

**SINTESIS DAN UJI SITOTOKSISITAS TURUNAN KHALKON DAN
FLAVON TERHADAP SEL KANKER HeLa, T47D DAN WiDr
SECARA IN VITRO**

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INTISARI

Senyawa turunan khalkon dan flavon telah berhasil disintesis dan diuji sitotoksitas terhadap sel kanker. Senyawa khalkon (**1-2**) disintesis melalui reaksi kondensasi Claisen-Schmidt dari 2-hidroksiasetofenon atau 2,4-dihidroksiasetofenon dengan 6-bromoveratraldehida. Sintesis khalkon (**1-2**) dilakukan dengan refluks (65°C) selama 24 jam menggunakan metanol sebagai pelarut dan NaOH sebagai katalis. Senyawa flavon (**3-4**) disintesis melalui siklisasi oksidatif senyawa khalkon (**1-2**) dengan katalis iodin dalam pelarut dimetilsulfoksida (DMSO) pada kondisi refluks (120°C) selama 4 jam. Senyawa hasil sintesis dianalisis strukturnya dengan FTIR, GC-MS/MS-*direct*, KLT-*scanner*, ^1H - dan ^{13}C -NMR. Senyawa hasil sintesis diuji sitotoksitas terhadap sel kanker HeLa, T47D, dan WiDr dengan metode MTT assay.

Hasil penelitian menunjukkan bahwa senyawa **1** (2'-hidroksi-2-bromo-4,5-dimetoksikhalkon), senyawa **2** (2',4'-dihidroksi-2-bromo-4,5-dimetoksikhalkon), senyawa **3** (2'-bromo-4',5'-dimetoksiflavon) dan senyawa **4** (7-hidroksi-2'-bromo-4',5'-dimetoksiflavon) diperoleh dengan rendemen secara berurutan sebesar 78; 72; 28 dan 30%. Hasil uji sitotoksitas menunjukkan bahwa senyawa **1** memiliki aktifitas sedang dalam menghambat sel kanker WiDr dengan nilai IC_{50} sebesar $87,09\text{ }\mu\text{g/mL}$ sedangkan senyawa **2** memiliki aktifitas sedang dalam menghambat sel kanker HeLa dan WiDr dengan nilai IC_{50} sebesar 48,97 dan $47,86\text{ }\mu\text{g/mL}$. Uji sitotoksitas terhadap senyawa **3** dan **4** dinyatakan tidak aktif dalam menghambat sel kanker HeLa, T47D dan WiDr dengan nilai IC_{50} lebih dari $100\text{ }\mu\text{g/mL}$.

Kata kunci: khalkon, flavon, uji sitotoksitas, antikanker

SYNTHESIS AND IN VITRO CYTOTOXICITY TEST OF CHALCONE AND FLAVONE DERIVATIVES AGAINST HeLa, T47D AND WiDr CANCER CELLS

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ABSTRACT

Synthesis and cytotoxicity test against cancer cell lines of chalcone and flavone derivatives have been investigated. The chalcone derivatives (**1-2**) was synthesized by Claisen-Schmidt condensation of 2-hydroxyacetophenone or 2,4-dihydroxyacetophenone with 6-bromoveratraldehyde. The synthesis of chalcones (**1-2**) was carried out by reflux condition (65°C) for 24 hours using methanol as a solvent and NaOH as a catalyst. The flavone derivatives (**3-4**) were synthesized by oxidative cyclization of chalcone (**1-2**) with iodine as a catalyst in reflux condition (120°C) for 4 hours. The structures of synthesized compounds were characterized by FTIR, GC-MS/MS-direct, TLC-scanner, ¹H- and ¹³C-NMR. The cytotoxicity of compound (**1-4**) was tested against HeLa, T47D and WiDr cancer cell lines by MTT assay.

The results showed that 2'-hydroxy-2-bromo-4,5-dimethoxychalcone (**1**), 2',4'-dihydroxy-2-bromo-4,5-dimethoxychalcone (**2**), 2'-bromo-4',5'-dimethoxy flavone (**3**) and 7-hydroxy-2'-bromo-4',5'-dimethoxyflavone (**4**) produced 78, 72, 28 and 30,33% in yield, respectively. The cytotoxicity test indicated that compound **1** had a moderate activity for inhibiting the growth of WiDr cancer cells with IC₅₀ 87,09 µg/mL, while compound **2** had a moderate activity for inhibiting the growth of HeLa and WiDr cancer cells with IC₅₀ 48,97 and 47,86 µg/mL. The cytotoxicity test indicated that compound **3** and **4** were not active for inhibiting the growth of HeLa, T47D and WiDr cancer cells with IC₅₀ value more than 100 µg/mL.

Keyword: chalcone, flavone, cytotoxicity, anticancer