

A Retrospective Epidemiological Analysis of Fungal Infections of Skin and Soft Tissue in a Health Care Setup in Delhi, India

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ABSTRACT

Introduction: Fungal infections of the skin and soft tissue can be an important cause of morbidity and mortality in both immunocompromised and immunocompetent patients and usually pose a major diagnostic challenge.

Aim: The present study was done to study the type of fungal isolates from pus and soft tissue specimens and to clinically correlate these fungal isolates.

Materials and Methods: A retrospective analysis of fungal isolates from pus and soft tissue specimens was done over a period of one year and their clinical history was studied from medical records department. The identification of yeast isolates was done by automated (MALDI-TOF-Vitek MS and Vitek-2) and conventional methods and mycelial fungi were identified by lactophenol cotton blue mount of growth and/or slide culture.

Results: Out of 288 pus/tissue specimens received for fungal culture during the study period, 37 showed growth of fungi. Out

of 37 fungi, 24 were yeast isolates and 13 were mycelial fungi. *Candida albicans* (24.3%) constituted the majority of the fungal infections followed by *Candida glabrata* (21.6%), *Aspergillus* sp. (18.9%) and *Dematiaceous* fungi (13.5%). All the *Dematiaceous* fungi were isolated from post-transplant patients. Out of 37 patients, 12 were immunocompromised and 22 had history of surgery. Antifungal resistance was not seen in *C. albicans* while *Candida haemulonii/auris* showed 100% resistance to amphotericin B and fluconazole. All the patients in our study had history of prior antibiotic intake for one or other reasons. In contradiction to leucopenia, many of the patients in our study had leucocytosis at the time of isolation of fungi.

Conclusion: A variety of fungi may cause pus and soft tissue infections in both immunocompromised and immunocompetent individuals. The results signify the importance of clinicomycological awareness, correlation, fungal speciation and trend of antifungal susceptibility pattern for the optimum treatment of the patients.

Keywords: Abdominal pus, *Aspergillus*, *Candida*, MALDI-TOF, Necrotic tissue

INTRODUCTION

The incidence of fungal infections has markedly increased in recent years particularly due to greater use of immunosuppressive drugs; prolonged use of broad-spectrum antibiotics; widespread use of indwelling catheters; and the Acquired Immunodeficiency Syndrome (AIDS). Fungal infections have emerged as a major cause of death among cancer and transplant recipients [1]. In addition, immunocompromised patients often experience more frequent and severe fungal infections. Most fungi that are pathogenic for humans are saprophytes in nature; they cause infection when airborne spores reach the lung or paranasal sinus or when hyphae or spores are accidentally inoculated into the skin or cornea. Most fungi infect hosts preferentially by one route and only infrequently by another routes. *Candida albicans*, a normal commensal in the mouth and intestine, reaches deeper tissues only when mucosal or cutaneous barriers are breached by disease, surgery, trauma or catheterization [2]. Some fungi, such as *Aspergillus* are said to be opportunists in that they usually infect hosts with compromised immunity [3]. Skin and soft tissue infections are the common sites of infection in immunocompromised hosts and usually pose a major diagnostic challenge. It is important to differentiate between bacterial and fungal infection in such cases because the successful management of such cases depend upon early recognition, timely surgical debridement or drainage and appropriate antifungal therapy. So, keeping all these factors in mind, the aim of the present study is: 1) to study the type of fungal isolates from pus and soft tissue specimens; 2) to clinically correlate these fungal isolates; 3) to study antifungal susceptibility profile of various *Candida* species.

MATERIALS AND METHODS

The study was conducted at Indraprastha Apollo Hospitals, New Delhi, India. A retrospective analysis of the fungal isolates from pus/tissue specimens was done over a time period of one year i.e., September 2013 to August 2014. All the clinical data of the patient was collected from Medical Record department of the hospital with respect to demographic details, immune status, underlying co-morbid conditions, any surgical procedure, treatment and follow-up. All the pus/tissue specimens received in the Department of Microbiology for fungal culture were processed as per standard protocol. The tissue specimens comprised of any type of skin and subcutaneous tissue, nasal polyps/paranasal sinus tissue, other soft tissue specimens from visceral organs like Pancreas. The details of the type of specimen and underlying condition have been described in [Table/Fig-1]. Direct microscopic examination (KOH mount) of the specimens was done and the fungal culture was put on two types of Sabouraud's Dextrose Agar (SDA) slants (SDA with and without Gentamicin- Chloramphenicol- cycloheximide) in duplicate and incubation of both types of SDA slants was done at 25°C and 37°C. The two types of SDA were prepared in house from dehydrated culture powder (HiMedia, Mumbai, India). Gentamicin (HiMedia, Mumbai, India), chloramphenicol (HiMedia, Mumbai, India) and cycloheximide (Sigma Aldrich, US) were added in SDA in concentration of 5 µg/mL, 16 µg/mL and 0.5 µg/mL respectively to inhibit the growth of bacteria and environmental moulds. The significance of the fungal growth was determined by clinical history and correlation with direct microscopic examination. Only that growth was considered to be significant in which direct microscopic examination showed presence of fungal elements (hyphae/yeast) along with clinical correlation. The fungal growth was identified

Fungal isolates	Number (%)	Site of specimen	Underlying disease/Diagnosis	Treatment	Outcome
<i>Candida albicans</i>	09 (24.3)	Abdominal pus	Post cholecystectomy-subdiaphragmatic collection	Ultrasound guided tube drainage of pus and Fluconazole	Discharged in stable condition
		Wound discharge post Coronary Artery Bypass Grafting (CABG)	Coronary artery disease, Chronic Renal disease	Fluconazole	Cured
		Ulcers on glans penis	Pyrexia of unknown origin with autoimmune disease	Voriconazole	Cured
		Aspirate from retropharyngeal mass	Skull base osteomyelitis	Amphotericin B followed by Voriconazole	Discharged in stable condition
		Oral ulcer	Dermatomyositis with facial cellulitis	Local application of clotrimazole	Recurrence
		Abdominal wound	Ileal stricture and perforation peritonitis	Amphotericin B	Expired
		Right lung empyema	Chronic renal disease, right lower lobe consolidation	Ultrasound guided pleural fluid drainage and Amphotericin B followed by Voriconazole	Cured
		Pus from groin and buttocks	Bilateral lower limb cellulitis on antibiotics	Daily dressing and local application of clotrimazole	
		Trophic ulcer right foot	Ulcer right foot, Coronary artery disease	Debridement	Discharged in stable condition
<i>Candida glabrata</i>	08 (21.6)	Ulcerative blisters all over the body	Chronic renal disease on Haemodialysis	Diagnosed as Toxic Epidermal Necrolysis, Local application of Fluticasone and oral Voriconazole	Expired
		Abdominal wound	Post operative case of bowel ischaemia with Faecal peritonitis and anastomotic leak	Anidulafungin	Discharged on request (Poor prognosis predicted)
		Necrotic Pancreatic tissue	Traumatic Pancreatitis	Pancreatic necrosectomy with FJ	Discharged in stable condition
		Abdominal wound discharge	Adenocarcinoma duodenum with whipple's procedure done	Caspofungin	Cured
		Intraabdominal abscess	Decompensated chronic liver disease	Drainage and Caspofungin	Expired
		Tongue ulcer	Post chemotherapy mucositis post renal transplant, metastatic carcinoma	Local application of Clotrimazole	Ulcers subsided
		Abdominal pus	Follow up case of carcinoma colon, Peripheral vascular disease	Drainage and Caspofungin	Discharged in stable condition
		Abdominal wound discharge	Disseminated tubercular perforation peritonitis	Caspofungin	Discharged
<i>Candida tropicalis</i>	03 (8.1)	Liver abscess	Gall stone disease with choledocholithiasis, hepatic abscess	Ultrasound guided drainage of hepatic abscess followed by Voriconazole, Common bile duct exploration	Cured
		Burn wound swab	Burns with sepsis, acute kidney and lung injury	Daily dressing	Discharged in stable condition
		Wound sinus on face	Road traffic accident with facial injury	Voriconazole	Discharged in stable condition
<i>Candida haemulonii/auris</i>	03 (8.1)	Subglial granulation tissue	Post cranioplasty infected bone implant	Re-exploration of left fronto-parieto-temporo skin flap with removal of infected brain cement and granulation tissue	Wound healthy at the time of discharge
		Wound bilateral cheek	Road traffic accident with head injury with polytrauma	Daily dressing	Cured
		Slough on under surface of tibia and femur	Infected Total knee replacement (TKR)	Revision TKR Voriconazole/ Amphotericin B	Recurrence till date with isolation of <i>Candida haemulonii/auris</i>
<i>Candida pulcherrima</i>	01 (2.7)	Multiple ulcers over left leg	Peripheral vascular disease	Voriconazole	Improved
Dematiaceous fungi	05 (13.5)	Left ankle infected cyst (<i>Exophiala</i> sp.)	Post Renal transplant	Excision of cyst followed by Itraconazole	Improved
		Abscess right foot (<i>Exophiala</i> sp.)	Post Renal transplant	Incision and drainage followed by Ketoconazole	Improved
		Left parieto-occipital abscess (<i>Cladophialophora</i> sp.)	Post Liver transplant	Craniotomy followed by Itraconazole	Improved and discharged in stable condition
		Right ankle swelling (<i>Exophiala</i> sp.)	Renal transplant	Incision and drainage followed by Voriconazole	Improved
		Intracranial abscess (<i>Cladophialophora</i> sp.)	Post renal transplant	Craniotomy and decompression followed by Itraconazole	Expired

<i>Aspergillus</i> spp.	07 (18.9)	Nasal mucosa (<i>Aspergillus flavus</i>)	Rhinosinusitis	Functional endoscopy Sinus Surgery (FESS)	Improved
		Wound pus on face (<i>Aspergillus fumigatus</i>)	Gunshot injury	Daily dressing and Voriconazole	Improved
		Ethmoidal cell mucin (<i>Aspergillus flavus</i>)	Rhinosinusitis	FESS	Improved
		Nasal polyp (<i>Aspergillus flavus</i>)	Nasal Polyposis	Polypectomy	Improved with no recurrence in one year follow up
		Scrotal abscess sinus (<i>Aspergillus fumigatus</i>)	Scrotal abscess	Excision of Scrotal sinus	Improved
		Nasal polyp (<i>Aspergillus flavus</i>)	Rhinosinusitis	Polypectomy	Improved
		Tissue from maxillary sinus (<i>Aspergillus flavus</i>)	Rhinosinusitis	FESS followed by Voriconazole	Improved
<i>Rhizopus</i> spp.	01 (2.7)	Subcutaneous abdominal swelling	Post renal transplant	Debridement followed by Amphotericin B and Posaconazole	Cured

[Table/Fig-1]: Frequency of isolation of fungal isolates from pus/tissue specimens with respect to clinical details of the patients.

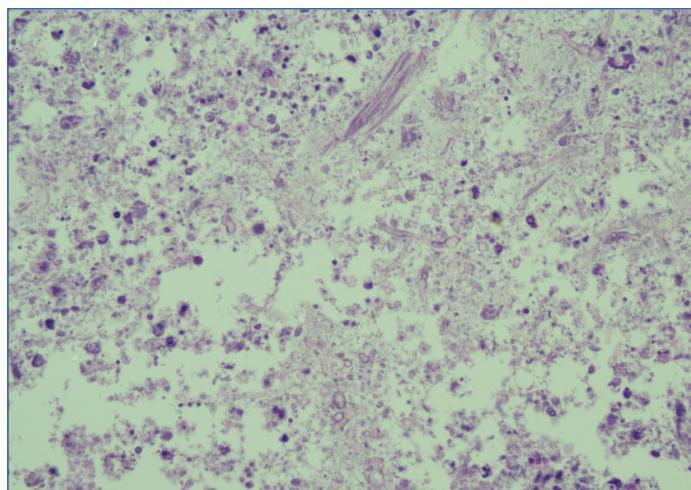
by macroscopic and microscopic examination using Lactophenol Cotton Blue Mount (LPCB) examination. The yeast identification was done by automated methods using Vitek-2 compact (software version 7.01, Biomerieux, France) or MALDI-TOF Vitek-MS (software version V2, Biomerieux, France) and correlated with conventional methods like germ tube test, Chrome agar (Hi Media, India) and cornmeal agar. For Germ tube test, yeast growth was inoculated in 0.5 ml of pooled human serum followed by incubation at 37°C for 2-3 hours followed by microscopic examination of wet mount at 40X objective. The yeast growth was deeply streaked in three parallel lines 1.5 cm long and about 1 cm apart on Corn meal Tween agar culture plate and a sterile cover slip was placed on it. The incubation was done at 25°C and was examined microscopically for chlamydospore formation and growth morphology after 48-72 hours along the edge of the coverslip in low and high power objectives. The mycelial fungi were identified by LPCB mount examination of the fungal growth on SDA slants and/or slide culture. The antifungal susceptibility testing of the yeast isolates were done by determining Minimum Inhibitory Concentration (MIC) for various drugs using Vitek-2 AST-YS06 cards (Biomerieux, France) and/or ATB FUNGUS 3 strips (Biomerieux, France) and/or E-test (Biomerieux, France) using RPMI agar. These methods of antifungal susceptibility testing had already been validated at our centre by comparing it with microbroth dilution technique. So, the interpretation of the results was done as per CLSI guidelines on broth dilution antifungal susceptibility testing of yeasts [4]. For Quality assurance of antifungal susceptibility testing methods, *Candida glabrata* ATCC 6258 and *Candida parapsilosis* ATCC 22019 were used as quality control strains. It was also tried to correlate the fungal culture growth with fungal KOH mount, and histopathological examination. A clinical correlation was also done with respect to age, gender, underlying disease, immune status, treatment and outcome.

RESULTS

A total of 288 pus/tissue non duplicate specimens were received for fungal culture during the study period. Out of these 37 showed growth of fungi. Out of 37 fungi, 24 were yeasts and 13 were mycelial fungi. Direct microscopic examination (KOH mount) of all the specimens with growth of mycelial fungi showed presence of fungal hyphae and those with growth of yeast showed presence of yeast in the specimen. [Table/Fig-2] shows the presence of septate, branching pigmented fungal hyphae in pus specimen from infected ankle cyst of post renal transplant patient. Few of these specimens were also sent for histopathological examination which also corroborated well with KOH mount examination and culture growth. [Table/Fig-3,4] shows the histopathological examination of left ankle infected cyst. The distribution of type of fungal isolates is given in [Table/Fig-1].

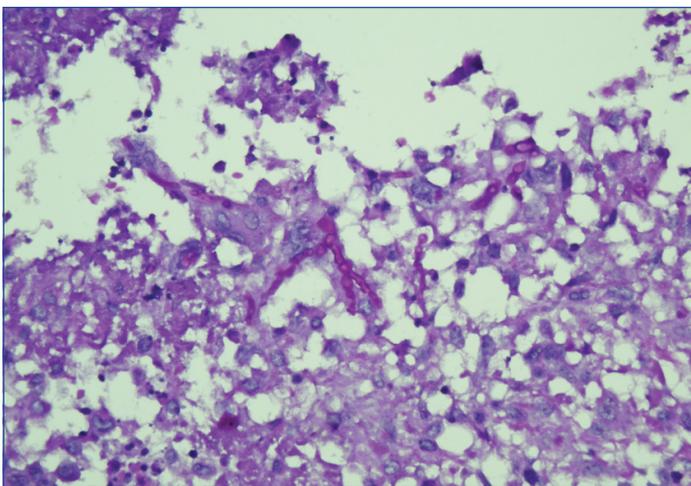


[Table/Fig-2]: KOH wet mount of pus from infected ankle cyst- shows black pigmented, septate fungal hyphae suggestive of Phaeoid fungi (Magnification 400X).

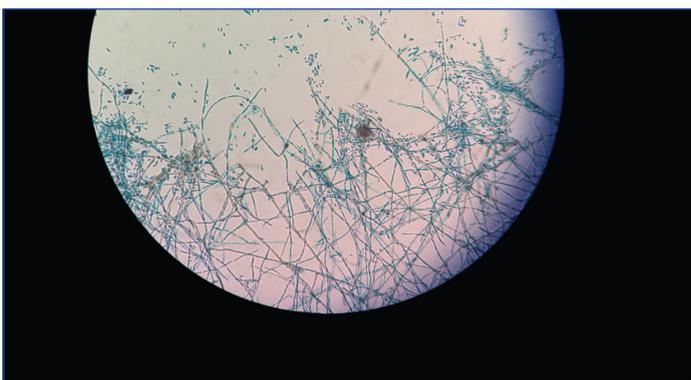


[Table/Fig-3]: Haematoxylin and Eosin stain- central necrotic area shows fungal hyphae and spore like structure enmeshed with necrotic cell debris (Magnification 400X).

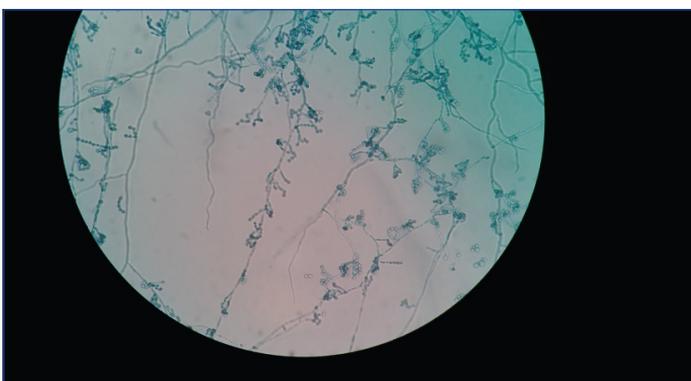
The yeast identification results by Vitek-2 and/or Vitek-MS correlated with conventional methods like germ tube test, Chrome agar and cornmeal agar for the common *Candida* species like *C. albicans*, *C. tropicalis* and *C. glabrata*. Germ tube test was positive in *C. albicans* and negative in other species. On Chrome agar (HiChrome *Candida* differential agar, HiMedia, Mumbai, India), *C. albicans* showed light green coloured smooth colonies, *C. tropicalis* showed blue coloured raised colonies and *C. glabrata* showed cream coloured smooth colonies. On corn meal agar, *C. albicans* showed pseudohyphae, clusters of blastoconidia and chlamydospores. However, *C. glabrata* did not show pseudohyphae formation and compactly arranged small yeast cells were seen. *C. tropicalis* showed presence of single or small cluster of blastoconidia along pseudohyphae. The identification of mycelial fungi was done by LPCB mount of slide culture. [Table/Fig-5,6] shows *Exophiala* sp. and *Cladophialophora* sp. on slide culture.



[Table/Fig-4]: PAS stain- Fungal structures ingested by multinucleated giant cells highlighted on PAS stain (Magnification 400X).



[Table/Fig-5]: LPCB mount of slide culture-*Exophiala* sp.- Conidia are one-celled, subhyaline, smooth, thin-walled, sub-globose to ellipsoidal, arising laterally or on tips and aggregating in clusters. Conidiophores are showing brown pigmented walls (Magnification 400X).



[Table/Fig-6]: LPCB mount of slide culture-*Cladophialophora* sp.-Conidia are formed in long, sparsely branched, chains from undifferentiated conidiophores. Conidia are one-celled, pale brown, smooth-walled, ellipsoid to oblong-ellipsoid (Magnification 400X).

The age range of the patient population showing growth of fungi varied from 12 years to 76 years and the median age was 44 years. Out of 37 patients 33 were males and four were females; 12 were immunocompromised (Post transplant and/or on immunosuppressant); 22 had history of surgery (including eight transplant surgeries) and sixteen were diabetic. Out of 37 patients, 18 patients had leucocytes within normal limits, 17 had leucocytosis and two had leucopenia. The distribution of fungal isolates with respect to the site of the specimen, clinical diagnosis, treatment and outcome is depicted in [Table/Fig-1]. The MIC values for various *Candida* species is depicted in [Table/Fig-7].

DISCUSSION

The frequency and occurrence of candidiasis has been well described [5,6]. More than 80% of high-risk patients in critical care units and those who are neutropenic may be colonized with *Candida* species,

Antifungal drug	<i>Candida albicans</i> (09)	<i>Candida glabrata</i> (08)	<i>Candida haemulonii/auris</i> (03)	<i>Candida tropicalis</i> (03)
M.I.C Values (µg/mL) range				
Amphotericin B	0.5-1	0.5-1	≥8	0.5-1
5 Flucytosine	≤1	≤1	≤1	≤1
Fluconazole	≤1	4-≥64	≥16-64	≤1
Voriconazole	≤0.12	≤0.12-1	≤0.12	≤0.12
Caspofungin	≤0.25	≤0.25	≤0.25	≤0.25
Micafungin	≤0.06-0.12	≤0.06-0.12	0.12	≤0.06

[Table/Fig-7]: Minimum inhibitory concentration (MIC Values) of various *Candida* species.



[Table/Fig-8]: Sinus formation at the surgical site (Total knee replacement)- caused by *Candida haemulonii/auris* (a) 3 incisions; (b) Middle incision showing dissection of Extensor Pollicis Longus tendon (EPL) and Dynamic Compression Plate (DCP); (C) Plate with screws in lateral view



[Table/Fig-9]: Healed sinus site after revision Total knee replacement followed by oral Voriconazole for three months

and superficial mucosal and cutaneous infections are common [7,8]. These non-invasive infections can be effectively treated with improved skin care and a topical antifungal agent or with a short course systemic azole antibiotic (e.g., fluconazole). In our study, out of 24 yeast isolates, *C. albicans* was isolated maximally (37.5%), followed by *C. glabrata* (33.3%), *C. tropicalis* (12.5%), *C. haemulonii/auris* (12.5%) and *C. pulcherrima* (4.1%). Few of these isolates

were present as colonizer and were not necessarily associated with the disease condition. So, the treatment in such cases was started only after clinical correlation especially in view of underlying disease and immune status of the patient. Amongst seven immunosuppressed/immunocompromised patients five (71.4%) showed growth of non albicans *Candida*. However, amongst 17 immunocompetent patients, 07 (41.1%) showed growth of *C. albicans*. So, the present study shows the predominance of non albicans *Candida* amongst immunocompromised patients in comparison to immunocompetent patients. We have seen the epidemiology of fungal infections in pus/tissue specimens only. In other studies, where the authors have studied the epidemiology of yeast isolates in all the specimens, the isolation of non albicans *Candida* have been found to be more than *C. albicans*.

In our study, amongst non albicans *Candida* the isolation of *C. glabrata* was found to be the maximum. Out of eight *C. glabrata* isolates, six were isolated from abdominal/intraabdominal abscess/tissue. *Candida glabrata* was also isolated from a case of pancreatic necrosis. Pancreatic infections usually occur due to enteric bacteria. Amongst yeast, the isolation of *C. glabrata* is rare; *C. albicans* being reported maximally [9,10]. Chakrabarti A et al., noted *Candida tropicalis* to be the most common fungal species grown from the samples collected from the pancreatic tissues of 335 patients with acute pancreatitis [11]. For all the isolates of *C. glabrata* the MIC value for Fluconazole was found to be ≥ 4 $\mu\text{g/mL}$. All the isolates of *C. glabrata* were found to be susceptible to Amphotericin B, 5 Flucytosine, Voriconazole, Caspofungin and Micafungin.

C. haemulonii complex/auris is now emerging as an invasive fungal pathogen especially amongst immunocompromised patients. *C. haemulonii* complex consists of three genotypically distinguishable species i.e., *C. haemulonii*, *Candida duobushaemulonii* and *C. haemulonii* var. *vulnera*. There are two other species which are related to *C. haemulonii* complex and can be misidentified as *C. haemulonii*. These species are *C. auris* and *C. pseudohaemulonii* [12]. *C. haemulonii* has been reported to be resistant with Amphotericin B and Fluconazole [13,14]. So, it is pertinent to identify these species because antifungal resistance is a great concern with this species. In this study all the three isolates of *C. haemulonii/auris* were from patients following surgery. In the present study we have also found that the isolates were resistant to amphotericin B and Fluconazole and sensitive to 5-Flucytosine, Voriconazole and Caspofungin. There are reports of misidentification of *Candida haemulonii* by Vitek-2 YST cards [15]. It has been seen that Vitek-2 system identifies *C. auris*, *C. haemulonii* and *C. pseudohaemulonii* as *C. haemulonii* [16]. The sequence analysis of the Internal Transcribed Spacer (ITS) region and the D1/D2 domain of the large subunit ribosomal RNA gene identify these isolates as *C. auris*. As mentioned in [Table/Fig-1], a case of Total Knee Replacement (TKR) is presenting till date with recurrent *C. haemulonii/auris* infection. The patient has taken several course of Voriconazole with intermittent response. Patient was a long standing diabetic with occasional periods of poor control. Patient also underwent revision TKR followed by Voriconazole for three months. [Table/Fig-8] shows the discharging wound Post TKR and [Table/Fig-9] shows healed wound after revision TKR. In literature, post surgical osteomyelitis by *Candida* species have been seen but orthopaedic infections due to *Candida haemulonii/auris* are not reported. This can be due to misidentification or non-identification [17].

In the present study, *Candida tropicalis* was isolated from hepatic abscess in a 36-year-old male patient who was a case of Gall stone disease with choledocholithiasis. Most of the cases of hepatic abscess have been reported in patients with haematological malignancies and the most common species have been found to be *C. albicans* [18]. However, Chen CY et al., reported the maximum isolation of *Candida tropicalis* in cases of hepatosplenic abscess in patients with acute leukaemia [19]. This patient was found to be immunocompetent and non-neutropenic. Like our case, Menachery J et al., reported liver abscess in a 30-year-old immunocompetent male patient due to

C. albicans [18]. The other two isolates of *C. tropicalis* were isolated from open wound. All the *Candida tropicalis* isolates in our study were found to be susceptible to all the drugs tested.

Candida pulcherrima was isolated from ulcers on leg in a patient of peripheral vascular disease. This species of *Candida* has earlier been reported to be involved in nail infections but now there are reports of fungaemia caused by this fungi [20,21].

For the identification of the yeasts, in comparison to conventional techniques, we found use of MALDI-TOF Vitek MS to be time saving, less labour intensive, cost-effective and accurate. The time required for the identification of the yeasts is at least 48 hours using conventional methods while by using MALDI-TOF-MS, the identification of the yeasts can be done in 10 minutes which is very helpful in the initiation of the appropriate antifungal therapy because there are certain species which are inherently resistant to certain antifungal drugs and few species usually show less susceptibility to particular antifungal drugs. MALDI-TOF-MS was also found to be very cost-effective and less labour intensive because the preparation and inoculation of a battery of biochemical tests are not required for the identification. A limited number of reagents and consumables (Target slide, matrix, formic acid, pipette tips, disposable loop or tooth pick) are required for the identification by MALDI-TOF-MS. Moreover, the interpretation of the results is also more specific and accurate and removes any confounding error.

Although, the isolation of Phaeoid fungi has been reported from both immunocompromised and immunocompetent patients but they are increasingly becoming opportunistic pathogens. In our study, all the five Phaeohyphomycetes (*Dematiaceous* fungi) were isolated from immunocompromised i.e., post-transplant patients. Out of five, two were isolated from brain abscess (cerebral phaeohyphomycosis) and three were isolated from lower limb lesions (subcutaneous phaeohyphomycosis). These fungi are generally found in soil or associated with plants and distributed worldwide. Exposure is thought to be from inhalation or minor trauma. Central nervous system infections can be due to haematogenous spread from an initial, presumably subclinical pulmonary focus. These cases were managed primarily surgically with wide excision and/or debridement of wound. After fungal culture report, Itraconazole was administered in most of the cases.

Aspergillus spp. was isolated mainly from cases of paranasal sinusitis. *Aspergillus* related sinus syndromes include allergic sinusitis, sinus aspergilloma, chronic granulomatous sinusitis, chronic invasive sinusitis and acute invasive sinusitis [22]. In the present study the patients of paranasal sinusitis were categorised as cases of allergic rhinosinusitis or nasal polyposis. These patients were immunocompetent and showed allergic manifestations like allergic rhinitis and/or asthma. All the cases of *Aspergillus* rhinosinusitis/polyposis were caused by *A. flavus*. Prateek S et al., also reported most common isolation of *A. flavus* amongst cases of both fungal rhinosinusitis and allergic fungal rhinosinusitis [23]. These patients were treated mainly surgically with or without antifungal therapy with good clinical response. Telmesani LM found 12.1% prevalence of Allergic Fungal Sinusitis (AFS) among patients with nasal polyps. They demonstrated *Aspergillus* sp. in 08 out of 11 cases of AFS [24].

In the present study, one case of cutaneous zygomycosis due to *Rhizopus* sp. was seen in post renal transplant patient. Bojdy A et al., also reported a case of cutaneous zygomycosis in renal transplant recipient [25].

All the patients in the study had history of prior antibiotic intake for one or other reasons. In contradiction to leucopenia, many of the patients in our study had leucocytosis at the time of isolation of fungi. Fungal infections in 16 patients were associated with history of previous surgery. Majority of these surgeries were performed outside the hospital for one or another reasons like wound repair in case of road traffic accidents, abdominal surgeries etc. Diabetes

was seen in 16 (43.2%) patients with fungal infections.

LIMITATION

The limitation of the study was that, we cannot exactly correlate poor outcome of the patient to the fungal infection or underlying co-morbid condition in immunocompromised patients. The prospective study on this concept especially with emphasis on long term follow-up will help in the optimum management of the patient.

CONCLUSION

In the present study, we have compiled all the clinically significant data of pus and soft tissue infections caused by fungi in our tertiary health care set-up. In literature, the studies pertaining to fungal isolates specifically from pus and tissue are very scarce. It is important to have such type of data in a clinical set-up because epidemiological profile of fungal infections varies with the type of specimen and patient population.

A variety of fungi may cause pus and soft tissue infections in both immunocompromised and immunocompetent individuals. So, it is important to diagnose fungal infections, identify the fungal species and know its antifungal susceptibility pattern to avoid unnecessary usage of antibiotics and appropriate antifungal therapy. A clinico-mycological awareness and correlation is required for optimum management.

REFERENCES

- [1] Person AK, Kontoyiannis DP, Alexander BD. Fungal infections in transplant and oncology patients. *Infect Dis Clin North Am.* 2010;24:439-59.
- [2] Conti HR, Huppler AR, Whibley N, Gaffen SL. Animal Models for Candidiasis. *Curr Protoc Immunol.* 2014;105:19.6.1-19.6.17.
- [3] Bennett JE. Diagnosis and treatment of fungal infections. In: Fauci AS, Braunwald E, Isselbacher KJ, Wilson JD, Martin JB, Kasper DL, Hauser SL, Longo DL, eds. *Principles of Internal Medicine, United States of America, 14th edn.* McGraw-Hill, 1998:1148-1150.
- [4] CLSI. Reference Method for Broth Dilution Antifungal Susceptibility Testing of Yeasts; Approved Standard-Third Edition. CLSI document M27-A3. Wayne, PA: Clinical and Laboratory Standards Institute; 2008.
- [5] Yapar N. Epidemiology and risk factors for invasive candidiasis. *Ther Clin Risk Manag.* 2014;10:95-105.
- [6] Rex JH, Walsh TJ, Sobel JD, Filler SG, Pappas PG, Dismukes WE, et al. Practice guidelines for the treatment of candidiasis. *Infectious Diseases Society of America. Clin Infect Dis.* 2000;30:662-78.
- [7] Eggimann P, Pittet D. Candida colonization index and subsequent infection in critically ill surgical patients: 20 years later. *Intensive Care Med.* 2014;40:1429-48.
- [8] Youngster I, Sharma TS, Duncan CN, McAdam AJ. Yield of fungal surveillance cultures in pediatric hematopoietic stem cell transplant patients: A retrospective analysis and survey of current practice. *Clin Infect Dis.* 2014;58:365-71.
- [9] Shanmugam N, Isenmann R, Barkin JS, Beger HG. Pancreatic fungal infection. *Pancreas.* 2003;27(2):133-38.
- [10] Kochhar R, Ahammed SK, Chakrabarti A, Ray P, Sinha SK, Dutta U, et al. Prevalence and outcome of fungal infection in patients with severe acute pancreatitis. *J Gastroenterol Hepatol.* 2009;24(5):742-47.
- [11] Chakrabarti A, Rao P, Tarai B, Shivaprakash MR, Wig J. Candida in acute pancreatitis. *Surg Today.* 2007;37:207-11.
- [12] Ramos LS, Figueiredo-Carvalho MH, Barbedo LS, Ziccardi M, Chaves AL, Zancopé-Oliveira RM, et al. Candida haemulonii complex: species identification and antifungal susceptibility profiles of clinical isolates from Brazil. *J Antimicrob Chemother.* 2015;70:111-15.
- [13] Khan ZU, Al-Sweih NA, Ahmad S, Al-Kazemi N, Khan S, Joseph L, et al. Outbreak of fungaemia among neonates caused by Candida haemulonii resistant to amphotericin B, itraconazole and fluconazole. *J Clin Microbiol.* 2007;45:2025-27.
- [14] Rodero L, Cuenca-Estrella M, Cordoba S, Cahn P, Davel G, Kaufman S, et al. Transient fungemia caused by an amphotericin B-resistant isolate of Candida haemulonii. *J Clin Microbiol.* 2002;40:2266-69.
- [15] Lee WG, Shin JH, Uh Y, Kang MG, Kim SH, Park KH, et al. First three reported cases of nosocomial fungemia caused by Candida auris. *J Clin Microbiol* 2011;49:3139-42.
- [16] Kim MN, Shin JH, Sung H, Lee K, Kim EC, Ryoo N, et al. Candida haemulonii and closely related species at 5 university hospitals in Korea: identification, antifungal susceptibility, and clinical features. *Clin Infect Dis.* 2009;48:e57-61.
- [17] Gamaletsou MN, Kontoyiannis DP, Sipsas NV, Moriyama B, Alexander E, Roilides E, et al. Candida Osteomyelitis: Analysis of 207 Pediatric and Adult Cases (1970-2011). *Clin Infect Dis.* 2012;55:1338-51.
- [18] Menachery J, Chawla Y, Chakrabarti A, Duseja A, Dhiman R, Kalra N. Fungal liver abscess in an immunocompetent individual. *Trop Gastroenterol.* 2012;33:232-33.
- [19] Chen CY, Chen YC, Tang JL, Yao M, Huang SY, Tsai W, et al. Hepatosplenic fungal infection in patients with acute leukemia in Taiwan: incidence, treatment, and prognosis. *Ann Hematol.* 2003;82(2):93-97.
- [20] Deconinck L, Meybeck A, Pradier M, Patoz P, Melliez H, Senneville E. Community acquired fungemia caused by Candida pulcherrima: diagnostic contribution of MALDI-TOF mass spectrometry. *Ann Clin Microbiol Antimicrob.* 2016;15:14.
- [21] Bereczki L, Bartha N, Kocsubé S, Sóki J, Lengyel G, Tálosi G, et al. Fungaemia caused by Candida pulcherrima. *Med Mycol.* 2012;50:522-24.
- [22] Richardson MD, Hope W. *Aspergillus*. In: Anaissie EJ, McGinnis MR, Pfaller MA, eds. *Clinical Mycology, United States of America, 2nd edn.* Churchill Livingstone, 2009:271-276.
- [23] Prateek S, Banerjee G, Gupta P, Singh M, Goel MM, Verma V. Fungal rhinosinusitis: A prospective study in a University hospital of Uttar Pradesh. *Indian J Med Microbiol.* 2013;31:266-69.
- [24] Telmesani LM. Prevalence of allergic fungal sinusitis among patients with nasal polyps. *Ann Saudi Med.* 2009;29:212-14.
- [25] Bojdy A, Shojaja SRH, Farid GA, Sarvghad MR, Naderi HR, Salehnia N, et al. Cutaneous mucormycosis (zygomycosis) in a kidney transplant recipient: recovery after amphotericin therapy. *Ann Biol Res.* 2013;4:275-79.

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