

Evaluation of the Protective Role of Date (*Phoenix dactylifera*) Against Flusilazole Fungicide Toxicity in Male Albino Rats

A thesis

Submitted for the Award of the Ph. D. Degree of Science in Zoology (Physiology)

 $\mathbf{B}\mathbf{y}$

Soad Ahmed Ebrahim Khwanes

M.Sc. (Zoology). 2007

Supervisors

Prof. Dr. Nadia Mohamed Abd El-Aziz El-Beih

Prof. of Physiology-Faculty of Science
Ain Shams University

Prof. Dr. Magdy Mohammed El-Said Gad

Department of Mammalian and Aquatic Toxicology

Central Agricultural Pesticides Lab

Agricultural Research Center

I dedicate this work to my family especially my mother, my husband, my dear daughter and my kind son.

THE THE THE POLICE

وَلْيَخْشَ لِلَّذِينَ لَوْ تَرَكُولِ مِنْ خَلْفِهِمْ ذُرِيَّةً فَلْيَتَقُولِ لِللَّهِ وَلْيَقُولُولِ ضِعَافًا خَافُولِ عَلَيْهِمْ فَلْيَتَقُولِ لِللَّهِ وَلْيَقُولُولِ فَعُرِيدًا لِللَّهِ وَلَيْقُولُولِ قَوْلُ اسْدِيدًا لِ

Acknowledgement

Acknowledgments

First and for the most, cordial thanks due to **Allah** who helped me to accomplish this work, and enabled me to overcome all the problems, which faced me throughout the work. Then I would like to express sincere thanks and gratitude to my committee for their guidance and support through this project.

I am deeply grateful and indebted to **Prof. Dr. Nadia Mohamed Abd El-Aziz El-Beih**, Prof of physiology, Department of Zoology, Faculty of Science, Ain Shams University, for her kind supervision, planning of the study, her valuable criticism and suggestion, motivation during this work and critical reading of the manuscript.

Thanks due to **Prof. Dr. Magdy Mohamed El-Said,** Prof of Toxicology, Department of Mammalian and Aquatic Toxicology, Central Agricultural Pesticides Laboratory (CAPL), Agricultural Research Center.

Appreciation and deep thanks are due to **Prof. Dr. Mohamed Abdel-Razek** Director of Central Agricultural Pesticides Laboratory (formerly), for fruitful advices and practical help.

I would like also to express my gratitude and appreciation to **Prof. Dr. Monir Almaz** Director of Central Agricultural Pesticides Laboratory (formerly), for his practical help during this thesis.

Special thanks are due to **Prof. Dr. Mamdouh El-Sherif** Director of Central Agricultural Pesticides Laboratory for his great aid.

Also, I would like to express my deepest and sincere appreciation to **Prof. Dr. Islam Noeman Nasr** head of Residual Department (formerly) for his practiced help.

Thanks are also extended to head and all members of Zoology Department, Faculty of Science, Ain Shams University.

(Late) Prof. Dr. Al-Huseiny Naguib, words cannot express my feelings nor my thanks for all his help. If the world had more people like him it would be a better place.

Thanks very much to **Prof. Dr. Mohamed Farid** head of Mammalian and Aquatic Toxicology Department (formerly), and everybody at Central Agricultural Pesticides Laboratory. Special thanks to head and all my co-workers at the Mammalian Toxicology Department that have been like part of my family and have helped me in my weak moments. This project would not have been possible without their help.

Lastly, I would like to extend great thanks to my family especially my mom, my sisters, my brother, my husband, my dear daughter, Noha and my kind son, Mohamed. They encouraged me to continue moving forward when I thought I would fail. Thanks for all their support, confidence and patience which helped me achieve what I am today.

List of Abbreviations

A/G ratio Albumin/ Globulin ratio
ALAT Alanine aminotransferase
ANOVA Analysis of variance

ASAT Aspartate aminotransferase ATP Adenosine triphosphate BAT Brown adipose tissue

b. w. Body Weight

Cal. Calorie

CAPL Central Agricultural Pesticides Lab.

CAT Catalase

CCL₄ Carbon tetra-chloride

CDNB 1-chloro-2,4-dinitrobezene

Cl⁻ Chloride ion

CYP Cytochrome P450
DNA Deoxyribonucleic acid

DTNB 5-5'-dithio-bis(2-nitrobenzoic acid)

EC Emulsifiable concentration

EDCs Endocrine-disrupting Chemicals
EDTA Ethylene diamine tetra acetic acid

EPC Erythropoietin

Fig Figure

FL Femtolitre, 10^{-15}

g gram

GGT Gamma glutamyl transferase

GI Glycemic index

GR Glutathione reductase
GS Glutamin synthetase
GSH Total glutathione

GSSG Glutathione Disulfied

GST Glutathione S-Transferase GPx Glutathione peroxidase HAT Hour after treatment

Hb Haemoglobin concentration

HCl Hydrochloric acid

HDL-C High density lipoprotein-Cholesterol

HPLC High Performance Liquid Chromotography

H₂O₂ Hydrogen Peroxide

hr Hour

i.e. Identified example Ig Immunoglobulin

LC Liquid Chromatography LD₅₀ Median Lethal dose

LDL-C Low density lipoprotein-Cholesterol

LP Lipid Peroxidation

MCV Mean Corpuscular Volume

MCH Mean Corpuscular Haemoglobin

MCHC Mean Corpuscular Haemoglobin concentration

MDA Malondialdehyde

mg Milligram

MTD Maximum tolerated dose

M. W. Molecular Weight NaCl Sodium chloride

NAD Nicotinamide adenine dinucleotide

NADPH Nicotinamide adenine dinucleotide phosphate

ND Not Detected nmol Nanomole

OD Optical Density
OH Hydroxyl radical

 O_2 Oxygen

O₂ Superoxide anion
PCV Packed Cell Volume
PF Pair-Fed control
Pg Picogram, 10⁻¹²
ppb Pert per Billion
ppm Part per million

PROTOX Protoporphyrinogen Oxidase PUFA Polyunsaturated Fatty Acids

RBCs Red Blood Corpuscles

RDA Recommended Dietary Allowance

ROOH Reduction of variety of hydroperoxides

ROS Reactive Oxygen Species

rpm Round Per Minute

SDS Sodium Dodecyl Sulphate

SE Standard Error

SOD Superoxide dismutase T₃ Tri-iodothyronine

T₄ Thyroxine

TAG Triacylglycerol
TBA Thiobarbituric acid

TBARS Thiobarbituric acid reactive substance

TC Total cholesterol
TCA Trichloroacetic Acid
THs Thyroid Hormones

TMP 1,1,3,3- tetramethoxy propane

TNB 5-Thio- 2- Nitrobezoate

T.P Total Protein

TSH Thyroid stimulating hormone

UDP-GT uridine-diphosphate glucuronosyl transferase.

U/l Unite/ Litre
UV Ultra-Violet

VLDL Very low density lipoprotein

V/V Volume/Volume

Vol. Volume

WBCs White Blood Cells W/V Weight/Volume

 $\begin{array}{ccc} xg & & Gravity \\ \mu M & & Micro \ mole \end{array}$

Contents

Abstract	1
Introduction	3
Literature Review	6
Materials and methods	44
Results	76
Discussion	214
Summary and conclusion	254
References	259
Arabic summary	•••••