Study of Candida albicans Isolated from Wounds of Type 2 Diabetes Patients after Plastic Surgery

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ABSTRACT This study included 65 wo Din General Hospital in Ti determine the prevalence DM patients after plastic three and six days of burn inoculated on culture med Chrome agar) and incubate incubation at 37°C for 4 preformed based on a co differential release of chro substrates by <i>Candida</i> s activity. In this study, 30° were obese, although the s infection but still referred that maximum rate of patie	bund patients were admission in Salah krit during November 2017- May 2019, of <i>C. albicans</i> causing wound infectior surgery. Wound swabs were taken at patients from the pus of the wound are dia (sabouraud dextrose agar and Cand d aerobically for (24-48) hours at 37C ² . At 8 hours the identification of yeast v lony colour. The method is based on mogenic breakdown products from varia species following differential excenzy % of patients were overweight and 44 study didn't include the relation of BMI w to its association. The study also view ents were within the age group 37-46 ye the age group 47%-56%. Additionally, s	to fugal cause of wound (30%). The present appeared on saboura circular colonies and the predominant isc patients who admitte Keywords: T2DM; the infections Correspondence: Abdulrazzaq Abbas V 0% M.B.Ch.BFICM (Pla ith Iraq e-mail: abdulrazzaq2t ars

INTRODUCTION

Article History

During the past 10 years, fungal infections have gained considerable medical importance, particularly in patients with severe underlying diseases. While Candida species were previously the predominant causative agents of invasive mycoses, in recent years opportunistic moulds, such as Aspergillus species or other hyphomycetes, and zygomycetes, have increasingly been implicated in human infection. In contrast to the yeasts, several of these moulds show a high degree of resistance to most antifungal drugs ⁽¹⁾. Candida albicans occurs naturally as a commensal of mucous membranes of the oral cavity, gastrointestinal tract and genitourinary tract of healthy human⁽²⁾. On sabouraud dextrose agar colonies are white to cream colored, smooth and glabrous in appearance. Microscopic morphology shows spherical to subspherical budding yeast-like cells or blastoconidia, Gram positive. It is much larger than bacteria, reproduced by budding ⁽³⁾. The genus *Candida* includes about 150 different species, however, only a few are known to cause human infections. C. albicans is the most significant pathogenic species; at least 70% of all human Candida infections are caused by C. albicans. Other Candida spp. pathogenic in humans include C. tropicalis, C. glabrata, C. krusei, C. parapsilosis, C. dubliniensis, and C. lusitaniae⁽⁴⁾. Several Candidal infections are a serious problem, especially in individual whose immune defense mechanism has been weakened. C. albicans cause infection of the skin, oral cavity and esophagus, gastrointestinal tract, vagina and vascular system of humans⁽⁵⁾. Virulence factors of *C. albicans* include host recognition biomolecules (adhesins), morphogenesis (the reversible transition between unicellular yeast cells and filamentous, growth forms), secreted hydrolytic enzymes includes proteases and phospholipases. There are a limited number of antifungal drugs, especially compared with the number of antibacterial drugs ⁽⁶³⁾. So the aim of this study was

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re females. The *C. albicans* was the most isolated d infection (70%) and Aspergillus was the second ting study revealed that C. albicans colonies aud dextrose agar as white to cream, smooth and green colonies on CHROM agar. C. albicans was olated fungal cause of wound infection in DM ed because plastic surgery.

Plastic surgery; Candida albicans; Wound

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to determine the prevalence of C. albicans causing wound infection in DM patients after plastic surgery.

MATERIAL AND METHOD

This study included 65 wound patients were admission in Salah ad-Din General Hospital in Tikrit during the period from November 2017 to May 2019 with age ranged between (17-70) years. They were suffering from wound infection after different types of surgical plastic surgery were done for them. Wound swabs were taken after three and six days of burn patients from the pus of the wound area in the morning before the bathing of the affected area (before hydrotherapy). Each swab was placed in a sterile tube with transport media till reaching the laboratory to be inoculated on culture media (sabouraud dextrose agar and Candida Chrome agar) and incubated aerobically for (24-48) hours at 37C° (1). CHROMagar Candida Staniszewska et al., 2011b is a selective fungal medium that includes chromogenic substances allowing for guick identification of several different Candida spp. based on a colour change. It can be used the for identification of individual non-albicans species, as well as C. albicans, if germ tube test was not characteristic. After incubation at 37°C for 48 hours the identification of yeast was preformed based on a colony colour. Using this method, we were able to identify the following individual non-albicans species: C. glabrata (dark pink colonies, wet), C. tropicalis (blue colonies, wet), C. krusei (light pink colonies, dry), and C. albicans (green colonies, wet). , which also facilitates the detection of mixed infections with more than one species of Candida. The method is based on the differential release of chromogenic breakdown products from various substrates by Candida species following differential exoenzyme activity.

FINDINGS

In this study, 30% of patients were overweight and 40% were obese, although the study didn't include the relation of BMI with infection but still referred to its association. The study

also viewed that maximum rate of patients were within the age group 37-46 years (40%) followed by 30% in the age group 47%-56%. Additionally, the study concluded that 60% of patients were females

Parameters		n(%)
BMI (kg/ m²)	Non-Obese (18.5 24.9)	30
	Overweight (25-29.9)	30
	$Obese \ge 30$	40
Age (years)	17-26	5
	27-36	20
	37-46	40
	47-56	30
	57-66	3
	67-76	2
	Range: 17-50	
	Mean: 43.01±3.4	
Sex	Males	40
	Females	60

Table 1: Distribution of the general characteristics Diabetes patients

The study demonstrated that C. albicans was the most isolated fugal cause of wound infection (70%) and Aspergillus was the second (30%)

Table 2: Isolated fungal gerera		
	Present study	
Total DM patients	65	
Fungal isolates	22%	
First line	Candida (70%)	
2 nd line	Aspergillus (30%)	

The presenting study revealed that *C. albicans* colonies appeared on sabouraud dextrose agar as white to cream, smooth and circular colonies and green colonies on CHROM agar Fig. (1-4). The chromogen mix consists of artificial substrates (chromogens), which release different colored compounds upon degradation by specific enzymes (Horvath et al., 2008). The results of microscopic examination show that the Gram negative bacteria produced rod colonies and Gram positive bacteria produced cluster-like colonies. Regarding to the *C. albicans*, it produced round blastoconidia single or in small cluster under microscope (Salyers and Whitt, 2002; Kayser et al., 2005). Candida albicans cultures on Sabouraud's (SDA) medium was white to cream smooth, glabrous,were is on CHROMagar medium was green, wet Figure 1 and 2. Microscopic examination showed that Spherical to sub spherical budding blastoconidia Figure 3. Germ tube positive.. Figure 1.



Figure 1: Candida albicans on Sabouraud's (SDA) agar (left) and CHROMagar (right)



Figure 2: CHROMagar plate showing C. albicans



Figure 3: Microscopy of *C. albicans* showing Budding and yeast cells (40x)



Figure 4: Germ tube of *Candida albicans* after three hours incubation (40x)

DISCUSSION

In this study, 30% of patients were overweight and 40% were obese, although the study did not include a BMI relationship to infection but still indicated its association. The study also showed that the maximum rate of patients were within the age group 37-46 years (40%) followed by 30% in the age group 47% -56%. In addition, the study concluded that 60% of patients are females. This finding is consistent with the results of previous studies. A previous study found that middle-aged young people were the most visited by plastic surgery clinics ⁽⁷⁾. Another study showed that the age above 25 years are the most vulnerable to cosmetics because of the requirements of modern life as well as the increase of cosmetic clinics and the development of cosmetic procedures in the world, and social media has actually contributed to increase the number of visitors to these clinics. Skin colonization and the development of several skin

manifestations in patients with poor glycemic control and elevated glycosylated hemoglobin, who have abnormal carbohydrate metabolism, relative insulin availability, neuron degeneration, deranged collagen production, microvascular complications and impaired wound healing, seems to be related to a multitude of skin disorders ⁽⁸⁻¹¹⁾. In addition, the prevalence of cutaneous disease have relation to number of risk factors including age, male gender, family history of onychomycosis and intake of immunosuppressive agents (12,13). C. albicans was completely 100% sensitive to Amphotericin B and Clotrimazole and 25% to Fluconazole. In this study, this result fitted to the result detected by several studies done earlier ⁽¹⁴⁻¹⁶⁾. The most common mechanisms responsible for drug resistance in Candida spp. are lower accumulation of drugs into the cells due various drug efflux proteins. Another possibility is the mutations or overexpression of the target gene, that leads to change in the structure of target enzyme which may result in alteration of the target and hence in resistance towards Azole drugs ^(17,18). Diabetes mellitus is firmly connected with the advancement of parasitic diseases and dermatoses associated with skin appearances. Two third of diabetic subjects have expanded recurrence of contaminations brought about by Candida or dermatophytes (tinea pedis and onychomycosis) during the course of the malady procedure ^(5,7). Dermatologic issues happen all the more frequently in patients with type 2 diabetes mellitus, by whom irresistible inclusion of the skin is all the more frequently (immune system skin sores are progressively normal in type 1 diabetes) ⁽⁸⁾. The most visit contagious illnesses are: tinea pedis (interdigitalis), onychomycosis, candidal intertrigo and candidal vaginitis ^(4,9). The regiment of medications relies upon culture distinguishing proof, influenced some portion of body, reality of disease, prior administration and invulnerable circumstance of the patient ⁽⁴⁾. Routinely, azoles, terbinafine are endorsed. Terbinafine (inhibitor of parasitic squalene epoxidase) cream and sertaconazole cream is the prescribed first line treatment for shallow skin diseases. Onychomycosis is a sign for fundamental antimycotics (for instance terbinafine 250 mg/d for 2-12 weeks). The foundational treatment is associated with unfavorable impacts: sickness, stomach torment, retching, loose bowels, dazedness, rash and pruritus ⁽¹⁸⁻²¹⁾. In patients with mucormycosis, amphotericin B with debridement is utilized (22). Other studies involving trauma patients reported a rate of candidemia <10%.3,5,13 For example, the incidence of Candida infection in our population was similar to that reported by our findings. This difference between the rate of Candida infection in trauma patients and the rate among other non-trauma critically ill surgical patients might indicate the relatively low initial inoculum of Candida, prompt surgical attention and the relatively younger age of trauma patients.

CONCLUSION

It was concluded that *C. albicans* was the predominant isolated fungal cause of wound infection in DM patients who admitted because plastic surgery.

CONFLICT OF INTEREST none SOURCE OF FINDINGS self-findings.

ETHICAL CLEARANCE

This research was carried out with the patient's verbal and analytical approval before the sample was taken. According to this approval, all the samples were collected and the tests were carried out. A copy of the results of the tests was then given to the patients

REFERENCES

- 1. Powers AC. Diabetes mellitus. Harrison's principles of internal medicine. 19th Ed. 2008:2275-305.
- 2. Sahay BK. API Textbook of Medicine. 8th Ed. Diabetology. Vol II. Sec 18-, 2009:1042.
- 3. Sridhar GR, Rao PV and Ahuja MMS. RSSDI Textbook of Diabetes. Hyderabad, RSSDI; 2002:95-112.
- Gadepalli R, Dhawan B, Sreenivas V, Kapil A, Ammini AC, Chaudhry R. A clinico-microbiological study of diabetic foot ulcers in an Indian tertiary care hospital. Diabetes care. 2006;29(8):1727-32.
- Norbury W, Herndon DN, Tanksley J, Jeschke MG, Finnerty CC, Scientific Study Committee of the Surgical Infection Society. Infection in burns. Surgical infections. 2016 Apr 1;17(2):250-5.
- Fan C, Tian Q, Huang G, Zhang L, Wu Q, Zhang K. Candida tropicalis burn wound sepsis: a series of histopathology-confirmed cases. Intensive and Critical Care Nursing. 2018 Jun 1;46:6-9.
- Mahadik K, Yadav P, Bhatt B, Shah RA, Balaji KN. Deregulated AUF1 Assists BMP-EZH2–Mediated Delayed Wound Healing during Candida albicans Infection. The Journal of Immunology. 2018 Dec 15;201(12):3617-29.
- Manolakaki D, Velmahos G, Kourkoumpetis T, Chang Y, Alam HB, De Moya MM, Mylonakis E. Candida infection and colonization among trauma patients. Virulence. 2010 Sep 1;1(5):367-75.
- Dai T, de Arce VJ, Tegos GP, Hamblin MR. Blue dye and red light, a dynamic combination for prophylaxis and treatment of cutaneous Candida albicans infections in mice. Antimicrobial agents and chemotherapy. 2011 Dec 1;55(12):5710-7.
- St. Denis TG, Dai T, Izikson L, Astrakas C, Anderson RR, Hamblin MR, Tegos GP. All you need is light: antimicrobial photoinactivation as an evolving and emerging discovery strategy against infectious disease. Virulence. 2011 Nov 1;2(6):509-20.
- Bassetti M, Marchetti M, Chakrabarti A, Colizza S, Garnacho-Montero J, Kett DH, Munoz P, Cristini F, Andoniadou A, Viale P, Della Rocca G. A research agenda on the management of intra-abdominal candidiasis: results from a consensus of multinational experts. Intensive care medicine. 2013 Dec 1;39(12):2092-106.
- 12. Carneiro HA, Mavrakis A, Mylonakis E. Candida peritonitis: an update on the latest research and

treatments. World journal of surgery. 2011 Dec 1;35(12):2650-9.

- 13. Pittet D, Wenzel RP. Nosocomial bloodstream infections. Secular trends in rates, mortality and contribution to total hospital deaths. Arch Intern Med 1995; 155:1177-84.
- Eggimann P, Garbino J, Pittet D. Epidemiology of Candida species infections in critically ill nonimmunosuppressed patients. Lancet Infect Dis 2003; 3:685-702.
- 15. Ziegler TR. Parenteral nutrition in the critically ill patient. N Engl J Med 2009; 361:1088-97.
- Hostetter MK. Handicaps to host defense. Effects of hyperglycemia on C3 and *Candida albicans*. Diabetes 1990; 39:271-5.
- 17. Holley A, Dulhunty J, Blot S, Lipman J, Lobo S, Dancer C, et al. Temporal trends, risk factors and outcomes in albicans and non-albicans candidaemia: an international epidemiological study in four multidisciplinary intensive care units. Int J Antimicrob Agents 2009; 33:5541-7.
- Harris AD, Castro J, Sheppard DC, Carmeli Y, Samore MH. Risk factors for nosocomial candiduria due to Candida glabrata and *Candida albicans*. Clin Infect Dis 1999; 29:926-8.
- Goldacre MJ, Watt B, Loudon N, Milne LJ, Loudon JD, Vessey MP. Vaginal microbial flora in normal young women. Br Med J 1979; 1:1450-3.
- 20. Oriel JD, Waterworth PM. Effects of minocycline and tetracycline on the vaginal yeast flora. J Clin Pathol 1975; 28:403-6.
- 21. Miller LG, Hajjeh RA, Edwards JE Jr. Estimating the cost of nosocomial candidemia in the united states. Clin Infect Dis 2001; 32:1110.
- 22. Cruciani M, de Lalla F, Mengoli C. Prophylaxis of Candida infections in adult trauma and surgical intensive care patients: a systematic review and metaanalysis. Intensive Car