Investigation of ß-glucuronidase inhibitor from the secondary

metabolites of the root of Neolitsea acuminatissima

<u>Hsiao-Jung Chou (周孝容)</u>^{1,a}, Chih-Chi Chang (張芷綺)^{1,a}, Chu-Hung Lin (林居宏)², Ih-Sheng Chen (陳益昇)², Hsun-Shuo Chang (張訓碩)², Tian-Lu Cheng (鄭添祿)³, **Horng-Huey Ko (柯宏慧)**^{1,*}

¹Department of Fragrance and Cosmetic Science, Kaohsiung Medical University ²School of Pharmacy, Kaohsiung Medical University ³Department of Biomedical Science and Environmental Biology, Kaohsiung Medical University

According to literature, the main factor induced diarrhea in chemotherapy is β glucuronidase of Escherichia coli (eßG). The authors desire to investigate eßG inhibitor, for the release of side-effect induced by chemotherapy. In the preliminary screen data, the methnolic extract of the root of Neolitsea acuminatissima (NA) showed the anti-eßG activity and without affecting human B-glucuronidase (hBG) activity. The results indicated that NA is a valuable source from which natural product-based medicinal products can be derived. In fact, the phytochemistry and the anti-eßG activity of the root of NA has not been investigated. Thus, the root of NA was selected as the candidate. The methanolic extract of NA (NARM) was partitioned with dichloromethane (NARD), ethyl acetate (NARE), n-butanol (NARB), and water (NARW) to obtain different soluble fractions. Bioassay-guided fractionation of the root of NARE and NARD, one new eudesmanolide type sesquiterpenes, methoxyneolitacumone A (8), one new alkaloid, demethoxydaibucarboline A (5), along with 8 known compounds, namely as zeorin (1), neolitacumone C (2), quercetin (3), dihydroquercetin (4), β -sitosterol (6), neolitacumone A (7), oplopanone (9), and epicatechin (11), were isolated and the structures were characterized spectroscopically. Among them, compounds 3 and 5 showed potent antieβG activity with inhibition ratio of 70-80%.

^aThese authors contributed equally in the study.

Key Words: *Neolitsea acuminatissima*, β-glucuronidase, diarrhea.