

Potential utility of single and concurrent administration of a partial peroxisome proliferator activated receptor-γ agonist and a protein tyrosine phosphatase inhibitor in treating type II diabetic rats

A thesis submitted for the partial fulfillment of Master's degree in Pharmaceutical Sciences (Pharmacology and Toxicology)

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Besides the work presented in the thesis, the candidate has attended prerequisite postgraduate courses including the following topics:

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- Computer skills
- Statistics

Special Courses:

- Chemotherapy
- Clinical Toxicology
- Neuropharmacology
- Pharmacology
- Selected topics in Pharmacology and Toxicology

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List of Abbreviations

ACC Acetyl coenzyme-A carboxylase ACE Angiotensin-converting enzyme

AdipoR Adiponectin receptor ADP Adenosine diphosphate

AGEs Advanced glycation end products
AMP Adenosine monophosphate

AMPK Adenosine monophosphate activated protein kinase

ANOVA Analysis of variance
Ang II Angiotensin II
apoB Apolipoprotein B
AR Androgen receptor

AT1 Angiotensin II-type1 receptor
AT2 Angiotensin II-type 2 receptor
ATM Ataxia telangiectasia mutated gene

ATP Adenosine triphosphate
AUC Area under curve
BSA Bovine serum albumin
CAP Cbl associated protein

Cbl Casitas B-lineage lymphoma CD36 Cluster of differentiation 36

CE Cholesterol esterase COx Cholesterol oxidase

CPT-1 Carnitine palmitoyltransferase-1

CVD Cardiovascular disease

DAG Diacyl glycerol

DAP Dihydroxyacetone phosphate DIR Diabetic/Insulin-resistant

DM Diabetes mellitus

DTNB 5,5'- dithiobis-2-nitrobenzoic acid
EARPG European Arterial Risk Policy Group
EDTA Ethylene diamine tetra acetic acid
ELISA Enzyme-linked immunosorbent assay
eNOS Endothelial nitric oxide synthase

FAS Fatty acid synthase

FCPD Fibrocalculous pancreatic diabetes

FFA Free fatty acids

FTO fat mass- and obesity-associated gene

G-3-P Glyerol-3-phosphate
G6P Glucose-6-phosphate
G6Pase Glucose-6-phosphatase

G6PDH Glucose-6-phosphate-dehydrogenase

GAD Glutamic acid decarboxylase GDM Gestational diabetes mellitus

GIGT Gestational impaired glucose tolerance

GK Glycerol kinase GLUT Glucose transporter GO Glucose oxidase

GPO Glycerylphosphate oxidase
GPx Glutathione peroxidase
GSH Reduced glutathione
H₂O₂ Hydrogen peroxide
HDL High density lipoprotein

HFD High-fat diet

HFFD High-fat, high-fructose diet

HK Hexokinase

HMG-CoA 3-hydroxy-3-methyl-glutaryl-Coenzyme A

HOMA Homeostasis model assessment

HPLC High performance liquid chromatography

HRP Horseradish peroxidase

IDDM Insulin-dependent diabetes mellitus

IFG Impaired fasting glycemia IgG Immunoglobulin G

IGT Impaired glucose tolerance

IL-6 Interleukin-6

iNOS Inducible nitric oxide synthase

i.p. Intra-peritoneal IR Insulin resistance

IRS Insulin receptor substrate

IV Intravenous

LDL Low density lipoprotein
LSD Least significant difference
MCP-1 Monocyte chemotactic protein-1

MTP Microsomal triglyceride transfer protein MRDM Malnutrition-related diabetes mellitus

mRNA Messenger RNA

NADH Nicotinamide adenine dinucleotide NCEP National Cholesterol Education Program

NEFA Non-esterified fatty acids NF-KB Nuclear factor-kappa B

NIDDM Non-insulin-dependent diabetes mellitus

NO Nitric oxide
NPD Normal pellet diet
NSB Non-specific binding
OCT Organic cation transporter

OD Optical density

OGTT Oral glucose tolerance test
PAI-1 Plasminogen activator inhibitor-1

PBS Phosphate buffered saline PCOS Polycystic ovary syndrome PDH Pyruvate dehydrogenase

PDK Phosphoinositide dependent kinase PDPD Protein–deficient pancreatic diabetes

PEG Polyethylene glycol

PEPCK Phosphoenol pyruvate carboxykinase

PFK Phosphofructokinase

p-HBS p-hydroxybenzene sulfonate

PI3K Phosphatidyl-inositol 3-kinase

PIP2 Phosphatidyl-inositol-3,4-biphosphate PIP3 Phosphatidyl-inositol-3,4,5-triphosphate

PKA Protein kinase A
PKB Protein kinase B
PKC Protein kinase C
PO Peroxidase
p.o. Per Os (oral)

PPAR Peroxisome-proliferator activated receptor

PPRE PPAR responsive elements
PTP Protein tyrosine phosphatase

RAGE Receptor for advanced glycation end products

RAS Renin-angiotensin system

RIA Radioimmunoassay
ROS Reactive oxygen species
rpm Round per minute
RXR Retinoid X receptor

S.E.M. Standard error of the mean SHBG Sex hormone–binding globulin

SOD Superoxide dismutase

SNP Single nucleotide polymorphism SRE Sterol responsive elements

SREBP Sterol regulatory element binding protein

STZ Streptozotocin

T1DM Type 1 diabetes mellitus T2DM Type 2 diabetes mellitus

TG Triglycerides

TMB 3, 3', 5, 5'-Tetramethyl benzidine TNF-α Tumor necrosis factor alpha

TZD Thiazolidinedione UCP Uncoupling proteins

VLDL Very low density lipoprotein

WAT White adipose tissue

WHO World Health Organization

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