

CANDIDA AURIS: EMERGENCE AND EPIDEMIOLOGY OF A HIGHLY PATHOGENIC YEAST

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ABSTRACT

Candida auris is a multidrug-resistant emerging yeast, which was responsible for healthcare-associated infection outbreaks, and was cataloged as a new species in 2009, after being isolated from a patient's ear canal secretion in Japan. Since the notification of this first occurrence, numerous cases have been reported throughout the world, including Brazil. *C. auris* affects mainly inpatients, patients in intensive care units, exposed to broad-spectrum antifungal medications and who make use of vascular catheters. Currently, this yeast is one of the main responsible for invasive infections in hospitals and has been cause of concern by authorities and organs due to its rapid dissemination and difficult treatment caused by its low susceptibility to antifungal agents traditionally used in clinical practice. As a contributor to the severity of infections associated with *C. auris*, the transmission mechanism is still unknown, which implies in a lack of control of the microorganism and high mortality rates. Thus, this literature review presents relevant information in order to alert the importance of *C. auris* as an etiological agent of systemic infections, as well as its epidemiology and the real challenges of the treatment.

Keywords: *Candida auris*; candidiasis; candidemia; multidrug-resistance; biofilm; epidemiology; diagnosis

Infections caused by *Candida* genus yeasts are known as candidiasis or candidosis¹. Candidiasis, among other clinical forms, can also be considered a sexually transmitted disease (SDT) and is clinically manifested through lesions, which can be classified as superficial with cutaneous and mucosal involvement, or systemic, disseminated, and of high severity². This type of mycosis has the mouth, the throat, the tongue, scalp, genitals, fingers, nails and internal organs as infection sites¹. Its origin can be exogenous or endogenous, being the latter the main transmission mechanism, in which the *Candida* species that are part of the microbiota in tissues and organs, when faced with the vulnerability of the host, for example in immunocompromised situations, become opportunistic. The exogenous way is a transmission mechanism that occurs mainly through direct contact of health professionals with patients or contaminated medical-hospital issues².

The main virulence factors of yeasts that induce infection are³:

- a) secretion of extracellular enzymes such as phospholipases and proteinases, which degrade the host tissue leading to tissue invasion³;
- b) Production of toxic substances that damage the cells³;
- c) Creation of biofilms on cells and inert surfaces³;
- d) Formation of pseudohyphae by certain species of *Candida* spp.³;
- e) Production of hemolysins²;
- f) The ability to adhere to medical and hospital materials and host cells³.

The sum of these mechanisms with the weakness of the host response may lead to candidiasis³.

Microorganisms usually live naturally in communities, through the formation of biofilm; this is due to the advantages of being in association, among which

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the main one is a greater protection against the action of antifungal agents and the immune system defenses of the host².

The epidemiology of *Candida* spp. can be variable, according to the virulence of the isolate, parasitic load and predisposition of the host, and all these factors make them pathogenic yeasts¹. *Candida* species are responsible for bloodstream infections (BSIs) in hospitalized patients, and affect mainly those who are in intensive care units, exposed to broad spectrum antifungal agents, internal vascular catheters, parenteral nutrition, abdominal surgery and immunosuppressants⁴. The BSIs represent a challenge to public health due to their severity and intensity, leading to an increase in the length of hospital stay, increasing hospitalization costs, as well as morbidity and mortality rates in infected patients. In studies conducted throughout 3 years in 49 hospitals in the United States, *Candida* spp. was indicated as one of the main etiological agents of BSIs, responsible for 7.6% of the cases².

Among the most common causes of BSIs is candidemia, which is the syndrome most frequently associated with invasive candidiasis⁵ and high mortality rates (40% to 60%)². Invasive candidiasis is one of the major clinical manifestations of the *Candida* genus caused by *C. albicans* and non-*albicans*⁶ *Candida*, these, in turn, respond to at least 50% of the invasive infections by *Candida* spp.³, and can be fluconazole-resistant or difficult to extinguish due to the fact that they produce biofilm⁷. The *Candida* yeast has more than 200 existing species, among which are exemplified especially - *C. albicans*, *C. glabrata*, *C. parapsilosis*, *C. krusei*, *C. tropicalis*, *C. haemulonii*, *C. guilliermondii*, and the most recently identified species, *C. auris*^{1,6}. The latter has been classified as a highly pathogenic new species and it is multiresistant to antifungal agents traditionally used in clinical practice, being the cause of several systemic infections, which can be fatal⁶. In 2013, a study reported 33% of lethality in cases of BSIs attributed to *C. auris* in all patients and 57% in the subgroup of patients admitted to intensive care therapy units; however these rates can be attributed to the severity of underlying diseases in these patients⁸.

The rapid emergence of *C. auris* and the resistance to the three major classes of antifungal drugs (azoles, echinocandins and polyenes)⁹, the horizontal transmission among hospitalized patients, leading to healthcare-associated infection outbreaks, and the high mortality rates associated, make *C. auris* one of the most current causes of invasive infections in hospitals and reason of concern due to its evolution and worldwide spread⁴. Furthermore, fungal otomastoiditis, which is a rare and possibly

fatal disease for immunosuppressed patients, has *C. auris*^{10,11} as its etiologic agent. There have been increasing cases of otologic infection caused by *C. auris*¹⁰. Although the origin of the infections is inconclusive, it is not yet known whether these isolates are in the hospital environment or are of endogenous origin¹². Evidence indicates that the dissemination of *C. auris* in the hospital environment may occur through contact with contaminated surfaces and devices and also in an interpersonal way, thus alerting services to this opportunistic pathogen⁹.

Given the above, this review aims to address epidemiological aspects of *C. auris* candidiasis, evidencing cases already described in Brazil as well as frequency of cases by region. Additionally, we present the methods most used for the identification of *C. auris*, the data of microbial susceptibility against the most varied classes of antifungal drugs, and therapeutic challenges. Therefore, we intend to contribute to the elucidation of the clinical importance of the theme, gathering relevant information for a better understanding of which are the factors that promote the expansion and opportunism of the disease, which generates an alert about this noticeable species among the non-*albicans* *Candida* species currently, in regard to high virulence and pathogenicity.

METHODS

A systematic bibliographic review study was conducted in databases and online collections of the Elsevier Publisher (ScienceDirect) and the National Institutes of Health (PubMed) virtual health library from April to June 2017.

The terms used in the electronic searches were “*Candida auris*”, “candidíase/candidiasis”, “candidemia”, “*Candida auris* diagnóstico/diagnosis”, “*Candida auris* epidemiologia/epidemiology”, “*Candida auris* multirresistente/multidrug-resistant”, “*Candida auris* biofilme/biofilm” and “*Candida auris* tratamento/ treatment”.

In addition, an online survey was conducted to access documents of national (ANVISA) and international organizations, which could contribute with data to compose the present study.

The purpose of this research was to address the most relevant and current aspects of identification, epidemiology, clinical manifestations, treatment, and prevention of infections by *C. auris*, a species still little known in Brazil, whose information is still controversial between official records and scholars, indicating the demand for care and attention.

The inclusion criteria were scientific papers that explored the proposed theme, published between the 2003 and 2017. We excluded studies that did

not have adequate bibliographic references, which were incomplete or tertiary sources. In addition, it we decided not to include theses, dissertations, monographs and books.

Identification

In order for the dissemination of *C. auris* to be prevented and controlled in a hospital environment, it is of the utmost importance that the identification of the isolate be done quickly. When the identification of the species is suspected in an isolate, or even after confirmation, it is the responsibility of the laboratory to immediately inform the Hospital Infection Control Committee (CCH) of the health service⁹.

Infections caused by this new species of *Candida* are diagnosed by blood culture or culture of other body fluids. Its specific identification requires molecular laboratory methods, such as D1-D2 or Internal Transcribed Spacer (ITS)^{9,12,13}, or desorption/ionization with mass spectrometry, and the Matrix-Assisted Laser Desorption Ionization – Time of Flight (MALDI-TOF), the fastest and most suitable method described in the literature for the identification of *C. auris*¹⁴, making it possible to create reference spectrum and the care to be taken when adopting this approach.

The laboratory that isolates by the phenotypical methods one of the following species: *C. haemulonii*, *C. famata*, *C. sake*, *C. catenulata*, *C. lusitaniae*/*C. guilliermondii*, *Saccharomyces cerevisiae* and *Rhodotorula glutinis* and has the MALDI-TOF equipment or the genome sequencing should confirm whether the isolate is *C. auris*. It is imperative that it is made sure that MALDI-TOF contains protein profiles in its database that enable the detection of *C. auris* isolates. Currently, in Brazil, only *Research Use Only* (RUO) Bruker libraries of reference spectra have profiles of *C. auris*⁹.

In the case of laboratories that do not have the equipment, but are able to carry out screening tests for *C. auris*, they should conduct a culture examination and direct research with India ink to discard the presence of the encapsulated yeast. Subsequently, screening tests for possible identification of *C. auris* should be carried out with the aim of discarding the other non-*albicans* *Candida* species, among them filamentation tests on Tween-80 corn meal agar and/or colony color analysis on Sabouraud dextrose agar or other agar and/or color analysis on chromogenic agar and/or presence of germ tube (PGT). Isolates that do not present carotenoid pigments, negative PGT, and/or pseudohyphae formation on corn meal agar, and pink to purple coloration on chromogenic agar should be separated for further confirmation and, if positive for *C. auris*, the Commission for Hospital Infection Control (CHIC) should be notified⁹.

In the case of positive screening tests, the competent organ should perform biochemical tests to screen species phenotypically similar to *C. auris*; determine the susceptibility of the yeast to antifungal agents, to verify the resistance of the suspected isolate; to perform sequencing of the D1-D2 or ITS1-ITS4 regions and/or MALDI-TOF for confirmation of *C. auris*, or refer directly to reference laboratories⁹.

Therefore, criteria for considering an isolate suspect of *C. auris* are those phenotypically identified as *C. haemulonii*, *C. famata*, *C. sake*, *C. catenulata*, *C. lusitaniae*/*C. guilliermondii*, *S. cerevisiae* and *R. glutinis* and that presented high minimal inhibitory concentrations (MIC) for fluconazole (FCZ; MIC \geq 32 μ g/mL), voriconazole (VCZ; MIC \geq 2 μ g/mL), amphotericin B (APHB; MIC \geq 2 μ g/mL), and echinocandins (ECHs), such as anidulafungin (ADA; \geq 2 μ g/mL)⁹.

This new species can be erroneously identified as those mentioned above^{9,14}, if identification is made through classical methods^{6,9}. Gaitan et al.⁶ initially described isolates identified as *S. cerevisiae*, *C. sake*, *C. lusitaniae*, *C. haemulonii*, and eight inconclusive isolates, and 8 (eight) were isolated from the blood and 4 (four) from the tip of the catheter, from patients hospitalized in an ICU in a European hospital. Confirmation of *C. auris* from all isolates was conducted by molecular methods.⁶

In Colombia, isolates were initially identified as *C. haemulonii*, *C. famata*, *C. albicans* or *C. tropicalis*. After observing the unusual prevalence and the micromorphological discordance, the strains were cultured in CHROMagar *Candida* medium using MALDI-TOF. Pink colonies were observed in the CHROMagar *Candida* medium and there was subsequently molecular confirmation of *C. auris* in all isolates¹⁴.

According to Chowdhary et al., growth at 40 °C may differentiate the isolates of *C. auris* that are mistakenly identified as *C. haemulonii* by VITEK, considering that the species *Candida haemulonii* does not grow at 40 °C.^{8,12} Kumar et al. reported a rapid and inexpensive method using the CHROMagar *Candida* supplemented with Pal's medium to differentiate *C. auris* from isolates identified as *C. haemulonii* by VITEK2. All isolates of *C. auris* showed smooth, white-to-cream colonies at 37°C and at 42 °C, after 24 and 48 hours of incubation and did not produce pseudohyphae. The isolates of *C. haemulonii* had a weak growth of smooth, light-pink colonies, in 24 and 48 hours, and there was no pseudohyphae production either. The *C. haemulonii* yeast did not grow at 42 °C^{8,15}.

Thus, it is observed in the literature that the identification and the diagnoses of *C. auris* are definitely carried out by molecular methods^{6,8,9,12-21}.

Routine methods (direct, culture, biochemical tests) are used only as screening for initial detection of yeast in clinical samples.

As a recommendation to hospitals that are not able to routinely detect this new *Candida* species, it is advisable to monitor the monthly number of positive blood cultures with non-*albicans* *Candida* species, as any increase may be indicative of a potential outbreak, possibly related to *C. auris*⁸.

Epidemiology

Due to the severity of the outbreaks that it has been causing in hospitals around the planet, *C. auris* shows a real and eminent health risk and should be observed with great caution. It was isolated for the first time from the external auditory canal of a patient in Japan and was described as a new species in 2009¹⁶. Since that episode, several other countries have reported infections, including Japan, South Korea, India, Pakistan, South Africa, Kenya, Kuwait, Brazil, Israel, Venezuela, Colombia, United Kingdom, United States, and Canada. These findings prove that the species is not limited to a particular region^{9,12}.

The epidemiology of *C. auris* specifically in Brazil is still inconclusive. While studies conducted in 2016^{12,17,18} already indicate the presence of the yeast in the country, ANVISA⁹, through an official statement, reported that up to the present moment there have been no occurrences of infections by *C. auris* in our country⁹. However, this declaration does not exclude the possibility that the species is already present in the region, since its prevalence is not sufficiently known due to difficulties in its identification and diagnosis⁹.

In America, the first outbreak occurred in Venezuela between March 2012 and July 2013, in 18 affected patients, among which 13 were pediatric. The isolates were initially identified as *C. haemulonii* and after sequencing of the ITS region and analysis by Amplified Fragment Length Polymorphism (AFLP) it was identified that the microorganism involved was *C. auris*⁹.

Between February and July 2016 in Colombia, the outbreak occurred in 6 different hospitals totaling 17 inpatients. Among the 17 patients, 9 were male; the age group ranged between 0-77 years, 15 were hospitalized in ICUs and 2 in medical units. Of the total, 13 patients showed fungemia and in the other four, *C. auris* was isolated from peritoneal fluid, cerebrospinal fluid, bone, or urine. Most of the patients had central venous catheter, urinary catheter, and mechanical ventilation. Some had risk factors for

candidemia: red blood cell transfusion, parenteral nutrition, abdominal surgery, hemodialysis, diabetes, pancreatitis, cancer, and HIV infection¹⁴.

In India, the highest incidence of *C. auris* infection has occurred in public hospitals. In 27 ICUs 1400 cases of *Candida* infection were reported, among which 74 (5.3%) were isolated and confirmed as *C. auris*. In patients with diagnosis of candidemia, those infected with *C. auris* had an average period of hospital stay of 25 days, which was higher than those infected with non-*auris* *Candida*, with a mean of 15 days²². In the city of New Delhi, in two hospitals, 12 *C. auris* were isolated, collected between 2009 and 2011. The isolates from these hospitals were clonal, indicating an inter-hospital transmission. Most patients presented permanent urinary catheter and persistent candidemia - and the mortality rate was of 33%²³.

Between October 2012 and October 2013, 4 isolates were sent to the National Institute of Communicable Diseases in Johannesburg, South Africa, from 4 patients presenting candidemia and hospitalized in different institutions. The isolates were initially identified as *C. haemulonii* and *R. glutinis* by tests commonly used and later *C. auris* was correctly identified by genome sequencing²⁴.

The first and largest outbreak of *C. auris* in Europe occurred in a cardiac center in London between April 2015 and July 2016, totaling 50 cases. Among these, 44% (n=22/50) developed *C. auris* candidemia. Through an environmental sampling the persistent presence of the yeast was detected around the spaces between the beds²⁵.

In Spain, between April and June 2016, eight isolates from four patients (two per patient), at the *Hospital Universitario e Politécnico La Fe* in Valencia, were obtained from blood cultures and catheter tips. All patients were adults and were hospitalized in the ICU. The four cases were identified as *C. auris* after confirmation by sequencing the ITS region⁶.

Lockhart et al.²⁶, carried out a study throughout 3 years (2012 to 2015), to understand the emergence and epidemiology of *C. auris* from isolates of 54 patients from Pakistan, India, South Africa, Venezuela and Japan. From 41 isolates, which had information regarding the patients, antifungal susceptibility testing and complete genome sequencing (WGS) testing were performed. The clinical condition of patients were 41% diabetes mellitus, 51% had undergone surgery recently, 73% had central venous catheter, and 41% were receiving systemic antifungal therapy, since *C. auris* had been isolated. The mean time since admission until infection was of 19 days, where 61% of patients had blood infection and 59% died. Among the isolates, 93% were fluconazole-resistant, 35% were resistant to amphotericin B and 7% to

echinocandins, 41% were resistant to 2 antifungal classes and 4% to 3 classes²⁶.

Also, isolated cases and outbreaks have been reported in five different continents, including Europe, Asia, North America, South America, and Africa. A case in Norway, probably from another country outside the continent, confirms that there is a risk of transmission of yeast through the hospital transfer of patients. In a recent study, *C. auris* isolates present in the United Kingdom were shown to have diverse geographic origins, suggesting multiple introductions in the country⁸.

Considering the above, it was observed that *C. auris* is a species of high prevalence in various regions and climatic conditions around the world, being associated with high mortality rates in hospitalized patients. Due to the inherent difficulties of detection and diagnosis, its epidemiology is still controversial in Brazil.

Susceptibility and treatment challenges

The indiscriminate use of antifungals may have induced *C. auris* to become a multidrug-resistant pathogen, limiting treatment efficacy because of its resistance to fluconazole (FCZ), and variable susceptibility to other azoles, amphotericin B (APHB) and echinocandins (ECHs)^{12,27,28}, as it can be observed in Table 1²¹.

The ability to produce biofilms, the survival and dissemination in hospital settings, besides the risk of outbreaks, contribute to the pathogenicity of *C. auris*. According to the literature, *C. auris* may be as pathogenic as *C. albicans*. Despite the fact that it forms less biofilm when compared to *C. albicans*, this new species has demonstrated a greater virulence, but still needs to be further investigated. These factors, together with the resistance of *C. auris* to most antifungal agents, may explain why it is considered one of the most risky pathogens currently¹³.

According to Sherry et al.¹³, *C. albicans* presented a higher biofilm mass, fact that is directly related to its pathogenicity. *C. auris*, in turn, produced significantly less biofilm. However, the two species had a higher biofilm production compared to *C. glabrata*. The *C. albicans* biofilms were densely packed with pseudohyphae, *C. glabrata* formed a sparse biofilm with yeast cells, without extracellular matrix, and *C. auris* presented biofilm containing budding yeast and pseudohyphae. Still according to Sherry et al.¹³, broth microdilution susceptibility testing was also performed with FCZ, voriconazole (VCZ), caspofungin (CPA), micafungin (MCA), amphotericin B liposomal (APHB lipo), APHB, and chlorhexidine (CXA). For planktonic and sessile cells of *C. auris* showing MIC > 32 µg/mL, at FCZ and VCZ, it presented minimal activity against *C. auris* planktonic cells (Table 1). APHB lipo was active against planktonic cells (0.25-1.0 µg/mL) and decreased metabolic activity of *C. auris* requiring much higher concentrations of the same antifungal (16 µg/mL). APHB was more effective, requiring 4 µg/mL to be more active against biofilm. The MCA was the most active ECHs, with < 0.5 µg/mL to inhibit planktonic cells, compared to 2-32 µg/mL for CPA. However, these two antifungal agents, MCA and CPA were ineffective against biofilm (> 32 µg/mL). The CXA presented higher activity, with < 0.02% to inhibit, in fact, planktonic and sessile cells (Table 1). Among the antifungals analyzed, CPA was inactive against *C. auris* biofilm, which is noteworthy, because this antifungal agent is effective against *Candida* spp. biofilms. These factors may influence the virulence and survival of *C. auris*, and contribute to the outbreaks reported in hospital environments.

Susceptibility studies conducted by Chowdhary et al.¹² demonstrated that *C. auris* presented high MIC to FCZ (CIM90 > 64 µg/mL), VCZ, APHB (2 µg/mL) and the ECHs (Table 1). The CXA presented efficacy as disinfectant in the prevention and control of infection

Table 1: Susceptibility of clinical isolates of *C. auris* to *in vitro* treatment with various antifungal agents.

MIC (µg/mL) and MIC ₉₀												
FLZ	VCZ	ICZ	PSZ	ECHs	MCA	CPA	ADA	APHB lipo	APHB	FCA	CXA	References
> 32					<0.5	2-32		0.25-1.0 e 16*	4		<0.02%	13
>64	2								2			12
>256	2	RV	RV	RV					RV			6
		≥ 2										15
									>1			27
≥ 32	≥ 2			≥ 8					≥ 2	≥128		26
≥32	≥ 2						≥ 2		≥ 2			9

MIC (minimum inhibitory concentration); MIC₉₀ (minimum inhibitory concentration capable of inhibiting 90% of isolates); FCZ (fluconazole); VCZ (voriconazole); ICZ (itraconazole); PSZ (posaconazole); ECHs (echinocandins); MCA (micafungin); CPA (caspofungin); ADA (anidulafungin); APHB lipo (amphotericin B liposomal); APHB (amphotericin B); FCA (flucytosine); CXA (chlorhexidine). *MIC related to biofilm cells.

by *C. auris*, through hand hygiene, oral gargling and skin disinfection, as well as topical use of nystatin and terbinafine¹².

Gaitán et al.⁶ observed that *C. auris* is resistant to FCZ and VCZ and has MIC variable to ECHs and APHB. The isolates were resistant to FCZ (MIC > 256 µg/mL), VCZ (MIC 2 µg/mL) and to PSZ, ICZ, ECHs and APHB (Table 1)⁶.

Kumar et al.¹⁵, observed that *C. auris* was ICZ-resistant (MIC ≥ 2 µg/mL), indicating expressive virulence factors, including phospholipase, proteinase and hemolysin activity (Table 1).

Calvo et al.²⁷, between March 2012 and July 2013, analyzed *C. auris* in 18 patients with candidemia, from a medical center in Maracaibo, 13 pediatric patients, with men age of 26 days and 5 adults. All had received antibiotics and invasive procedures, at the onset of treatment with antifungal drugs the catheter was removed. Patients' survival was of up to 30 days. Through AFLP fingerprinting the isolates were genotyped and suggested a clonal outbreak. The isolates were azole-resistant, susceptible to anidulafungin (ADA) and MIC > 1 µg/ml for APHB (Table 1)²⁷.

Lockhart et al.²⁶, found MICs to *C. auris* with the following antifungal drugs: FCZ (≥ 32 µg / mL), VCZ (≥ 2 µg / mL), ECHs (≥ 8 µg/mL), flucytosine (FCA) (≥128 µg / mL) and APHB (≥ 2 µg / mL), the values found, in the present study, were similar to the MICs published by ANVISA (2017), FCZ (≥ 32 µg/mL), VCZ (≥ 2 µg/mL), APHB (≥2 µg/mL), and ECHs (ADA≥2 µg/mL) (Table 1)⁹.

In the United Kingdom, a hospital verified the utility of chlorine in hydrogen peroxide products and the *in vitro* activity of chlorhexidine against *C. auris*. The use of chlorhexidine disposable table cloths for surface cleaning and hand hygiene as well as chlorhexidine alcohol prior to intravenous catheter manipulation resulted in a 95% decrease in the incidence of BSIs by *C. auris* after application of these measures⁸.

Thus, it is observed that *C. auris* presents in general a very variable susceptibility to different classes of antifungal and other antimicrobial agents analyzed, with resistance in the majority of cases, which makes treatment difficult. For each new isolate of *C. auris*, it is recommended the use of antifungigram tests to determine MIC, as this should help to select the most appropriate medicine and accelerate the patient's clinical response, who is at serious risk of death when affected by this yeast.

CONCLUSION

C. auris is a multidrug-resistant, pathogenic yeast that can be a source of healthcare-associated infections in hospitals. It has a high potential for horizontal nosocomial transmission. In order for health services to be able to fight these infections, implementing control and prevention measures, it is necessary for laboratories to accurately identify *C. auris*. Thus, the competent authorities will be notified of possible outbreaks, and may adopt appropriate measures, controlling the epidemic.

Routine laboratory methods are insufficient for detection of *C. auris*, which may be erroneously identified as other species. The correct identification of the yeast requires sophisticated molecular methods for proper confirmation at the species level. In suspect cases the isolate should be sent to a reference laboratory capable of identifying the *C. auris* species.

Its epidemiology is variable and *C. auris* has no specific location, being widely found around the world. In Brazil, it is still a controversial issue, because although there are no reports of infection by the species in the country, its incidence is not reliable due to the obstacles found in its identification and diagnosis, a fact that compromises the effective fight against the spread of the yeast.

Chlorhexidine should be seen as a possible strategic solution in preventing *C. auris* infections, serving as a skin disinfectant. Its treatment is still limited, due to the antifungal resistance, an intrinsic species' trait. Currently, there is no evidence that point to an effective standard treatment. This fact, coupled with the high association with mortality, makes *C. auris* an emerging global threat.

Finally, *C. auris* is a species currently responsible for many outbreaks, which has been generating public health alert in Brazil and other countries. Surveillance and Monitoring Agencies in Health Services have issued notes of numerous reports of cases of infections by this yeast, which hardly responds to conventional treatment. Thus, further research should be developed to elucidate the risk factors as well as the transmission mechanism, allowing accurate and effective guidelines for the population to take the necessary actions and precautions to avoid the spread of this pathogen.

Conflict of interest

The authors declare no conflicts of interest.

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