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BIOACTIVE CHEMICAL CONSTITUENTS FROM  
*CHISOCHETON PENDULIFLORUS* AND *CF. AGLAIA ERYTHROSPERMA*

Miss Jarinporn Phongmaykin

A Dissertation Submitted in Partial Fulfillment of the Requirements  
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Thesis Title                    BIOACTIVE CHEMICAL CONSTITUENTS FROM  
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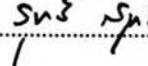
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จรินพร ผ่องเมญินทร์: องค์ประกอบทางเคมีที่มีฤทธิ์ทางชีวภาพจากหนาน奴ขี้ช้างและกล้อง  
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การศึกษาองค์ประกอบทางเคมีจากใบ เนื้อไม้และเปลือกต้นของหนาน奴ขี้ช้าง (วงศ์ Meliaceae) สามารถแยกสารในกลุ่ม aromadendrane sesquiterpene จำนวน 3 ชนิด คือ 14-hydroxyviridiflorol, (-)-10 $\beta$ ,13,14-trihydroxy-*allo*-aromadendrane และ 14-hydroxyepiviridiflorol ซึ่งเป็นสารชนิดใหม่ พร้อมกับ สารในกลุ่ม dammarane triterpene จำนวน 5 ชนิด คือ dammaradienone, cabraleadiol, eichlerialactone, cabralealactone และ cabraleahydroxylactone พนสารในกลุ่ม hexanortriterpene 1 ชนิด คือ hollongdione สารในกลุ่ม coumarin 2 ชนิด คือ scoparone และ scopoletin สารในกลุ่ม chromone 1 ชนิด คือ 5-hydroxy-7-methoxy-2-pentylchromone นอกจากนี้ยังพบสาร vanillic acid, ethyl orsellinate และ  $\beta$ -sitosterol glucoside สำหรับการศึกษาองค์ประกอบทางเคมีจากใบ เนื้อผล และเมล็ดของต้นกล้องซึ่งเป็นพืชอีกชนิดหนึ่ง ในวงศ์ Meliaceae สามารถแยกสารในกลุ่ม dammarane triterpene จำนวน 2 ชนิด คือ ethyl eichlerianoate และ aglinin A สารในกลุ่ม coumarin 2 ชนิด คือ scoparone และ scopoletin พนสารในกลุ่ม aglalactone 1 ชนิด คือ 5,6-desmethylenedioxy-5-methoxy-aglalactone นอกจากนี้ยังพบสารในกลุ่ม flavagline 1 ชนิด คือ 4'-demethoxy-3',4'-methylenedioxy-methyl rotaglate การพิสูจน์โครงสร้างทางเคมีของสารทั้งหมดที่สกัดแยกได้โดยอาศัยการวิเคราะห์เชิงสเปกตรัมของ UV, IR, MS และ NMR ร่วมกับการเปรียบเทียบข้อมูลของสารที่เคยมีการรายงานมาแล้ว พนว่าสาร cabraleadiol, cabraleahydroxylactone, cabralealactone, eichlerialactone, 14-hydroxyviridiflorol, (-)-10 $\beta$ ,13,14-trihydroxy-*allo*-aromadendrane, สารผอมระหว่าง 14-hydroxyviridiflorol และ 14-hydroxyepiviridiflorol ในอัตราส่วน 2:3, 5,6-desmethylenedioxy-5-methoxy-aglalactone, 4'-demethoxy-3',4'-methylenedioxy-methyl rotaglate และ ethyl eichlerianoate และ aglinin A และ aglinin A แสดงฤทธิ์ความเป็นพิษต่อเซลล์มะเร็ง ขณะที่สาร cabraleahydroxylactone, 5,6-desmethylenedioxy-5-methoxy-aglalactone, 4'-demethoxy-3',4'-methylenedioxy-methyl rotaglate, ethyl eichlerianoate และ aglinin A แสดงฤทธิ์ด้านเชื้อไวรัสเริโน และยังพบว่า 4'-demethoxy-3',4'-methylenedioxy-methyl rotaglate สามารถแสดงฤทธิ์ด้านเชื้ومากาเรียได้

สาขาวิชา: เภสัชเคมีและผลิตภัณฑ์ธรรมชาติ  
ปีการศึกษา 2549

ลายมือชื่อนิสิต..... จรินพร ผ่องเมญินทร์ .....  
ลายมือชื่ออาจารย์ที่ปรึกษา.....   
ลายมือชื่ออาจารย์ที่ปรึกษาร่วม.....  รองศาสตราจารย์ 

# # 4476979533:MAJOR PHARMACEUTICAL CHEMISTRY AND NATURAL PRODUCTS

KEY WORDS:*CHISOCHETON PENDULIFLORUS/CF. AGLAIA ERYTHROSPERMA/AROMADENDRANE-TYPE SESQUITERPENES/ DAMMARANE TRITERPENES/HEXANORTRITERPENES/ AGLALACTONES/ FLAVAGLINES*

JARINPORN PHONGMAYKIN: BIOACTIVE CHEMICAL CONSTITUENTS FROM *CHISOCHETON PENDULIFLORUS* AND CF. *AGLAIA ERYTHROSPERMA* THESIS  
ADVISOR: ASSOC. PROF. EKARIN SAIFAH, Ph.D., THESIS CO-ADVISOR: ASSOC.  
PROF. RUTT SUTTISRI, Ph.D., 335 pp.

Chemical investigation of the leaves, wood and stem bark of *Chisocheton penduliflorus* Planch. ex Hiern (family Meliaceae) led to the isolation of three aromadendrane sesquiterpenes, namely 14-hydroxyviridiflorol, (-)-10 $\beta$ ,13,14-trihydroxy-*allo*-aromadendrane and a new derivative, five dammarane triterpenes: dammaradienone, cabraleadiol, eichlerialactone, cabralealactone and cabraleahydroxylactone, a hexanortriterpene, hollongdione, two coumarins, scoparone and scopoletin, a chromone, 5-hydroxy-7-methoxy-2-pentylchromone, along with vanillic acid, ethyl orsellinate and  $\beta$ -sitosterol glucoside. Investigation of the leaves, fruits and seeds of another meliaceous plant, cf. *Aglaia erythrosperma* C.M. Pannell yielded two dammarane triterpenes, ethyl eichlerianoate and aglinin A, and two coumarins, scoparone and scopoletin. Furthermore, an aglalactone, 5,6-desmethylenedioxy-5-methoxy-aglalactone, and a flavagline, 4'-demethoxy-3',4'-methylenedioxy-methyl rotaglate, were also isolated. The structure determination of these compounds was accomplished by spectroscopic methods, including UV, IR, MS and NMR, and comparison with previously reported data. Cabraleadiol, cabraleahydroxylactone; cabralealactone, eichlerialactone, 14-hydroxyviridiflorol, (-)-10 $\beta$ ,13,14-trihydroxy-*allo*-aromadendrane, a 3:2 mixture of 14-hydroxyviridiflorol and 14-hydroxyepiviridiflorol, 5,6-desmethylenedioxy-5-methoxy-aglalactone, 4'-demethoxy-3',4'-methylenedioxy-methyl rotaglate and ethyl eichlerianoate exhibited antituberculosis activity. Cabraleadiol, cabraleahydroxylactone, cabralealactone, eichlerialactone, 5,6-desmethylenedioxy-5-methoxy-aglalactone, 4'-demethoxy-3',4'-methylenedioxy-methyl rotaglate, ethyl eichlerianoate and aglinin A displayed cytotoxic activity, whereas cabraleahydroxylactone, 5,6-desmethylenedioxy-5-methoxy-aglalactone, 4'-demethoxy-3',4'-methylenedioxy-methyl rotaglate and aglinin A also showed anti HSV-1 activity. Furthermore, 4'-demethoxy-3',4'-methylenedioxy-methyl rotaglate exhibited antimalarial activity.

Field of study Pharmaceutical Chemistry  
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## LIST OF ABBREVIATIONS AND SYMBOLS

$[\alpha]^{25}_D$	=	Specific rotation at 25° and sodium D line (589 nm)
<i>br s</i>	=	Broad singlet (for NMR spectra)
BSA	=	Bovine serum albumin
BuOH	=	Butanol
c	=	Concentration
°C	=	Degree Celsius
calcd	=	Calculated
CDCl <sub>3</sub>	=	Deuterated chloroform
cf.	=	Compare or consult
CFU	=	Colony forming unit
CHCl <sub>3</sub>	=	Chloroform
CH <sub>2</sub> Cl <sub>2</sub>	=	Dichloromethane
CD <sub>3</sub> OD	=	Deuterated methanol
cm	=	Centimetre
cm <sup>-1</sup>	=	Reciprocal centimeter (unit of wave number)
<sup>13</sup> C NMR	=	Carbon-13 Nuclear Magnetic Resonance
2D NMR	=	Two dimensional Nuclear Magnetic Resonance
<i>d</i>	=	Doublet (for NMR spectra)
<i>dd</i>	=	Doublet of doublets (for NMR spectra)
<i>ddd</i>	=	Doublet of doublets of doublets (for NMR spectra)
DEPT	=	Distortionless Enhancement by Polarization Transfer
DMSO	=	Dimethyl sulfoxide
DMSO- <i>d</i> <sub>6</sub>	=	Deuterated dimethyl sulfoxide
$\delta$	=	Chemical shift
ED <sub>50</sub>	=	Median effective dose
EIMS	=	Electron Impact Mass Spectrometry
ESIMS	=	Electrospray Ionization Mass Spectrometry
ESITOFMS	=	Electrospray Ionization Time of Flight Mass Spectrometry
EtOAc	=	Ethyl acetate
em.	=	emission
ex aff.	=	of affinity
ex.	=	excitation

**LIST OF ABBREVIATIONS AND SYMBOLS (continued)**

g	=	Gram
hr	=	Hour
$^1\text{H}$ NMR	=	Proton Nuclear Magnetic Resonance
$^1\text{H}$ - $^1\text{H}$ COSY	=	Homonuclear (Proton-Proton) Correlation Spectroscopy
HMBC	=	$^1\text{H}$ -detected Heteronuclear Multiple Bond Coherence
HMQC	=	$^1\text{H}$ -detected Heteronuclear Multiple Quantum Coherence
$\text{H}_2\text{O}$	=	Water
HREIMS	=	High Resolution Electron Impact Mass Spectrometry
HRESIMS	=	High Resolution Electrospray Ionization Mass Spectrometry
HSQC	=	Heteronuclear Single Quantum Correlation
Hz	=	Hertz
$\text{IC}_{50}$	=	Median Inhibitory Concentration
IR	=	Infrared Spectrum
$J$	=	Coupling constant
KBr	=	Potassium bromide
Kg	=	Kilogram
L	=	Liter
$\mu\text{g}$	=	Microgram
$\mu\text{Ci}$	=	MicroCurie
$\mu\text{l}$	=	Microliter
$\mu\text{M}$	=	Micromolar
$\lambda_{\max}$	=	Wavelength at maximal absorption
$\epsilon$	=	Molar absorptivity
$[\text{M}]^+$	=	Molecular ion
m	=	Metre
$m$	=	Multiplet (for NMR spectra)
$[\text{M}+\text{H}]^+$	=	Protonated molecular ion
MeOH	=	Methanol
mg	=	Milligram
MHz	=	Megahertz
MIC	=	Minimum inhibitory concentration
min	=	Minute

**LIST OF ABBREVIATIONS AND SYMBOLS (continued)**

ml	=	Milliliter
mM	=	Millimolar
mm	=	Millimeter
mp	=	Meling point
MS	=	Mass Spectrometry
MW	=	Molecular weight
<i>m/z</i>	=	Mass to charge ratio
NaCl	=	Sodium chloride
nm	=	Nanometer
NMR	=	Nuclear Magnetic Resonance
NOESY	=	Nuclear Overhauser Enhancement Spectroscopy
OD	=	Optical Density
PBS	=	Phosphate Buffer Saline
ppm	=	Part-per-million
<i>q</i>	=	Quartet (for NMR spectra)
<i>s</i>	=	Singlet (for NMR spectra)
$\nu_{\text{max}}$	=	Wave number at maximal absorption
rpm	=	round per minute
spp.	=	Species
TFA	=	Trifluoroacetic acid
<i>t</i>	=	Triplet (for NMR spectrum)
<i>tdd</i>	=	Triplet of doublet of doublets (for NMR spectrum)
THF	=	Tetrahydrofuran
TLC	=	Thin Layer Chromatography
UV	=	Ultraviolet