

UNIVERSITI TEKNOLOGI MARA

**THE CYTOTOXIC EFFECT OF RED AND GREEN
MALAYSIAN SEAWEEDS AGAINST LIVER CANCER
CELL LINES (HepG2)**

SYUHADA BINTI MOHD JALIL

Dissertation submitted in partial fulfilment of the requirement for
the degree of **Bachelor of Pharmacy (Hons)**

BACHELOR OF PHARMACY (Hons)

2014

ACKNOWLEDGEMENT

Alhamdulillah and praised to Allah s.w.t for His blessing, I am able to accomplish this thesis at the right given times. There were many people officially and unofficially being involved to help me for this accomplishment. It is quite impossible for me to acknowledge every one of them. However, some of them do really deserve for appreciation.

First and foremost, deepest gratitude and big thanks toward my final year research project's supervisor, Dr Siti Alwani binti Ariffin for accepting me as her student. She did teach me a lot of valuable things along this research project. I am really appreciated her continuous and nonstop encouragement, patience and guidance toward me during this project. All of her encouragements were very helpful in motivating me to work harder than everyone else without comparing the amount of the work in order to obtain extra knowledge throughout this research project. Besides, she also gave me ideas on how actually doing a research appropriately by demonstrating it. Apart from that, she did develop me to be beyond from who I am right now and realizing me, there is no shortcut in order to achieve success.

I also would like to express my appreciation to my inspirational colleague, Nur Fatihah binti Zainal Abdim who always be with me over the entire periods of making this thesis and also the other entire current and former member of Marine Pharmaceutical Research Group (MAREGs) Laboratory. Not forgetting, special thanks toward both of my parents as they have been very supportive in term of both encouragement and financially supported for the whole periods to make this project in reality.

TABLE OF CONTENT

TITLE PAGE	PAGE
ACKNOWLEDGEMENTS	
LIST OF TABLES	i
LIST OF FIGURES	ii
LIST OF ABBREVIATIONS	iii
ABSTRACT	v
CHAPTER I (INTRODUCTION)	
1.1 Background of study	1
1.2 Problem statement	4
1.3 Objectives	5
CHAPTER II (LITERATURE REVIEW)	
2.0 Cancer	6
2.1 Liver cancer	8
2.1.1 Risk factor for cancer development	9
2.2 Chemotherapy	12
2.3 Drug discovery from natural products	14
2.4 Drug discovery from marine sources	15
2.5 Seaweeds	17
2.5.1 <i>Acanthoporphyspiciera</i> (M. Vahl) Borgesen	20
2.5.2 <i>Chaetomorpha</i> genus	22
2.6 Endophytes	23
2.6.1 Metabolites produced by marine endophytic fungi	25

ABSTRACT

In recent years, generally about 30,000 new compounds from marine natural products have been explored and identified. This number of findings reflected the great potential of marine natural products as a source of novel chemicals classes. Due to the efficacy limitation of the available anticancer drugs in liver cancer treatment, there is an urge to seek new candidate for alternatives drugs. In search for new source of active metabolites, marine endophytic fungi isolated from seaweeds were chosen to be further studied. Hence, in this study, three marine endophytic fungi (ED1, ED2 and UF) isolated from red (*Acanthophora spicifera* (M. Vahl) Borgesen) and green Malaysian seaweeds (*Chaetomorpha minima* F.S Collins & Hervey 1917) were grown on two types of media, PDA with 1% and 3% of artificial sea salt (PDA 1% and PDA 3%). Marine endophytic fungi extracts were investigated for their cytotoxic effect against HepG2 cells by using MTT assay. In this presence study, the lowest IC₅₀ values were observed at 72 hours incubation times. Extracts with the IC₅₀ values less than 50µg/ml [UF 1% (16.5± 1.5µg/ml), ED1 1% (36 ± 16.00µg/ml) and UF 3% (42.5± 12.5µg/ml)] were considered to have promising cytotoxic activity. Only one extract [UF 1% (16.5± 1.5µg/ml)] showed active cytotoxic activity whereas two extracts [(ED1 1% (36 ± 16.00µg/ml) and UF 3% (42.5± 12.5µg/ml)] had moderate cytotoxic activity. As for salinity, endophytic fungi media supplemented with 1% artificial sea salt was sufficient in providing cytotoxic effect against HepG2 cells as compared to 3%. Therefore, the cytotoxic effects exhibited by these three extracts against HepG2 cells could be considered as suitable candidates for further studies as anticancer agents.

Keywords: Cytotoxic, seaweeds, marine endophytic fungi, HepG2.

CHAPTER ONE

INTRODUCTION

1.1 Background of study

Cancer is known as a second global leading cause of death after the cardiovascular disease (Bae et al., 2010). In year 2008, about 12.66 million people were diagnosed with cancer which reflects to 188 cases for every 100,000 people. Out of 12.66 million of cancer patients, about 7.56 million died due to the cancer complication. Lung, breast, colorectum, liver and stomach are the major cancer site that mostly occurred in Asian region for both sexes(UK cancer research, 2013). The cancer deaths are believed to be continuously increased with estimation, 13.1 million of deaths occur in 2030 (World Health Organization, WHO, 2013).

Cancer is a class of disease which characterized by the uncontrollable growth a group of cells, invasion and metastasis. These three properties help in differentiating them from benign tumors, which are self-limited and do not invade or metastasized (Dhorajiya et al., 2012). Currently, many treatments have been implemented to control the uncontrollable growth of cancer cells. The recommended treatments for cancer patients are depending on the types and stages of cancers. Chemotherapy is