

# Vasculoprotective Effect of Pioglitazone on Cyclosporine-Induced Vascular Damage in Rats

#### A Thesis

Submitted to the Faculty of Pharmacy
University of Alexandria
In Partial Fulfillment of the Requirements for the Degree

of **Master of Science** 

in
Pharmaceutical Sciences
(Pharmacology)

by

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B.Sc. in Pharmaceutical Sciences, 2006 Faculty of Pharmacy University of Alexandria

Department of Pharmacology and Toxicology
Faculty of Pharmacy
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#### LIST OF ABBREVATIONS

AT<sub>1</sub> Angiotensin receptor-1

**ATP** Adenosine triphosphate

**ATPase** Adenosine triphosphatase

AUC Area under the curve

BH<sub>4</sub> Tetrahydrobiopterin

**BK**<sub>Ca</sub> - **MaxiK** Large-conductance calcium activated potassium

channels

**cAMP** Cyclic adenosine monophosphate

**CAT** Catalase

**cGMP** Cyclic guanosine monophosphate

**CO** Carbon monoxide

**CORM** Carbon monoxide releasing molecule

**CSA** Cyclosporine-A

**Cu/Zn-SOD** Cupper zinc-containing superoxide dismutase

**DAG** Diacylglycerol

**DMSO** Dimethylsulphoxide

**DNA** Desoxyribonucleic acid

**DPTA** Diethylenetriamine pentacetic acid

EC<sub>50</sub> Half maximal effective concentration

**EDRF** Endotheilium derived relaxing factor

**Emax** Maximal response

**eNOS** Endothelial nitric oxide synthease

**ET** Endothelin

**FFA** Free fatty acids

**GC** Guanylate cyclase

GLUT Glucose transporter

**GPx** Glutathione peroxidase

**GR** Glutathione reductase

**GS** Thiyl radical

**GSH** Reduced glutathione

**GSSG** Oxidized glutathione

**GTP** Guanosine triphosphate

GW9662 Chloro-5-nitro-N-phenylbenzamide

**H&E** Hematoxylin and Eosin stain

H<sub>2</sub>O<sub>2</sub> Hydrogen peroxide

**HbO** Oxidized haemoglobin

**HDL-C** High density lipoprotein-Cholesterol

**HMG-CoA** Hydroxymethyl-glutaryl-coenzyme A

**HO** Heme oxygenase

ICAM Intracellular adhesion molecule

IL Interleukin

**iNOS** Inducible nitric oxide synthease

**IP3** Inositol triphosphate

LDL-C Low density lipoprotein-Cholesterol

MDA Malondialdehyde

MHC Major histocompatibility complex

MMP Metalloproteinase

mRNA Messenger ribonucleic acid

 $N_2O_4$  Dinitrogen tetraoxide

**NADP** Nicotinamide adenine dinucleotide phosphate

**L-NAME**  $N_{\omega}$ -Nitro-L-arginine methyl ester hydrochloride

**NED** N-1-naphthylethylene diamine dihydrochloride

**NFAT** Nuclear factor activated T-Cell

**NF-KB** Nuclear factor kappa B

**nNOS** Neuronal nitric oxide synthease

NO Nitric oxide

NOx NO metabolites (nitrite/nitrate)

 $O_2^-$  Superoxide anion

**OH** Hydroxyl radical

**ONOO** Peroxynitrite

Ox-LDL Oxidized-Low density lipoprotein

**PDE** Phosphodiesterase

**PE** Phenylephrine

PGI<sub>2</sub> Prostacyclin

**15d-PGJ(2)** 15-deoxy-Δ12,14-prostaglandin J2

PI3 Phosphoinositide 3-kinases

**PIO** Pioglitazone

**PIPE** piperazine-N,N'-bis(2-ethanesulfonic acid)

**PKA** Protein kinase A

**PPAR** Peroxisome proliferator activated receptors

PUFA Polyunsaturated fatty acids

**RO**<sup>.</sup> Alkoxyl radical

**ROO** Peroxyl radical

**ROS** Reactive oxygen species

**RPP** Renal perfusion pressure

**S.E.M** Standard error of mean

Ser Serine

**-SH** Thiol

**SNP** Sodium nitroprusside

**SOD** Superoxide dismutase

**TBA** Thiobarbituric acid

**TBARs** Thiobarbituric acid reactive substances

TC Total cholesterol

TCA Trichloroacteic acid

**TG** Triglycerides

T<sub>H</sub> Helper T-Cell

**TMP** 1,1,3,3-Tetramethoxypropane

TNF-α Tumor necrosis factor alpha

**TZD** Thiozolidinedione

VCAM Vascular cell adhesion molecule

**VEGF** Vascular endothelium growth factor

**ZnPPIX** Zinc protoprophyrin IX

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