ALPHA PARTICLES DEPOSITION AND ITS EFFECTS ON LUNGS, MALE INFERTILITY AND BLOOD COMPONENTS

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by

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This effort dedicated

То

Soul of the messenger of Allah "Muhammad –Allah peace upon him" Soul of my father "Haj Hamid Ismail-may Allah rest his soul in peace "

My lovely mother, brothers and sisters

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"All praises and thanks to ALLAH"

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ACRONYMS AND ABBREVIATIONS

Alveoli	Air sacs of the lungs
Amyloid	Is a substance which can be found in all tissue pathology
BALF	Broncho-Alveolar Lavage Fluid
BEIR	Biological Effects of Ionizing Radiation
Breathing	Process of inhaling and exhaling air
Bronchi	Largest branch of the bronchial tree
	between the trachea and bronchioles.
Bronchial tree	Entire system of air passageways within the lungs formed by the branching of bronchial tubes.
Bronchioles	Smallest of the air passageways within the lungs.
C	Celsius, centigrade
C.S.	cross section (cut perpendicular to the axis, or across the structure)
c.t.	connective tissue
CBC	Complete blood count
CBRN	chemical, biological, radiological, nuclear
e.	Epithelium
EPA	Environmental Protection Agency (United States)
Epiglottis	Flaplike piece of tissue at the top of the larynx that covers its opening when swallowing is occurring.
Exhalation	Also known as expiration, the movement of air out of the lungs.

GM	Geiger-Mueller
Gy	Gray
H&E stain, HE stain or hematoxylin and eosin stain,	Popular staining method in histology. It is the most widely used stain in medical diagnosis; for example when a pathologist looks at a biopsy of a suspected cancer, the histological section is likely to be stained with H&E and termed H&E section, H+E section, or HE section.
HAZMAT	Hazardous materials
Hemoglobin	Iron-containing protein pigment in red blood cells that can combine with oxygen and carbon dioxide.
HPA	Health Protection Agency
IAEA	International Atomic Energy Agency
ICRP	International Commission on Radiological Protection
Inhalation	known as inspiration, the movement of air into the lungs.
keV	kilo electron volts
kg	Kilogram
km	Kilometer
l.s.	longitudinal section (cut parallel to the axis, or along the structure)
Larynx	Organ between the pharynx and trachea that contains the vocal cords.
Lungs	Paired breathing organs.
magnification	On most tissues and organs we used the same three magnifications that students use in lab: 40X (scanning objective lens), 100X (low power objective lens) and 400X (high power objective lens).

Nasal cavity	Air cavity in the skull through which air passes from the nostrils to the upper part of the pharynx.
NCRP	National Council on Radiation Protection & Measurements
NTDs	Nuclear Track Detectors
Pleura	Membrane sac covering and protecting each lung.
ppm	parts per million
Rad	Radiation absorbed dose
RBE	Relative biologic effectiveness
RED	Radiological exposure device
Rem	Roentgen Equivalent Man (dose equivalent)
RERF	Radiation Effects Research Foundation
Respiration	Exchange of gases (oxygen and carbon dioxide) between living cells and the environment.
SRIM	The stopping and range of ions in matter
Sv	Sievert
t.s.	transverse section (cut perpendicular to the axis or across the structure)
Trachea	Also known as the windpipe, the respiratory tube extending from the larynx to the bronchi. The nasal cavity is lined by mucous membrane containing microscopic hairlike structures called cilia.
UNSCEAR	United Nations Scientific Committee on the Effects of Atomic Radiation
w.m.	whole mount (the tissue was not cut before it was mounted on the slide)
WHO	World Health Organization

LIST OF SYMBOLS

A	Alpha particle
В	Beta
μ	Micro
Γ	Gamma ray
θc	Critical Angle
ρ	Density
η	Efficiency of the CR-39 NTDs
V _B	Velocity of bulk etch rate
V _T	Velocity of track etch rate
K	Calibration factor
F	Equilibrium factor between radon and its daughter
RBC	Red Blood Cell
WBC	White Blood Cell
PLT	Blood Platelet
C _{Rn222}	Radon concentration
²²² Rn	Radon Gas
²²⁰ Th	Thoron Gas

TABLE OF CONTENTS

Acknowledgments	iv
Acronyms and Abbreviations	vi
List of Symbols	ix
Table of Contents	X
List of Tables	xvi
List of Figures	xvii
List of Appendices	xxiii
Abstrak	xxvi
Abstract	xxviii
CHAPTER 1- Introduction and an Overview	
1.1 Background and Overview	1
1.2 Terminology	3
1.3 Problem Statements and the Contributions	4
1.4 Objectives	5
1.5 Scope of Research	6
1.5.1 User Interface (UI) Elements	6
1.5.2 Locations of UI Elements	7
1.5.2 (a) Iraqi Kurdistan region	7
1.5.2 (b) Penang Island	7
1.6 Outline of Thesis	8
CHAPTER 2- Literature Review & Previous Research	
2.1 Introduction	9
2.2 Literature Review	11
2.2.1 Alpha Particle and the Radon Gas	11

2.2.2 Radon Inhalation and its Effects on the Lungs	13
2.2.3 Occupational (Physical) Exposures and Male Infertility	16
2.2.4 Deposition of Alpha Particles and the Blood Ionization	22
2.2.5 Nuclear Track Detectors (NTDs)	30
CHAPTER 3- Principle of Alpha Track Formation in Nuclear Tr Detectors	ack
3.1 Introduction	38
3.2 Characteristics of NTDs	38
3.3 Mechanism Formation of Latent Track	40
3.3.1 Defect Creation	40
3.3.2 Defect Relaxation	41
3.4 Models of Track Formation	42
3.5 Calculations for Track Parameters	45
3.5.1 Bulk and Track Etch Rates	47
3.5.1(a) Method of Calculating Bulk Etch Rate (μ m/h)	48
3.5.1(b) Method of Calculating Track Etch Rate (μ m/h)	49
3.5.2 Critical Angle and the Detector Efficiency	50
3.5.2(a) Critical angle	50
3.5.2(b) Detector efficiency (η)	51
3.6 Determining the Calibration Factor of Radon and its Progeny	52
3.7 Design of the measuring devices	54
3.8 Simulation of Radon Measurements with NTD in Diffusion Chamber	57
CHAPTER 4- Mechanism of Interaction of Alpha Particles with	Fissues
4.1 Introduction	59
4.2 Free Radicals	60
4.3 Ionization of Alpha Particles	62

4.4 Possibility of the Effects of Radiation on Cells	65
4.5 Interaction of Alpha Particles on Human Blood	66
4.6 Effects of Radon Inhalation Dose on Lung Damage (Cumulative exposure)	68
4.7 Radon and Genomic Damage (Cause of Infertility)	72
CHAPTER 5- Fabrication and Calibration Process of Alpha Radiation Collimator and Radon Dosimeter	
5.1 Introduction	78
5.2 Materials and the Equipments	78
5.2.1 Materials	78
5.2.1(a) CR-39 NTDs	78
5.2.1(b) Pen tubes	79
5.2.1(c) Radium (²²⁶ Ra)	79
5.2.1(d) Other materials	80
5.2.2 Equipments	81
5.2.2 (a) Water bath	81
5.2.2 (b) Radiation dosimeters	82
5.2.2 (c) Radon monitor model 1027	83
5.2.2 (d) Scanning System of Tracks of Alpha Particles	83
5.3 Methodology	84
5.3.1 Calibration Process of CR-39 NTDs	84
5.3.1(a) Optimum time of etching	84
5.3.1(b) Evaluation of the efficiency of the CR-39 NTDs	86
5.3.2 Design and Constructs of Alpha Irradiation Collimators	88
5.3.3 Fabricate of an Optimum Radon Dosimeter.	91
5.4 Results and Discussion	94
5.5 Summary	101

Human Dangs and Mate Interently	
6.1 Introduction	102
6.2 Materials and the Equipments	103
6.2.1 Materials	103
6.2.1(a) An electronic balance	103
6.2.1(b) Tools to collect soil samples	103
6.2.1(c) An Electric oven (Humidity chamber HCP)	104
6.2.2 Equipments	105
6.3 Area under Study	105
6.4 Methodology	107
6.4.1 Indoor radon	107
6.4.1(a) Experimental procedures (Distribute and exposure of the dosimeters)	107
6.4.1(b) Measurements	111
6.4.2 Radon in the Soil Samples	114
6.4.2(a) Experimental procedures (Collection and exposing of the soil samples)	114
6.4.2(b) Measurements	116
6.4.3 Radon in Drinking Water	119
6.4.3(a) Experimental procedure	119
6.4.3(b) Measurements	122
6.5 Results and Discussion	126
CULADTED 7 Disk Evolution of Aluba Doutiels Doute the state	T

CHAPTER 6- Evaluation of the Risks of Deposition of Alpha Particles on Human Lungs and Male Infertility

CHAPTER 7- Risk Evaluation of Alpha Particle Deposition on the Human Blood Samples: *In vitro*

7.1 Introduction	145
7.2 Materials and the Equipments	146

7.2.1 Materials	146
7.2.1(a) Human blood samples	147
7.2.1(d) TPP tissue culture test plate	148
7.2.1(e) Device of a blood analysis	149
7.3 Research Methodology	149
7.3.1 Fabrication of an alpha irradiation technique and procedures of human blood irradiation	149
7.3.1(a) Procedures of the fabrication of alpha irradiation technique using ²²⁶ Ra	150
7.3.1(b) Experimental procedures to irradiate a human blood samples	151
7.3.2 Fabrication of a radon exposure technique and procedures of deposition of radon's progenies on the surface of human blood	153
7.4 Results and Discussion	158
CHAPTER 8- Preliminary Study on the Effects of Radon Inhalation Lungs, Trachea and Testes of the Male Rabbits	1 on the
8.1 Introduction	174
8.1 Introduction8.2 Materials and the Equipments	174 175
8.1 Introduction8.2 Materials and the Equipments8.2.1 Materials	174 175 175
 8.1 Introduction 8.2 Materials and the Equipments 8.2.1 Materials 8.2.2 Equipments 	174 175 175 178
 8.1 Introduction 8.2 Materials and the Equipments 8.2.1 Materials 8.2.2 Equipments 8.3 Research Methodology 	174 175 175 178 178
 8.1 Introduction 8.2 Materials and the Equipments 8.2.1 Materials 8.2.2 Equipments 8.3 Research Methodology 8.3.1 Install and Fabrication a Radon Exposure Technique 	174 175 175 178 178 178
 8.1 Introduction 8.2 Materials and the Equipments 8.2.1 Materials 8.2.2 Equipments 8.3 Research Methodology 8.3.1 Install and Fabrication a Radon Exposure Technique 8.3.2 Procedures of the Exposure 	174 175 175 178 178 178 178
 8.1 Introduction 8.2 Materials and the Equipments 8.2.1 Materials 8.2.2 Equipments 8.3 Research Methodology 8.3.1 Install and Fabrication a Radon Exposure Technique 8.3.2 Procedures of the Exposure 8.3.3 Histological techniques 	174 175 175 178 178 178 178 179 181
 8.1 Introduction 8.2 Materials and the Equipments 8.2.1 Materials 8.2.2 Equipments 8.3 Research Methodology 8.3.1 Install and Fabrication a Radon Exposure Technique 8.3.2 Procedures of the Exposure 8.3.3 Histological techniques 8.3.3 (a) Anaesthetization process of the rabbits 	174 175 175 178 178 178 178 179 181 181
 8.1 Introduction 8.2 Materials and the Equipments 8.2.1 Materials 8.2.2 Equipments 8.3 Research Methodology 8.3.1 Install and Fabrication a Radon Exposure Technique 8.3.2 Procedures of the Exposure 8.3.3 Histological techniques 8.3.3 (a) Anaesthetization process of the rabbits 8.3.3 (b) Removal of the organs 	174 175 175 178 178 178 179 181 181 181
 8.1 Introduction 8.2 Materials and the Equipments 8.2.1 Materials 8.2.2 Equipments 8.3 Research Methodology 8.3.1 Install and Fabrication a Radon Exposure Technique 8.3.2 Procedures of the Exposure 8.3.3 Histological techniques 8.3.3 (a) Anaesthetization process of the rabbits 8.3.3 (b) Removal of the organs 8.3.3 (c) Fixation process (24 hours) 	174 175 175 178 178 178 178 179 181 181 181 182 183

8.3.3(e) Clearing process	185
8.3.3(f) Impregnation (infiltration) process	186
8.3.3(g) Procedure of blocking in wax (Embedding)	187
8.3.3(h) Trimming process	188
8.3.3(i) Sectioning process	188
8.3.3(j) Dehydration and the staining procedures of the slides	189
8.3.3(k) Mounting process	190
8.3.4 Imaging Process of the Tissue Slides	191
8.4 Result and Discussion	191
CHAPTER 9- Conclusions and Future Works	
9.1 Introduction	205
9.2 Conclusions	205
9.3 Suggestions for Future works	207
9.4 Recommendations	208
References	210
Appendix A	227
Appendix B	239
Appendix C	240
LIST OF PUBLICATIONS	243

LIST OF TABLES

Page

Table 5.1	The comparison between present results and the theoretical results of the other references	101
Table 5.2	Optimum parameters for the irradiation collimator and the radon dosimeter	101
Table 6.1	Number of poor, partial, and good dwellings for each location. For each location, there are 4 dwelling. The definition of poor, partial and	109
Table 6.2	Summary values of the used parameters in this section and main formulas	113
Table 6.3	Summary values of the used parameters in this section and main formulas	119
Table 6.4	Summary values of the used parameters in this section and main formulas	126
Table 6.5	Data analysis based on the type of ventilation	129
Table 6.6	Results of indoor radon concentration and the risk factor inside dwellings in selected locations in Iraqi Kurdistan.	134
Table 6.7	Average radon concentration, radon exhalation rate and the effective radium content in soil samples using CR-39 NTDs and RAD 7.	136
Table 6.8	Radon concentration in drinking water and the equilibrium factors with the source of drinking water.	140
Table 6.9	Results of the radon concentration and it's contribute indoor radon concentration using CR-39 NTDs in drinking water, and its dose equivalent to the bronchial epithelium	143
Table 6.10	Results of the annual dose equivalent to the stomach and the whole body, with the life time cancer risks for stomach and whole body.	144
Table 7.1	More details about the Test plate version F-face and 96 wells.	149
Table 7.2	Results of the analysis of the human blood sample without irradiation (background of indoor radon = 210 Bq/m^3) for 20 minutes	161
Table 8.1	Change of Bouin's in each stage of the running	184

Table 8.2	Change of alcohol concentration in each stage of the running	185
Table 8.3	Staining for animal sample	186
Table 8.4	Types of solutions and the specific periods for dehydrating the slides	189

LIST OF FIGURES

		Page
Figure 3.1	Formation of etch-able track	40
Figure 3.2	Axially sections through a particle track in a polymer (Durrani <i>et al.</i> 1975).	42
Figure 3.3	Radial section-chain breaks allow preferential etching at a lower damage density (Ilic <i>et al.</i> 1990, Durrani <i>et al.</i> 1975).	42
Figure 3.4	The ion-explosion spike mechanism for track formation in inorganic solids (Fleischer <i>et al.</i> 1965, 1975)	43
Figure 3.5	Track constructions for normal incident particle at constant V_T (Nikezic and Yu 2004)	46
Figure 3.6	Geometrical forms for the latent tracks dependence on the incident angle (a) $\theta < \delta$ (b) $\theta = \delta$ and (c) $\theta > \delta$ (Illic & Durrani 2004, Nikezic et al. 2004).	51
Figure 3.7	Open (bare) detector samples: different sizes and shapes(Nikolaev and Ilic 1999).	55
Figure 3.8	Open-chamber samples: different sizes and shapes (Nikolaev and Ilic 1999).	55
Figure 3.9	Closed diffusion chambers for the radon gas; (a) radiometer with an inlet filter, (b) radiometer with improved parameters, (c) Multipurpose parameters, (d) combination of etch track detector and electrostatic field, (e) etch track detector and activated charcoal (Nikolaev and Ilic, 1999).	57
Figure 4.1	Schema of creation free radicals of oxygen atom.	60
Figure 4.2	In-direct and direct route of radiation on DNA.	61
Figure 4.3	Helium atom becoming helium ion, as a result of loss of an electron.	62
Figure 4.4	Process of producing positive and negative ions due to alpha irradiation.	63
Figure 4.5	Ratio of component of human blood: mixed and unmixed	67
Figure 4.6	Biomicroscope shapes of the blood parameter: (a) red blood cells, (b) white blood cells, and (c) platelet (IAEA 1997).	67
Figure 4.7	Normal red blood cells move easily through blood vessels, taking oxygen to every part of your body	68

Figure 4.8	Respiratory structures in the human body (WHO 2009).	69
Figure 4.9	Alpha-particle tracks in the tissue;	71
	•: Ionized molecule & o: Normal molecule	
Figure 4.10	Reactive Oxygen Speckles and Cellular Damage	77
Figure 5.1	Laser-engraved code of the CR-39 NTDs; Intercast Europe SRL.	80
Figure 5.2	Samples of the stainless steel and plastic pen tubes used at an irradiation collimator.	80
Figure 5.3	Decay schema of Uranium (238 U) to Radon (222 Rn) gas (Guilmette <i>et al</i> . 1991).	81
Figure 5.4	Shape of the source of Radium (²²⁶ Ra).	81
Figure 5.5	Some of the equipments and materials used in the selected work; A) PVC tub, B) Past solution to contact of the materials, C) Soft paper, D) Digital balance to measure the mass of NaOH, E) Sensitive balance, F) Air pump (push and pull), G) Wire connections, H) Small electric fan, I) Pieces of sponge used as a barrier, and J) Plastic dispenser (container)	82
Figure 5.6	Water bath "GOTECH TESTING MACHINES INC.	82
Figure 5.7	Radiation dosimeters: type RAMS DA3-2000 (A) and VICTOREEN (B).	83
Figure 5.8	Professional continuous radon monitor version 1027	84
Figure 5.9	An optical microscope equipped with CCD camera, connected to a PC-based image analyzer	85
Figure 5.10	Samples of the fixable tubes to store NaOH solution	86
Figure 5.11	Method of irradiation of CR-39 NTDs (2π geometry)	86
Figure 5.12	Process of the chemical etching of the CR-39 NTDs.	87
Figure 5.13	Window for the SRIM-2000 program: ion stopping and range of the incident of alpha particles inside CR-39 NTDs.	89
Figure 5.14	Variation of the radiation dose with the material type of the alpha irradiation collimators for different energy of alpha particles.	90
Figure 5.15	Irradiation collimators for alpha particles (informal incident	90

particles=90°)

Figure 5.16	Sample of the PVC chamber used as a radon dosimeters	92
Figure 5.17	Sample of the PVC chamber whose radius was reduced by the paper	92
Figure 5.18	Two-PVC diffusion radon chambers; with and without a piece of sponge	92
Figure 5.19	Cubic Perspex chamber; A) Mechanism of attached radon dosimeter and B) mechanism of recode radon concentration using radon monitor.	92
Figure 5.20	AR1 is the area of the scanning (=23466.483 $\mu m^2)$ for alpha track registration.	93
Figure 5.21	Track registration from different densities for different times of exposure	94
Figure 5.22	Change of track density on the surface of CR-39 NTDs with the etching time.	95
Figure 5.23	Running of the TRACK_TEST program: Graphic plot of a normal incident alpha particle (5.49 MeV).	95
Figure 5.24	Energy loss of alpha particle with electrons and nucleus of CR-39 NTDs. In addition, the range of the alpha particle in the CR-39 NTDs depends on its energy.	96
Figure 5.25	Linear relationship between residual energy and the source-detector distance	97
Figure 5.26	Images of the alpha tracks for the energies 1, 2, 3 and 4 MeV(Normal incident =90 ^{0}).	98
Figure 5.27	Experimental response of radon as a function of radius for different heights	98
Figure 5.28	Variation of the calibration factor with the dimension of the detection system on the base of the volume	99
Figure 5.29	Photo of the optimum radon dosimeter ($r=3$ cm; $h=7$ cm).	99
Figure 5.30	Exponential relationship of K (cm) with the efficiency of CR-39 NTDs for optimum radon dosimeter.	99
Figure 6.1	Wire mesh garden, soil ogre and electronic balance	102
Figure 6.2	Drying processes of the soil samples inside the electric oven	103

Figure 6.3	Radon detector version 7 (RAD 7)	103
Figure 6.4	Sketch map of the Iraq (A) and Iraqi Kurdistan region (B): Red spots represent locations of study.	104
Figure 6.5	Exponential relationships between the rate of men infertility and the location under study	106
Figure 6.6	S1, S2, S3, &S4 are the locations of the dwellings, the angle between (S2, S3, S4) them \approx 120 °, and we get average indoor radon in each location.	108
Figure 6.7	Schematic diagram of the optimum radon dosimeter	109
Figure 6.8	Long-term measurements' detection techniques; Two radon dosimeters; with and without filter.	112
Figure 6.9	Soil sample under exposure using long-term measurements technique	112
Figure 6.10	Short -term measurement detection techniques to evaluate a radon exhalation rate of soil samples, using RAD7.	112
Figure 6.11	Cylindrical volume tubes (scale tubes)	114
Figure 6.12	Plastic bottle to keep the water samples	117
Figure 6.13	The apparatus used to study the radon dosimeters for drinking water; (a) without a barrier, (b) with a barrier for the passive detection technique.	117
Figure 6.14	An active detection technique of airborne radon gas in drinking water.	118
Figure 6.15	Average indoor radon concentration in Iraqi Kurdistan; histogram of frequency distribution, lognormal and percentages.	122
Figure 6.16	Variation of the average radon concentration for the locations under study	123
Figure 6.17	Correlation between men infertility, annual effective dose and the locations under study.	124
Figure 6.18	Defects variation in sperm concentration with locations under study: Online version for the red colures is high and low values	125
Figure 6.19	Abnormal sperm shapes found at the locations under study: Online version for the red colures is in high values	125
Figure 6.20	Variation in sperm activity with locations under study: Online version	126

for the red colures is in low values

- Figure 6.21 Exponential relation between annual effective dose (mSv/y) and the 127 rate of men infertility
- Figure 6.22 Correlation ship between ventilation rate, average of an indoor radon 129 concentration and average ratio of men infertility.
- Figure 6.23 Un-uniform distribution of soil radon concentration in the selected 130 locations using CR-39 NTDs.
- Figure 6.24 Un-uniform distribution of soil radon concentration in the selected 130 locations using RAD7.
- Figure 6.25 Un-uniform distribution of radium content in soil samples using CR- 131 39 NTDs detection method.
- Figure 6.26 Linear relationship between radon concentration, radon exhalation 132 rate, and ratio of effective radium in the soil sample.
- Figure 6.27 A relation between the ratio (=Soil radon concentration ($C_{Soil Rn222}$) / 133 Indoor radon concentration ($C_{indoor Rn222}$)) and the effective radium content in soil samples, using CR-39 NTDs.
- Figure 6.28A Un-uniform distribution of radon concentration emanation from 135 drinking water for selected locations, using active detecting methods.
- Figure 6.28B Un-uniform distribution of radon concentration emanation from 135 drinking water for selected locations, using passive detecting methods.
- Figure 6.29 Correlation between active and passive detecting technology for the 135 values of average radon concentration
- Figure 6.30 Annual doses equivalent to the bronchial epithelium and the radon 136 concentration in drinking water
- Figure 7.1 A requirement of materials, tools to fabricate, and exposure 141 techniques process.
- Figure 7.2Plastic test plate forms142
- Figure 7.3 Sysmex, K-Series range KX-21 equipped with a computer to keep 143 patient information
- Figure 7.4 Blood irradiations, exposure, withdraw dropper, and other tools of the 144 blood kept inside an electric oven.
- Figure 7.5 Process of evaluating radiation dose rate 145

Figure 7.6	Irradiation of human blood samples by ²²⁶ Ra.	146
Figure 7.7	Windows of SRIM-2008 program shows ion stopping and range tables of the incident of alpha particles into the human blood.	147
Figure 7.8	Alpha track registration for 226 Rn (dose=40±1.6 µSv/h) per 4 minutes ; (a) Magnification=50X, (b) Magnification=1000X	147
Figure 7.9	Tools and stages of preparing a chamber of radon gas	148
Figure 7.10	Human blood exposure to radon gas.	149
Figure 7.11	Installed of the radon exposure technique	150
Figure 7.12	Schema diagram of human blood exposed <i>in vitro</i> to radon gas (irradiation technique).	150
Figure 7.13	Alpha track register by CR-39 NTDs (Area 0.101912 mm ²)	151
Figure 7.14	Reverse changes for each of the range and energy loss of alpha particle inside the CR-39 NTDs.	152
Figure 7.15	Reverse changes for each of the range and energy loss of alpha particle inside the CR-39 NTDs.	153
Figure 7.16	Comparative study of alpha particle energy, radiation dose and alpha particle registration on CR-39 NTDs.	154
Figure 7.17	Relatively change in the main blood component with the different time of irradiation (1and 2 indications to before and after irradiation for each parameter).	156
Figure 7.18	Contour plot of radiation doses from the radium source vs ratio of PLT and ratio of WBC	156
Figure 7.19	Contour plot of radiation doses from the radium source vs ratio of PLT and ratio of RBC	156
Figure 7.20	Relative change in the main blood components (before and after irradiation) with a different radiation dose	157
Figure 7.21	Main effects plot for the ratio (after/ before irradiation) of PLT with the age and gender of the samples.	158
Figure 7.22	Main effects plot for the ratio (after/ before irradiation) of WBC with the age and gender of the samples.	159
Figure 7.23	Main effects plot for the ratio (after/ before irradiation) of RBC with the age and gender of the samples.	159

Figure 7.24	Percentage decreases of radon concentration with the stages of exposure during exposure.	160
Figure 7.25	Average loss ratio of radon concentration depends on time of exposure	160
Figure 7.26	Changes of radon concentration with the time of exposure	161
Figure 7.27	Rate and percentage occupation of absorption dose into blood samples for various exposure times	162
Figure 7.28	Increasing the deposition of alpha particles with the time of exposure	162
Figure 7.29	Relative changes in reduced of platelet count (PLT) with rate of alpha particle deposition on the blood samples.	163
Figure 7.30	Average absorbed dose increased at increasing time of exposure	164
Figure 7.31	Most change (reduced) of PLT is at 20minut exposure of radon gas	165
Figure 7.32	Increase of the RBC and PLT with the time of exposure to radon gas	166
Figure 8.1	Electric fan, stands, and clipper	169
Figure 8.2	Part of the Perspex box, which is used to an exposure technique of radon gas.	169
Figure 8.3	Food, drinking water and the bed of the Rabbits	169
Figure 8.4	Thermo scientific used to cover slipping	169
Figure 8.5	Some of the tools have been used within histological process of the Rabbits	170
Figure 8.6	Radon exposure techniques	171
Figure 8.7	Place of the radon exposure technique; Rabbits are checked by the researcher and put it inside the technique to exposure, location is inside the underground lab of Biophysics lab, USM.	173
Figure 8.8	Radon exposure technique and the animals are ready for exposure process expose.	173
Figure 8.9	Anaesthetize tools; A) is the box of exposure rabbits and B) is the information board of the anaesthetize matter (Chloroform)	174
Figure 8.10	Process of the removable of the organs	175
Figure 8.11	Specimen dispenser included dumped organs	175

Figure 8.12	Shows specimen tubes of the slide pieces of the organs.	176
Figure 8.13	Specimen bottles containing suitable amount of Alcohol concentrations	177
Figure 8.14	Xylene is a clearing agent	178
Figure 8.15	Tools of impregnation process	179
Figure 8.16	Cold plate for the equipment of Embedding.	179
Figure 8.17	Place of the filled mould on the ice plate	180
Figure 8.18	Shape of blocking in wax	180
Figure 8.19	Microtome section; Microtome blades, holders, and supplies.	181
Figure 8.20	A) Water bath, alcohol and dropper used to expending of the slid tissue, and B) is a device of the Thermostat.	181
Figure 8.21	Stages of the infiltration process for the rabbit's organs.	182
Figure 8.22	Equipments for keeping the slides A: wood dispenser slide, B) fracture for keep the slides for 1 day	183
Figure 8.23	Scanning and image process for the tissue slides	183
Figure 8.24	AR1= 23734.435 Micron, for 30 days	185
Figure 8.25	AR1= 11367.3 Micron , for 60 days	185
Figure 8.26	AR1= 23884.022 Micron, for 90 days	185
Figure 8.27	Microscopic view of a histological specimen of rabbit lung (left: L, and right: R) tissue stained rabbits for the magnification of 5X.	187
Figure 8.28	Microscopic view of a histological specimen of rabbit lung (left: L, and right: R) tissue stained rabbits for the magnification of 10X.	188
Figure 8.29	Microscopic view of a histological specimen of rabbit lung (left: L, and right: R) tissue stained rabbits for the magnification of 20X.	189
Figure 8.30	Microscopic view of a histological specimen of rabbit trachea tissue stained rabbits for 30 days of exposure and different magnifications 5, 10, & 20X.	190
Figure 8.31	Microscopic view of a histological specimen of rabbit trachea tissue stained rabbits for 60 days of exposure and different magnifications 5, 10, & 20X.	191

- Figure 8.32 Microscopic view of a histological specimen of rabbit trachea tissue 192 stained rabbits for 90 days of exposure and different magnifications 5, 10, & 20X.
- Figure 8.33 Microscopic view of a histological specimen of rabbit tests tissue 194 stained rabbits at magnifications 5X.
- Figure 8.34 Microscopic view of a histological specimen of rabbit tests tissue 195 stained rabbits at magnifications 10X.
- Figure 8.35 Microscopic view of a histological specimen of rabbit tests tissue 196 stained rabbits at magnifications 20X.

PENGENDAPAN ZARAH ALFA DAN KESANNYA KE ATAS PARU-PARU, KETIDAKSUBURAN LELAKI DAN KOMPONEN DARAH

ABSTRAK

Pengendapan zarah alfa pada paru-paru, ketidaksuburan laki-laki dan komponen darah manusia (platelet, PLT; sel darah putih, WBC dan sel darah merah, RBC) telah dilakukan dengan menggunakan teknik pendedahan baru CR-39 NTDs. Penyelidikan ini meliputi lima bahagian utama. Pada bahagian pertama, proses penentukuran CR-39 NTD telah dilakukan. Didapati bahawa kecekapan dan masa optimum etsa CR-39 NTD adalah $80.3 \pm 1.23\%$ dan 9 jam, masing-masing.

Pada bahagian kedua, bergantung kepada faktor penentukuran gas radon, dosimeter radon optimum telah direkabentuk. Dimensi optimum dosimeter radon adalah 7 cm panjang dan 6 cm diameter pada faktor penentukuran 2.68 ± 0.03 cm.

Pada bahagian ketiga, kepekatan radon dalam udara, tanah dan air minum telah dinilai di 124 kediaman yang mana ketaksuburan adalah ketara di 31 lokasi berbeza di Kurdistan Iraq, dengan menggunakan pengesan pasif (pengesan plastik CR-39) dan aktif (RAD7). Suatu hubungan ditemui antara kepekatan radon dalam bilik, kadar pengudaraan, dan kadar ketaksuburan lelaki. Kadar tinggi kepekatan radon dalam bilik dan ketaksuburan lelaki adalah realtifnya berkorelasi dengan pergularan rendah atau buruk. Selanjutnya, kadar eskhalasi dari tanah dan air minum menyumbang kepada kadar peningkatan radon dalam bilik di sebahagian besar lokasi. Menurut faktor risiko anggaran, radon disebabkan oleh risiko kanser paru-paru untuk kediaman di beberapa lokasi terpilih adalah berubah dari 43.17 hingga 108.79 dengan purata lebih kurang 65.22 ± 20.93 per juta orang. Kadar kepekatan radon mempunyai nilai rendah (139.11 \pm 17.26 Bq/m³) untuk pengudaraan yang baik (13.7%), dan mempunyai nilai tinggi (189.78 \pm 55.91 Bq/m³) untuk pengudaraan yang buruk (56.45%). Oleh kerana itu,

pengendapan zarah alfa, yang dipancarkan daripada progeni radon, didapati meningkatkan kanser paru-paru dan ketaksuburan lelaki.

Pada bahagian keempat, pengendapan zarah alfa pada sampel darah manusia telah diukur dengan menfabrikasikan teknik penyinaran dan pendedahan. Salah satunya adalah pengkolimat penyinaran alfa dari sumber radium, digunakan untuk mengukur penyinaran dalam sampel darah manusia dengan tenaga berbeza zarah alfa. Kaedah pengagihan zarah alfa pada permukaan CR-39 NTDs telah digunakan untuk menganggarkan ketumpatan zarah alfa yang terkumpul pada permukaan sampel darah. PLT, RBC dan WBC adalah didapati agak terjejas. Namun, yang paling terjejas adalah bilangan PLT, yang menurun dengan peningkatan dos sinaran zarah alfa. Teknik kedua adalah teknik pendedahan gas radon bagi mendedahkan sampel darah manusia. Kajian perbandingan CR-39 NTD dan sampel darah manusia adalah suatu teknik baru untuk kajian *in vitro* pengionan darah. Dalam teknik pendedahan ini, kepekatan radon dikurangkan menjadi sekitar 4.9 %, dan dengan demikian, sekitar 95% dari kepekatan radon diselamatkan. Nisbah pengendapan zarah alpha didapati mencukupi untuk mengubah bilangan PLT bagi kedua-dua lelaki dan perempuan.

Dalam bahagian akhir, suatu teknik pendedahan gas radon (553.20 \pm 26.87 Bq/m³) telah direkabentuk bagi mengkaji kesan *in vivo* penyedutan radon ke atas paru-paru, trakea, dan testis arnab jantan. Pertumbuhan sel abnormal tidak kelihatan dalam paru-paru dalam masa 30 hari pendedahan tetapi muncul selepas 60 hari. Selanjutnya, didapati penyedutan gas radon purata (553.20 \pm 26.87 Bq/m³) tidak menjejaskan trakea dalam masa 30 dan 60 hari. Namun, kesannya bermula selepas 90 hari pendedahan. Di sisi lain, penyedutan kepekatan radon purata tidak menjejaskan testis dalam masa 30, 60, dan 90 hari.

ALPHA PARTICLES DEPOSITION AND ITS EFFECTS ON LUNGS, MALE INFERTILITY AND BLOOD COMPONENTS

ABSTRACT

Alpha particles deposition and its effects on lungs, male infertility and blood components (platelets, PLT; white blood cells, WBC; and red blood cells, RBC) have been performed by using new exposure techniques of CR-39 Nuclear Track Detectors (NTDs). The present research includes five main parts. In the first part, the calibration process of the CR-39 NTDs has been carried out. It was found that the efficiency and the optimum time of etching of CR-39 NTDs are 80.3 ± 1.23 % and 9 h, respectively.

In the second part, depending on the calibration factor of the radon gas, optimum radon dosimeter has been fabricated. Optimum dimensions of the radon dosimeter were 7 cm length and 6 cm diameter at the calibration factor of 2.68 ± 0.03 cm.

In the third part, the concentration of radon in air, soil and drinking water has been evaluated in 124 dwellings where infertility was prevalent at 31 different locations in Iraqi Kurdistan, using passive (CR-39 plastic detector) and active (RAD 7) detectors. A relationship was found between indoor radon concentration, ventilation rate, and rate of male infertility. High rate of indoor radon concentration and male infertility was relatively correlated with low or bad ventilation. Furthermore, radon exhalation rate from the soil and drinking water contributed to increased rate of indoor radon in most of the locations. According to the estimation risks factor, the radon-induced lung cancer risks for dwellings in selected locations varies from 43.17 to 108.79 with the average of about 65.22 + 20.93 per million person. Radon concentration rate has low value (139.11 \pm 17.26 Bq/m³) for good ventilation (13.7%), and has high value (189.78 \pm 55.91 Bq/m³) for poor ventilation (56.45%). Therefore, deposition of the alpha particles, which are

emitted from radon's progenies, has been found to increase of lung cancer and male infertility.

In the fourth part, deposition of the alpha particles on the human blood samples has been estimated by fabricating irradiation and exposure techniques. One was the alpha irradiation collimator of the radium source, used to measure the irradiation in human blood samples with different energies of alpha particles. The method of distribution of alpha particles on the surface of CR-39 NTDs was used to estimate the density of alpha particles that accumulate on the surface of the blood samples. PLT, RBC and WBC counts were found to be relatively affected. However, the most was the PLT count, which decreased with increasing radiation dose of alpha particles. The second technique was the exposure technique of radon gas to expose human blood samples. Comparative study of CR-39 NTDs and the human blood samples is a new technique for *in vitro* studies of ionization of blood. In the present exposure technique, the radon concentration was reduced to around 4.9 %, and thus, around 95% of the radon concentration was saved. A ratio of the deposition of alpha particles was found to be adequate to change the PLT count in both males and females.

In the final part, an exposure technique of radon gas $(553.20 \pm 26.87 \text{ Bq/m}^3)$ has been fabricated to study *in vivo* effects of radon inhalation on the lungs, trachea and testes of the male rabbits. Abnormal growth of cells was not observed in the lungs within 30 days of exposure but it appeared after 60 days. Furthermore, it was found that inhalation of the average radon gas $(553.20 \pm 26.87 \text{ Bq/m}^3)$ did not affect the trachea within 30 and 60 days. However, the effects started after 90 days of exposure. On the other hand, inhalation of average radon concentration did not affect the testes within 30, 60, and 90 days.

CHAPTER 1

Introduction and an Overview

1.2 Background and Overview

Development of the techniques of irradiation and exposures to nuclear radiation in the field of medical physics started from the discovery of radioactivity. Ionizing radiation is a portion of the electromagnetic spectrum with sufficient energy to pass through matter and physically dislodge orbital electrons to form ions. These ions, in turn, can produce biological changes when introduced into the tissues, and can exist in two forms, i.e., electromagnetic wave, such as an X-ray or gamma ray, or as a particle, in the form of an alpha or beta particle, neutron, or proton (ATSDR 2005). Different forms of ionizing radiation have various abilities to generate biological damages (Hada *et al.* 2011).

Natural sources inherent to life on earth are considered to be major source of human exposure to ionizing radiation. Radon gas, gamma rays, cosmic (natural sources) radiations, and internal radiations constitute 2.4 mSv/y of the absorbed radiation dose. In addition, artificial and other sources contribute to 2.8 mSv/y (ICRP 1984, Shankarnarayanan 1998) of the absorbed radiation dose. People may be exposed to external and internal radiations by inhalation and ingestion due to background radiations that exist in the environment. Radon exposure occupies 50% of the average annual dose contribution of population radiation exposure; thus, most of the risks are from the inhalation of radon gas (Mehta 2005, Somlai *et al.* 2009).

The most significant characteristic of the radon-222 gas is the four short-lived progeny products from polonium-218 (²¹⁸Po) to polonium-214 (²¹⁴Po), which, shortly

after their formation get attached themselves to aerosol particles. However, a small fraction of these particles remains in an unattached form, depending on the movement of the air mass, which in turn depends on the installed ventilation systems (Somlai *et al.* 2009). Deposition of the radon's progeny on the lungs can cause cancer, and may result in male sterility producing undirected effects, via aberration of the DNA (Shankarnarayanan 1998, Abo-Emagd *et al.* 2008, Rajamanickam 2010). The interaction mechanism of the alpha particles with the tissues has been explained in Chapter 4.

Designing and fabricating of an optimum dosimeter to detect and measure radon density in the air is attracting great interest. Furthermore, estimation of the contribution of radon in soil and water in increasing the indoor radon density is also gaining interest among researchers.

Deposition of alpha particles emitted from radon daughter particles may influence the reduction in the number of platelets in both the genders at different rates, depending on the energy of the alpha particles. Hematology studies in the field of radiation have played an active role in estimating the exposure to ionizing radiation, as it increases the number of chromosomal aberrations in human blood lymphocytes (IAEA 1997).

Nuclear track detectors (NTDs) are nuclear detectors commonly made from polyallyl diglycol carbonate (PADC), and the most common NTD material is CR-39 (Hepburn and Windle 1980, Talat and Elsayed 2010). The principle of detection using NTDs is as follows; heavily charged particles cause damage to the materials along their path due to the excitation and ionization of atoms with which they interact. This damaged region is very narrow (30-100 °A) around the particle trajectory, and consists of disordered, but continuous, damage trails in a higher energy state. These damaged

regions could be visualized in the form of "tracks," using special techniques either through direct electron microscopy at a very high magnification or by optical microscopes after selective chemical etching (Dorschel *et al.* 1999). However, their size is changeable, which makes them only as a practical system in *in vitro* studies of human blood samples.

In the present study, a new irradiation and exposure techniques has been developed to evaluate the risks of the deposition of alpha particles on human lungs and its risks on male infertility, through measurement of radon concentration in indoor air, soil samples, and drinking water in the Iraqi Kurdistan. An irradiation and exposure technique to evaluate the risks of alpha particle deposition on the surface of the human blood samples was fabricated, and its methodology and results have been explained in Chapter 7.

Furthermore, a new exposure technique has been fabricated to expose male rabbits to the inhalation of radon gas (*in vivo*) and carry out the histological study of the rabbits to investigate the risks of radon inhalation (deposition of the alpha particles onto the lungs, trachea, and testis). CR-39 NTDs have been used as an essential detector, and the methodology and the results have been explained in Chapter 8.

1.2 Terminology

The main terminologies used in this investigation are as follows:

- 1. Evaluation of an alpha particle density (track/cm² per time of exposure), emitted from the radium-226 source into the human blood samples and the CR-39 NTDs.
- 2. Etching parameters; Bulk etch rate (V_B), track etch rate (V_T), critical angle for track registration (θ c), detector efficiency (η), and the optimum time of etching.

- Range and restricted energy loss of alpha particles in CR-39 NTDs and human blood samples.
- 4. Efficiency of the CR-39 NTDs (%).
- 5. Calibration factor between radon and its daughter.
- 6. An optimum dimension of the radon dosimeters (cm).
- 7. Track density
- 8. Concentrations of the radon gas (Bq/m^3) .
- 9. Potential alpha particle concentration (mWL).
- 10. Annual effective dose (μ Sv /y).
- 11. Inhalation and ingestion doses (μ Sv).
- 12. Average annual dose equivalent to the bronchial epithelium, stomach, and whole body (μ Sv).
- 13. Concentrations of the blood parameters: White blood cell (WBC) count, red blood cell (RBC) count, and platelet (PLT) count.
- 14. The histological study of some organs of the male rabbits

1.3 Problem Statements and the Contributions

In this project, the problem statements comprise the following elements:

- 1. No relationships have been published between the equilibrium factor for radon and the efficiency of the NTDs. This study is the first to demonstrate this relationship.
- 2. Experimentally, the optimum dimension of the dosimeter of radon gas has not been improved. Therefore, some problems related to the environmental factors (humidity and storage of the detector inside the store or lab) affected track density. Thus,

depending on the calibration factor of the radon gas, a new dosimeter of the radon gas has been fabricated experimentally.

- 3. Relationships between radon concentration, ventilation rate, and male infertility had not established. In this study, this relationship has been established.
- 4. Risks of radon inhalation and the effects of the deposition of radon's progenies on lungs and trachea have not been authenticated histologically. This study presented the effect of inhalation of radon on lungs, trachea, and testes of male rabbits, histologically.
- 5. Old exposure techniques had problem in detecting blood exposure to radon gas (Hamza and Mohankumar 2009), because the loss of radon concentration could not be estimated and the density of the deposition of the alpha particles could not be measured. In addition, detection of blood irradiation sample by the normal incident alpha particles was impossible. Therefore, in the present work, two different techniques have been fabricated; one for normal irradiation of the alpha particles for human blood samples, and other for exposing human blood samples by the radon exposure technique.
- 6. Human blood has not been irradiated directly by different energies of alpha particles to determine its effects on human blood components. In this study, the theory of track registration by CR-39 NTDs has been employed to detect and measure alpha track density in the human blood samples (*in vitro*). This technique will be beneficial for blood sterilizations.

1.4 Objectives

This study comprises four overall objectives in four main stages as follows:

- To evaluate the risk of accumulation of alpha particles (that produced from decay radon gas) on the lungs and infertility in men, via the design an optimum radon dosimeter, using CR-39 NTDs; Case study in Iraqi Kurdistan.
- 2. To fabricate radium (²²⁶Ra) irradiation technique to evaluate risks of the normal incident of alpha particles onto the surface of human blood samples (*in vitro*) on the blood components (PLT, WBC & RBC).
- 3. To fabricate a suitable radon exposure technique to evaluate risks of the accumulation of the ²¹⁸Po, ²¹⁴Po and ²¹⁰Po (progenies of radon) onto the surface of human blood samples (*in vitro*) on the concentration of the blood components.
- 4. Evaluate the risk of accumulation of the alpha particles on the structure of the Rabbit's tissues (lungs, trachea and testes) via histologically study (*in vivo*) and fabricating a comfortable technique of the radon exposure.

1.5 Scope of Research

1.5.1 Interface of the Elements

The types of user interface elements that are the subject of this study are individual terms, symbols, measurements, and indicators CR-39 NTDs in the field of radon gas. Furthermore, measurement techniques to investigate the deposition of alpha particles on human blood using this type of detector are the interface elements of this study. In addition, inhalation of low radon dose by male rabbits is another scope of this study. Thus, the scope of this study is limited to the following three interface elements:

1. Concentration of the radon gas, annual effective dose and the lifetime risks of the radon's progenies in air, soil, and drinking water, and their risks on human health (lung cancer and men infertility).

- 2. Hematological studies of the alpha particles exposure (in vitro) using CR-39 NTDs.
- 3. *In vivo* histological studies of the inhalation of low doses of indoor radon. The male rabbits are the case study, and the CR-39 NTDs with RAD7 are the detection techniques used.

1.5.2 Locations of Interface Elements

Locations of the present study depended on the case study, and were as follows.

1.5.2 (a) Iraqi Kurdistan region

Iraqi Kurdistan region was the area under study for measuring the radon concentration inside the houses of infertile males (depending on the Erbil center for the infertility). In addition, samples of soil and drinking water were also collected from this region. Iraqi Kurdistan is located in the north of Iraq, with the numerous cases of infertility and cancers (blood, breast and prostate cancer) prevail. This region is largely mountainous with a current population of around six million, and covers approximately 40,643 square kilometers. The map of the Iraqi Kurdistan and the details of the data collection have been presented in Chapter 6.

1.5.2 (b) Penang Island

Some of the official locations in the Penang Island have been used for analytical laboratories for the following purposes:

(a) Medical Physics and Biophysics laboratories in the School of Physics, Universiti Sains Malaysia (USM) were used to design all the techniques employed in this study. In addition, procedures of blood irradiation and the exposure of rabbits were carried out in theses laboratories.

- (b) Human blood samples were collected and analyzed from the Wellness Center of USM main campus.
- (c) Histological examination of the rabbits was carried out in the Histological laboratory, School of Biological Science, USM.

1.6 Outline of the Thesis

The thesis covers risk evaluation of alpha particle deposition on lung cancer, male infertility, and human blood using new irradiation and exposure techniques. It comprises of nine chapters, classified according to the subjects.

Chapter 1 presents an introduction and overview of this thesis. Chapter 2 presents the literature review and previous research on radon inhalation, CR-39 NTDs, deposition of alpha particles on blood, and the effects of deposition of radon's daughter ions in rabbit's lungs. Chapter 3 describes the mechanism of track formation of CR-39 NTDs, while Chapter 4 presents the mechanism of the interaction of alpha particles with the tissues. Chapter 5 presents the calibration of CR-39 NTDs, fabrication of an alpha irradiation collimator, and selection of an optimum radon dosimeter. Chapter 6 describes the evaluation of the risks of deposition of alpha particles on human lungs and male infertility. Chapter 7 presents the risk evaluation of alpha particle deposition on the surface of human blood samples. Chapter 8 describes the preliminary study on the effects of radon inhalation on lungs, trachea, and testes of male rabbits. Finally, Chapter 9 draws the conclusion and future works.

CHAPTER 2

Literature Review & Previous Research

2.1 Introduction

The aim of this chapter is to present a literature review and previous research on the effects of alpha particle deposition onto the surface of lungs, trachea, and blood samples. The focus point will be the detection techniques of nuclear track detectors (NTDs) to determine the inhalation and injection doses of alpha particles emitted from radon's progenies. The next two chapters will discuss on the principle of alpha track formation in nuclear track detectors and mechanism of interaction of alpha particles with tissues.

The risks of deposition of alpha particles (interaction) on lungs, blood, and male fertility are included in the literature review. Deposition of the alpha particles on the tissues and cells may cause damage (temporary and permanent), depending on the time of exposure, energy of the alpha particles, and quantity of the absorbed dose. Thus, studies on the risks of radiation have been carried out for a long time and research on the techniques is novel.

Alpha particles have two protons bound with two neutrons to make a helium nucleus. It is a heavy nuclear particle and is denoted by the Greek alphabet, α . It has a net spin of zero and has a highly ionization form of particle radiation, and exhibits low penetration and low velocity into the matter. Thus, it loses most of its energy in a short distance (Quseph and Nostovych 1978, NRCNA 2006).

The sources of alpha particles are the decay of radioactive nuclei, such as uranium, thorium, actinium, radium, radon and thoron, as well as, the transuranic elements

(chemical elements with atomic number \geq 92). In other words, spontaneous emission of the alpha particles occurs in elements with a mass number of about 150 (NRCNA 2006).

In addition, alpha decay can generate radon gas ${}^{222}_{88}Rn$ from radium ${}^{226}_{88}Ra$. The decay of alpha particles as a process must have sufficient atomic nucleus to support it. Thus, sometimes, during the process of emission of alpha particles, the nucleus may be left in an excited state, and hence, to remove the excess energy, gamma rays will be emitted from the nucleus. The production of alpha particles is through the mechanism of Coulomb repulsion between an alpha particle and the rest of the nucleus, both having same electric charge (positive) (Maher *et al.* 2006).

The speed of the alpha particles, along with its positive charge, easily removes the electrons from the atom's orbits causing ionization of that atom. Thus, its energy of motion will be transferred to the medium and this transfer energy slows down into the target (medium), and the rate of slow down depends on the type of medium.

Alpha particles can penetrate up to 7.5 cm in air, and have a high linear energy transfer (LET). Thus, they lose most of their energy in a small range, which makes them a radiation hazard if ingested (NRCNA 2006, Maher 2006).

Health risks of the deposition of alpha particles occur from ingestion and inhalation of the radionuclide elements that emit alpha particles. Radon (²²²Rn) and thoron (²²⁰Th) are two natural radioactive gases that are colorless, odorless and tasteless. As they are radioactive gases, they will decay to their progenies by emitting alpha particles in lungs, trachea, and stomach, which get deposited onto these organs. As a result, they become a health hazard.

Techniques of an irradiation and exposure are based on their ability to detect and maintain a radiation dose during periods of exposure, as much as possible. Moreover, the rate of energy loss, the purpose of the technique, and the fields of study (hematology, histology, and environmental study) are important parameters, and has been considered in this chapter.

2.2 Literature Review

2.2.1 Alpha Particle and the Radon Gas

Radium $\binom{226}{88}Ra$) is a source of radon gas, which decays by emitting an alpha particle to the radon gas. Henri first discovered radium in 1896 as mentioned by Christie in 1909. In the years of 1899 and 1900, Paul Villard and Ernest Rutherford have separated radiation into three types: alpha, beta, and gamma, depending on their ability to penetrate objects and cause ionization. Alpha rays were defined by Rutherford based on their lowest penetration of ordinary objects (Rutherford 1900, Pohl and Pohl-Ru 1977).

Pierre and Marie Curie n 1899 observed that a radioactive gas emitted by radium remained radioactive for a month (Del and Regato 1979, Mazeron and Gerbaulet 1998, Diamantis *et al.* 2008). At that same year, Rutherford discovered variations during the measurement of radiation from thorium oxide. Rutherford noticed that the radioactive gas that was continuously emitting from compounds of thorium retained its radioactivity for several minutes. He called this gas "emanation," and later renamed it as Thorium Emanation (Th Em) (Rutherford 1900).

Friedrich from Germany was the first to discover of radon in 1900, as mentioned by Rutherford (1900). Friedrich placed radon as the fifth radioactive element, next to

uranium, thorium, radium and polonium. In the same year, during his experiments on radium, he discovered an emanation from the radioactive gas from the radium compounds, and named it as Radium Emanation (Ra Em) (Rutherford 1900).

In 1901, Rutherford proved that the emanations were also radioactive (Rutherford 1901&1902, Del and Regato 1979). Radcliffe and Lond (1909) reported that in 1903, André-Louis Debierne observed similar emanations from actinium and called it as Actinium Emanation (Ac Em). Several names have been suggested for the above mentioned gases (Th Em, Ra Em & Ac Em) during the period of 1904-1920 by various scientists, as follows: in 1904 (exradio, exthorio, and exactinio), in 1918 (radon, thoron, and akton), in 1919 (radeon, thoreon, and actineon), and eventually in 1920 (radon, thoron, and action) (Kathren , 1998). In same year, Sir William Ramsay suggested that the emanations (radon, thoron, and actineon) might be an element of the noble gas family (argon, krypton, and xenon) (Radcliffe and Lond 1909, Moore 1918).

In 1910, radon gas was isolated by Sir William Ramsay and Robert Whytlaw-Gray and its density was determined (Moore 1918). They suggested that radon could be the heaviest gas known. Thus, they suggested a new name called niton (Nt). In 1912, the International Commission accepted it for Atomic Weights. In 1913, the International Committee for Chemical Elements and International Union of Pure and Applied Chemistry (IUPAC) selected the names of radon (Rn), thoron (Tn), and actinon (An), as reported by Aston et al.,(1923). Furthermore, the names of the isotopes were denoted with the number, with the most stable isotopes (radon; ²²²Rn) taking the element name, however, thoron (tn) and action (An) became (²²⁰Rn) and (²¹⁹Rn), respectively (Aston *et al.* 1923). Short-term tests to test the level of radon by remaining in home for 2-90 days, depending on the device have been employed. "Charcoal canisters," "alpha track," "electric ion chamber," "continuous monitors," and "charcoal liquid scintillation" detectors are most commonly used for short-term testing. As radon levels tend to vary from day to day and season to season, a short-term test is less likely to detect the annual average radon level than long-term tests to determine year round average radon level.

2.2.2 Radon Inhalation and its Effects on the Lungs

As an inert gas, radon has a low solubility in body fluids, which leads to a uniform distribution of the gas throughout the body. Exposure to this gas, and its solid decay product, polonium-218, and -214, also result in health risks such as cancer. Once the decay products are inhaled into the lung, they undergo further radioactive decay and release small burst of energy in the form of alpha particles that cause DNA breakage or production of free radicals. Radon not only causes lung cancer, but is also likely to have toxic effects related to the health and survivability of an embryo or fetuses (NRCNA 2006).

When radon and its short-lived decay products are inhaled, the alpha particles emitted by the deposited decay products dominate the radiation dose in the lung tissues, and these products, especially those attached to small size aerosols or those which remain in an unattached form, cause damage to sensitive lung cells, thereby increasing the probability of developing cancer (Field 2011, Mole et al. 1990). The World Health Organization first drew attention to the health effects of residential radon exposures in 1979 (WHO 2009), through a European working group on indoor air quality. Further, radon was classified as a human carcinogen by IARC (1988).

Historical roots of the risk of inhalation of radon on lung cancer started from the discovery of radium by Henri in 1896, and polonium (²¹⁰Po) by Marie and Pierre Curie in 1898. In 1901, Elster and Geitel measured the radon concentration for the first time (Jacobi 1993). The relationship between inhalation of radon and lung cancer was first noted from the incidence of workers who died of lung cancer in the mines of Schneeberg (small city in Saxony/Germany at the northern slope of the "Erzgebirge"). Schneeberg and Jachymov (Jacobi 1993) observed that the high ratio of radon concentration in the air of mines was related to the high ratio of lung cancer among workers in the mines of Schneeberger, based on some findings that have been assumed.

More precise radon measurements carried out in the 1920s in the Schneeberg and Jachymov mines supported this hypothesis. However, the role of radon as a causative factor for the Schneeberger lung cancer was not generally accepted. In a pathological summary report from Dresden in 1926, the cancer was reported to have been caused by the inhalation of toxic dusts (Walsh 1970, Jacobi 1993).

A research program in Germany provided more clarification on the relation between radon concentration and lung cancer. This was a comprehensive study and included measurements of radon concentration in the mines near Schneeberg. In addition, measurements of the alpha activity in tissue samples via histopathological analysis of lung tissues of miners who had died from lung cancer were analyzed.

It was found that the average radon concentration in most mines at Schneeberg was within the range of 70-120 kBq/m³. It was demonstrated that most of the workers in this mine died from lung cancer, and termed it as "death mine." Because of the observations and supporting biological studies, it was concluded that the inhalation of radon must be

regarded as a possible cause for the high ratio of lung cancer among miners in Schneeberg region. These results were summarized in 1945 and the data are available from Schneeberg and Jachymov, but without the mention of the possible role of inhaled short-lived decay products of radon (Walsh 1970, Jacobi 1993).

In 1988, Schiittmann considered that Miiller was the first person to recognize the causal link, as reported by Durrani (1993). Miller concluded that the Schneeberger lung cancer incident was a specific occupational disease, caused by the high radon content in the air of these mines, and when inhaled, it initiated a carcinogenic process in the airways of the lungs (Durrani 1993, Jacobi 1993).

Crameri and Burkart (1989) submitted a report on the importance of radon. They concluded that indoor exposure to radon and radon daughters particles amounted to about 40% of the total effective dose to which the population was exposed to, both from natural and manmade sources.

Studies by National Cancer Institute in 1995 presented a report on radon exposed underground miners. It was found that 40% of the 2700 lung cancer deaths that occurred in 65,000 miners were due to radon (Lubin *et al.* 1995). The Environmental Protection Agency (EPA) recommended that the level of indoor radon concentration should be 4 pCi/l, which had a 6.2% lifetime chance of lung cancer death for cigarette smokers, 2.3% for the general population, and 0.7% for those who never smoked (EPA 2003).

In addition, it provided a reliable document about the effect of smoking on the increased risk of lung cancer in parallel with the inhalation of radon gas. Among non-smokers, 70% of the lung cancer deaths are believed to be due to radon. Thus, the

houses of people who smoke may have high radon levels and their risk of lung cancer is especially high (EPA 1992, Mendez 1998).

EPA 2009 and 2010 recommended that with today's technology, radon levels in most homes could be reduced to 2 pCi/l ($0.02WL=74Bq/m^3$) or below. Thus, it considered an optimum of indoor radon between 2 and 4 pCi/l. In fact, the emanation rate of radon gas has been found to vary from month to month, season to season and year to year, depending on the geological formation, ventilation rate, building material, rate of porosity, soil permeability, etc. (Garakani *et al.* 1988, Hubbard and Hagberg 1996, Gillmore *et al.* 2005, Ismail and Hussyin 2007, Prasad *et al.* 2009, Bochicchio *et al.* 2009, Groves-Kirkby *et al.* 2010, Binesh *et al.* 2011). As a result, long-term measurements of the radon gas are suitable.

2.2.3 Occupational (Physical) Exposures and Male Infertility

In general, occupational exposures are classified into physical exposures (heat and radiation), chemical exposures (solvents and pesticides), psychological exposures (distress), and exposure to metals and welding. Radiation exposure for the occupation of male is under present literature review.

Inhalation of radon and its short-lived decay products is considered to be the most common human exposure. One-third of the deposited decay products from radon are transported from the lungs into the bloodstream, causing ionization of the cells. Thus, a random deposition of the energy in the cells will produce ionization and mutation causing genetic risk. Accordingly, the effects caused by radiation are related to the dose of radiation received, regardless of whether the radiation is of natural or artificial origin. In addition, most of this exposure comes from radon in the air $(1300 \ \mu Sv)$ (Sankaranarayana1999).

The point of importance in the context of this study is that the mutations induced by radiation of human germ cells will cause an increase in the frequency of the genetic material of the spermatozoa, in an increase in sterility, which can be identified by their respective phenotypes. Thus, human fertility is reduced due to pollution, which may be caused by gene mutation, lowered sperm counts, impairment of sperm motility, or many other reasons. Neither the average mutation rate nor the numbers of loci capable of mutating to dominant detrimental form, as well as mutations that cause sterility are known (Sutton 1975).

Review of the effects of radiation resulting in infertility began from study on the effects of ionizing radiation causing genetic damage in human cells. Numerous occupational exposures have been linked to impairment of male fertility (Sheiner *et al.* 2003). However, studies have been limited by inadequate sample sizes, inappropriate study designs, and/or selection bias. Additionally, the use of semen measures as surrogates for male fertility has been problematic, because there is considerable intra individual variability, substantial overlap between infertile men and fertile men, and poor correlation between fertility and decrements in semen measures.

Salvin (1956) submitted a report on his research under the title 'Effect of Atomic Radiation on the Incidence of Sterility and Mutation.' He carried out his research after the atomic bombings in Hiroshima and Nagasaki, with the cooperation of the United States Atomic Energy Commission. This committee supplied funds to the Atomic Bomb Casualty Commission, which sponsored investigations of the results of the bombings. However, owing to the following reasons, he could not obtain satisfactory results: no accurate birthrates were maintained by the Japanese authorities before the bombings took place, and it was difficult to ascertain the damage to fertility by the bombs alone, because during the war, there were other factors affecting fertility, such as fear, anxiety, and malnutrition.

1. There was the great difficulty in obtaining statistics in Japan with reference to the incidence of sterility and mutation rates related to atomic bombing.

2. There was no apparent increase in the incidence of sterility as determined by the birth rates in bombed and unbombed cities.

3. The sterility dose of radiation was approximately the same as the lethal dose of whole body radiation.

4. The fertility of men who survived the bombings returned, as a rule, after several months.

5. The number of stillbirths and abnormalities was greatly increased as the result of atomic radiation of pregnant women. Most of the abnormalities were microcephalics, which is consistent with the current known effects of X-rays on fetus in uterus.

6. There has been no appreciable effect till date on the mutation rate due to the bombings. However, the great majority of geneticists feel that there is an analogy between the mutations produced in animals and humans by radiations, and that the effects of radiation will ultimately express themselves. They also feel that we must balance the patient well at the concealed illness, and use radiation only when one would seem to outweigh the other. There can be nothing but the genetic disadvantage for man in artificially raising his mutation rate above that which sufficed for his evolution till

date. The effects of atomic radiation are destructive by increasing the incidence of sterility temporarily.

In 1959, Oakberg submitted his experiment about irradiation of mouse by the doses of 20 rads of gamma rays and 100, 300, and 600 rad of X-rays. He concluded that the killing of cells, rather than inhibition of mitosis, was the primary factor responsible for radiation-induced depletion of spermatogonia.

In 1960, Heller and Rowley investigated some excellent cell data available with regard to the men testes. They studied the radiation effects on human spermatogenesis in biopsies and ejaculates from a group of 67 volunteers administered with testicular doses of 8-600 rad of X-rays. They found that the spermatogonia in the mouse were most radiosensitive cells for doses under 50 rad (Rowley *et al.* 1974, Gaulden 1983).

In 1971, Léonard submitted his research entitled "Radiation induced translocations in spermatogonia of mice." He found that the many years of radiation translocation would cause heritable semi-sterility.

In 1974 Rowley *et al.*, carried out made an experiment on uniform irradiation of the human testes. A portable unit was developed to provide uniform irradiation of the human testes. The device had built-in radiological protection and provided a dosage independent of the subject geometry, uniform to within $\pm 5\%$. Single doses, between 8 and 600 rad were administered to the testes of human subjects. Dose-response relationships and recovery times were determined for each dose range studied. In 1975 Sutton (1975) found that mutations will cause sterility.

In 1979, Evans *et al*, explained that the nuclear radiation in nuclear dockyard had induced aberrations of the chromosome of the worker. In 1986, Germai submitted his

paper about the situation of radioactive patients. He found that the given therapeutic amounts of radionuclide represent sources of exposure and contamination to personnel providing care. Thus, the workers inside those hospitals were found to receive enough dose of radiation.

In 1987, Searle in his report titled "Review: Radiation and the genetic risk, trends in Genetics," concluded that the ionizing radiation can induce mutations and chromosome structural changes, leading to lethality. In 1988, a number of new reports have appeared which dealt with health risks of radon and other internally deposited alpha emitters (NRC 1988).

In 1994, Roger *et al.*, submitted their article, "Relative biological effectiveness of alpha-particle emitters in vivo at low doses." They found that the therapeutic potential of radionuclides that emit alpha particles and their associated health hazards have attracted considerable attention. In addition, they considered that the above mentioned relationships are based on *in vivo* experimental data, and could be valuable in predicting the biological effects of alpha particle emitters.

In 1995, Hagelström *et al*, submitted their results about ionizing radiation and the chromosomal damage in workers occupationally exposed to chronic low-level ionizing radiation. They found a 4-fold increase in the level of chromosomal aberrations between the exposed and control groups, without qualitative or quantitative cytogenetic differences between X-rays and nuclear medicine exposed workers.

In 2005, Gracia *et al*, submitted their results of their experiments on the effects of occupational exposures on male infertility. They determined the association between male occupational exposures and infertility. They performed their experiments with the