

Accepted Manuscript

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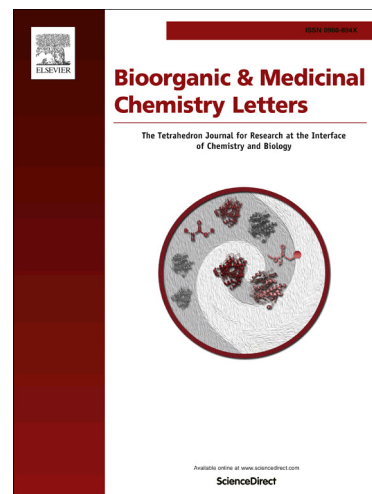
PII: S0960-894X(15)00396-0
DOI: <http://dx.doi.org/10.1016/j.bmcl.2015.04.064>
Reference: BMCL 22647

To appear in: *Bioorganic & Medicinal Chemistry Letters*

Received Date: 24 February 2015
Revised Date: 17 April 2015
Accepted Date: 20 April 2015

Please cite this article as: Mutai, P., Pavadai, E., Wiid, I., Ngwane, A., Baker, B., Chibale, K., Synthesis, antimycobacterial evaluation and pharmacophore modeling of analogues of the natural product formononetin, *Bioorganic & Medicinal Chemistry Letters* (2015), doi: <http://dx.doi.org/10.1016/j.bmcl.2015.04.064>

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Synthesis, antimycobacterial evaluation and pharmacophore modeling of analogues of the natural product formononetin

Peggoty Mutai^a, Elumalai Pavadai^a, Ian Wiid^b, Andile Ngwane^b, Bienyameen Baker^b, Kelly Chibale^{a,c,d*}

^aDepartment of Chemistry, University of Cape Town, Rondebosch 7701, South Africa

^bDST-NRF Centre of Excellence for Biomedical Tuberculosis Research, SAMRC Centre for Tuberculosis Research, Division of Molecular Biology and Human Genetics, Faculty of Medicine and Health Sciences, Stellenbosch University, PO Box 19063, Tygerberg 7505, South Africa

^cSouth African Medical Research Council Drug Discovery and Development Research Unit, University of Cape Town, Rondebosch 7701, South Africa

^dInstitute of Infectious Disease and Molecular Medicine, University of Cape Town, Rondebosch 7701, South Africa

*Corresponding author

Email address: Kelly.Chibale@uct.ac.za (K. Chibale)

Abstract

The synthesis and antimycobacterial activity of formononetin analogues is hereby reported. Formononetin and its analogue **11E** showed 88% and 95% growth inhibition, respectively, against the H37Rv strain of *Mycobacterium tuberculosis*. Pharmacophore modeling studies indicated that the presence of a hydroxyl group in formononetin and its analogues, is crucial for maintaining activity.

Keywords Antimycobacterial, Formononetin, Pharmacophore, Structure, Activity

Tuberculosis (TB) is a leading cause of mortality and morbidity, being estimated to infect about one third of the world's population¹. The Global tuberculosis report by WHO in 2013 reported that there were an estimated 8.6 million incidents of TB in 2012 and 1.3 million TB related deaths². The HIV scourge has made the TB situation worse, especially in sub-Saharan Africa, with the co-epidemic being particularly concentrated in the Southern African countries where HIV prevalence is high³. Increasing rates of drug-resistant tuberculosis are a significant concern and pose serious implications for current and future treatment of the disease⁴. Resistance has been reported for new drugs such as bedaquiline⁵, indicating that there is an urgent need for more new drug candidates to raise the probability of developing a novel, short-course and safe 'universal' regimen applicable to drug-susceptible and all forms of drug-resistant TB⁶.

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