BEAM OPTIMIZATION FOR BORON NEUTRON CAPTURE THERAPY AT THERMAL COLUMN OF TRIGA MARK-II RESEARCH REACTOR

SAFWAN BIN SHALBI

A thesis submitted in fulfilment of the requirements for the award of the degree of Master of Philosophy

School of Chemical and Energy Engineering Faculty of Engineering Universiti Teknologi Malaysia

DECEMBER 2018

ACKNOWLEDGEMENT

"In the name of Allah, the most Gracious and the most Merciful"

Alhamdulillah and thank you to Allah s.w.t., the Almighty, for providing me His showers of blessings. It also drives my effort, inspiration and a lot of forbearance throughout my research to finish it successfully. My greatest profound gratitude goes to my supervisor Dr. Wan Norharyati binti Wan Salleh, from Faculty of Chemical and Energy Engineering, Universiti Teknologi Malaysia (UTM), for her sincere help, full cooperative support and determination has deeply motivated me at every level of this research. Thank you for sharing a lot of progressive critics and ideas throughout the experimental phase, and thesis writing that resulting in the favorable outcomes of this research. My greatest recognition also goes to my co-supervisor, Dr. Faridah binti Idris from Malaysian Nuclear Agency (MNA) for her tremendous effort, encouragement, and critical propositions. Most essentially, thank you so much to my parent and siblings for their ultimate love, endless encouragement and patience with continuous support throughout my journey; without them, this thesis would not be completed. Finally, my appreciation goes to all unnamed personals who help and encourages my journey to finish this research. May Allah s.w.t bless the good deed that has been done and bless us. Amin.

ABSTRACT

The thermal column at TRIGA PUSPATI research reactor has an ability to produce thermal neutron. However, the optimization on the thermal neutron flux produced should be performed in order to gain a sufficient thermal neutron for boron neutron capture therapy purpose. Thus, the objective of this study is to optimize the thermal neutron flux by designing the collimator with different materials at the thermal column. In order to fulfil the requirement, set by the IAEA standard, the collimator was designed using Monte Carlo N-Particle simulation. Initially, the measurement of the thermal neutron flux was conducted along the thermal column at 250 kW. The thermal column was divided into 3 phases (Phase 1, Phase 2 and Phase 3) so that an accurate measurement can be obtained by using gold foil activation method and thermoluminescent dosimeter detector. This value was used as a benchmark for the neutron flux produced from the thermal column. The collimator was designed using different types of materials and their characteristic towards gamma and neutron flux was investigated. The results demonstrated that the final thermal neutron flux produced was significantly depends on the shielding thickness, aperture size and collimator conditions (uncovered or fully covered). The collimator design using thickness shielding of 5+10 cm of lead, aperture size of 3 cm and operated using uncovered condition has produced the optimum thermal neutron flux. The sufficient amount of thermal neutron flux of 3.28 x 10⁸ neutron.cm⁻²s⁻¹ at 250 kW of TRIGA PUSPATI research reactor power produced from the designated collimator was achieved.

ABSTRAK

Turus terma di reaktor penyelidikan TRIGA PUSPATI mempunyai keupayaan untuk menghasilkan neutron terma. Walaubagaimanapun, penghasilan neutron fluks terma yang optima perlu untuk memperolehi neutron terma yang cukup untuk tujuan terapi penangkapan neutron boron. Oleh itu, objektif kajian ini adalah untuk mengoptimumkan fluks neutron terma di turus terma dengan merekabentuk kolimator dengan menggunapakai bahan yang berbeza. Bagi memenuhi kriteria yang ditetapkan oleh piawaian IAEA, kolimator direka menggunakan simulasi Monte Carlo N-Particle. Kajian ini dimulakan dengan mengukur fluks neutron di turus terma pada kadar kuasa reaktor 250 kW. Bagi memudahkan pengukuran, turus terma telah dibahagikan kepada 3 fasa (Fasa 1, Fasa 2 dan Fasa 3) supaya pengukuran menggunakan keranjang emas dan pengesan meter dos pendarkilau haba lebih mudah dan tepat. Nilai fluks tersebut digunakan sebagai penanda aras bagi penyelidikan ini. Kolimator direkabentuk menggunakan bahan yang berbeza dan ciri-cirinya terhadap gama dan fluks neutron dikaji. Hasil penyelidikan ini mendapati fluks neutron terma yang terhasil sangat bergantung terhadap ketebalan perisai, saiz apertur dan jenis kolimator (jenis bertapis dan tidak bertapis). Rekabentuk kolimator yang menggunakan ketebalan pelindung 5 + 10 cm plumbum, saiz apertur 3 cm dan beroperasi menggunakan jenis tidak bertapis telah menghasilkan fluks neutron terma yang optimum. Jumlah neutron terma dihasilkan mencukupi iaitu 3.28 x 10⁸ neutron.cm⁻²s⁻¹ pada 250 kW kuasa reaktor penyelidikan TRIGA PUSPATI daripada kolimator yang direkabentuk telah tercapai.

TABLE OF CONTENTS

CHAPTER	TITLE	PAGE
	DECLARATION	ii
	ACKNOWLEDGEMENT	iii
	ABSTRACT	iv
	ABSTRAK	V
	TABLE OF CONTENTS	vi
	LIST OF TABLES	ix
	LIST OF FIGURES	xi
	LIST OF ABBREVIATIONS	xiv
	LIST OF APPENDICES	XV
1	INTRODUCTION	1
	1.1 Research Background	1
	1.2 Problem Statement	3
	1.3 Objectives	4
	1.4 Scopes of Study	5
	1.5 Significance of Research	5
2	LITERATURE REVIEW	7
	2.1 BNCT Historical and Review	7
	2.2 Neutron Properties	13
	2.2.1 Neutron Absorption	14

	2.2.2 Neutron Shielding	15
	2.2.3 Neutron Collimator	16
	2.2.4 Neutron Moderator	17
	2.3 BNCT Design and Modifications	18
3	RESEARCH METHODOLOGY	24
	3.1 Research Framework	24
	3.2 Thermal Column	27
	3.3 Measurement of Neutron Spectrum and Gamma-ray	29
	3.4 Gold Foil Activation Method	31
	3.5 Characterization of Material	35
	3.5.1 Thermoluminescent Dosimeter	35
	3.5.2 Microspec 6 N Probe Spectrometer	36
	3.5.3 Background Radiation Measurement	38
	3.6 Fluxes of Neutron and Gamma Estimation	39
	3.7 Simulation software using MCNP	40
	3.8 Collimator Design	48
	3.9 Collimator Fabrication	50
4	RESULT AND DISCUSSION	52
	4.1 Measurement of neutron and gamma rays	52
	4.1.1 Phase 1 Measurement	53
	4.1.2 Phase 2 and Phase 3 measurement	56
	4.1.3 Thermal and Epithermal Neutron flux Across	58
	A 1 A TI D-600 and TI D-700 measurements	50 60
	4.2 Characterization of material for BNCT collimator	61
	4.2.1.1 Characterization of material using	01
	4.2.1.1 Characterization of material using Microspec-6	61
	4.2.1.2 Neutron Spectrum Profiles	61
	4.2.1.3 Neutron and Gamma Attenuation Coefficient	64

	4.2.2 Thermoluminescence Detector (TLD)	67
4.3 Design of BNCT collimator using MCNP simulation		71
	4.3.1 Optimization of Collimator Design	76
	4.3.2 Shielding Thickness	76
	4.3.3 Aperture Size	78
	4.3.4 The Different Between Uncovered and Fully	
	Covered Collimator	79
	4.4 Collimator Performance	82
5	CONCLUSIONS	85
	5.1 Conclusions	85
	5.2 Limitation and Recommendation	87
REFERENCES		88

97-104

LIST OF TABLES

TABLE NO.	TITLE	
2.1	The recommended value for BNCT by IAEA	18
2.2	Neutron shielding material studied by a few researchers for BNCT purpose	
2.3	Neutron collimator material studied by a few researchers for BNCT purpose	23
2.4	Neutron moderator material studied by a few researchers for BNCT purpose	23
3.1	Stringers and holes use in experiments	32
3.2	The sample of bare gold foil and cadmium-covered gold foil for the Phase 2 and Phase 3 with distance from G7	32
3.3	Differences between MCNP5 and MCNPX	42
4.1	Thermal and epithermal neutron flux measured using the gold foil activation method for Phase 1	54
4.2	Cadmium ratio from stringer at the thermal column	55
4.3	The measurement of thermal and epithermal by using gold foil activation method from this research and by Munem (2007)	56

4.4	Thermal neutron and epithermal neutron measured using the gold foil activation method for Phase 2 and Phase 3	57
4.5	Thermal neutron and epithermal neutron measured using gold foil activation method across the beam line for Phase 1, Phase 2 and Phase 3	59
4.6	The comparison between the measurement of neutron and gamma using TLD and gold foil activation method	60
4.7	Thermal peaks recorded	64
4.8	Neutron intensity of each material with neutron attenuation coefficient	66
4.9	The measurement of neutron and gamma dose using TLD-600 and TLD-700	67
4.10	Summarization of materials used in collimator	73
4.11	Simulation results of neutron flux across the beam line of BNCT collimator using different material	74
4.12	Simulation results of gamma flux across the beam line of BNCT collimator using different material	76
4.13	The lead thickness result on gamma and thermal from centre and outer	77

LIST OF FIGURES

FIGURE NO.	TITLE	PAGE
1.1	Neutron capture by the boron and subsequent nuclear reaction and nuclear fissions	2
2.1	Illustration view of FiR-1 BNCT facility	10
2.2	Geometric characteristic of Polyethylene/Cerrobend Collimator	11
2.3	Schematic diagram of thermal column before modifications	19
2.4	Schematic diagram of thermal column after modifications	19
2.5	MCNP geometry of BSA for epithermal neutron beam BNCT	21
3.1	Flowchart of the research framework	26
3.2	The outer part of the thermal column	28
3.3	The position and drawing of the stringer at the thermal column	29
3.4	Phase 1, Phase 2 and Phase 3 at the thermal column of RTP	30
3.5	Illustration of sample position for the Phase 2 and Phase 3	33
3.6	The MICROSPEC 6N-probe spectrometer	37

3.7	The Microspec-6N-probe spectrometer setup	38
3.8	Portable Survey Meter	39
3.9	The general flowchart process of MCNP	41
3.10	The cell card for MCNPX input file	44
3.11	The surface card for MCNPX input file	45
3.12	The data card for MCNPX input file	45
3.13	The source card for MCNPX input file	46
3.14	The input file and output file command for MCNPX	47
3.15	Collimator design using simulation	49
3.16	The material cut into pieces for collimator fabricate	50
3.17	The straight test collimator	51
4.1	The neutron flux measurement across the beam line for the Phase 1, Phase 2 and Phase 3	59
4.2	Neutron spectrum profile obtained from the open beam source	62
4.3	Neutron spectrum profile obtained from Microspec-6 for lead, HDPE, 30% borated polyethylene, polyethylene, cadmium, 5% borated polyethylene	63
4.4	The percentage of gamma and fast neutron shield by material based on linear attenuation coefficient	66
4.5	The percentage of neutron produce by using Microspec-6 and TLD detector	68
4.6	The percentage of gamma produce by using Microspec-6 and TLD detector	69

4.7	The collimator design D1-D10	72
4.8	Collimator cell card	73
4.9	Graph of neutron flux across the beam line of collimator with different material	74
4.10	The different lead thickness result on gamma and thermal from centre and outer	77
4.11	The graphical comparison between thermal neutron flux obtain from different size of aperture along the distance	78
4.12	The different between thermal neutron produce over distance by fully covered collimator and uncovered collimator	80
4.13	Contour plot of thermal neutron from uncovered collimator	81
4.14	The geometry of collimator at the thermal column of RTP	82
4.15	The graphical different of thermal neutron flux along the thermal column by simulation (collimator) and experiment	83
4.16	The graphical different of epithermal neutron flux along the thermal column by simulation (collimator) and experiment	84

xiii

LIST OF ABBREVIATIONS

Al	-	Aluminium
BNCT	-	Boron Neutron Capture Therapy
BTI	-	Bubble Technology Industries
BPE	-	Borated Polyethylene
Cd	-	Cadmium
CRANE	-	Centre for Application of Nuclear Energy
IAEA	-	International Atomic Energy Agency
LANL	-	Los Alamos Nuclear Laboratory
LET	-	Linear Energy Transfer
Li	-	Lithium
MCNP	-	Monte Carlo N-Particle
MINT	-	Malaysia Institute for Nuclear Technology Research
MOSTI	-	Ministry of Science, Technology and Environment
Pb	-	Lead
PUSPATI	-	Tun Ismail Atomic Research Centre
RTP	-	Reactor TRIGA Mark II PUSPATI
SANS	-	Small Angle Neutron Scattering
TLD	-	Thermoluminescence Detector

LIST OF APPENDIXES

APPENDIX	TITLE	PAGE	
А	Neutron flux measurement at thermal column stringers	97	
В	Example of Mobile Microspec-6 data	99	
С	TLD-600 and TLD-700 data	102	

CHAPTER 1

INTRODUCTION

1.1 Research Background

Cancer is the most leading cause of death based on 172 countries in the world (Bray *et al.*, 2018). In fact, cancer had caused death over 8.2 million each year, which was roughly estimated about 13% around the world (Organization, World Health, 2003). There are more than 100 types of cancer exist and each requiring different techniques of diagnosis.

BNCT is one of the promising methods in order to cure cancer by using neutron source either from nuclear research reactor or neutron generator such as an accelerator. BNCT method used the combination of low energy of neutron (slow neutron) irradiation and the targeting of a tumour site injected with a proper boron containing compound. Basically, the tumour cell was not directly destructed by neutron, but indirectly destroyed by the results of nuclear reaction between neutron and boron. In theory, the BNCT also is one of the forms of radiotherapy which is selectively kills the cancer cells and less effect on other normal cell that used photon that will selectively deposited in tumour cell of boron carriers such as BPA and BSH. The standard boron carries compound must be enriched by Boron-10 about 20% natural abundance. The patient will be irradiated with slow energy of neutron to reach the ratio of high concentration of boron in the tumour cell (Durisi, 2007). The irradiation with slow neutron points out of nuclear reaction that form the products of Boron-10 capture the thermal neutron and as a result of production Boron-10 convert into Boron-11 which decay from the emission of an alpha particle (Faião-Flores *et al.*, 2011).

Initially in BNCT, thermal neutron is captured by Boron-10 and becomes unstable because of its properties towards neutron. Neutron, especially thermal neutron was well known for its neutral properties (no electrical charge) and could be absorbed by atomic nuclei which collide with it to create heavier isotope. The unstable reaction of Boron-10 and thermal neutron results of emitting both high LET (Linear Energy Transfer) α and γ before it changes to Lithium-7. The LET is a term that used to measure the force acting on a charged particle travelling through matter (Solleh, 2016). There are three factors affecting LET which is charge, velocity and the mass of the particle. (Kraft, 2000). BNCT emitted both high LET α and γ which have deeper penetration in the range of ~10µm which is approximately to the diameter of mammalian cell and promptly attenuating the radiation (Valda *et al.*, 2005). Hence, the BNCT give a highly localized treatment to the tumour cell compare to other treatment. The basic nuclear reaction is illustrated in Figure 1.1:



Figure 1.1 Neutron capture by the boron and subsequent nuclear reaction and nuclear fission (Gohil *et al.*, 2015)

The neutron produce from the research reactor was mostly use in the studies of BNCT around the world (Gohil *et al.*, 2015). This kind of research are aim to develop a BNCT facility for the cancer treatment studies with safe and controlled from radiation and practically using slow neutron emitted from the Malaysia TRIGA MARK II research reactor (RTP). Thus, the first steps to achieve the long-term goal is to establish the suitability neutron fluxes produce from the research reactor. Commonly, most of the TRIGA reactor has a thermal column, which is the special place design to produce thermal neutron which can be utilized in BNCT facility. Therefore, the thermal column of RTP was identified to produce thermal neutron source in the investigation. In contemplation of developing the facility at the thermal column of RTP, the thermal neutron beam needs to be optimized by designing suitable ideal neutron collimator.

1.2 Problem Statement

The development of the BNCT facility in Malaysia can be performed at the thermal column of the Malaysia research reactor. TRIGA MARK II is one of the facilities that can provide neutron souce for BNCT facility. The specification of neutron flux and the gamma dose rate must be considered for the development of the BNCT facility as a safety precaution for this research. Based on previous research, the thermal column is identified as a suitable place for BNCT facility.

RTP was believed to supply a proportionate of neutron beam in developing the BNCT facility (Munem, 2007). The thermal neutron flux measurement at inside the thermal column show that the thermal neutron flux inside the thermal column was 1.17×10^{10} neutron.cm⁻²s⁻¹ which is higher than the IAEA requirement for BNCT (1.0×10^{9} neutron.cm⁻²s⁻¹) at 1000 kW based on research by Solleh (2016).

Unfortunately, those thermal neutron fluxes were decreased into 8.58×10^8 neutron.cm⁻²s⁻¹ as measurement at the thermal column door (Solleh, 2016). Thus, the optimization on the thermal neutron flux at outside of the thermal column to get the equivalent result as thermal neutron flux inside the thermal column for BNCT facility by designing neutron collimator was carried out.

In order to design the neutron collimator for BNCT purpose, the characterization of material towards thermal neutron flux and gamma was explored and the optimization of collimator was performed in terms of shielding thickness, aperture size of collimator and types of collimator design in order to produce high thermal neutron flux.

1.3 Objectives

The main aims of this study are:

- To measure the neutron flux at thermal column using two technique (gold foil activation analysis and TLD detector).
- 2. To design the collimator for BNCT research facility at the thermal column of TRIGA MARK II.
- To simulate the desired neutron flux for the BNCT purpose by using MCNPX.

1.4 Scopes of Study

In order to achieve the objectives of this study, the following scopes of study have been carried out:

- a) Collecting data of thermal neutron flux and gamma flux from experimental work at thermal column using TLD detectors and gold foil activation method analysis.
- b) Characterizing material behaviour towards neutron and gamma using TLD detector and Microspec-6 N-probe for designing collimator.
- c) Designing collimator, the BNCT research facility and simulate from the reactor core using MCNPX software.
- d) Estimating the thermal neutron flux and gamma flux produce through the BNCT facility design using MCNPX.
- e) Maximizing the production of thermal neutron passing and minimize the unnecessary gamma and fast neutron by optimizing the aperture size, shielding thickness and collimator conditions.
- f) Comparing the data obtained from simulation and experimental work.
- g) Fabricating process for the BNCT collimator as the result of thermal neutron flux and gamma flux produce from the MCNPX was sufficient for BNCT.

1.5 Significance of Research

There is one and only research reactor in Malaysia that was identified as an available neutron source for BNCT. The thermal column of RTP are determine to be a suitable place for BNCT purpose based on the early studied of BNCT. The measurement with the verification by MCNP for the thermal neutron at the thermal column was clear that the availability of the thermal column of Malaysia RTP was

sufficient enough for the development of BNCT as the quality and intensity of the neutron beam produced was well within standard requirement for BNCT facility. Research of BNCT at thermal column are being continued by Solleh (2016) in order to produce the data of gamma and neutron outside of the thermal column and as well to develop basic neutron collimator for these purposes. The result produce by Solleh (2016) show that the neutron flux at the end of thermal column door was 8.58 x 10⁸ neutron.cm⁻²s⁻¹ which is lower than IAEA standard (was 1.00 x 10⁹ neutron.cm⁻²s⁻¹) for BNCT facility. Thus, aim of this research is to optimize the thermal neutron beam by designing the neutron collimator for the BNCT purpose so that the neutron flux obtained is within the range that set by the IAEA standard. Furthermore, in the previous research done by Solleh (2016), only single phase was considered which is refer to the Phase 1 (in this work). In this study, three phases were involved in collimator design.

REFERENCES

Agency (2016). History. Available from:

<<u>http://www.nuclearmalaysia.gov.my/new/profile/history.php</u>>.[28 November 2016]

- Abdi, M., Rezaee, K., Shayan, P., & Farzaneh, A. (2012). Collimator Design for Neutron Radiography Systems Using a Reactor Flux. World Applied Sciences Journal, 20(10), 1439-1442.
- Aihara, Teruhito, Hiratsuka, Junichi, Morita, Norimasa, Uno, Masako, Sakurai,
 Yoshinori, Maruhashi, Akira, . . . Harada, Tamotsu. (2006). First clinical case
 of boron neutron capture therapy for head and neck malignancies using 18FBPA PET. *Head & neck*, 28(9), 850-855.
- A. J. B. John R. Lamarsh, Introduction to Nuclear Enginnering, Third Edition ed.
 United States of America: Prentice-Hall, 2001
- Akan, Zafer, Türkmen, Mehmet, Çakır, Tahir, Reyhancan, İskender A, Çolak, Üner, Okka, Muhittin, & Kızıltaş, Sahip. (2015). Modification of the radial beam port of ITU TRIGA Mark II research reactor for BNCT applications. *Applied Radiation and Isotopes*, 99, 110-116.
- Allen BJ, Ralston A, Boron dose enhancement for ²⁵²Cf Brachytherapy. In: Larsson B, Crawford J, Weinrich R,eds. Advances in Neutron Capture Therapy, Vol. 1 Medicine And Physics. Amsterdam: Elsevier, 1998: 271-4.
- Arif, Study of Moderator (Collimator) Materials for BNCT, 2016
- Auterinen, Iiro, Kotiluoto, Petri, Hippeläinen, E, Kortesniemi, Mika, Seppälä, Tiina, Seren, Tom, Collan, Juhani. (2004). Design and construction of shoulder recesses into the beam aperture shields for improved patient positioning at the FiR 1 BNCT facility. *Applied radiation and isotopes*, *61*(5), 799-803.

- Bray, F., Ferlay, J., Soerjomataram, I., Siegel, R. L., Torre, L. A., & Jemal, A. (2018). Global cancer statistics 2018: GLOBOCAN estimates of incidence and mortality worldwide for 36 cancers in 185 countries. *CA: a cancer journal for clinicians*.
- Beddoe, A. H. (1997). Boron Neutron Capture Therapy. British Journal of Radiology. Vol. 70, Pp.665-667.
- Bilski, P., and Obryk, B. (2006). Thermoluminescent dosimetry at the IFJ Krakow.Institute of Nuclear Physics (IFJ), Krakow, Poland.
- Bonora, M., Corti, M., Borsa, F., Bortolussi, S., Protti, N., Santoro, D., Altieri,
 S.Zonta, C., Clerici, A. M., Cansolino, L., Ferrari, C., Dionigi, P., Marchetti,
 A., Zanoni, G., and Vidari, G. (2011). 1H and 10B NMR and MRI
 investigation of boron- and gadolinium–boron compounds in boron neutron
 capture therapy. *Applied Radiation and Isotopes*. Vol. 69, no. 12, pp. 1702-1705.
- Bosko, Andrey. (2005). General Electric PETtrace cyclotron as a neutron source for boron neutron capture therapy. Texas A&M University.
- BTI. (2011). Spectroscopic Neutron Probe. Available from: <<u>http://bubbletech.ca/wp-content/uploads/2014/02/BTI_MICRO_Neutron-</u> <u>Probe_2011-Feb-26.pdf</u> >. [12 December 2016]
- BTI. (2013). Portable Microscopic Survey System (Microspec-6). Available from: <<u>http://bubbletech.ca/wp-content/uploads/2014/02/BTI_MICROSPEC-6_2013-Jun-14_rev2.pdf</u>>. [12 December 2016]
- Burton, J. L., Choksy, S., Edwards, S. J. L., Gallagher, J. A., Glennon, P., Hall, P.
 A.,Howes, T. Q., Jobling, J. C., Laniado, M. E., Lepping, P., Manji, H.,
 McFerran, D.J., Patel, A., Ford, H. E. P., Ford, T. R. P., Rahemtulla, A.,
 Shafiq, A., Sizer, B. F.,Stanaway, S., and Willocks, C. (2007). *Concise Colour Medical Dictionary*.Oxford University Press Inc., New York.
- Cacuci, Dan Gabriel. (2010). Handbook of Nuclear Engineering: Vol. 1: Nuclear Engineering Fundamentals; Vol. 2: Reactor Design; Vol. 3: Reactor Analysis; Vol. 4: Reactors of Generations III and IV; Vol. 5: Fuel Cycles, Decommissioning, Waste Disposal and Safeguards (Vol. 2): Springer Science & Business Media.
- Capture Therapy 2010. Proceedings of 14th International Congress on Neutron Capture Therapy. Buenos Aires, Argentina.

Chandler (2016). Aluminum used in nuclear reactors and other harsh environments may last longer with new treatment. from <u>https://phys.org/news/2016-03-</u> <u>aluminum-nuclear-reactors-harsh-environments.html (</u>2 Febuary 2017)

Chemicool (2015), Available from :<<u>http://www.chemicool.com/elements/boron.html</u>>.[23March 2017].

Chen, A. Y., Liu, Y. W. H., and Sheu, R. J. (2008). Radiation shielding evaluation of the BNCT treatment room at THOR: A TORT-coupled MCNP Monte Carlo simulation study. *Applied Radiation and Isotopes*. Vol. 66, pp. 28–38.

Chen, N. S. (2007). Neutrons. *Encyclopedia of Cancer and Society*. SAGE Publications, Inc. Pp.613-615.

- Chiragkumar J. Gohil and Malleshappa N. Noolvi. (2015). Selective cancer treatment by Boron Neutron Capture Therapy (BNCT) – a review. *International Journal* of Pharmaceutical Chemistry and Analysis; July - September 2015;2(3):136-138
- Devine, R. T. (2002). Evaluation of spectrum measurement devices for operational use. Nuclear Instruments and Methods in Physics Research A, 476 (416–422).
- Da Silva, A. X., & Crispim, V. R. (2001). Moderator–collimator-shielding design for neutron radiography systems using 252Cf. *Applied Radiation and Isotopes*, 54(2) (217-225).
- Durisi, E, Zanini, A, Manfredotti, C, Palamara, F, Sarotto, M, Visca, L, & Nastasi, U. (2007). Design of an epithermal column for BNCT based on D–D fusion neutron facility. *Nuclear Instruments and Methods in Physics Research Section A: Accelerators, Spectrometers, Detectors and Associated Equipment, 574*(2), 363-369.
- Faghihi, F, & Khalili, S. (2013). Beam shaping assembly of a D–T neutron source for BNCT and its dosimetry simulation in deeply-seated tumor. *Radiation Physics* and Chemistry, 89, 1-13.
- Faião-Flores, F, Coelho, PRP, Muniz, ROR, Souza, GS, Arruda-Neto, J, & Maria, Durvanei A. (2011). Antitumor potential induction and free radicals production in melanoma cells by Boron Neutron Capture Therapy. *Applied Radiation and Isotopes*, 69(12), 1748-1751.
- Fauziah, Nina, Widiharto, Andang, & Sardjono, Yohannes. (2015). A Conceptual Design Of Neutron Collimator In The Thermal Column Of Kartini Research

Reactor For In Vitro And In Vivo Test Of Boron Neutron Capture Therapy. *Tri* Dasa Mega-Jurnal Teknologi Reaktor Nuklir, 15(2).

- Ferlay, Jacques, Shin, Hai-Rim, Bray, Freddie, Forman, David, Mathers, Colin, & Parkin, Donald Maxwell. (2010). Estimates of worldwide burden of cancer in 2008: GLOBOCAN 2008. *International journal of cancer*, 127(12), 2893-2917.
- Floberg, John. (2005). The physics of boron neutron capture therapy: an emerging and innovative treatment for glioblastoma and melanoma. *Physics and Astronomy Comps Papers*.
- Geng, C., Tang, X., Guan, F., Johns, J., Vasudevan, L., Gong, C., . . . Chen, D. (2015). GEANT4 calculations of neutron dose in radiation protection using a homogeneous phantom and a Chinese hybrid male phantom. Radiation protection dosimetry, 168(4), 433-440.
- Gohil, Chiragkumar J, & Noolvi, Malleshappa N. (2015). Selective cancer treatment by Boron Neutron Capture Therapy (BNCT)–a review. *International Journal* of Pharmaceutical Chemistry and Analysis, 2(3), 136-138.
- Harris, D. R. (2017). Neutron Fluctuations in a Reactor of Finite Size. Nuclear Science and Engineering, 21:3(369-381).
- Hatanaka, H, Sweet, WH, Sano, K, & Ellis, F. (1991). The present status of boronneutron capture therapy for tumors. *Pure and applied chemistry*, 63(3), 373-374.
- Hawk, Andrew E, Blue, TE, & Woollard, JE. (2004). A shielding design for an accelerator-based neutron source for boron neutron capture therapy. *Applied radiation and isotopes*, 61(5), 1027-1031
- Hopewell, JW, Gorlia, T, Pellettieri, L, Giusti, V, H-Stenstam, B, & Sköld, K. (2011). Boron neutron capture therapy for newly diagnosed glioblastoma multiforme: an assessment of clinical potential. *Applied Radiation and Isotopes*, 69(12), 1737-1740.
- IAEA, International Atomic Energy Agency. (2001).Current status of neutron capture therapy. IAEA-TECDOC-1223.
- Ibrahim, T. N. S. (2004). National Cancer Control Programme. *Malaysia's Health* 2004. Ministry of Health Malaysia.
- Joensuu, Heikki, Kankaanranta, Leena, Seppälä, Tiina, Auterinen, Iiro, Kallio, Merja, Kulvik, Martti, Kotiluoto, Petri. (2003). Boron neutron capture therapy

of brain tumors: clinical trials at the Finnish facility using boronophenylalanine. *Journal of neuro-oncology*, *62*(1), 123-134.

- Kotiluoto, P., Auterinen, I., Savolainen, S., Kouri, M., and Joensuu, H. (2007).Boron neutron capture therapy in the treatment of locally recurred head and neck cancer. *Int. J. Radiat. Oncol. Biol. Phys.* Vol. 69, pp.475.
- Kankaanranta, L, Koivunoro, H, Seppälä, T, Atula, T, Mäkitie, A, Kortesniemi, M, .
 Savolainen, S. (2008). Boron Neutron Capture Therapy (BNCT) In The Treatment Of Locally Recurred Head And Neck Cancer. *Clinical Otolaryngology*, *33*(4), 385.
- Kato, Itsuro, Fujita, Yusei, Maruhashi, Akira, Kumada, Hiroaki, Ohmae, Masatoshi, Kirihata, Mitsunori, . . . Sumi, Tetsuro. (2009). Effectiveness of boron neutron capture therapy for recurrent head and neck malignancies. *Applied Radiation* and Isotopes, 67(7), S37-S42.
- Kapusta, B, Sainte-Catherine, C, Averty, X, Scibetta, M, & Rommens, M. (2003).
 Present status on the mechanical characterization of aluminum alloys 5754-NET-O and 6061-T6 irradiated at high fluences. Paper presented at the 9th Meeting of the International Group on Research Reactors, IGORR, Sydney, Australia.
- Kasesaz, Y., Khala, H., Rahmani, F., Ezzati, A., Keyvani, M., Hossnirokh, A., Shamami, M. A., and Amini, S. (2014). Design and construction of a thermal neutron beamfor BNCT at Tehran Research Reactor. *Applied Radiation and Isotopes*. Vol. 94,pp.149–151
- Kortov, V. S., Milman, I. I., Nikiforov, S. V., and Gorelova, E. A. (2000). The Use of Thermoluminescent Detectors for Radiation Monitoring on Territories of Atomic Power Plants. Journal of International Research Publication. Issue 1 -2000/01 ISSN, pp.1311-8978
- Kraft, Gerhard. (2000). Tumor therapy with heavy charged particles. *Progress in Particle and Nuclear Physics*, 45, S473-S544.
- Kreiner, AJ, Baldo, M, Bergueiro, JR, Cartelli, D, Castell, W, Vento, V Thatar, Sandin, JC Suarez. (2014). Accelerator-based BNCT. *Applied Radiation and Isotopes*, 88, 185-189.
- Lehtinen, P. M. (2007). A new radiation therapy treatment developed for head and neck cancer patients in Finland. *Press Releases of University of Helsinki*. Vol. 13, pp.21.

- M., Debray, M. E., Somacal, H. R., Capoulat, M. E., Herrera, M. S., Grosso, M. F., Gagetti, L., Anzorena, M. S., Canepa, N., Real, N., Gun, M., and Tacca, H. (2014).Present status of Accelerator-Based BNCT. *Reports of Practical Oncology and Radiotherapy*. Greater Poland Cancer Centre. Pp.414-420.
- Mallick, I. (2014). Overall Survival (OS), Available from: http://lymphoma.about.com/od/glossary/g/overallsurvival.htm.[29 November 2016]
- Masunaga, S., Nagasawa, H., Hiraoka, M., Sakurai, Y., Uto, Y., Hori, H., Nagata, K.,Suzuki, M., Maruhashi, A., Kinashi, Y., and Ono, K. (2004). The usefulness of 2-nitroimidazole-sodium borocaptate-10B conjugates as 10B-carriers in boron neutron capture therapy. *Applied Radiation and Isotopes*. Vol. 61, pp.953–958.
- Minsky, D. M., Valda, A. A., Kreiner, A. J., Green, S., Wojnecki, C., and Ghani, Z. (2011). First tomographic image of neutron capture rate in a BNCT facility.
- Mitsumoto, T., Yajima, S., Tsutsui, H., Ogasawara, T., Fujita, K., Tanaka, H., Sakurai, Y., and Maruhashi, A. (2013). Cyclotron-Based Neutron Source for BNCT. AIP Conf. Proc. Vol. 1525, pp. 319.
- Mitchell, Hannah Elizabeth. (1996). An accelerator-based epithermal photoneutron source for boron neutron capture therapy: School of Mechanical Engineering, Georgia Institute of Technol
- Mohamed, A. A., Muhammad, A., Idris, F., Bokhari, A., and Yunus, M. N. (2003). Neutron Beam Applications Using Low Power Research Reactor Malaysia Perspectives
- Monshizadeh, M, Kasesaz, Y, Khalafi, H, & Hamidi, S. (2015). MCNP design of thermal and epithermal neutron beam for BNCT at the Isfahan MNSR. *Progress in Nuclear Energy*, 83, 427-432.
- Munem, Eid Mahmoud Eid Abdel. (2007). Neutron flux measurements with Monte Carlo verification at the thermal column of a Triga mark II reactor: Feasibility study for a BNCT facility
- Normey-Rico, J. E. (2007). Control of dead-time processes: Springer Science & Business Media.
- Nigg, D. W. (2006). Neutron Sources and Applications in Radiotherapy-A Brief History, and Current Trends. United States. Department of Energy.

- Organization, World Health. (2003). *The world health report 2003: shaping the future*: World Health Organization.
- Paiva, Siqueira, P. T., & Cavalieri, T. A. Comparing the responses of TLD 100, TLD 600, TLD 700 and TLD 400 in mixed neutron-gamma fields.
- Payudan, A., Haryadi, A., & Abdullatif, F. (2017). Optimization of Collimator Neutron Design for Boron Neutron-Capture Cancer Therapy (BNCT) Based Cyclotron 30 MeV. Indonesian Journal of Physics and Nuclear Applications, 2(3), 128-136.
- Pohjola, M. (2009). Biologically targeted treatment. Available from: ">http://www.hightech.fi/direct.aspx?area=htf&prm1=748&prm2=article>">http://www.hightech.fi/direct.aspx?area=htf&prm1=748&prm2=article>">http://www.hightech.fi/direct.aspx?area=htf&prm1=748&prm2=article>">http://www.hightech.fi/direct.aspx?area=htf&prm1=748&prm2=article>">http://www.hightech.fi/direct.aspx?area=htf&prm1=748&prm2=article>">>">">
- Pouryavi, Mehdi, Masoudi, S Farhad, & Rahmani, Faezeh. (2015). Radiation shielding design of BNCT treatment room for DT neutron source. *Applied Radiation and Isotopes*, 99, 90-96.
- Pozzi, Emiliano CC, Trivillin, Verónica A, Colombo, Lucas L, Hughes, Andrea Monti, Thorp, Silvia I, Cardoso, Jorge E, Curotto, Paula. (2013). Boron Neutron Capture Therapy (BNCT) For Liver Metastasis In An Experimental Model: Dose–Response At Five-Week Follow-Up Based On Retrospective Dose Assessment In Individual Rats. *Radiation and environmental biophysics*, 52(4), 481-491.
- Rafi (2015). Moderator, Collimator and Shielding Studies for BNCT Research at Malaysian Nuclear Agency. PhD Thesis. Universiti Sains Malaysia.
- RSICC. (2008-2013). Monte Carlo N–Particle Transport Code System MCNP6.1. Available from:

<<u>https://rsicc.ornl.gov/codes/ccc/ccc8/ccc-810.html</u>>. [30 December 2016].

- Robinson, J. A., Hartman, M. R., and Reese, S. R. (2010). Design, Construction and Characterization of a Prompt Gamma Activation Analysis Facility at the Oregon State University TRIGA® Reactor. *Journal of Radio analytical and Nuclear Chemistry*. 283(2): 359-369
- Rossini, Andrés E, Dagrosa, Maria A, Portu, Agustina, Saint Martin, Giselle, Thorp, Silvia, Casal, Mariana, Pisarev, Mario A. (2015). Assessment of biological effectiveness of boron neutron capture therapy in primary and metastatic melanoma cell lines. *International journal of radiation biology*, *91*(1), 81-89.

- Valda, A, Minsky, DM, Kreiner, AJ, Burlon, AA, & Somacal, H. (2005).
 Development of a tomographic system for online dose measurements in BNCT (Boron neutron capture therapy). *Brazilian journal of physics*, 35(3B), 785-788.
- Shalbi, S., Salleh, W. N. W., Idris, F. M., Rosdi, M. A. A., Sarkawi, M. S., Jamsari, N. L., & Nasir, N. A. M. (2018). Neutron measurement at the thermal column of the Malaysian Triga Mark II reactor using gold foil activation method and TLD. In *IOP Conference Series: Materials Science and Engineering* (Vol. 298, No. 1, p. 012031). IOP Publishing.
- Sakurai, Yoshinori, Sasaki, Akira, & Kobayashi, Tooru. (2004). Development of neutron shielding material using metathesis-polymer matrix. Nuclear Instruments and Methods in Physics Research Section A: Accelerators, Spectrometers, Detectors and Associated Equipment, 522(3), 455-461.
- Savolainen, S., and Joensuu, H. (2012). Boron neutron capture therapy in the treatment of locally recurred head-and-neck cancer: final analysis of a phase I/II trial. *International Journal of Radiation Oncology/Biology/Physics*. Vol. 82, Issue 1, pp.67–75.
- Shaaban, Ismail, & Albarhoum, Mohamad. (2015). Design calculation of an epithermal neutronic beam for BNCT at the Syrian MNSR using the MCNP4C code. *Progress in Nuclear Energy*, 78, 297-302.
- Shirakawa, Makoto, Yamamto, Tetsuya, Nakai, Kei, Aburai, Kenichi, Kawatobi, Sho, Tsurubuchi, Takao, Matsumura, Akira. (2009). Synthesis and evaluation of a novel liposome containing BPA–peptide conjugate for BNCT. *Applied Radiation and Isotopes*, 67(7), S88-S90.
- Shukri, A. (2005). Boron Neutron Capture Therapy (BNCT). Research Reactor for Sustainable Development. *Prosiding Bengkel dan Seminar Reactor Interest Group (RIG)*. RIG, Institut Penyelidikan Teknologi Nuklear Malaysia (MINT).
- Shultis, J Kenneth, & Faw, Richard E. (2011). An MCNP primer. *Kansas State University, Manhattan*.
- Solleh, Mohd, & Rafi, Mohd. (2016). *Moderator, Collimator And Shielding Studies* For Bnct Research At Malaysian Nuclear Agency. Universiti Sains Malaysia.

Solleh M. R. M., Tajudin, A. A., Mohamed, A. A., Munem, E. M. E.

- A., Rabir, M. H., Yoshiaki, K. (2011). Collimator And Shielding Design For
 BNCT Research At TRIGA MARK II Reactor. *Proceeding of Nuclear Science, Technology & Engineering Conference (NUSTEC2011). Malaysian Nuclear Society*
- Turgay Korkut , A. K., Gokhan Budak, Bunyamin Ayg, Osman Gencel, Aybaba Hanc-erliogulları (2012). Investigation of neutron shielding properties depending on number of boron atoms for colemanite, ulexite and tincal ores by experiments and FLUKA Monte Carlo simulations. *Applied Radiation and Isotopes*, 70(1), 341–345. doi: <u>https://doi.org/10.1016/j.apradiso.2011.09.006</u>
- Walker, Simon J. (1998). Boron neutron capture therapy: principles and prospects. *Radiography*, 4(3), 211-219.

Wagiran, H. (1997). Prinsip Asas Sinaran dan Reaktor Nuklear. Penerbit UTM.

- Weisberg, D. E. (2006). Autodesk and AutoCAD.
- Winter, M. (2015) Boron: the essentials, Available from: <<u>http://www.webelements.com/boron/</u>>.[23 March 2017].
- X-5 Monte Carlo Team (2003). MCNP A General Monte Carlo: N-particle Transport Code, version 5. University of California, Los Alamos National Laboratory
- Zafiropoulos, D., and Scarabottolo, G. (2001). Neutron Spectra using a portable Neutron Spectrometer. *Radiation protection Group, INFN, Laboratory Nazionali di Legnaro*.
- Zolghadri, S., Yousefnia, H., Afarideh, H., Bahrami-Samani, A., Jalilian, A., & Ghannadi-Maragheh, M. (2013). Measurement of thermal neutron capture cross-section and resonance integral for the 165Ho (n, γ) 166gHo reaction by the activation method. Nuclear Instruments and Methods in Physics Research Section B: Beam Interactions with Materials and Atoms, 295, 94-98.