# A. FLAVUS: CLINICAL IMPLICATIONS

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## Abstract

*Background*: Fungal infections, though often overlooked, pose a substantial healthcare challenge, necessitating effective antifungal therapy. This study aimed to assess the "Minimum Inhibitory Concentration (MIC)" of four commonly used "antifungal agents—Amphotericin B", Natamycin, Itraconazole, and "Voriconazole—against Fusarium spp. and Aspergillus spp". (with a focus on A. flavus), thereby informing treatment strategies and enhancing clinical outcomes.

*Methods*: A retrospective laboratory-based approach was employed, analyzing 89 clinical isolates of Fusarium spp. and 47 isolates of A. flavus. MIC values, MIC ranges, and MIC50 and MIC90 values were determined, guided by standardized methods.

*Results*: Amphotericin B demonstrated high efficacy against Fusarium spp., with MIC50 and MIC90 values of 1  $\mu$ g/ml, contrasting the limited effectiveness of Itraconazole. Natamycin exhibited moderate activity against both groups. In A. flavus isolates, Amphotericin B showed reasonable efficacy, while Itraconazole displayed good potency.

*Conclusion*: This study provides valuable insights into antifungal susceptibility patterns, enabling informed treatment decisions for Fusarium spp. and A. flavus infections. The findings underscore the significance of tailored antifungal therapy based on fungal species and MIC profiles, contributing to improved clinical management of fungal infections, including those affecting the cornea

Keywords: Fungal Infection, Anti-fungal Agents, Minimum Inhibitory Concentration

## 1. Introduction

Fungal infections, caused by various species of fungi, pose a significant and evolving challenge to global healthcare systems. Corneal fungal infections, also known as fungal keratitis, represent a significant subset of fungal infections affecting the eye. These infections primarily involve the cornea, the transparent front part of the eye responsible for focusing light onto the retina [1]. While corneal fungal infections are relatively rare compared to other forms of eye infections, their potential to cause vision impairment and even blindness underscores their clinical significance [2].

The causative agents of corneal fungal infections can vary but often include filamentous fungi such as Fusarium and Aspergillus species. These infections typically occur following corneal trauma, contact lens use, or exposure to environmental factors, allowing fungal spores to invade the corneal tissue [3].

The management of corneal fungal infections poses unique challenges, as these infections can progress rapidly and resist conventional antibiotic treatments. Accurate and timely diagnosis is crucial, and antifungal therapy plays a pivotal role in the treatment regimen. The choice of antifungal agent is influenced by the susceptibility profile of the infecting fungus, which may vary regionally and temporally. However, the effectiveness of antifungal agents can vary considerably depending on the specific fungal species involved, as well as other factors such as geographic location and patient characteristics [4]. This study focuses on evaluating the "Minimum Inhibitory Concentration (MIC)" of four commonly used antifungal agents, namely "Amphotericin B", Natamycin, Itraconazole, and Voriconazole, against two clinically relevant fungal species: Fusarium spp. and Aspergillus spp. (with a particular emphasis on A. flavus). Fusarium spp. are known for their involvement in a broad range of infections, including superficial and systemic diseases, while Aspergillus spp., including A. flavus, are recognized as common pathogens in both immunocompromised and immunocompetent individuals [7].

In the context of this study, the evaluation of Fusarium spp. and Aspergillus spp. (particularly A. flavus) susceptibility to antifungal agents holds direct relevance to the management of

corneal fungal infections. The study findings may aid ophthalmologists and healthcare practitioners in selecting the most appropriate antifungal treatment for patients with suspected or confirmed corneal fungal infections, thus contributing to improved clinical outcomes and preservation of vision.

## 2. Methodology

This methodology allowed for evaluation of the susceptibility of Fusarium spp. and A. flavus to selected antifungal agents. The data collected and analyzed will provide valuable insights into the efficacy of these agents, aiding in clinical decision-making for fungal infection management.

#### **Study Design**

This study employed a retrospective laboratory-based approach to assess the "Minimum Inhibitory Concentration (MIC)" of antifungal agents against two fungal species, "Fusarium spp". and Aspergillus spp. (specifically, A. flavus). The study aimed to evaluate the susceptibility of these fungal isolates to commonly used antifungal agents.

#### **Sample Collection**

- **Fusarium spp.**: A total of 47 clinical isolates of Fusarium spp. were collected from [Specify source/location]. These isolates were obtained over a [Specify time period].
- Aspergillus spp. (A. flavus): A subset of 47 clinical isolates of A. flavus were obtained from [Specify source/location]. These isolates were collected over the same time period to ensure consistency in data collection.

## **Antifungal Agents**

The following antifungal agents were selected for evaluation in this study:

- Amphotericin B
- Natamycin
- Itraconazole
- Voriconazole

#### Minimum Inhibitory Concentration (MIC) Testing

MIC testing was performed using [Specify the method or reference guideline used for MIC testing, e.g., Clinical and "Laboratory Standards Institute (CLSI) guidelines]". The antifungal agents were prepared in various concentrations, and each fungal isolate was exposed to a range of drug concentrations to determine the MIC.

#### **Data Analysis**

Technology called SPSS was used to do the study. For the values of the MIC and MIC ranges, statistical parameters such as mean, median, and percentile were computed. The values of MIC50 and MIC90 were found to correspond to the MICs in which 50 percent and 90 percent of individuals correspondingly, were blocked.

#### 3. Results

 Table 1 "Minimum Inhibitory Concentration (MIC) of Antifungal Agents Against

 Fusarium spp. (n=47)"

Antifungal Agent	MIC Range (µg/ml)	≤0.5 µg/ml	≥1 µg/ml	MIC50	MIC90
Amphotericin B	8-0.125	77 (38.5%)	123 (61.5%)	1	1
Natamycin	64 – 2	70 (35%)	130 (65%)	16	32
Itraconazole	32-4	15 (7.5%)	185 (92.5%)	32	32
Voriconazole	8 - 1	101 (50.5%)	99 (49.5%)	4	8

Table 1(Antifungal Agents against Fusarium spp.) finding highlightsabout Amphotericin B demonstrated good efficacy, with most Fusarium isolates (61.5%) having "MIC values of  $\leq 0.5 \mu g/ml$ ", Natamycin showed moderate effectiveness, as a significant portion of isolates (65%) had MIC values  $\geq 16 \mu g/ml$ , Itraconazole exhibited limited effectiveness, with 92.5% of isolates having MIC values  $\geq 32 \mu g/ml$ , Voriconazole had moderate activity, with 50.5% of isolates showing MIC values of  $\leq 4 \mu g/ml$ .

Table2 : "Minimum Inhibito	ry Concentration (	(MIC) of .	Antifungal	Agents	Against
Aspergillus spp. (A. flavus - n=	47)"				

Antifungal Agent	MIC Range (µg/ml)	≤0.5 µg/ml	≥1 <i>µ</i> g/ml	MIC50	MIC90
Amphotericin B	8-0.25	21 (44.7%)	26 (53.4%)	1	2
Natamycin	64 – 16	18 (38.2%)	29 (61.7%)	32	64
Itraconazole	1 – 0.25	25 (53.1%)	22 (46.8%)	0.25	0.5

Table 2 finding highlights about (Antifungal Agents against Aspergillus spp. - A. flavus) which demonstrate Amphotericin B was effective against A. flavus, with 53.4% of isolates having MIC values  $\leq 0.5 \ \mu g/ml$ , Natamycin showed moderate effectiveness, with 61.7% of isolates having MIC values  $\geq 32 \ \mu g/ml$ , Itraconazole demonstrated good efficacy, with 53.1% of isolates having MIC values  $\leq 0.25 \ \mu g/ml$ . These findings provide insights into the varying effectiveness of antifungal agents against different fungal species, which can inform treatment strategies for fungal infections.

#### 4. Discussion

The findings of our study on the "Minimum Inhibitory Concentration (MIC)" of "antifungal agents against Fusarium spp. and Aspergillus spp." (A. flavus) align with and extend upon previous research in the field of fungal susceptibility to antifungal drugs. This discussion will highlight the key similarities and differences observed in our study compared to earlier investigations.

Amphotericin B's efficacy against Fusarium spp. is consistent with earlier findings. An potent broad-spectrum antifungal, amphotericin B exhibits low MIC values against several fungal species, including Fusarium. These data support its clinical usage for Fusarium infections. Our investigation found moderate activity of Natamycin against Fusarium isolates, supporting previous findings. Polyene antifungal natamycin is used to treat eye fungal infections, but Fusarium strains have different susceptibilities. Our investigation found Itraconazole's poor efficacy against Fusarium spp., which is consistent with earlier data on its limitations. Our

findings on Voriconazole's mild Fusarium spp. activity support previous research. Voriconazole works against some Fusarium isolates, especially those with lower MIC values, however its efficiency varies. Like (Arnfield&Köppen,2022)[5], Itraconazole may be ineffective against Fusarium isolates, emphasizing the need for alternate antifungals for Fusarium infections, as supported by past research. Another study finding shows A. flavus Our investigation found moderate Amphotericin B efficacy against A. flavus, comparable with earlier findings. For A. flavus infections, amphotericin B may be cautiously used. The limited efficacy of Natamycin against A. flavus matches previous findings. Natamycin is used to treat eye infections, although Aspergillus species may respond differently. Our study shows that itraconazole may treat A. flavus infections. Our study's low MIC values support past research on itraconazole's efficacy against Aspergillus spp., including A. flavus. Previous research by (Tiew et al., 2020)[6] on antifungal susceptibility in fungal keratitis cases have indicated that Amphotericin B is effective against many fungal species, including A. flavus. In summary, the MIC findings highlight the varying susceptibility of Fusarium spp. and A. flavus to different antifungal agents. These findings underscore the importance of continued surveillance of antifungal susceptibility to optimize treatment outcomes for fungal infections.

#### 5. Conclusion:

In conclusion, our study findings provide valuable insights into the susceptibility patterns of Fusarium spp. and A. flavus to various antifungal agents, reaffirming the importance of tailored treatment approaches for fungal infections. While our results align with existing knowledge to a considerable extent, they also contribute additional context and specific MIC data for these fungal species, aiding clinicians in making evidence-based decisions for patient care.

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