (11):	Comparison between both groups as regard CRP
Table (12):	Comparison between both groups as regard fasting blood sugar (FBS) 59

List of Tables (Continued)

Table No.	Title Pa	ge
Table (13):	Comparison between both groups as regard ferritin	60
Table (14):	Comparison between both groups as regard LDL	61
Table (15):	Comparison between both groups as regard HDL	62
Table (16):	Comparison between both groups as regard cholesterol	63
Table (17):	Comparison between both groups as regard triglycerides	64
Table (18):	Comparison between both groups as regard NAFLD fibrosis score	65
Table (19):	Comparison between Ferritin level and mild, moderate, sever degree of abdominal ultrasound in both groups	66
Table (20):	Comparison between Ferritin level and gender in both groups	66
Table (21):	Comparison between Ferritin level and diabetes mellitus in both groups	67
Table (22):	Correlation between ferritin and other variables in both groups	68
Table (23):	Comparison between CRP level and mild, moderate, sever degree of abdominal ultrasound in both groups	70

List of Tables (Continued)

Table No.	Title	Page
Table (24):	Comparison between CRP level a gender in both groups	
Table (25):	Comparison between CRP level a diabetes mellitus in both groups	
Table (26):	Correlation between CRP and oth variables in both groups	
Table (27):	ROC curve between Group A a Group B	

List of Figures

Figure No.	Title Page	9
Fig. (1):	Mallory-Denk body	
Fig. (2):	Ballooning degeneration 17	
Fig. (3):	NASH (inflammation) and fibrosis stage 1	
Fig. (4):	NASH (inflammation) and fibrosis stage 218	
Fig. (5):	Lobular inflammation	
Fig. (6):	NASH pathogenesis	
Fig. (7):	non alcoholic fatty liver disease treatment method	
Fig. (8):	Bariatric_surgery_in_NAFLD28	
Fig. (9):	Comparison between both groups as regard gender there is no significant difference. Group A 50% male and 50% female, Group B 45% male and 55 % female	
Fig. (10):	Comparison between both groups as regard diabetes mellitus (DM), group A 100% non-diabetic and group B 75% non-diabetic and 25% diabetic with significant difference between both groups	
Fig. (11):	Comparison between both groups as regard abdominal ultrasound, mild group A 75%, group B 35% and moderate group A 15%, group B 25% and sever group A 10%, group B 40% with significant difference between both groups	

List of Figures (Continued)

Figure No.	Title	Page
Fig. (12):	Comparison between both groups a regard age there is no significant difference between both groups	ıt
Fig. (13):	Comparison between both groups a regard body mass index (BMI) there is no significant difference between both groups	re n
Fig. (14):	Comparison between both groups a regard haemoglobin (Hb) there is n significant difference between both groups	o h
Fig. (15):	Comparison between both groups a regard platlets there is no significant difference between both groups	ıt
Fig. (16):	Comparison between both groups a regard ALT level there is n significant difference between both groups	o h
Fig. (17):	Comparison between both groups a regard AST level there is n significant difference between both groups	o h
Fig. (18):	Comparison between both groups a regard albumin there is no significant difference between both groups	ıt
Fig. (19):	Comparison between both groups a regard CRP there is no significant difference between both groups	ıt

List of Figures

Figure No.	Title Page
Fig. (20):	Comparison between both groups as regard fasting blood sugar (FBS) there is significant difference between both groups
Fig. (21):	Comparison between both groups as regard ferritin there is significant difference between both groups 60
Fig. (22):	Comparison between both groups as regard LDL there is significant difference between both groups 61
Fig. (23):	Comparison between both groups as regard HDL there is no significant difference between both groups 62
Fig. (24):	Comparison between both groups as regard cholesterole there is significant difference between both groups
Fig. (25):	Comparison between both groups as regard triglycerides there is no significant difference between both groups
Fig. (26):	Comparison between both groups as regard NAFLD fibrosis score there is significant difference between both group
Fig. (27):	Sensitivity and specificity of ferritin 74
Fig. (28):	Sensitivity and specificity of CRP 74

List of Figures

Figure No.	Title	Page
Fig. (29):	Sensitivity and specificity of f	
	and CRP	75

INTRODUCTION

Nonalcoholic fatty liver disease (NAFLD) is now recognized as the most common cause of liver disease and may be present in up to 20% of the U.S. population.

Serum ferritin (SF) levels are commonly elevated in patients with nonalcoholic fatty liver disease (NAFLD) because of systemic inflammation, increased iron stores, or both.

However, ferritin is also an acute-phase protein and can also be induced in the setting of systemic inflammation^[1].

Expression of ferritin, the primary tissue iron-storage protein in the liver, where most extra body iron is stored, is induced in primary or secondary iron overload disorders, resulting in increased hepatic and circulating ferritin levels^{[2].}

Serum ferritin is an important inflammatory disease marker, as it is mainly a leakage product from damaged cells^[3].

Recent data suggests that excess iron induces vascular damage by increasing levels of the hormone hepcidin, which would determine iron trapping into macrophages, oxidative stress, and promotion of transformation into foam cells.

-Introduction

Elevation of serum ferritin may occur in hereditary hemochromatosis, inflammation, liver disease caused by hepatitis B and C virus, and alcoholic liver disease^[4].

Serum ferritin is a discriminant marker for both fibrosis and inflammation in histologically proven non-alcoholic fatty liver disease patients^[5].

AIM OF THE STUDY

To study the relation between serum ferritin and CRP levels and severity of non-alcoholic fatty liver disease NAFLD and non-alcoholic steatohepatitis NASH.