

Rotenoid compounds, isolated from dried root of plant and their inhibitory activities on cancer cells

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Background: 6-Deoxyclitriacetal (6-DA), isolated from dried roots of *Stemona collinsae* Craib, and derivatives of 6-DA modified at C-11 position with morpholine showed relative cytotoxic activities to doxorubin, a commercial drug, against various types of human carcinoma cell lines such as breast carcinoma cell (BT474), colon carcinoma (SW260), gastric carcinoma (KATO3), hepato carcinoma (HEP-G2), and lung carcinoma cell (CHAGO) [Sangthong, S., et al, 2011], therefore, in this work, we present the cytotoxicity of derivatives of stemone, stemonal compounds.

Objective: To enhance the cytotoxicity of rotenoid compounds

Methods: 6-Deoxyclitriacetal (6-DA), stemone and stemonal are rotenoid and its derivative were extracted from the dried roots of *Stemona collinsae* Craib using CH₂Cl₂, isolated by silica gel column chromatography and analyzed by TLC visualize using UV light. Tosylated stemone (1), 6-Deoxyclitriacetal di(4-dimethylamino) benzoate (3) and Stemone diamine (4) were synthesized. The structure of rotenoid compounds and its derivative as showed in figure 1. were confirmed by ¹H, ¹³C NMR spectroscopy and Mass spectroscopy. The cytotoxic activity was evaluated by the MTT assay against human cancer cell lines of; lung carcinoma (CHAGO), hepato carcinoma (Hep-G2), gastric carcinoma (KATO), colon carcinoma (SW260), and human breast carcinoma (BT474)

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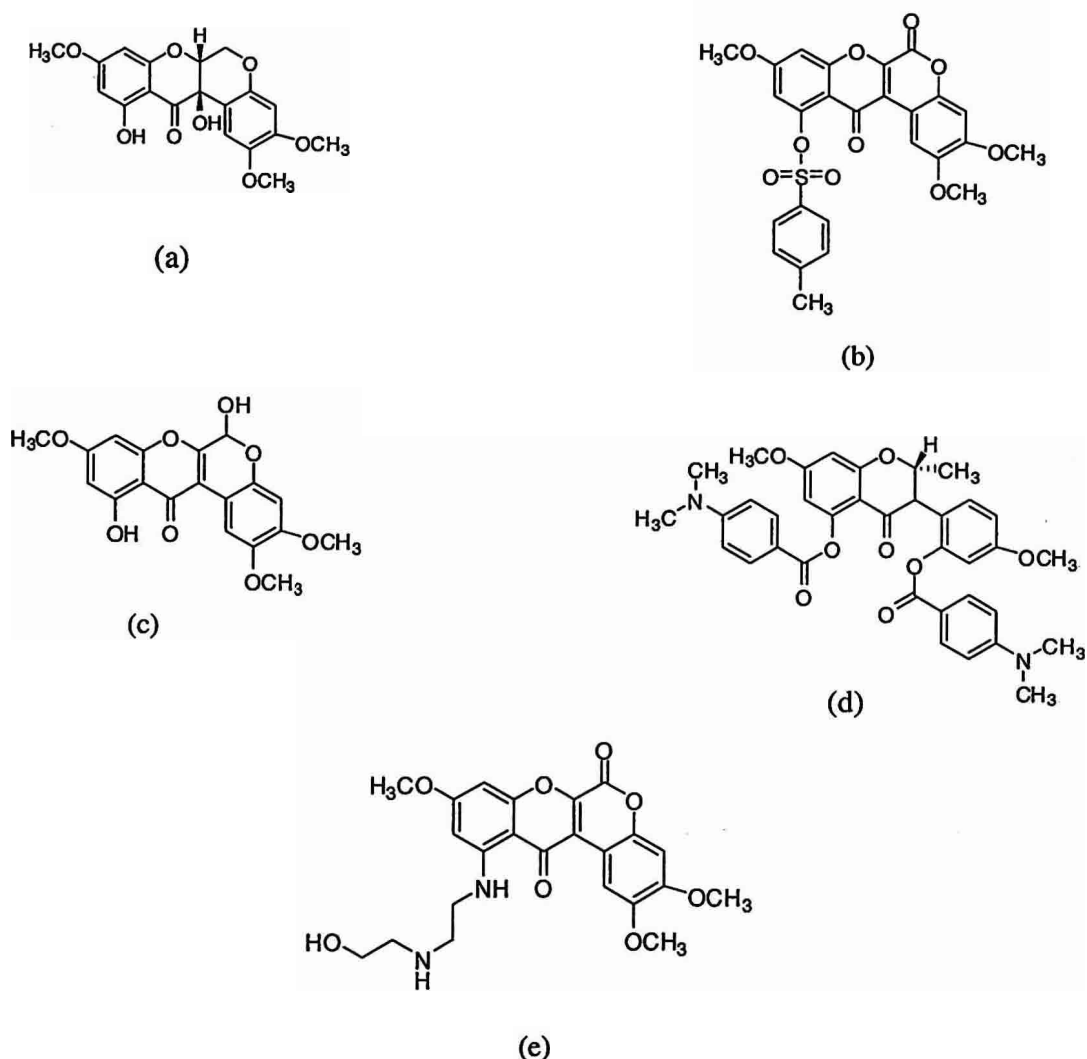


Figure 1. Representation the structure of (a) 6-DA, (b) tosylated stemonone, (c) stemonal , (d) 6-deoxyclitriacetal-di-(4-dimethylamino) benzoate and (e) stemonone diamine.

Results: 6-DA showed IC_{50} value against cancer cell lines; CHAGO and HEP-G2 at 5.098 and 4.980 μM , respectively, but are inactive in BT474, KATO and SW260 cell lines. Compound 1 and 3 are inactive against all tested cancer cell lines, while Compound 2 showed IC_{50} values of ??? against cancer cell lines; BT474, CHAGO and HEP-G2 but inactive cytotoxicity against KATO and SW260. For compound 4 showed active cytotoxicity against only CHAGO cancer cell line at IC_{50} value 3.687 μM .

Table 1. The inhibitory of activity (IC_{50}) of test compounds

Compound	IC_{50} (μM)				
	BT474	CHAGO	HEP-G2	KATO	SW260
Doxorubicin	1.53±0.45	2±0.32	1.23±0.39	Inactive	1.22±0.01
6-Deoxyclitriacetal	Inactive	5.1± 1.23	4.98±0.87	Inactive	Inactive
Compound 1	Inactive	Inactive	Inactive	Inactive	Inactive
Compound 2	4.84±0.22	4.96±0.19	4.37±0.73	Inactive	4.61±0.50
Compound 3	Inactive	Inactive	Inactive	Inactive	Inactive
Compound 4	Inactive	3.69±1.46	Inactive	Inactive	Inactive

Conclusion: Pre- study of this work demonstrated that stemonal (2) has a strong cytotoxic activity and against various type of cancer cell lines such as BT474, CHAGO, SW260 and HEP-G2. While 6-DA showed cytotoxicity to CHAGO and HEP-G2 cancer cell line less than stemonal. For tosylated stemonone (1), which is stemonone derivative, and 6-deoxyclitriacetal-di-(4-dimethylamino) benzoate (3) which is 6-deoxyclitriacetal derivative has no cytotoxic activity to cancer cell lines. For stemonone diamine (4) is the one of stemonone derivative showed strong cytotoxicity to CHAGO cancer cell line than its parent compound.

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