



Effect of Manganese Bionanoparticles and /or Low Dose of Gamma Radiation on Rheumatoid Arthritis in Rats

A thesis

Submitted in partial fulfillment of the requirements for the Master degree of Science
In Biochemistry

By

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﴿قَالُواْ سُبْحَــانَكَ لاَ عِلْمَ لَنَا إِلاَّ مَا عَلَّمْتَنَا إِنَّكَ أَنتَ الْعَلِيمُ الْحَكِيمُ﴾

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Declaration

I declare that this thesis has been composed by me and that the work which is recorded herein after has been done by me. It has not been submitted for a degree at this or any other university.

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I dedicate this work with all my love to the spirit of my dear father, deep thanks and appreciation to my beloved father for his kindness and continuous advice; god bless his soul, to all my family and my husband Ramy who helped me during my entire thesis and to all those who contributed towards my scientific knowledge, whenever and wherever they are.

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List of abbreviations

ACD Anemia of chronic disease

AD Atopic dermatitis

ADP Adenosine diphosphate
AFCs Antibody-forming cells

AI Arthritic index

AIA Adjuvant-induced arthritis

AKT Protein kinase B

ALT Alanine transaminase (aminotransferase)

ANOVA One-way analysis of variance
Anti-CCP Anti-cyclic citrullinated peptide
Acute respiratory distress syndrome

AST Aspartate transaminase (aminotransferase)

B-cells B stand for maturation in the bone marrow

CAMs Cell adhesion molecules

CCL20 Chemokine (C-C motif) ligand 20

CD Cluster of differentiation (cell surface molecule)
CD25+ Cluster of differentiation 25 (surface marker protein

on regulatory T-cells)

CD36 Thrombospondin receptor CD4+ T-cells Helper T-cells (T-lymphocyte)

CD8+ T-cells Cytotoxic T-cells

CFA Complete Freund's adjuvant

COXs Cyclooxygenases CRP C-reactive protein

Cu-Zn SOD Copper-zinc superoxide dismutase

Cyt c Cytochrome c DCs Dendritic cells

DLS Dynamic light scattering

DMARDs Disease-modifying anti-rheumatic drugs

DTNB 5, 5` dithiobis-2-nitrobenzoic acid

e aq Hydrated electron

EAEA Egyptian atomic energy authority
EC-SOD Extracellular superoxide dismutase
EDTA Ethylenediaminetetraacetic acid
Extracellular superoxide in and othelial call

E-selectin Selectin expressed in endothelial cells

ETC Electron transport chain

Fc Fragment crystallizable region of the antibody

Fcγ Fragment crystallizable gamma receptors

FMN Flavonoids mediated nanomaterial

Fox p3+ Forkhead box p3 (surface marker protein on

regulatory T-cells)

FTIR Fourier transform infrared analysis

GFR Glomerular filtration rate
GIT Gastrointestinal tract
GPx Glutathione peroxidase
GS Glutamine synthetase
GSH Reduced Glutathione

GSSG Glutathione disulfide (oxidized glutathione)

HCl Hydrochloric acidHDR High-dose radiationHO-1 Hemoxygenase-1

HRP Horseradish peroxidaseHT Hashimoto's thyroiditis

ICAMs Intracellular adhesion molecules

IFN-γ Interferon gammaIg ImmunoglobulinIkB Inhibitor of kappa B

IL Interleukin

IL-12p70 Inteleukin-12 protein of 70 kDa

IL-1β Interleukin-1beta

ILD Interstitial lung disease

iNOS Inducible nitric oxide synthase

IRR Irradiation

ISO International organization for standardization

LD₅₀ Median lethal dose
 LDH Lactate dehydrogenase
 LDI Low-dose irradiation
 LDR Low dose radiation
 LD-RT Low-dose radiotherapy

LOXs Lipoxygenases

L-selectins Selectin expressed in lymphocytes
 M1 Pro-inflammatory macrophages
 M2 Anti-inflammatory macrophages

MC Mononuclear cells

MCP Monocyte chemoattractant protein

MDH Malate dehydrogenase

MIP-α Macrophage inflammatory protein-α

Matrix metalloproteases **MMPs MnBNPs** Manganese bionanoparticles

MnCl₂ Manganese chloride **MnNPs** Manganese nanoparticles **MNPs** Metal nanoparticles

MnSOD Manganese superoxide dismutase

MPCs Mononuclear phagocytes

MS Multiple sclerosis

Nicotinamide adenine dinucleotide phosphate **NADPH**

(Reduced form)

NBT Nitroblue tetrazolium

National center for radiation research and technology **NCRRT** NF-kB p65 65 kDa protein subunit of nuclear factor kappa B

NF-kB Nuclear factor- kappa B NK Natural killer cells

NO Nitric oxide **NPs Nanoparticles**

NAD(P)H quinone dehydrogenase-1 **NOO-1** Nrf-2 Nuclear factor E2-related factor-2 **NSAIDs** Non-steroidal anti-inflammatory drugs **OECD**

Organization for economic co-operation and

development

ONOO. Peroxynitrite ion OS Oxidative stress

p38 MAPK Class of mitogen-activated protein kinases (MAPK) p450 Cytochrome protein shows maximum absorption at

450nm

PBMC Peripheral blood mononuclear cells

PG E2 Prostaglandin E2

PMNs Polymorphonuclear neutrophils (cells)

P-selectins Selectin expressed in platelets

Rheumatoid arthritis $\mathbf{R}\mathbf{A}$

RANKL Receptor activator of nuclear factor kappa-B ligand

RDA The recommended dietary allowances

RF Rheumatoid factor **RNS** Reactive nitrogen species

ROO Peroxyl radical

ROS Reactive oxygen species

SE Standard error

SLE Systemic lupus erythematosus

SLs Sesquiterpene lactones
SOD Superoxide dismutase

SOD-1 Superoxide dismutase-1 (Copper-zinc superoxide

dismutase)

TAOC Total antioxidant capacity

TBA Thiobarbituric acid
TCA Tricarboxylic acid

T-cells T stand for cells maturation in the thymus

TEM Transmission electron microscopy

TEs Trace elements

TGF-β Transforming growth factor-beta

TLC Total leukocytic count
TMB Tetramethyl-benzidine

TNFR1 Tumor necrosis factor receptor 1
TNF-α Tumor necrosis factor-alpha

Tregs Regulatory T-cells

UDP Up-and-Down procedure

UV-Vis Ultraviolet-visible spectral analysisVEGF Vascular endothelial growth factor

VFs Vertebral fractures WBC White blood cells

WHO World health organization

β-NADH Beta nicotinamide adenine dinucleotide (Reduced

form)

γ Gamma

γ-GCS Gamma-glutamylcysteine synthetase