## **Exploring Quinoxaline Derivatives as Potential Antifungal Agents: A Comprehensive Study on Candida and Aspergillus Species Susceptibility**

Abdelbagi Alfadil, MD, Ph.D\*,\*\*\* Hamoud Alsamhan, Ph.D\*\* Ahmed Ali, Ph.D\*\* Huda Alkreathy, MD, Ph.D\*\* Mohammad W. Alrabia, MD, Ph.D\*,\*\*\* Bandar Hasan Saleh, MD Ph.D\* Sameer E.M. Alharthi, Msc, PhD\* Hani Zakareya Asfour, Msc, PhD\* Nabil A. Alhakamy, Msc,Ph. D\*\*,\*\*\*\* Karem Ibrahem, MD, PhD\*

## ABSTRACT

The escalating prevalence of fungal diseases presents a significant global health concern, impacting millions of individuals annually. Fungal infections encompass a spectrum of conditions ranging from superficial to life-threatening systemic diseases, with Candida, Aspergillus, and Cryptococcus species posing substantial threats. Particularly, Candida albicans is a leading cause of invasive fungal infections, including candidiasis and candidemia, with significant mortality rates and economic burdens. Addressing the challenges posed by fungal infections requires innovative therapeutic approaches, especially in the face of emerging drug-resistant strains such as Candida auris. In this context, the antifungal potential of quinoxaline derivatives, exemplified by 2,3-dimethylquinoxaline (DMQ), has garnered attention for its efficacy against fungal infections and wound healing properties. Building on previous research, our study aims to explore the antifungal activity of various quinoxaline compounds against Candida and Aspergillus species. Notably, compounds such as 2-Methyldibenzo[f,h] quinoxaline and 4-Chloro-7-fluoropyrrolo[1,2-a]quinoxaline demonstrated significant activity against specific Candida strains, while 2-Chloro-6-(trifluoromethyl)quinoxaline and 2-Chloro-7-(trifluoromethyl)quinoxaline exhibited efficacy against multiple Candida species. However, none of the tested compounds showed activity against certain Candida and Aspergillus strains, highlighting the diverse responses of different fungal species to these compounds. The observed variations in antifungal activity can be attributed to differences in chemical structure, mode of action, and target specificity among the compounds. While some compounds effectively target fungal vulnerabilities, others may lack the necessary biochemical properties for interaction with fungal cells. In conclusion, our study underscores the need for continued research to develop effective antifungal therapies capable of combating a broad spectrum of fungal infections. By elucidating the molecular mechanisms underlying compound efficacy and exploring novel therapeutic strategies, we can address the evolving challenges posed by fungal diseases and improve patient outcomes.

Keywords: Candida, Aspergillus, Quinoxaline derivative, drug resistance

Bahrain Med Bull 2025; 47 (1): 2671-2675

*	Department of Clinical Microbiology and Immunology,
	Faculty of Medicine, King Abdulaziz University, Jeddah
	Saudi Arabia.
	E-mail: kaibrahem@kau.edu.sa
**	Department of Pharmacology, Faculty of Medicine
	King Abdulaziz University, Jeddah, Saudi Arabia.
***	Center of Research Excellence for Drug Research
	and Pharmaceutical Industries
	King Abdulaziz University, Saudi Arabia.
****	Department of Pharmaceutics
	Faculty of Pharmacy, King Abdulaziz University, Jeddah, Saudi Arabia
****	Mohamed Saeed Tamer Chair for Pharmaceutical Industries
	King Abdulaziz University, Jeddah 21589, Saudi Arabia.