ISOLATION OF SOME BIOACTIVE COMPOUNDS FROM HELIOTROPIUM INDICUM LINN. AND ASYMMETRIC SYNTHESIS OF (+)-CASTANOSPERMINE

THEERAPHAN MACHAN

A THESIS SUBMITTED TO THE GRADUATE SCHOOL IN PARTIAL FULFILLMENT OF THE REQUIREMENTS FOR THE DEGREE OF DOCTOR OF PHILOSOPHY IN PHARMACY

ลิขสิทธิ์มหาวิทยาลัยเชียงใหม่ Copyright[©] by Chiang Mai University All rights reserved

> GRADUATE SCHOOL CHIANG MAI UNIVERSITY MAY 2007

ISOLATION OF SOME BIOACTIVE COMPOUNDS FROM HELIOTROPIUM INDICUM LINN. AND ASYMMETRIC SYNTHESIS OF (+)-CASTANOSPERMINE

THEERAPHAN MACHAN

THIS THESIS HAS BEEN APPROVED TO BE A PARTIAL FULFILLMENT OF THE REQUIREMENTS FOR THE DEGREE OF DOCTOR OF PHILOSOPHY IN PHARMACY

EXAMINING COMMITTEE B. Janving ath CHAIRPERSON Assoc.Prof. Dr. Boonsom Liawruangrath S. Lawruangrath MEMBER Assoc.Prof. Dr. Saisunee Liawruangrath MEMBER Dr. Apiwat Baramee MEMBER Prof. Dr. Stephen Geoffrey Pyne MEMBER Assoc.Prof. Dr. Duang—Buddhasukh MEMBER Dr. Pakawan Puangsombut MEMBER

11 May 2007 © Copyright by Chiang Mai University

ACKNOWLEDGEMENTS

I wish to express my sincere gratitude and appreciation to Associate Professor Dr. Boonsom Liawruangrath, my supervisor, for her very kind supervision, encouragement and inspiring discussion throughout this research work. I am also grateful for the opportunity she offered me to study in Ph.D program at the Faculty of Pharmacy, Chiang Mai University.

I would like to sincerely thank Professor Dr. Stephen G. Pyne, my co-supervisor, for his wisdom, kindness and patience throughout my period of Ph.D project at the Department of Chemistry, University of Wollongong, Australia. I am also thankful for the opportunity he offered me to learn organic synthesis and to attend Australian conferences and have fun life in Australia.

I would like to express my sincere thanks to my co-supervisors, Associate Professor Dr. Saisunee Liawruangrath and Lecturer Dr. Apiwat Baramee, for their guidance, discussions and support. I am also grateful to Lecturer Dr. Pakawan Puangsombut and Associate Professor Dr. Duang Buddhasukh, the external examiners, for their helpful comments and valuable suggestions.

I gratefully acknowledge to the Thailand Research Fund (TRF) under the Royal Golden Jubilee (RGJ) project for the financial support. Thanks are express to the University of Wollongong, Australia, and Graduate School, Chiang Mai University as well as the Commission on Higher Education, Ministry of Education, Thailand for their partial financial supports. The Department of Pharmaceutical Science, Faculty of Pharmacy, Chiang Mai University is also gratefully acknowledged for providing all laboratory facilities.

I would like to thank all the technical support staff in the Department of Chemistry, University of Wollongong, particularly Dr. John Korth for teaching me GC-MS techniques and helping my GC-MS experiments, Dr. Wiford Lie and Sandra Chapman for teaching me NMR techniques and helping my NMR experiments. Thanks are express to Larry Hick, and Roger Kanitz for helping me with MS experiments. I would like to thank Dr. Alison Ung for her very kind help and instruction in Pyne's lab.

My friends, who made me feel very warm and funny when I was in Australia, namely; Dr. Sarah Yong, Dr. Minyan Tang, Andrew Davis, Morwenna Williams, Dr. Joe Hartley, Dr. Johana Mberre, Dr. Uta Batenburg-Nguyen, Dr. Stephen Taylor, Joe Ambrus, Leena Burgess, Christopher Au, Qinyong Mao, Arife Yazici, and my Thai friends at the University of Wollongong are acknowledged for their help and friendships.

Special thanks are also grateful due to my lovely friends; namely Dr. Chalermporn Thongpoon, Piyaporn Srisom, Wisanu Thongchai, Suwanna Saisin, Kamolporn Panyakad, and all of graduated students in the Faculty of Pharmacy, for their useful help encouragement and happy time.

Finally, I wish to express my heartfelt gratitude to my father, my passed away mother, my sisters, my bothers-in-law, and my lovely nephews and niece for their source of inspiration, tender love care and endless encouragement.