

2nd International Conference on

PHARMACEUTICAL CHEMISTRY

October 02-04, 2017 Barcelona, Spain

Synthesis and molecular docking of novel non-Cytotoxic, anti-Angiogenic sulfonyl coumarin derivatives against hepatocellular carcinoma cells *In Vitro*

Eslam EL-SAWY¹, Manal EBAID¹, Hanaa RADY¹, Aziza SHALBY², Khadiga AHMED¹ and Heba ABO-SALEM¹
National Research Centre, Egypt

Resistance to conventional chemotherapy, leads to the need for development of novel safe and effective cancer therapies with new mechanism of action. Anti-angiogenic drugs are major example of such newly developed targeted therapeutics. In cancer drug development arena, coumarin-type compounds have been reported to possess marked cytotoxic activities, in addition act as novel angiogenesis inhibitors. In this respect, a new series of coumarin derivatives was synthesized starting from 2-oxo-2H-coumarin-6-sulfonyl chloride (**1**), 6-nitro-2-oxo-2H-coumarin-3-sulfonyl chloride (**10**) and 6-amino coumarin-2-one (**19**). Reaction of **1** or **10** with 2-cyanoacetohydrazide, 2'-acetyl-2-cyanoacetohydrazide or 3-amino-5-pyrazolone afforded pyrazoline derivatives. While reaction of **1** or **10** with malononitrile followed by reaction with hydrazine hydrate, urea, thiourea or guanidine led to the formation of pyrazole and pyrimidine derivatives. On the other hand, compound **19** on reaction with Vilsmeier-Haack reagent yielded the corresponding aldehyde **20**. Compound **20** under reaction with chlorosulfonylisocyanate afforded N-chlorosulfonyl amid which cyclized to give pyranobenzothiazine derivative **25**. The tested compounds **4**, **5**, **8**, **12**, **13** and **14** were non-cytotoxic against hepatocellular carcinoma cells (HepG2) using MMT. These non-cytotoxic compounds were evaluated as anti-angiogenic agent. Collectively, our results indicate that, coumarin molecules **4**, **5**, **8**, **13** and **14** can be utilized as lead compounds to develop potential non-toxic angiogenesis inhibitors and small molecular ligands to target (HepG2), which was in concomitant with molecular docking results. 1-Acetyl-5-amino-4-(2-oxo-2H-chromene-6-sulfonyl)-1,2-dihydro-pyrazol-3-one (**4**) considered a promising anti-angiogenic agent, where it exhibited MMP-dependent anti-migratory activity and down regulated CD105.

Biography

El-Sawy has completed her PhD at the age of 28 years from Al-Azher University and postdoctoral studies from National Research Centre, Cairo Egypt. She has published more than 34 papers in reputed journals. Current position: Professor Doctor of Organic Chemistry at Chemistry of Natural Compounds Department, National Research Centre (NRC).

eslamelsawy@gmail.com

Notes: