

ความเป็นพิษต่อเซลล์มะเร็งของสารประกอบไดเทอร์ปีนของเปลือกต้นเปล่าใหญ่

Croton oblongifolius Roxb. จังหวัดฉะเชิงเทรา



นางสาว พัชรา บุญญาณณี

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CYTOTOXITY TO CANCER CELL LINES OF DITERPENOID COMPOUNDS FROM STEM
BARKS OF *Croton oblongifolius* Roxb. FROM CHACHOENGSAO PROVINCE

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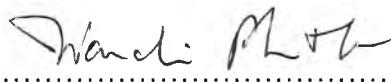
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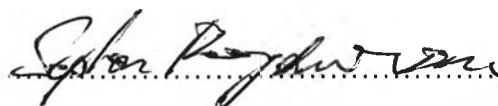
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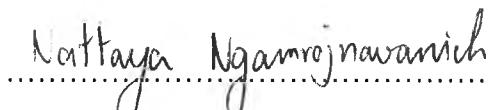
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ใหญ่ *Croton oblongifolius* Roxb. จังหวัดฉะเชิงเทรา (CYTOTOXICITY TO CANCER CELL
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การสกัดเปลือกต้นเปล้าใหญ่ (*Croton oblongifolius* Roxb.) จากจังหวัดฉะเชิงเทรา
ด้วยเยกเซนและเอธิลอะซิเดต สามารถแยกของผสมสเตอรอยด์ได้ 1 ชนิด และสารบิสุทธิ์อีก 6
ชนิด การหาสูตรโครงสร้างของสารเหล่านี้อาศัยคุณสมบัติทางกายภาพและเทคนิคทางสเปกต์
ร์สโคปี สารเหล่านี้คือ ของผสมของ stigmasterol, β -sitosterol และ campesterol (5), สาร
ประกอบเคลอโรเดนไดเทอร์ปีนอยด์ 2 ชนิดคือ (-)-hardwickiic acid (2) และ (-)-20-
benzyloxyhardwickiic acid (7) และสารประกอบแลบเดนไดเทอร์ปีนอยด์ 4 ชนิดคือ labda-
7,12(*E*),14-triene-17-oic acid (1), 2-acetoxy-labda-8(17),12(*E*),14-triene-3-ol (3), 3-
acetoxy-labda-8(17),12(*E*),14-triene-2-ol (4) และ labda-8(17),12(*E*),14-triene-2,3-diol
(6) และนำสารที่แยกได้มาทดสอบการยับยั้งเซลล์มะเร็ง Hep-G2 (ตับ), SW 620 (ลำไส้ใหญ่),
Chago (ปอด), Kato-3 (กระเพาะอาหาร) และ BT 474 (เต้านม) พบว่าสารประกอบ 3 และ 6 มี
ฤทธิ์ยับยั้งเซลล์มะเร็งทั้ง 5 ชนิด สารประกอบ 4 มีฤทธิ์ยับยั้งเซลล์มะเร็งตับและมะเร็งกระเพาะ
อาหารได้ในระดับปานกลาง และเป็นครั้งแรกในการรายงานผลการทดสอบการยับยั้งเซลล์
มะเร็งของ (-)-hardwickiic acid (2)

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| สาขาวิชา.....เคมีอินทรีย์..... | ลายมือชื่ออาจารย์ที่ปรึกษา..... |
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PATCHARA BUNYAMANEE : CYTOTOXICITY TO CANCER CELL LINES OF
DITERPENOID COMPOUNDS FROM STEM BARKS OF *Croton oblongifolius* Roxb.
FROM CHACHOENGSAO PROVINCE.

THESIS ADVISOR : Associate. Professor. AMORN PETSOM, Ph.D.

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Extraction of the stem barks of *Croton oblongifolius* Roxb. from Chachoengsao province with hexane and ethyl acetate gave a mixture of steroids and six compounds. The structures of these substances were established on the basis of physical properties and spectroscopic data. They were a mixture of stigmasterol, β -sitosterol and campesterol (5), two clerodane diterpenoids; (-)-hardwickiic acid (2) and (-)-20-benzyloxyhardwickiic acid (7) and four labdane diterpenoids; labda-7,12 (*E*),14-triene-17-oic acid (1), 2-acetoxy-labda-8(17),12(*E*),14-triene-3-ol (3), 3-acetoxy-labda-8(17),12(*E*),14-triene-2-ol (4) and labda-8(17),12(*E*),14-triene-2,3-diol (6). They were tested for their cytotoxicity against human tumor cell lines; Hep-G2 (hepatoma), SW 620 (colon), Chago (lung), Kato-3 (gastric) and BT 474 (breast). Compound 3 and 6 showed significant cytotoxicity against all cell lines. Compound 4 exhibited moderate cytotoxicity against Hep-G2 and Kato-3. Moreover, this is the first report of cytotoxicity test of (-)-hardwickiic acid (2).

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ABBREVIATIONS

| | | |
|---------------------------------|---|---|
| b.p. | = | Boiling point |
| br s | = | Broad singlet (for NMR spectra) |
| c | = | Concentration |
| ^o C | = | Degree Celsius |
| CDCl ₃ | = | Deuterated chloroform |
| CHCl ₃ | = | Chloroform |
| CH ₂ Cl ₂ | = | Dichloromethane |
| cm | = | Centimeter |
| ¹³ C-NMR | = | Carbon-13 nuclear magnetic resonance |
| COSY | = | Correlated Spectroscopy |
| d | = | Doublet (for NMR spectra) |
| dd | = | Doublet of doublet (for NMR spectra) |
| ddd | = | Doublet of doublet of doublet (for NMR spectra) |
| DEPT | = | Distortionless Enhancement by Polarization Transfer |
| DMSO | = | Dimethyl sulfoxide |
| δ | = | Chemical Shift |
| EI MS | = | Electron Impact Mass Spectrum |
| EtOAc | = | Ethyl acetate |
| g | = | Gram |
| ¹ H-NMR | = | Proton nuclear magnetic resonance |
| Hz | = | Hertz |
| HMBC | = | Heteromolecular Multiple Bond Correlation |
| HMQC | = | Heteromolecular Multiple Quantum Correlation |
| IR | = | Infrared spectrum |
| J | = | Coupling constant |

| | | |
|----------------|---|---|
| kg | = | Kilogram |
| L | = | Litre |
| M ⁺ | = | Molecular ion |
| mg | = | Milligram |
| MHz | = | Megahertz |
| ml | = | Millilitre |
| mm | = | Millimetre |
| m.p. | = | Melting point |
| MeOH | = | Methanol |
| M | = | Molar |
| <i>m/z</i> | = | Mass to charge ratio |
| M.W. | = | Molecular weight |
| MS | = | Mass spectrometry |
| No. | = | Number |
| NMR | = | Nuclear Magnetic Resonance |
| NOESY | = | Nuclear Overhauser Enhancement Spectroscopy |
| ppm | = | Part per million |
| q | = | Quartet (for NMR spectra) |
| s | = | Singlet (for NMR spectra) |
| t | = | Triplet (for NMR spectra) |
| TLC | = | Thin layer Chromatography |
| wt | = | Weight |
| R _f | = | Retention factor in chromatography |