

CHAPTER 4

SUPERFICIAL MYCOSES

Esra ADIŞEN¹
Özge KESEROĞLU²

Superficial fungal infections of the skin are a common presentation in clinical practice. They can affect any skin surface including the mucous membranes, hair follicles, nail plates, and nail beds. In this section common manifestations of *Malassezia* and *Candida* species are summarized.

4.1. *Malassezia*

Malassezia species are members of the normal commensal skin microflora of human body and many warm-blooded vertebrates. They are lipophilic and are associated with some skin diseases. They are found in the form of budding yeast in the stratum corneum of patients with various skin diseases (1). They may cause systemic disease in children and adults and sepsis in newborns and may play a role in certain dermatological diseases as well, due to differentiation of the skin structure, mostly in immunosuppressed patients (2).

4.1.1. Taxonomy and History

Malassezia is found in the *Cryptococcaceae* family of the Blastomycetes class. *Malassezia* species are dimorphic, which means that they can be found both in yeast and in mycelial form, and both forms have been regarded as separate breeds for many years. In order to emphasize these differentiations, *Pityrosporum* was used for yeast-shaped form and *Malassezia* for mycelial-shaped form. Since the yeast cells have different forms, they are separated into two different yeast-shaped species; *Pityrosporum orbicularula* with round cell and *Pityrosporum ovale* with oval cell (2-5).

Malassezia species were first described by Eichstedt in 1846. In 1873, Rivolta observed a psoriatic, double-walled, bud-like cell. In 1874, Malassez reported that

¹ Prof. Dr., Gazi University School of Medicine, Department of Dermatology, eozsoy@gazi.edu.tr

² Dr., Ankara Özel Drderm Polikliniği, ozgederm@yahoo.com

candidiasis is a condition that may be inherited autosomal recessively, autosomal dominantly or may occurs sporadically. It presents with different clinical manifestations. Deep hyperkeratotic granulomatous lesions, diffuse grayish white plaques in the oral and genital mucosa, thickening and dystrophy in the nails occur during the course of the disease (50, 51, 54, 60, 61).

Chronic mucocutaneous candidiasis (CMC) is clinically classified in four groups (50, 51, 54, 60, 61):

4.2.1.3.1. Early-onset CMC is the most severe and typical form of chronic mucocutaneous candidiasis. This type often begins in the mouth with moniliasis, then they spread to the face, head and nails. Candidal granulomas form thick crusts that bleed when removed. In half of the cases endocrinological disorders can be seen. Candidal granuloma is unique to patients with CMC. In this severe form of onychomycosis, *Candida* invades the full thickness of the nail. Advanced cases with the swelling of lateral and proximal nail folds may cause digit deformity called pseudoclubbing or “chicken drumstick” appearance.

4.2.1.3.2. Late-onset CMC is the mildest type of disease and is associated with nail disease and oral lesions. There are also publications showing that this disease, usually seen in the fourth decade of life or later, and is accompanied by underlying neoplasms such as timoma.

4.2.1.3.3. Familial CMC is usually transmitted autosomal recessively, and accompanying endocrinologic disorders are rare. It is characterized by recurrent paronychia and oral moniliasis that usually occur in the first year of the life. The disease usually runs a benign course. In some family members, only mild form of the disease is detected and immunological tests of the patients are found normal.

4.2.1.3.4. Juvenile familial polyendocrinopathy and candidiasis syndrome usually shows autosomal recessive transmission. Endocrinological disorders usually occur before the appearance of the candidiasis infections; parathyroid, adrenal, thyroid, pituitary gland or ovarian dysfunction may be found in these patients.

REFERENCES

1. Stein DH. Superficial fungal infections. *Pediatr Clin North Am* 1983; 30: 545-561.
2. Kantarcıoğlu AS, Yücel A. *Malassezia* türleri: taksonomi, mikoloji, immünoloji, patogenez, vücuttaki dağılımı ve ilişkili enfeksiyonlar, laboratuvar tanımı, antifungallere duyarlılığı. *Cerrahpaşa J Med.* 2005; 36: 134-154.
3. Ashbee HR, Evans EGV. Immunology of diseases associated with *Malassezia* species. *Clin Microbiol Rev* 2002; 15: 21-57.
4. Ljubojevic S, Skerlev M, Lipozencic J, Basta-Juzbasic A. The role of *Malassezia furfur* in dermatology. *Clin Dermatol* 2002; 20: 179-182.
5. Guého E, Midgley G, Guillot J. The genus *Malassezia* with description of four new species. *Antonie Leeuwenhoek* 1996; 69: 337-355.

6. Gaitanis G, Magiatis P, Hantschke M, Bassukas ID, Velegraki A. The *Malassezia* genus in skin and systemic diseases. *Clin Microbiol Rev* 2012; 25: 106-141.
7. Cabañes FJ, Vega S, Castellá G. *Malassezia cuniculi* sp. nov., a novel yeast species isolated from rabbit skin. *Med Mycol* 2011; 49: 40-48.
8. Crespo-Erchiga V, Gómez-Moyano E, Crespo M. Pityriasis versicolor and the yeasts of genus *Malassezia*. *Actas Dermosifiliogr* 2008; 99: 764-771.
9. Guého E, Boekhout T, Ashbee HR, Guillot J, Van Belkum A, Faergemann J. The role of *Malassezia* species in the ecology of human skin and as pathogens. *Med Mycol* 1998; 36: 220-229.
10. İkizoğlu G. *Malassezia* türleri ile ilişkili deri hastalıkları. *Türk Klin J Dermatol-Special Topics* 2008; 1: 25-35.
11. Kiremitçi Ü, Alyanak A, Tüzün Y. Pityriasis versicolor'da yeni eğilimler. *Dermatose* 2004; 3: 92-97.
12. Leeming JP, Notman FH. Improved methods for isolation and enumeration of *Malassezia furfur* from human skin. *J Clin Microbiol* 1987; 25: 2017-2019.
13. Ran Y, Yoshiike T, Ogawa H. Lipase of *Malassezia furfur*: some properties and their relationship to cell growth. *J Med Vet Mycol* 1993; 31: 77-85.
14. Mittag H. Fine structural investigation of *Malassezia furfur*. II. The envelope of the yeast cells. *Mycoses* 1995; 38: 13-21.
15. Kesavan S, Holland KT, Ingham E. The effects of lipid extraction on the immunomodulatory activity of *Malassezia* species in vitro. *Med Mycol*. 2000; 38: 239-247.
16. Belew PW, Rosenberg EW, Jennings BR. Activation of the alternative pathway of complement by *Malassezia ovalis* (*Pityrosporum ovale*). *Mycopathol* 1980; 70: 187-191.
17. Mayser P, Pickel M, Haze P, Erdmann F, Papavassilis C, Schmidt R. Different utilization of neutral lipids by *Malassezia furfur* and *Malassezia sympodialis*. *Med Mycol* 1998; 36: 7-14.
18. Sugita T, Takashima M, Shinoda T, Suto H, Unno T, Tsuboi R, Ogawa H, Nishikawa A. New yeast species, *Malassezia dermatis*, isolated from patients with atopic dermatitis. *J Clin Microbiol* 2002; 40: 1363-1367.
19. Nell A, James SA, Bond CJ, Hunt B, Herrtage ME. Identification and distribution of a novel *Malassezia* species yeast on normal equine skin. *Vet Rec* 2002; 150: 395-398.
20. Kim SC, Kim HU. The distribution of *Malassezia* species on the normal human skin according to body region. *Korean J Med Mycol* 2000; 5: 120-128.
21. Gupta AK, Kohli Y. Prevalence of *Malassezia* species on various body sites in clinically healthy subjects representing different age groups. *Med Mycol* 2004; 42: 35-42.
22. Leeming JP, Notman FH, Holland KT. The distribution and ecology of *Malassezia furfur* and cutaneous bacteria on human skin. *J Appl Bacteriol* 1989; 67: 47-52.
23. Jang SJ, Lim SH, Ko JH, Oh BH, Kim SM, Song YC, Yim SM, Lee YW, Choe YB, Ahn KJ. Investigation on the distribution of *Malassezia* yeasts on the normal Korean skin by 26S rDNA PCR-RFLP. *Ann Dermatol* 2009; 21: 18-26.
24. Savin R. Diagnosis and treatment of tinea versicolor. *J Fam Pract* 1996; 43: 127-132.
25. Borelli D, Jacobs PH, Nall L. Tinea versicolor: epidemiologic, clinical, and therapeutic aspects. *J Am Acad Dermatol* 1991; 25: 300-305.
26. Ertam İ, Aytimur D. *Malassezia* spp. ve dermatoloji'deki yeri. *Türkderm* 2006; 40: 7-10.
27. Gupta AK, Batra R, Bluhm R, Boekhout T, Dawson TL Jr. Skin diseases associated with *Malassezia* species. *J Am Acad Dermatol* 2004; 51: 785-798.
28. Difonzo EM, Faggi E. Skin diseases associated with *Malassezia* species in humans. Clinical features and diagnostic criteria. *Parasitol* 2008; 50: 69-71.
29. Gupta AK, Bluhm F, Summerbell R. Pityriasis versicolor. *J Eur Acad Dermatol Venereol* 2002; 16: 19-33.
30. Sei Y. *Malassezia* related diseases. *Nihon Ishinkin Gakkai Zasshi* 2006; 47: 75-80.

31. Gupta AK, Kohli Y, Summerbell RC, Faergemann J. Quantitative culture of *Malassezia* species from different body sites of individuals with or without dermatosis. *Med Mycol* 2001; 39: 243-251.
32. Gemmer CM, DeAngelis YM, Theelen B, Boekhout T, Dawson JT Jr. Fast, noninvasive method for molecular detection and differentiation of *Malassezia* yeast species on human skin and application of the method to dandruff microbiology. *J Clin Microbiol* 2002; 40: 3350-3357.
33. Gupta AK, Kohli Y, Summerbell RC. Molecular differentiation of seven *Malassezia* species. *J Clin Microbiol* 2000; 38: 1869-1875.
34. Theelen B, Silvestri M, Gueho E, Van Belkum A, Boekhout T. Identification and typing of *Malassezia* yeasts using Amplified Fragment Length Polymorphism (AFLP), Random Amplified Polymorphic DNA (RAPD) and Denaturing Gradient Gel Electrophoresis (DGGE). *FEMS Yeast Res* 2001; 1: 79-86.
35. Back O, Faergemann J, Hornqvist R. Pityrosporum folliculitis: a common disease of the young and middle-aged. *J Am Acad Dermatol* 1985; 12: 56-61.
36. Nakabayashi A, Sei Y, Guillot J. Identification of *Malassezia* species isolated from patients with seborrhoeic dermatitis, atopic dermatitis, pityriasis versicolor and normal subjects. *Med Mycol* 2000; 38: 337-341.
37. Sandström MH, Bartosik J, Bäck O, Scheynius A, Särnhult T, Tengvall Linder M, et al. The prevalence of the *Malassezia* yeasts in patients with atopic dermatitis, seborrhoeic dermatitis and healthy controls. *J Eur Acad Dermatol Venereol* 2001; 15: 104-274.
38. Tajima M. *Malassezia* species in patients with seborrhoeic dermatitis and atopic dermatitis. *Nihon Ishinkin Gakkai Zasshi*. 2005; 46: 163-167.
39. Prohic A. Distribution of *Malassezia* species in seborrhoeic dermatitis: correlation with patients' cellular immune status. *Mycoses* 2010; 53: 344-349.
40. Tajima M, Sugita T, Nishikawa A, Tsuboi R. Molecular analysis of *Malassezia* microflora in seborrhoeic dermatitis patients: comparison with other diseases and healthy subjects. *J Invest Dermatol* 2008; 128: 345-351.
41. Sugita T, Suto H, Unno T, Tsuboi R, Ogawa H, Shinoda T, Nishikawa A. Molecular analysis of *Malassezia* microflora on the skin of atopic dermatitis patients and healthy subjects. *J Clin Microbiol* 2001; 39: 3486-3490.
42. Sugita T, Takashima M, Shinoda T, Suto H, Unno T, Tsuboi R. New yeast species, *Malassezia dermatis*, isolated from patients with atopic dermatitis. *J Clin Microbiol* 2002; 40: 1363-1367.
43. Hay RJ, Moore MK. Mycology. In: Burns T, Breathnach S, Cox N, Griffiths C. eds. *Rook's Textbook of Dermatology*. 7th ed. Oxford, Blackwell Sci. Pub 2004: 31.1-31.101.
44. Kelly BP. Superficial fungal infections. *Pediatr Rev* 2012; 33: e22-37.
45. Bilgehan H. Candidaların tarihçesi, ekolojisi ve dağılımı. Tümbay E, editör. *Candida ve enfeksiyonları'da*. İzmir: Türk Mikrobiyoloji Yayınları No:6. Bilgehan Basımevi 1986; 1-9.
46. Seçkin D, Baba M. Kandidiyazisin Kliniği. *Turk Klin J Int Med Sci* 2005; 1: 16-22.
47. Grillot R. Epidemiological survey of candidemia in Europe. *ECMM / ECMM Mycology Newsletter* 2003; 1-6.
48. Yücel A. Medical mycology: Yesterday and Today. *Cerrahpasa J Med* 1999; 30: 191-198.
49. Hay RJ, Ashby HR. Fungal infections. In: Griffiths CE, Barker J, Bleiker T, Chalmers R, Creamer D, eds. *Rook's Textbook of Dermatology*. Chichester: Wiley Blackwell, 2016: 32.1e329643.
50. Zuber TJ, Baddam K. Superficial fungal infection of the skin. Where and how it appears help determine therapy. *Postgrad Med* 2001; 109: 117-20, 123-6, 131-2.
51. Hall JH Jr, Leshner JL Jr. Superficial fungal infections. *Pediatrician* 1991; 18: 224-232.
52. Edwards JE. *Candida* species. In: Bennett JE, Dolin R, Blaser MJ, eds. *Mandell, Douglas,*

- and Bennett's Principles and Practice of Infectious Diseases, Updated Edition. 8th ed. Philadelphia, PA: Elsevier Saunders; 2015.
53. James WD, Berger TG, Elston DM. Diseases resulting from fungi and yeasts. In: James WD, Berger TG, Elston DM, eds. *Andrews' Diseases of the Skin*. 12th ed. Philadelphia, PA: Elsevier; 2016.
 54. Brodell RT, Elewski B. Superficial fungal infections. Errors to avoid in diagnosis and treatment. *Postgrad Med* 1997; 101: 279-287.
 55. Sobel JD. Recurrent vulvovaginal candidiasis. *Am J Obstet Gynecol* 2016; 214: 15e21.
 56. Lisboa C, Costa A, Ricardo E, Santos A, Azevedo F, Pina-Vaz C, Rodrigues A. Genital candidosis in heterosexual couples. *J Eur Acad Dermatol Venerol* 2010; 25: 145-151.
 57. McMillan R, Young H, Ogilvie M, Scott G. *Clinical practice in sexually transmissible infections* (1 ed). London, Saunders, 2002.
 58. Wisdom A, Hawkins D. *Diagnosis in color: Sexually transmitted Diseases* (2 ed.). London, Mosby Ltd., 1997.
 59. Evans EG. Causative pathogens in onychomycosis and the possibility of treatment resistance: a review. *J Am Acad Dermatol* 1998; 38: S32-56.
 60. Palma-Carlos AG, Palma-Carlos ML. Chronic mucocutaneous candidiasis revisited. *Allerg Immunol (Paris)* 2001; 33: 229-232.
 61. Rybojad M, Melec P, Feuilhade M, Morel P, Bourrat E. Familial chronic mucocutaneous candidiasis associated with autoimmune polyendocrinopathy. Treatment with fluconazole: 3 cases. *Ann Dermatol Venereol* 1999; i26: 54-56.