## UNIVERSITI TEKNOLOGI MARA

# EFFECT OF SUPPLEMENTATION OF TOCOTRIENOL ON ALTERATION OF ESTROUS CYCLE IN FEMALE RATS SUBJECTED TO STRESS.

## **MARIA BINTI EMBONG**

Dissertation submitted in partial fulfillment of the requirement for the

**Faculty of Pharmacy** 

October 2008

## **ACKNOWLEDGEMENT**

First and foremost I would like to express my greatest attitude to Allah S.W.T for giving me the strength and spirit to complete this research project. The deepest appreciations are expressed to my supervisor, Pn. Masita Nordin for giving me supports and guide me throughout this research. Sincere thanks are also expressed to Mr. Aloysious, Mr Syed Ridzuan and Mr. Halim and those who contributed their help and support me either directly or indirectly. Lastly, I would like to thanks to my family and all my friends who always with me whenever I need them.

Thank you very much.

# TABLE OF CONTENT

			Page
TITLE PAGE APPROVAL ACKNOWLEDGE TABLE OF CONTILIST OF FIGURE LIST OF TABLE LIST OF PLATE ABSTRACT			ii iii v vi vii viii
CHAPTER 1	INTRODUC	TION	
1.1	Background of	of study	1
1.2	Objective		2
CHAPTER 2	LITERATU	RE REVIEW	
2.1	Introduction		3
2.2	The reproductive system of female rat		4
2.3	Rat estrous cycle		5
2.4	Vitamin E		
	2.4.1	The chemical structure	8
	2.4.2	Pharmacokinetic properties	8
	2.4.3	Mechanism of action	9
	2.4.4	Vitamin E and health related issue	10

#### **ABSTRACT**

Menstrual cycle can be altered by many factors including stress. Physiological stress such as exercise has been proved to cause menstrual irregularities. This may be due to the distruption at the hormone level or due to the production of free radical during intense exercise. Antioxidants such as tocotrienol could reduce the activity of free radical thus prevent the damaging effect on the biological structure in the body. Therefore, this study was carried out to observe the effect of supplementation of tocotrienol in female rat estrous cycle that had been put under forced swimming stress. Twenty Sprague Dawley rats were divided into four groups that consist of 5 rats. The first group was the control group where no treatment was given. The second group was given 0.2ml corn oil only while the third group was put under force swimming stress. The forth group also was put under force swimming stress but supplemented with 60mg/kg tocotrienol dissolved in 0.2 ml pure corn oil. The mean cycle length between control and treated groups were compared using independence T-test with 95% confidence interval. The result showed that mean cycle length of the group that was given 0.2ml corn oil was 4.55±0.17 days. For the group that was put under force swimming stress but supplemented with 60mg/kg tocotrienol had mean cycle length of 5.27±0.26 days. These result were statistically no significant different with the control group where the mean cycle length was 4.61±0.14 days. Group that have been put under force swimming stress had significantly longer estrous cycle length (6.85±0.40 days) compare with control group, p<0.05. This proved that force swimming stress can alter the estrous cycle by prolonging its length and this alteration can be reversed by supplementation of tocotrienol.

#### **CHAPTER 1**

## INTRODUCTION

## 1.1 Background of study

Stress is a psychological and physiological response to events that upset our personal balance in some way. Women's menstrual cycle disturbance can occur due to physiological stress such as strenuous exercise (Warren & Perlroth, 2001) and emotional stress (Dmitry et. al, 2005). Extended or repeated activation of the stress response takes a heavy toll on the body. Jenifer et. al (2007) reported that stressful life events in incarcerated women cause ammenorhea (9%) and menstrual irregularities (33%). Fenster et al. (1999) found that women who experienced high stress at work had higher risk of short cycle. Study by Susannah et. al (2004) also found that increase stress levels in perimonopousal women shorten the menstrual cycle and duration of bleeding. However, the result has no relationship with the age of the women. Menstrual irregularity that are due to strenuous exercise originates from dysfunction at hypothalamic level (Warren & Perlroth, 2001). Low energy availability during strenuous exercise also could lead to menstrual disturbances (Nancy et. al, 2001). Reactive oxygen species (ROS), a type of free radicals play a role in female reproductive function (Oyawoye et. al, 2003). Though ROS is essential in some process of reproduction,