

Anticancer Activities of Some Novel Synthesized Thiazole and Thiohydantoin Derivatives

Yaser M. ALAHMADI ¹, Sultan S. AL-THAGFAN ¹, Sultan O. ALOLAYAN ¹,
Asma S. AL-WASIDI ², Nawal M. AL-JAFSHAR ², Amal M. AL-ANAZI ²,
Ali A. MOHAMAD ³ & Ahmed M. NAGLAH ^{4,5} *

¹ Clinical and Hospital Pharmacy Department, College of Pharmacy,
Taibah University, Madina Munawarah, Saudi Arabia

² Department of Chemistry, College of Science, Princess Nourah bint Abdulrahman University,
Riyadh 11671, Saudi Arabia

³ Invertebrate and Parasitology Department, Faculty of Science, Zagazig University, Zagazig, Egypt

⁴ Department of Pharmaceutical Chemistry, Drug Exploration & Development Chair (DEDC),
College of Pharmacy, King Saud University, Riyadh 11451, Saudi Arabia

⁵ Peptide Chemistry Department, Chemical Industries Research Division,
National Research Centre, 12622-Dokki, Cairo, Egypt

SUMMARY. A series of new compounds carrying 1,3-thiazole and 2-thiohydantoin scaffold were synthesized and structurally confirmed by different spectroscopic methods such as IR, ¹H-NMR, ¹³C-NMR and mass spectroscopy along with elemental analyses. All the synthesized compounds were examined for their cytotoxic activity versus HepG2 cell line. Compound 4 and its acetylated derivative showed the least IC50 values within the tested compounds. Additionally, compounds 4 and 6 were evaluated against VEGFR-2. The results revealed that compound 4 showed good VEGFR-2 inhibitory activity. Moreover, cell cycle analysis of compound 4 appeared cell cycle arrest at both G1 and G2/M phase of cell cycle profile of HepG2 cells.

RESUMEN. Se sintetizaron y confirmaron estructuralmente una serie de nuevos compuestos con andamiaje de 1,3-tiazol y 2-tiohidantoína mediante diferentes métodos espectroscópicos como IR, ¹H-NMR, ¹³C-NMR y espectroscopía de masas junto con análisis elementales. Todos los compuestos sintetizados fueron examinados por su actividad citotóxica versus la línea celular HepG2. El compuesto 4 y su derivado acetilado mostraron los menores valores de CI50 dentro de los compuestos probados. Además, los compuestos 4 y 6 se evaluaron frente a VEGFR-2. Los resultados revelaron que el compuesto 4 mostró una buena actividad inhibidora de VEGFR-2. Además, el análisis del ciclo celular del compuesto 4 pareció detener el ciclo celular tanto en la fase G1 como en la G2/M del perfil del ciclo celular de las células HepG2.

KEY WORDS: anticancer activities, cytotoxic activity, kinase assay, thiazole, thiohydantoin.

* Author to whom correspondence should be addressed. E-mail: amnaglah@gmail.com