UNIVERSITI TEKNOLOGI MARA

TIMING OF ACUTE CORONARY SYDROME ONSET WITH OBSTRUCTIVE SLEEP APNOEA RISK STUDY (TACOS)

MUHAMMAD HANIS BIN MUHMAD HAMIDI

MMed

May 2018

AUTHOR'S DECLARATION

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

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

Name of Student : Muhammad Hanis Bin Muhmad Hamidi

Student I.D. No. : 2013974251

Programme : Master of Internal Medicine – MD774

Faculty : Medicine

Thesis : Timing of Acute Coronary Syndrome Onset with

Obstructive Sleep Apnoea Risk Study (TACOS)

Signature of Student :

Date : May 2018

ABSTRACT

Introduction: Acute coronary syndrome (ACS) remains the principal cause of death in Malaysia. It is estimated about 20-23% of ACS occurs at night-time between 12am to 6am (nocturnal ACS). Factors associated with nocturnal ACS are unknown. Acute nocturnal pathophysiological response to obstructive sleep apnoea (OSA) may increase risk of nocturnal ACS. This study aims to determine the prevalence of nocturnal ACS and establish any risk factors in particular OSA risk which may be associated with nocturnal ACS. Methodology: This was a cross-sectional study of ACS patients undergoing coronary angiogram in CTC UiTM Sg Buloh between 1st August 2017 and 28th February 2018. Combination of STOP-BANG and Epworth Sleepiness Scale (ESS) questionnaires was used to determine OSA risk. The timing of ACS onset of each subject was established from interview and divided into two groups; nocturnal ACS (12am to 559am) and non-nocturnal ACS (6am to 1159pm). Twenty high risk OSA subjects were selected to undergo level I polysomnography (PSG). All subjects timing of ACS onset, OSA risk, demography, anthropometric measurements, comorbidities, echocardiography and PSG characteristics were analysed. Result: A total of 200 subjects were recruited. The prevalence of nocturnal ACS among ACS patients was 19% (38/200). The proportion of patients who were found to be high risk OSA was significantly larger in nocturnal ACS group compared to non-nocturnal ACS group, 36/38 (95%) vs 49/162 (30%), respectively (p<0.001). Nocturnal ACS patients were significantly younger, had higher BMI, waist circumference and neck circumference compared to non-nocturnal ACS patients; 50.1 $\pm 8.7 \text{ vs } 60.4 \pm 9.6 \text{ years}, 33.9 \pm 4.3 \text{ vs } 27.1 \pm 5.3 \text{ kg/m}^2, 106.7 \pm 10.3 \text{ vs } 96.8 \pm 11.3 \text{cm},$ 44.6±3.3 vs 39.5±3.3cm, respectively (p<0.05). Based on multiple logistic regression analysis, significant predictors for nocturnal ACS were age, OSA risk and neck circumference. There was no significant difference in PSG characteristics in high risk OSA patients between those with nocturnal ACS compared to non-nocturnal ACS. The positive predictive value of combined STOP-BANG and ESS scoring in determining OSA risk was 1. Conclusion: The prevalence of nocturnal ACS was 19%. The prevalence of high risk OSA among nocturnal ACS was very high compared to non-nocturnal ACS. There may be a strong association between OSA risk and nocturnal ACS. The significant predictors for nocturnal ACS were age, OSA risk and neck circumference.

ACKNOWLEDGEMENT

Alhamdulillah, Praise to the Almighty and the Most Merciful for blessing me with the strength, health and wisdom for enduring this challenging yet satisfyingly enriched journey of Masters in Internal Medicine. My highest gratitude goes to my supervisor Dr. Muhd Amin Ibrahim and my co-supervisors Associate Professor Dr Izuanuddin Ismail, Dr Mohd Arif M.Zim and Dr Johan Rizwal Ismail who have inspired me and have provided endless support, ideas and assistance at every step of this project.

I would also like to express my gratefulness to the professors and lecturers of Internal Medicine department UiTM with special thanks to Prof. Dr Mohd Fauzi Abdul Rani, Assoc. Prof. Dr Rohana Abdul Ghani, Assoc. Prof. Dr Sazzli Shahlan Kassim, Dr Muhd Iqbal Abd Hafiz, Dr Aisya Natasya Musa, Dr Muhd Yazli Yuhana, Dr Rizmy Najme Khair, Dr Nicholas Chua, Dr Khairul Shafiq Ibrahim and Dr Rafiz Abd Rani. They have been more than willing to provide me holistic guidance, knowledge and support to ensure UiTM Masters of Internal Medicine candidates progress through this physically, mentally and emotionally demanding journey with triumph.

My humble appreciation also goes to the medical laboratory technicians and physiotherapist of UiTM Respiratory unit, namely Abdul Wafi Zakaria, Siti Noor Aishah Zahari, Hilmi Abdullah, Mohd Nazeri Kamaruddin and Muhamad Afandi Arippin who have provided me the facilities and assistance during my data collection. Without the collaboration and hardwork of these remarkable individuals, the project would not have been a success.

I have always believed that success is achieved not individually but merely with the collective effort and support between individuals. Therefore, my sincerest gratitude goes to all my superiors, colleagues and friends with special thanks to Dr Yusniza Yusoff, Dr E Chia Qi, Dr Sharimila Shanmugam, Dr Kausalyaa Krishnabalan, Dr Noor Shazana Mustaffa, Dr Idazureen Ismas Ismail, Dr Rohan Shanmuganathan, Dr Ng Ken Yaw, Dr Huzairi Sani and Dr Burhanuddin Salim who have provided me help in every way possible to ensure the success of this project.

Last but not least, I would like to express my love and greatest appreciation to my loving wife, Norhashimah Matulidi, my parents, Hj Muhd Hamidi Nazir and Hjh Norhalini Siew Abdullah and my family. They have been my pillars of strength that never give up praying for my success and have given me endless support in the best possible way to ensure that I advance through. This piece of work is not only a victory to myself but also specially dedicated to all of you.

TABLE OF CONTENT

			Page				
CONFIRMATION BY PANEL OF EXAMINERS			ii				
AUTHOR'S DECLARATION			iii				
ABSTRACT ACKNOWLEDGEMENT TABLE OF CONTENT LIST OF TABLES LIST OF FIGURES LIST OF SYMBOLS AND ABBREVIATIONS LIST OF APPENDICES			iv v vi ix x xi				
				CHA	APTER (ONE: INTRODUCTION	1
				1.1	Resea	rch Background	1
				1.2	Research Objectives		1
					1.2.1	Primary Objectives	1
					1.2.2	Secondary Objectives	1
				1.3	Research Hypotheses		2
1.4	Significance of the study		2				
1.5	Definition of Terms		3				
	1.5.1	Acute Coronary Syndrome	3				
	1.5.2	Timing of ACS onset	4				
	1.5.3	Obstructive Sleep Apnoea	4				
	1.5.4	Polysomnography	5				
	1.5.5	Obstructive Sleep Apnoea Risk	6				
CHA	APTER T	TWO: LITERATURE REVIEW	8				
2.1	Epidemiology		8				
	2.1.1	Circadian rhythm and Timing of ACS onset	8				
	2.1.2	OSA and ACS	9				
	2.1.3	OSA and Nocturnal ACS	10				