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#118: Frequency and molecular epidemiology of *Aspergillus* isolated from patients with

suspicion of respiratory fungal infection

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ABSTRACT

We retrospectively analyzed a cohort of 146 patients with suspicion of respiratory fungal infection. The prevalence of Aspergillus was calculated, as well as the frequency of each section and species of these fungi isolated from respiratory samples. Resistance to azoles was determined for the Aspergillus of the Fumigati section.

Fifty-seven (39.0%) of the patients studied were positive for Aspergillus. Six sections and 13 different species of Aspergillus were detected, of which 9 were cryptic species. The Nigri section and the species A. niger sensu stricto were the most frequent. The A. fumigatus sensu stricto isolates tested did not show resistance to the azoles, but an A. udagawae strain revealed low susceptibility to voriconazole and an A. pseudofelis strain showed reduced susceptibility to voriconazole and itraconazole.

The genera Aspergillus was frequently detected in the respiratory samples tested and a great variety of species were identified. Surveillance of Aspergillus resistance should persist, although we only detected a decrease in the susceptibility of two cryptic species.

INTRODUCTION

In Portugal, it is estimated that there are approximately 12 600 cases of allergic bronchopulmonary aspergillosis. 776 cases of chronic pulmonary aspergillosis and 240 cases of invasive aspergillosis per year [1], but data about the epidemiology of these diseases is lacking.

Posaconazole, voriconazole and itraconazole are the standardized drugs used for prophylaxis and treatment of aspergillosis [2,3]. However, the use of these antifungals is being compromised by intrinsic resistance of some cryptic species of Aspergillus, as well as by the emergence of secondary resistance in A. fumigatus sensu stricto.

OBJECTIVE

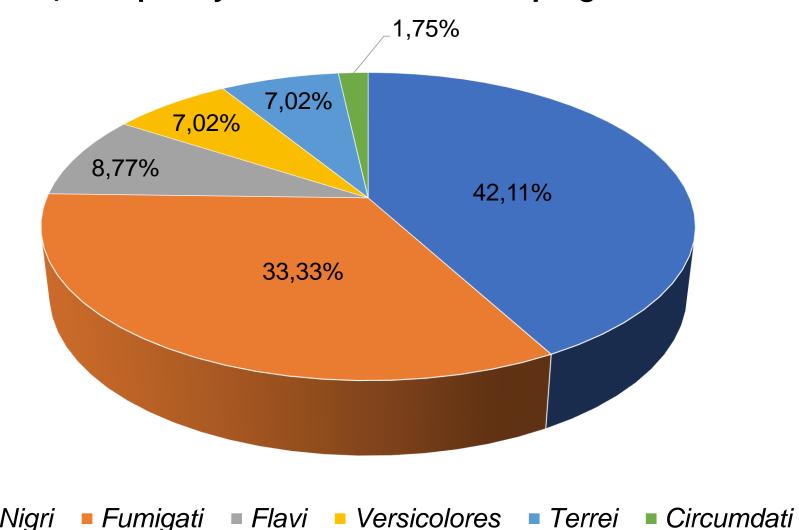
We aimed to contribute to the surveillance of Aspergillus in Portugal by determining the prevalence and frequency of these fungi in respiratory samples from a cohort of patients with symptoms of respiratory fungal infection, as well as by determining the susceptibility to azoles of Aspergillus of the Fumigati section

METHODS Collection of Determination of the Species identification respiratory samples Morphological **Detection of fungi by** susceptibility to trough sequencing of the (Nov. 2011 - Dec. 2017) identification as immunoenzimatic, Positive culture for DNA azoles of isolates calmodulin/β-tubulin in 16 hospitals from molecular and cultural filamentous fungi Aspergillus spp. extraction (n=52) from the genes different districts of techniques Fumigati section (n=58)**Portugal** Bronchoalveolar lavages **Bronchial secretions** Inoculation in Sabouraud **Bronchial aspirate** Calculation of the prevalence of dextrose agar media Expectoration Samson et al.[4]. Aspergillus spp. In the cohort. supplemented with voriitra- and posaconazole Calculation of the frequency of each section and species of Aspergillus detected.

RESULTS

A cohort of 146 patients with suspicion of respiratory fungal infection was studied and the presence of Aspergillus (n=187) in their respiratory samples was recorded (Figure 1A and B). Thirty-seven of these patients (39.0%) were positive for *Aspergillus* spp.

A) Frequency of each section of Aspergillus detected





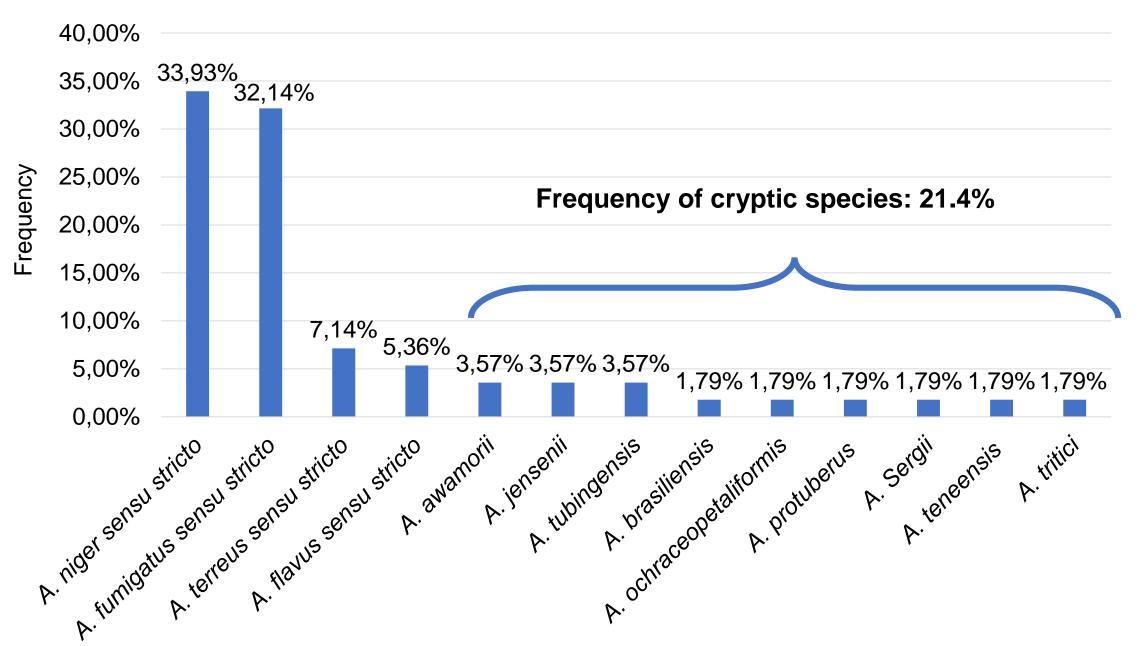


Figure 1 – Frequency of each section (A) and species (B) of Aspergillus identified.

Azole susceptibility of the 52 Fumigati isolates were tested by screening media. Obtained results are exemplified in Figure 2 and summarized in Table 1.

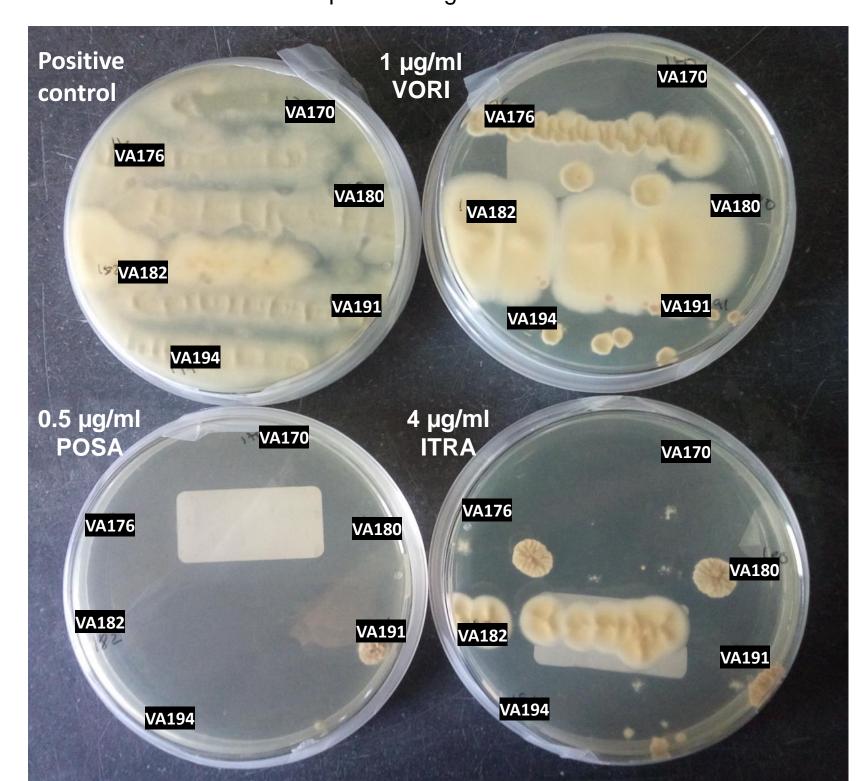


Figure 2 – Susceptibility screening of Aspergillus spp. The isolates VA170, VA176, VA180, VA191 and VA194 were identified as A. fumigatus sensu stricto. The isolate VA182 was identified as A. pseudofelis. VORI – voriconazole; POSA – posaconazole; ITRA –itraconazole.

Table 1 – Number of isolates of each species with significative (+), residual (±) or negative (-) growth on the screening medium supplemented with voriconazole, itraconazole and posaconazole.

	Voriconazole (4 μg/ml)			Itraconazole (4 μg/ml)			Posaconazole (0.5 μg/ml)		
	Growth on the screening media			Growth on the screening media			Growth on the screening media		
Species	+	±	-	+	±	-	+	±	-
A. fumigatus sensu stricto (n=45)	0	0	45	0	1	45	0	0	45
A. lentulus (n=4)	0	0	4	0	0	4	0	0	4
A. pseudofelis (n=1)	1	0	0	1	0	0	0	0	1
A. udagawae (n=2)	1	0	1	0	0	2	0	0	3

FINAL REMARKS

- ✓ There is a high prevalence and a great variety of *Aspergillus* spp. colonizing/infecting the respiratory tract of patients with symptoms of respiratory fungal infection.
- Contrary to what has been previously reported worldwide including in Portugal [5,6] -, the Nigri section and the species A. niger sensu stricto were the most frequent in the respiratory samples tested. This reveals the diversity and complexity of Aspergillus spp. distribution.
- Cryptic species were commonly isolated in the respiratory samples of this cohort and some of them A. pseudofelis and A. udagawae revealed a decrease in susceptibility to some of the azoles tested.
- ✓ A. fumigatus sensu stricto resistance to azoles was not described in this work, but surveillance should persist.

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