

AIN SHAMS UNIVERSITY
FACULTY OF SCIENCE

HISTOLOGICAL AND ULTRASTRUCTURAL STUDIES ON THE EFFECT OF CERTAIN NONSTEROIDAL ANTI - INFLAMMATORY DRUGS ON THE TESTES AND SPERMATOZOA OF MICE

A THESIS
SUBMITTED
BY

OLFAT MOHAMED HUSIEN YOUSSEF

(B. Sc., Ed. B. Sc. & M.Sc.)

ASSISTANT LECTURER OF ZOOLOGY,
FACULTY OF EDUCATION, AIN SHAMS UNIVERSITY



FOR

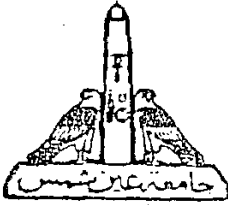
THE AWARD OF THE DEGREE OF
DOCTOR OF PHILOSOPHY IN ZOOLOGY

TO

THE FACULTY OF SCIENCE
AIN SHAMS UNIVERSITY



1999



AIN SHAMS UNIVERSITY
FACULTY OF SCIENCE

HISTOLOGICAL AND ULTRASTRUCTURAL STUDIES ON THE EFFECT OF CERTAIN NONSTEROIDAL ANTI - INFLAMMATORY DRUGS ON THE TESTES AND SPERMATOZOA OF MICE

A THESIS
SUBMITTED
BY

OLFAT MOHAMED HUSIEN YOUSSEF
(B. Sc., Ed. B. Sc. & M.Sc.)

*ASSISTANT LECTURER OF ZOOLOGY,
FACULTY OF EDUCATION, AIN SHAMS UNIVERSITY.*

FOR

THE AWARD OF THE DEGREE OF
DOCTOR OF PHILOSOPHY IN ZOOLOGY

TO

THE FACULTY OF SCIENCE
AIN SHAMS UNIVERSITY

1999

M. Shahn
Elham



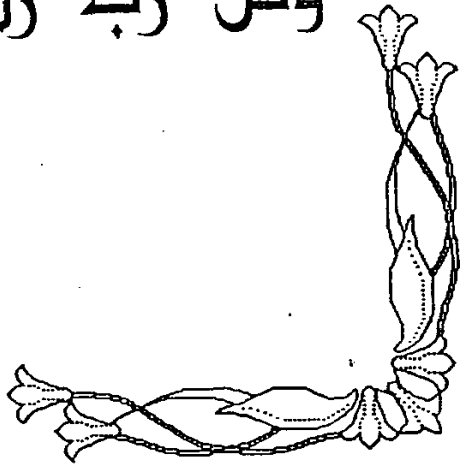


"بسم الله الرحمن الرحيم"

"وقل رب زدني علماً"

"صدق الله العظيم"

{سورة "طه" - الآية رقم ١١٤}



THIS DISSERTATION HAS NOT PREVIOUSLY
BEEN SUBMITTED FOR ANY DEGREE, AT THIS
OR ANY OTHER UNIVERSITY. THE
REFERENCES CITED, SHOW THE EXTENT TO
WHICH I HAVE AVAILED MYSELF OF THE
WORK OF OTHER AUTHORS.

OLFA7 HUS7EN

***We apologize for any errors we have
inadvertently allowed into print***

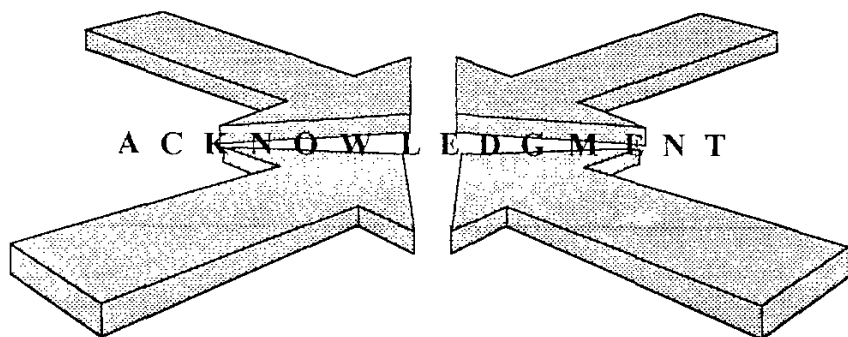


**A WORD OF THANKS,
A WORD OF PRAISE,
➤ FOR MY
“MOTHER, HUSBAND, SON AND
DAUGHTERS”
FOR BEING SO GREAT IN MANY WAYS
THAT PUSHED ME FORWARD.
THEY SKILLFULLY AWAKENING MY
MOTIVITY TO WORK.**

OLFAT

27/02/00





ACKNOWLEDGMENT

First of all, thanks to ALLAH, who enabled me to overcome all problems which faced me throughout the work.

I am grateful to Dr. Fahmy I. Kattab, Professor of Cell Biology, Department of Zoology, Faculty of Science, Ain Shams University and Dr. Mohamed A. Shahin, Professor of Vertebrates and Embryology, Department of Biological Sciences & Geology, Faculty of Education, Ain Shams University, for suggesting the point, supervising the work, helpful advice and reading the manuscript. Also deepest thanks for continuous guidance and encouragement throughout the work. The author wishes to express her deepest thanks to Dr. Ilham I. Seif, Professor of Pathology, Faculty of Medicine, Ain Shams University for helpful advice and reading the manuscript.

Sincere gratitude to Prof. Dr. Nahed Hussien, Head of Zoology Department, Faculty of Science, Ain Shams University, for providing valuable facilities and encouragement.

Sincere thanks are also afford to Prof. Dr. Mohamed Gharieb, Head of Biological Sciences & Geology Department, Faculty of Education, Ain Shams University, for his advice, providing valuable facilities and encouragement.

I wish to express my deepest appreciation to all members of the Biological Sciences & Geology Department, Faculty of Education and Zoology Department, Faculty of Science, Ain Shams University, for their help and cooperation.



ABSTRACT

HISTOLOGICAL AND ULTRASTRUCTURAL STUDIES ON THE EFFECT OF CERTAIN NONSTEROIDAL ANTI-INFLAMMATORY DRUGS ON THE TESTES AND SPERMATOZOA OF MICE.

Eight microscopical and ultrastructural studies were conducted to assess the effect of the two nonsteroidal anti-inflammatory drugs, voltaren (diclofenac sodium, 9.4 mg/kg b. wt.), and feldene (piroxicam, 2.5 mg/kg b. wt) on the testes and spermatozoa of mice. The two applied drugs caused a decrease in the diameter of the seminiferous tubules, disorganization of the germinal epithelium, maturation arrest at various stages of spermatogenesis, formation of multinucleate giant cells and various degrees of degeneration of spermatocytes and spermatids. Ultrastructural results showed spermatocytes, spermatids and Sertoli cells with disintegrating organelles, particularly mitochondria and endoplasmic reticulum, whereas the cytoplasm exhibited a vacuolated appearance. The drugs caused thickening and folding of the peritubular tissue. The common types of abnormalities observed in sperms of treated mice were amorphous heads and coiled tails. Diclofenac sodium caused a marked increase in the percentage of total sperm abnormalities as compared to piroxicam. Ultrastructural observations revealed malformed heads and degenerative changes of the acrosome, the connecting piece and the middle piece. Moreover, confused arrangement of the mitochondrial sheath and disorganization of the components of the axial filament complex of the middle piece were detected.

➤ **In conclusion**, diclofenac sodium and piroxicam induced variable time-dependent deleterious effects in the testicular tissue and spermatozoa of mice.

Key words:

Anti-inflammatory drugs, Diclofenac sodium, Piroxicam, Testes, Spermatozoa, Histopathology, Ultrastructure.

CONTENTS

ACKNOWLEDGMENT	I
ABBREVIATIONS	VI
LIST OF TABLES	IX
LIST OF FIGURES	X
ABSTRACT	i
INTRODUCTION	1
AIM OF THE WORK	4
REVIEW OF LITERATURE	6
MATERIALS AND METHODS	18
RESULTS AND OBSERVATIONS	26
HISTOLOGY AND HISTOPATHOLOGY OF THE TESTIS.	26
HISTOLOGY	26

(777)

Control mice	26
HISTOPATHOLOGY	42
Diclofenac sodium-treated mice	42
Examination after 14 days	42
Examination after 21 days	51
Piroxicam-treated mice	67
Examination after 14 days	67
Examination after 21 days	74
Recovering mice	87
ULTRASTRUCTURE OF THE TESTES	93
Control mice	93
Diclofenac sodium-treated mice	119
Examination after 14 days	119
Examination after 21 days	132
Piroxicam-treated mice	151
Examination after 14 days	151
Examination after 21 days	168
Recovering mice	179
LIGHT MICROSCOPIC OBSERVATIONS OF	
MICE SPERMATOZOA	187
Control mice	187
Diclofenac sodium-treated mice	189
Examination after 14 days	189
Examination after 21 days	189
Piroxicam-treated-mice	205
Examination after 14 days	205
Examination after 21 days	205
Recovering mice	206
ELECTRON MICROSCOPIC OBSERVATIONS	
OF MICE SPERMATOZOA	225
Control mice	225
Diclofenac sodium-treated mice	238
Piroxicam-treated mice	244

DISCUSSION	251
SUMMARY AND CONCLUSIONS	286
REFERENCES	291
ARABIC SUMMARY	v
ARABIC ABSTRACT	i

ABBREVIATIONS

AC	Acrosomal cap
AF	Axial filament
AG	Acrosomal granule
AN	Annulus
AT	Articular surface
AV	Acrosomal vesicle
BI	Basal lamina
BT	Boundary tissue
C	Centriol
CB	Chromatoid body
CD	Cytoplasmic droplet
CF	Central fibrils
CP	Connecting piece
CV	Coated vesicle
DB	Dense bodies
DC	Degenerated cells
DF	Dense fibers
Dg	Degenerative germ cell
DPP	Distal principal piece
EP	End piece
FS	Fibrous sheath
G	Golgi apparatus
Hd	Head
Hk	Hook

SS	Secondary spermatocyte
ST	Seminiferous tubule
TU	Tunica albuginea
V	Vacuole
Z	Spermatozoa