

Increasing trend of mucormycosis mixed with other invasive mycoses from a tertiary care centre in north India

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Introduction

- Mucormycosis is an angioinvasive, aggressively devastating and lifethreatening fungal infection essentially requiring timely intervention thereby prompt treatment.
- The incidence of mucormycosis, mixed with other invasive mycoses is increasing gradually with rise in the vulnerable population, especially among those presenting with uncontrolled diabetes mellitus.
- During recent times, we encountered mucormycosis mixed with other mycoses, whereby there was concomitant infection with hyaline or dematiaceous molds and/or yeasts.
- The present study intends to evaluate the increasing trend of mucormycosis along with such infections and the underlying significant risk factors with poor prognosis if timey intervention is not instituted.

Material and Methods

- A total of seventeen cases of mixed infection occurred during last five years from January 2013 to December 2017.
- Biopsy tissue from nasal cavity in rhino-orbital infection, necrotic tissue from cutaneous infection and sputum in case of pulmonary mucormycosis was processed for conventional mycological examination.
- Fungal etiology was established by direct examination of KOH wet mount, CFW staining and fungal culture on media like SDA and BHIA.
- The morphological identification of fungal isolates was done by LCB preparation.
- Histopathology study of tissue sections stained by hematoxylin and eosin (H&E), periodic acid-Schiff (PAS), Gomori methenamine silver (GMS) and Masson-Fontana (MF) stainings was done.
- The final diagnosis of isolates was established on the basis of molecular identification done by sequencing ITS region of isolates and compared with those of type strains.
- In vitro antifungal susceptibility testing of all mucoralean fungi was done by broth microdilution method according to the Clinical and Laboratory Standard Institute (CLSI) document M38-A2.

Results

- Thirteen cases were rhino-orbital clinical type, two each of pulmonary and cutaneous types of mucormycosis.
- Rhizopus arrhizus and Aspergillus flavus were isolated from six rhinoorbital cases and one from pulmonary case.
- Rhizopus arrhizus and Alternaria alternata were isolated from one rhino-orbital case while Lichtheimia corymbifera and Aspergillus *flavus* were isolated from one pulmonary case.
- Two patients showed Candida tropicalis and one showed Aspergillus fumigatus along with the direct positive findings of aseptate hyphae.
- The remaining cases turned out to be sterile on culture despite direct KOH/CFW showed mixed fungal etiology.
- Thirteen cases had underlying diabetes mellitus Type-II as the underlying risk factor.
- In addition, a significant risk factor in cutaneous type was the history of taking intramuscular injection at the affected site.
- AFST revealed all strains were sensitive to amphotericin B and posaconazole.
- All patients were treated with extensive surgical debridement and intravenous liposomal amphotericin B with good prognosis thereby eventually their survival.
- The results are also shown in tabulated form as well as figures as clinical photographs and photomicrographs.

Isolate	Site / clinical type	Risk factor	Treatment	Outcome of therapy
Rhizopus arrhizus Aspergillus flavus	Six rhino- orbital cases, one pulmonary case	DM type II, allergy to dust (one rhino- orbital case)	Surgical debridement + L-AMB	Five survived, one expired
Rhizopus arrhizus Alternaria alternata	One rhino- orbital case	DM type II	Surgical debridement + L-AMB	Survived
Lictheimia corymbifera Aspergillus flavus	One pulmonary case	DM type II	L-AMB	Survived
Rhizopus arrhizus Candida tropicalis	One rhino- orbital case	Chronic kidney disease	L-AMB	Survived
Aspergillus flavus Candida tropicalis	One rhino- orbital case	DM type II	Surgical debridement + L-AMB	Survived
Aspergillus fumigatus	One rhino- orbital case	DM type II	L-AMB	Survived
Sterile	Three rhino-orbital cases	DM type II	Surgical debridement + L-AMB	Survived
Sterile	Two Cutaneous cases	Intramuscular injection	Surgical debridement + L-AMB	One expired, one survived

Table Depicting details of mucormycosis with other invasive mycoses



Fig. 1. Bilateral ROCM with chemosis and proptosis



Fig. 2. KOH wet mount Fig. 3. CFW staining showing aseptate and septate hyphae



showing aseptate and septate hyphae

Discussion

- There has been an upsurge in cases of mixed fungal infection with a growing susceptible population.
- Diabetes mellitus appears to be a significant risk factor for the development of such infections to thrive and establish.
- Intramuscular injection has a major role in causation of cutaneous mixed infections.
- These mixed fungal infections carry poor prognosis, often leading to fatal consequences, if not timely managed.
- The aggressive nature of disease warrants a prompt and aggressive treatment regimen in order to save the life of patient.
- Amphotericin B is the mainstay therapeutic drug for treatment along with extensive surgical debridement/resection and control of risk factors.

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