UNIVERSITI TEKNOLOGI MARA

TOXIC EFFECTS OF MAGNESIUM OXIDE NANOPARTICLES ON HUMAN NEURONAL CELL LINES

NUR INTAN BINTI HASBULLAH

Thesis submitted in fulfilment of the requirements for the degree of **Master of Science**

Faculty of Applied Sciences

July 2015

AUTHOR'S DECLARATION

I declare that the work in this thesis was carried out in accordance with the regulations of Universiti Teknologi MARA. It is original and is the result of my own work, unless otherwise indicated or acknowledged as referenced work. This thesis has not been submitted to any other academic institution or non-academic institution for any degree or qualification.

I, hereby, acknowledge that I have been supplied with the Academic Rules and Regulations for Post Graduate, Universiti Teknologi MARA, regulating the conduct of my study and research.

Name of Student	:	Nur Intan Binti Hasbullah		
Student I.D. No.	:	2010804432		
Programme	:	Master of Science (AS780)		
Faculty	:	Faculty of Applied Sciences		
Thesis Title	:	Toxic Effects of Magnesium Oxide Nanoparticles on		
		Human Neuronal Cell Lines		
		$\langle \varphi \rangle$		
Signature of Student	:	·····		
Date	:	July 2015		
		,		

ABSTRACT

Rapid increase in the industrial application of nanomaterials such as MgO nanoparticles has great impact on the public. MgO nanomaterial is now used in medical and pharmaceutical products as well as industrial products such as paints. Thus, they are now widely used by the public, but their significant potential risks of exposure have not been systematically investigated. In this study, toxic effects of MgO nanoparticles which differ in terms of size and physical dimensions were investigated against cancer neuroblastoma SH-SY5Y cell lines and neuron-phenotypic cell lines. Viability of cells was measured by using MTS assay. Three types of MgO were used in the study, of which two of them were ultra-thin MgO sheets designated as MgO-5 (size: 500 µm, 5 nm thickness) and MgO-10 (size: 500 µm, 2 nm thickness). The other nanoparticle was spherical with average size of 10-100 nm (MgO-24). Oxidative stress was measured by the production of reactive oxygen species (ROS) using dicholofluorescein (DCF) assay. Beneficial effects of MgO nanoparticles were also evaluated for potential neuroprotection. Results indicated that the toxic effects of MgO nanoparticles on cancer neuroblastoma SH-SY5Y cell lines and neuron-phenotypic cell lines were dependent on shape, concentration and size of MgO nanoparticles. Particles with spherical shape (MgO-24) exerted more toxic effects compared to ultra-thin nanosheet particles (MgO-5 and MgO-10). MgO nanoparticles were not toxic to both cell lines at concentration $\leq 100 \ \mu$ g/ml and did not exert any potential for neuroprotection. However, once the concentration was increased to 1 mg/ml, MgO nanoparticles significantly reduced the viability of both cell lines tested due the production of ROS.

ACKNOWLEDGEMENT

It is an honor to thank people who have made this dream come true. Through this opportunity, I would like to express sincere thanks to my advisor Prof. Dr. Norlida Kamarulzaman for her guidance, suggestions, and support to finish this work. Her motivational words are worth remembering for me. To Madam Mazatulikhma Mat Zain, my heartfelt appreciation for the ideas, criticisms and patience you showed me throughout this course of study, and for the knowledge and expertise you shared with all your students. Most importantly, special thanks to my family, particularly my parents Hasbullah Mahmud and Siti Rahani Yahya for their love and support during my hard time in completing this thesis.

I would like to extend my gratitude to my lab mates, Sharil Sarman, Nordalilah Rosni, Mohd Hafiz Rothi, Mohd Syafiq Mohd Ridhuan, Nurulhuda Ismail, Ma'ati El-Moghoul and all members of tissue culture lab, for their unfailing support and valuable time to discuss and share knowledge in cell culture.

Lastly, I would like to thank my beloved friends, Muhamad Faidzal Razali, Norfaezah Mazalan, Siti Hajar Noor Alshurdin and Zainuddin Rossdi, who laughed and cried with me throughout my best and worst times of lab work.

After all, I would never have completed this research project without the financial support by FRGS fund (600-RMI/ST/FRGS 5/3/Fst (11/2008) and internal funds from Institute of Science, Universiti Teknologi Mara (UiTM), Shah Alam.

TABLE OF CONTENTS

		Page	
CO	NFIRMATION BY PANEL OF EXAMINERS	ii	
AU	THOR'S DECLARATION	iii	
AB	STRACT	iv	
AC	KNOWLEDGEMENTS	v	
TA	BLE OF CONTENTS	vi	
LIS	LIST OF TABLES		
LIS	ST OF FIGURES	x	
LIS	ST OF ABBREVIATIONS	xii	
СН	APTER ONE: INTRODUCTION	1	
1.1	Background of study	1	
1.2	Problem statement	4	
1.3	Significance of study	4	
1.4	Objectives of study	4	
1.5	Scope and limitation of the study	5	
СН	APTER TWO: LITERATURE REVIEW	6	
	An overview of nanotechnology and nanomaterials	6	
	Possible route of entry and potential harmful effects of nanoparticles on	7	
	biological system		
	2.2.1 Inhalation of nanoparticles	8	
	2.2.2 Ingestion of nanoparticles	9	
	2.2.3 Dermal exposure of nanoparticles	10	
2.3	Toxicity status of magnesium oxide nanoparticles	11	

2.4 Factors influencing toxicity of nanoparticles
2.4.1 Particle size
2.4.2 Shape
2.4.3 Dosage

2.4.4 Type of cells

13

13

14

14

15