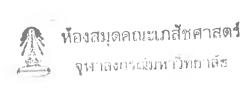
องค์ประกอบที่เป็นพิษต่อเชลล์จากเอื้องคำฝอย

นางสาวพรพรหม คล่องคำนวณการ



วิทยานิพนธ์นี้เป็นส่วนหนึ่งของการศึกษาตามหลักสูตรปริญญาเภสัชศาสตรมหาบัณฑิต สาขาวิชาเภสัชเวท ภาควิชาเภสัชเวทและเภสัชพฤกษศาสตร์ คณะเภสัชศาสตร์ จุฬาลงกรณ์มหาวิทยาลัย ปีการศึกษา 2556 ลิขสิทธิ์ของจุฬาลงกรณ์มหาวิทยาลัย





CYTOTOXIC CONSTITUENTS FROM DENDROBIUM BRYMERIANUM



Miss Pornprom Klongkumnuankarn

A Thesis Submitted in Partial Fulfillment of the Requirements

for the Degree of Master of Science in Pharmacy Program in Pharmacognosy

Department of Pharmacognosy and Pharmaceutical Botany

Faculty of Pharmaceutical Sciences

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Thesis Title CYTOTOXIC CONSTITUENTS FROM DENDROBIUM **BRYMFRIANUM** Miss Pornprom Klongkumnuankarn Ву Field of Study Pharmacognosy Thesis Advisor Associate Professor Boonchoo Sritularak, Ph.D. Thesis Co-Advisor Professor Kittisak Likhitwitayawuid, Ph.D. Accepted by the Faculty of Pharmaceutical Sciences, Chulalongkorn University in Partial Fulfillment of the Requirements for the Master's DegreeDean of the Faculty of Pharmaceutical Sciences (Assistant Professor Rungpetch Sakulbumrungsil, Ph.D.) THESIS COMMITTEE S. Amoypul. Chairman (Associate Professor Surattana Amnuoypol, Ph.D.) Bonchoo Snitularah Thesis Advisor (Associate Professor Boonchoo Sritularak, Ph.D.) C. Cikhit Thesis Co-Advisor (Professor Kittisak Likhitwitayawuid, Ph.D.) Pithi Chamorachotes Examiner (Assistant Professor Pithi Chanvorachote, Ph.D.) Superary Chami Examiner (Supakarn Chamni, Ph.D.) Veena Notealliam External Examiner

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การศึกษาทางพฤกษเคมีของสารสกัดหยาบจากเมทานอลของเอื้องคำฝอย สามารถแยก สารได้ทั้งสิ้น 8 ชนิด โดยสารที่แยกได้สามารถแบ่งออกได้เป็น 3 กลุ่ม คือ กลุ่ม bibenzyl (moscatilin gigantol และ tristin) กลุ่ม phenanthrene (flavanthrinin และ lusianthridin) และกลุ่ม fluorenone (nobilone dendroflorin และ denchrysan B) การพิสูจน์โครงสร้าง ทางเคมีของสารที่แยกได้ อาศัยการวิเคราะห์สเปกตรัมของ MS IR UV และ NMR ร่วมกับการ เปรียบเทียบกับข้อมลที่เคยมีรายงานมาก่อน และทดสอบฤทธิ์เป็นพิษต่อเชลล์ 2 ชนิด ได้แก่ เซลล์มะเร็งในช่องปากและเซลล์มะเร็งปอด พบว่า สารที่มีฤทธิ์เป็นพิษต่อเซลล์มะเร็งในช่องปาก ได้แก่ moscatilin flavanthrinin lusianthridin denchrysan B และ tristin โดยพบว่า moscatilin มีฤทธิ์ยับยั้งเซลล์มะเร็งชนิดนี้ได้แรงที่สุด โดยมีค่าความเข้มข้นที่สามารถทำให้เกิด การตายของเชลล์ได้ 50% (IC $_{50}$) คือ 2.62 μ M ขณะที่ flavanthrinin และ lusianthridin มีฤทธิ์ ปานกลาง โดยมีค่า IC50 79.67 และ 44.13 μ M ตามลำดับ ส่วน denchrysan B และ tristin มี ฤทธิ์ค่อนข้างอ่อน มีค่า IC₅₀ 158.91 และ 163.83 **µ**M ตามลำดับ โดยใช้ ellipticine (IC₅₀ 5.00 µM) และ doxorubicin (IC₅₀ 1.44 µM) เป็นชุดควบคุมผลบวก นอกจากนี้ เมื่อนำสารทั้ง 8 ชนิดมาทดสอบฤทธิ์เป็นพิษต่อเซลล์มะเร็งปอด พบว่า moscatilin gigantol lusianthridin และ dendroflorin สามารถยับยั้งการเจริญเติบโตของเซลล์มะเร็งปอดได้ดี โดยมีค่า IC_{50} เท่ากับ 674.04 85.40 268.60 และ 487.60 **µ**M ตามลำดับ เมื่อนำสารทั้งสี่ชนิดมาทดสอบฤทธิ์ antimigration ของเซลล์มะเร็งปอด พบว่า moscatilin และ dendroflorin แสดงฤทธิ์ antimigration ได้ดีที่สุด รองลงมาคือ lusianthridin และ gigantol ตามลำดับ

ภาควิชา เภสัชเวทและเภสัชพฤกษศาสตร์ สาขาวิชา เภสัชเวท ปีการศึกษา 2556

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> PORNPROM KLONGKUMNUANKARN: CYTOTOXIC CONSTITUENTS FROM DENDROBIUM BRYMERIANUM. ADVISOR: ASSOC. PROF. BOONCHOO SRITULARAK, Ph.D., CO-ADVISOR: PROF. KITTISAK- LIKHITWITAYAWUID, Ph.D., 189 pp.

Phytochemical study of the methanol extract from Dendrobium brymerianum (Orchidaceae) led to the isolation of eight compounds, which could be classified into 3 groups, consist of bibenzyls (moscatilin, gigantol and tristin), phenanthrenes (flavathrinin and lusianthridin) and fluorenones (nobilone, dendroflorin and denchrysan B). Their structure determinations were carried out by analysis of their spectroscopic data (UV, IR, MS ¹H NMR and ¹³C NMR) and comparison with previously reported data. These obtained compounds were evaluated for cytotoxic activity against KB (oral cavity cancer) and H460 (lung cancer) cell lines. The results suggested that, for the KB cells, moscatilin exhibited the strongest cytotoxic effect with an IC50 value of 2.62 μ M, whereas flavanthrinin, lusianthridin showed moderate activity (IC $_{50}$ 79.67 and 44.13 μ M, respectively), and denchrysan B and tristin showed weak activity (IC₅₀ 158.91 and 163.83 μ M, respectively). Ellipticine (IC₅₀ 5.00 μ M) and Doxorubicin (IC₅₀ 1.44 μ M) were used as a positive control. Moreover, for H460 cells, moscatilin, gigantol, lusianthridin and dendroflorin exhibited potential cytotoxic effect with IC₅₀ values of 674.04, 85.40, 268.60 and 487.60 \$\mu M\$, respectively. These compounds were further investigated for their anti-migration activity. The results indicated that at 48 hr moscatilin showed the strongest anti-migration effect, followed by dendroflorin, lusianthridin and gigantol.

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