Accepted Manuscript

Synthesis and evaluation of novel azoles as potent antifungal agents

Liangjing Li, Hao Ding, Baogang Wang, Shichong Yu, Yan Zou, Xiaoyun Chai, Qiuye Wu

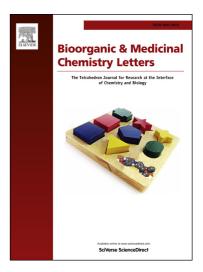
PII: S0960-894X(13)01316-4

DOI: http://dx.doi.org/10.1016/j.bmcl.2013.11.037

Reference: BMCL 21064

To appear in: Bioorganic & Medicinal Chemistry Letters

Received Date: 29 July 2013 Revised Date: 23 October 2013 Accepted Date: 16 November 2013



Please cite this article as: Li, L., Ding, H., Wang, B., Yu, S., Zou, Y., Chai, X., Wu, Q., Synthesis and evaluation of novel azoles as potent antifungal agents, *Bioorganic & Medicinal Chemistry Letters* (2013), doi: http://dx.doi.org/10.1016/j.bmcl.2013.11.037

This is a PDF file of an unedited manuscript that has been accepted for publication. As a service to our customers we are providing this early version of the manuscript. The manuscript will undergo copyediting, typesetting, and review of the resulting proof before it is published in its final form. Please note that during the production process errors may be discovered which could affect the content, and all legal disclaimers that apply to the journal pertain.

ACCEPTED MANUSCRIPT

Synthesis and evaluation of

novel azoles as potent antifungal agents

Liangjing Li a,b,† , Hao Ding a,† , Baogang Wang a , Shichong Yu a , Yan Zou a , Xiaoyun Chai a,* , Qiuye Wu a,*

^aDepartment of Organic Chemistry, School of Pharmacy, Second Military Medical University, 325 Guohe Road, Shanghai 200433, China

^bCadet Company 2nd, Faculty of Naval Medicine, Second Military Medical University, 800 Xiangyin Road, Shanghai 200433, China

Abstract: Using a rational approach to the design of antifungal agents, a series of azole agents with 1,3,4-oxadiazole side chains were designed and synthesised. The results of preliminary *in vitro* antifungal tests with eight human pathogenic compounds showed that all of the title compounds exhibited excellent activities against all of the tested fungi except *Aspergillus fumigatus*. Compounds **11e** and **11f** were found to be the most effective, with a minimum inhibitory concentration of 0.0039 μg/mL, followed by voriconazole, which has a MIC of 0.0625 μg/mL. The 1,3,4-oxadiazole side chain is not the major contributor but plays a role in eliciting the observed antifungal activity.

Fungal infections, which are an increasing healthcare concern worldwide, are associated with significant costs, morbidity, and mortality.¹⁻³ Although the number and types of antifungal drugs have markedly increased, only a few of the identified drugs can be used in the clinic.⁴ The commonly used antifungal agents are azoles, polyenes, and echinocandins.^{3,5} Among these antifungal drugs, the gold standard for

..

^{*}Corresponding authors. Tel: +86 21 81871228 (Xiaoyun Chai), Tel./fax: +86 21 81871225 (Qiuye Wu).

E-mail addresses: chaixy1207@163.com (Xiaoyun Chai), wuqy640319@163.com (Qiuye Wu).

[†] These two authors contributed equally to this work.

Download English Version:

https://daneshyari.com/en/article/10592793

Download Persian Version:

https://daneshyari.com/article/10592793

<u>Daneshyari.com</u>