



Synthesis of Piperidine and Morpholine Amides of Ferulic Acid and their Bioactivity against P-388 Leukemia Cells

Firdaus*¹, Dirayah Rauf Husain², Tajuddin Naid³, Seniwati¹,
Nunuk Hariani Soekamto¹, Sabir Sumarna¹, and Muhammad Fajar Islam¹

¹Departement of Chemistry, Faculty of Mathematic and Natural Sciences, Hasanuddin University, Makassar 90245, Indonesia

²Departement of Biology, Faculty of Mathematic and Natural Sciences, Hasanuddin University, Makassar 90245, Indonesia

³Faculty of Pharmacy, Hasanuddin University, Makassar 90245, Indonesia

Abstract: Synthesis of *N*-feruloylpiperidine (**5a**) and *N*-feruloylmorpholine (**5b**) from ferulic acid through acetylation, chlorination, amidation, and deacetylation reactions have been conducted. The acetylation was carried out using acetic anhydride reagent in pyridine solvent at room temperature for 6 hours. The chlorination was performed with thionyl chloride in benzene solvent by refluxing at 75°C for 4 hours, proceeded by *in situ* amidation using piperidine to synthesize of compound **1** and morpholine to synthesize of compound **2** in the presence of triethylamine and pyridine in dichloromethane solvent at room temperature. The deacetylation was performed using pyrrolidine reagent in ethyl acetate solvent at room temperature for 2 hours giving compounds **5a** and **5b** as yellowish crystalline solids with m.p. of 127-129°C and 151-153°C, respectively. Characterization of these compounds was committed by FTIR spectrophotometer and NMR spectrometer. The bioassay of the both compounds against P-388 leukemia cells gave IC₅₀ of 46.67 and 57.10 µg/mL, respectively.

Keywords: *N*-feruloylpiperidine, *N*-feruloylmorpholine, anticancer, ferulic acid, P-388 leukemia cell.