EMERGING INFECTIONS NETWORK LETTER

Recognition of Antifungal-Resistant Dermatophytosis by Infectious Diseases Specialists, United States

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Antifungal-resistant dermatophyte infections have recently emerged as a global public health concern. A survey of US infectious diseases specialists found that only 65% had heard of this issue and just 39% knew how to obtain testing to determine resistance. Increased clinician awareness and access to testing for antifungal-resistant dermatophytosis are needed.

Dermatophytosis (i.e., ringworm, tinea) is a common superficial fungal skin infection caused by dermatophyte molds (1). In the past decade, widespread outbreaks of antifungal-resistant dermatophytosis have occurred in South Asia because of the spread of *Trichophyton indotineae* fungus, which causes extensive pruritic plaques on the trunk, extremities, groin, and face in immunocompetent persons (2–4). Unlike typical dermatophyte skin infections, those involving *T. indotineae* fungus often do not resolve with over-the-counter topical antifungals or oral terbinafine (first-line systemic therapy) (2,3).

T. indotineae fungus has been detected in at least 11 US states, and recalcitrant dermatophytosis caused by antifungal-resistant T. rubrum fungus also has been reported (2,5,6). Affected patients have experienced diagnostic delays and received inappropriate therapies, including topical corticosteroids, which can worsen dermatophytosis (2). Antifungal-resistant dermatophytosis cases in the United States probably are underrecognized because this condition is not reportable to public health authorities in any state (https://www.cdc.gov/fungal/php/case-reporting) and most superficial fungal infections are not confirmed with diagnostic testing (7). Further, laboratory capacity to identify dermatophyte species and perform antifungal-susceptibility testing (AFST) may

be limited, given that definitive identification of certain species, such as *T. indotineae*, requires advanced molecular techniques (e.g., using internal transcribed spacer region sequencing) and most clinical laboratories do not perform AFST for molds (2,8).

Antifungal-resistant dermatophytosis cases might require infectious diseases (ID) clinician consultation both to manage antifungal use and because of the potential effect of these infections on highly immunocompromised patients. Understanding ID clinicians' level of awareness regarding antifungal-resistant dermatophytosis and access to laboratory testing could improve strategies to increase disease recognition and facilitate early appropriate treatment. Therefore, we surveyed ID clinicians by using the Emerging Infections Network (EIN) (https://ein.idsociety.org), a sentinel network of ID physicians and other ID specialists.

During December 2023, EIN distributed a survey link to ≈3,000 member subscribers on 3 separate occasions ≈1 week apart. EIN queries are designated as non-human subjects research by the institutional review board of the University of Iowa.

In total, we received 158 responses (Table). The most common practice setting was university hospital (47%), followed by community hospital (16%) and nonuniversity teaching hospital (12%). Most respondents were adult ID physicians (80%), followed by pediatric ID physicians (12%), pharmacists (4%), nurse practitioners, physician assistants or physician associates (1%), or other (3%). Overall, 103 (65%) respondents had heard of antifungal-resistant dermatophytosis before receiving the survey; among those, most (58%) had heard of it through previous EIN list-serv emails. Seventeen percent of respondents reported seeing or consulting on a patient who had dermatophytosis that was potentially resistant to treatment or concerning for resistance within the previous year.

Approximately half (47%) of respondents reported that if they saw a patient with potentially resistant dermatophytosis, they either would not be able to obtain laboratory testing to determine the species (16%) or were unsure whether they could obtain such testing (31%) (Table). Likewise, most respondents (61%) reported they would either not be able to obtain testing to determine if a dermatophyte was resistant to antifungals (16%) or were unsure whether they could obtain such testing (45%).

In summary, our survey found that approximately one third (35%) of ID clinicians had not heard of antifungal-resistant dermatophytosis. Only 53% of respondents reported knowing how to obtain access to dermatophyte speciation testing,

Table. Survey regarding 158 infectious diseases clinicians' awareness of antifungal-resistant dermatophyte infections and access to laboratory testing, United States, 2023

laboratory testing, United States, 2023	
Survey query	No. (%) respondents
Primary setting of clinical practice	
University hospital	75 (47)
Community hospital	26 (16)
Non-university teaching hospital	19 (12)
Veterans Affairs hospital or Department of Defense	13 (8)
Outpatient setting only	8 (5)
Free-standing children's hospital	5 (3)
City, county, or public hospital	4 (3)
Other	4 (3)
Question not answered	4 (3)
Type of clinical practice	
Adult infectious diseases physician	126 (80)
Pediatric infectious diseases physician	19 (12)
Pharmacist	6 (4)
Nurse practitioner, physician assistant, or physician associate	2 (1)
Other	4 (3)
Question not answered	1 (1)
How did you hear about the issue of antifungal-resistant dermatophyte infections?	(select all that apply)
I had not previously heard of this issue	55 (35)
Answers among those who had heard of this issue, n = 103	` '
Emerging Infections Network emails	60 (58)
CDC webpage or Morbidity and Mortality Weekly Report	46 (45)
Scientific publication	28 (27)
News reports	18 (17)
From clinical colleagues	18 (17)
Clinician education website (e.g., Doximity and Medscape)	6 (6)
Point-of-care medical resource (e.g., UpToDate)	4 (4)
Social media	2 (2)
Other	6 (6)
In the past year, I have seen or consulted on a patient with a dermatophyte infecti	
concerning for resistance	ion that had petermany redictant to treatment of
Yes	27 (17)
No	119 (75)
Unknown	12 (8)
If I saw a patient with a potentially resistant dermatophyte infection, I would be abl	
species	
Yes	83 (53)
No	26 (16)
Unknown	49 (31)
I would be able to obtain testing to determine if a dermatophyte is resistant to anti-	1 7
Yes	62 (39)
No	25 (16)
Unknown	71 (45)

and even fewer (39%) reported knowing how to obtain testing for dermatophyte resistance (Table). Those findings probably reflect the relatively recent recognition of these infections in the United States and the fact that testing to identify *T. indotineae* and other resistant dermatophyte species is limited to select US mycology reference centers and public health laboratories (2,8).

Dermatophyte species identification and AFST are critical to guiding antifungal treatment decisions for patients with potentially resistant dermatophytosis and for monitoring population-level trends in resistance profiles to inform treatment guidelines (8). National guidelines for treating antifungal-resistant dermatophytosis are lacking, but the azole antifungal itraconazole has been used successfully for patients

with dermatophytosis caused by T. indotineae and terbinafine-resistant T. rubrum fungi (2,9). However, clinicians should be aware of challenges with itraconazole, including pharmacokinetic variability (e.g., absorption), insurance coverage, drug-drug interactions, need for prolonged treatment (e.g., ≥ 6 weeks), and reports of emerging itraconazole-resistant T. indotineae and T. rubrum fungi (2,6,9,10).

One limitation of our survey is its low response rate and nonrepresentative nature. Furthermore, the survey might overestimate clinician awareness of antifungal-resistant dermatophytosis and laboratory testing access if clinicians experienced in this topic were likelier to respond. Despite those limitations, our survey highlights a need for increased awareness of antifungal-resistant dermatophytosis and

increased laboratory capacity to identify and perform susceptibility testing for dermatophytes to address this emerging public health concern.

Healthcare professionals can find information about recognizing, diagnosing, treating, and reporting emerging dermatophyte infections online (https://www.aad.org/member/clinical-quality/clinical-care/emerging-diseases/dermatophytes), including information about laboratories that can perform testing (https://www.aad.org/member/clinical-quality/clinical-care/emerging-diseases/dermatophytes/recognizing-trichophyton-indotineae#testing). Those websites were developed as a collaboration between the Centers for Disease Control and Prevention and the American Academy of Dermatology's Emerging Diseases Task Force.

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About the Author

Dr. Gold is a medical epidemiologist in the Mycotic Diseases Branch, Division of Foodborne, Waterborne, and Environmental Diseases, National Center for Emerging and Zoonotic Infectious Diseases, Centers for Disease Control and Prevention, Atlanta, Georgia, USA. His research interests include the epidemiology and prevention of fungal infections.

References

- Gupta AK, Ryder JE, Chow M, Cooper EA. Dermatophytosis: the management of fungal infections. Skinmed. 2005;4:305– 10. https://doi.org/10.1111/j.1540-9740.2005.03435.x
- 2. Caplan AS, Chaturvedi S, Zhu Y, Todd GC, Yin L, Lopez A, et al. Notes from the field: first reported U.S. cases of tinea caused by *Trichophyton indotineae* New York City,

- December 2021-March 2023. MMWR Morb Mortal Wkly Rep. 2023;72:536-7. https://doi.org/10.15585/mmwr.mm7219a4
- Verma SB, Panda S, Nenoff P, Singal A, Rudramurthy SM, Uhrlass S, et al. The unprecedented epidemic-like scenario of dermatophytosis in India: I. Epidemiology, risk factors and clinical features. Indian J Dermatol Venereol Leprol. 2021;87:154–75. https://doi.org/10.25259/IJDVL_301_20
- Gupta AK, Renaud HJ, Quinlan EM, Shear NH, Piguet V. The growing problem of antifungal resistance in onychomycosis and other superficial mycoses. Am J Clin Dermatol. 2021;22:149–57. https://doi.org/10.1007/ s40257-020-00580-6
- Cañete-Gibas CF, Mele J, Patterson HP, Sanders CJ, Ferrer D, Garcia V, et al. Terbinafine-resistant dermatophytes and the presence of *Trichophyton indotineae* in North America. J Clin Microbiol. 2023;61:e0056223. https://doi.org/10.1128/ jcm.00562-23
- 6. Gu D, Hatch M, Ghannoum M, Elewski BE. Treatment-resistant dermatophytosis: a representative case highlighting an emerging public health threat. JAAD Case Rep. 2020;6:1153–5. https://doi.org/10.1016/j.jdcr.2020.05.025
- Benedict K, Wu K, Gold JAW. Healthcare provider testing practices for tinea and familiarity with antifungal-drugresistant tinea – United States, 2022. J Fungi (Basel). 2022;8:831. https://doi.org/10.3390/jof8080831
- Lockhart SR, Smith DJ, Gold JAW. Trichophyton indotineae and other terbinafine-resistant dermatophytes in North America. J Clin Microbiol. 2023;61:e0090323. https://doi.org/ 10.1128/jcm.00903-23
- 9. Khurana A, Agarwal A, Agrawal D, Panesar S, Ghadlinge M, Sardana K, et al. Effect of different itraconazole dosing regimens on cure rates, treatment duration, safety, and relapse rates in adult patients with tinea corporis/cruris: a randomized clinical trial. JAMA Dermatol. 2022;158:1269–78. https://doi.org/10.1001/jamadermatol.2022.3745
- Burmester A, Hipler U-C, Uhrlaß S, Nenoff P, Singal A, Verma SB, et al. Indian *Trichophyton mentagrophytes* squalene epoxidase *erg1* double mutants show high proportion of combined fluconazole and terbinafine resistance. Mycoses. 2020;63:1175–80. https://doi.org/10.1111/myc.13150

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Corrections

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The name of author Glenn Patriquin was misspelled in Case Series of Jamestown Canyon Virus Infections with Neurologic Outcomes, Canada, 2011–2016 (V. Meier-Stephenson et al.). The article has been corrected online (https://wwwnc.cdc.gov/eid/article/30/5/22-1258_article).

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The name of author Carlos E. Sanz-Rodriguez was misspelled in Outbreak of Intermediate Species *Leptospira venezuelensis* Spread by Rodents to Cows and Humans in *L. interrogans*–Endemic Region, Venezuela (L. Caraballo et al.). The article has been corrected online (https://wwwnc.cdc.gov/eid/article/30/8/23-1562_article).