Accepted Manuscript

Anti-*Mycobacterium tuberculosis* activity of naphthoimidazoles combined with isoniazid and rifampicin

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Tuberculosis

Taberculosis

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PII: \$1472-9792(18)30131-8

DOI: 10.1016/j.tube.2018.06.015

Reference: YTUBE 1726

To appear in: Tuberculosis

Received Date: 26 March 2018

Revised Date: 4 June 2018
Accepted Date: 29 June 2018

Please cite this article as: Corrêa Barros LéPacheco, Del Rio KP, Carvalho TatianedosSantosConceiçã, Pinto MdCFR, de Moura KCG, Halicki PCB, Ramos DF, da Silva PEA, Anti-*Mycobacterium tuberculosis* activity of naphthoimidazoles combined with isoniazid and rifampicin, *Tuberculosis* (2018), doi: 10.1016/j.tube.2018.06.015.

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ABSTRACT

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Tuberculosis (TB) is the cause of more than one million deaths worldwide, and despite being a curable disease, some factors can make therapy difficult, emphasizing the need for the development of new drugs that may potentiate the action of the classic anti-TB antimicrobials. **Naphthoimidazoles** show broad spectrum of biological activities, including a antimycobacterial activity. The aim of this study was to evaluate the anti-Mycobacterium tuberculosis activity of nine naphthoimidazoles, alone and combined with isoniazid (INH) and rifampicin (RIF). We evaluated the minimum inhibitory concentration (MIC) of the compounds, the fractional inhibitory concentration of the combinations of the naphthoimidazoles with INH or RIF, and the cytotoxicity of these compounds. Eight compounds showed MICs ranging from 1.56 to 25 µg/mL and the presence of substituents on phenyl groups shown to be essential for antimycobacterial activity. Four compounds showed additivity with both INH and RIF and showed SI values higher than 10, indicating safety. Thus, considering the antimycobacterial activity and the absence of antagonism between naphthoimidazoles and the two main drugs for TB treatment, these compounds could be scaffolds for the development of new anti-TB drugs.

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Key words: Tuberculosis; new drugs; naphthoimidazoles.

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