



CASE REPORT

Laryngotracheal and pulmonary aspergillosis in an adolescent with acute lymphoblastic leukemia: case report

Gabriela De Martin Silva^{1*}, Fábio da Cunha Peixoto Ladeira¹, João Batista de Oliveira Andrade¹, Scheilla Torres de Oliveira¹, Joaquim Caetano de Aguirre Neto¹

Abstract

Patients undergoing chemotherapy for acute lymphoblastic leukemia are at greater risk for invasive fungal diseases that represent an important cause of morbidity and mortality. Aspergillosis is a fungal infection, usually pulmonary, that rarely affects the laryngeal and tracheal regions. This is a case report about an immunocompromised adolescent undergoing chemotherapy for acute lymphoblastic leukemia, who evolved with febrile neutropenia and pulmonary and laryngotracheal aspergillosis. After 3 weeks of antifungal treatment using voriconazole, a chest tomography showed the reduction of the multiple pulmonary nodules and the fibronasolaryngoscopy endorsed a complete response to the treatment of laryngotracheal lesions.

Keywords: acute lymphoblastic leukemia; invasive fungal disease; aspergillosis; adolescent.

¹Hospital Santa Casa de Misericórdia, Belo Horizonte, MG, Brasil

Financial support: None.
Conflicts of interest: No conflicts of interest declared concerning the publication of this article.
Submitted: July 8, 2021.
Accepted: July 31, 2021.

The study was carried out at Departamento de Cirurgia de Cabeça e Pescoço, Hospital Santa Casa de Misericórdia de Belo Horizonte, Minas Gerais, Brasil.

XXVIII Brazilian Congress of Head and Neck Surgery.

How to cite: Silva GM, Ladeira FCP, Oliveira JB, Oliveira ST, Aguirre Neto JC. Laryngotracheal and pulmonary aspergillosis in an adolescent with acute lymphoblastic leukemia: case report. Arch Head Neck Surg. 2021;50:e20215025. <https://doi.org/10.4322/ahns.2021.0006>

Introduction

Acute lymphoid leukemia (ALL) is the hematologic neoplasm with a greater incidence in children, currently with high survival rates^{1,2}. Due to the immunosuppression caused by the disease and the treatment, fungal infections are common in these patients and they are related to high rates of morbidity and mortality.

Aspergillosis is a disease caused by inhalation of the *Aspergillus* fungus, usually with pulmonary involvement. Extrapulmonary manifestations of aspergillosis are less common, and laryngotracheal aspergillosis is a rare entity, with few descriptions in the literature and with high mortality. Early diagnosis and proper treatment of aspergillosis are essential to improve outcomes in these patients³.



Copyright Silva et al. This is an Open Access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Case report

This is a 15-year-old female patient diagnosed with high-risk Acute Lymphoblastic Leukemia (ALL). The patient was undergoing chemotherapy treatment, when she evolved a febrile neutropenia, and the hospitalization for infectious treatment was indicated. Due to persistent dry cough, a computed tomography of the chest was performed, which showed the presence of pulmonary nodules indicating aspergillosis.

On the seventh day of hospitalization, the patient presented dysphonia and pharyngeal globus sensation, which evolved after 3 days with stridor and dyspnea, then she was referred to the Intensive Care Unit (ICU). An endoscopic evaluation of the larynx was performed, showing evidence of a semi-obstructive glottic lesion with apparent extension to the subglottis. The patient underwent a tracheostomy identifying a lesion on the tracheal wall. In the same procedure, laryngotracheoscopy was performed with evidence of a semi-obstructive vegetating lesion extending to the subglottis and trachea. A biopsy of the lesion showed the presence of hyphae characterized as *Aspergillus*. Serum Galactomannan – *Aspergillus* wall antigen – was also collected with a positive result, endorsing the diagnosis of laryngeal, tracheal and pulmonary aspergillosis. Treatment with voriconazole antifungal was started (Figure 1).

The patient had a significant improvement in symptoms, tolerating staying in ambient air, without fever, maintaining only residual cough and ventilatory-dependent pain, being discharged to the ward after 5 days in the intensive care unit. After 3 weeks of antifungal treatment, a chest tomography showed



Figure 1. Fibronasolaryngoscopy image showing a semi-obstructive vegetating lesion extending to the subglottis and trachea.



Figure 2. Chest tomography image at diagnosis and after three weeks of treatment.

a reduction in pulmonary nodules, associated with a reduction in serum galactomannan. Control fibronasolaryngoscopy was performed, showing a complete response to laryngotracheal treatment, with the disappearance of the lesions, enabling decannulation (Figure 2).

Discussion

In this report, we presented a rare case of an immunocompromised adolescent undergoing chemotherapy for ALL and diagnosed with aspergillosis in the larynx, trachea and lung. The use of voriconazole resulted in an improvement of the symptoms and a good pulmonary and tracheal response to treatment.

The invasive fungal infections are one of the main causes of morbidity and mortality in hematologic neoplasms patients, and also immunocompromised and in chemotherapy treatment ones^{4,5}. A severe and prolonged neutropenia, as in this patient, is a known risk factor for the development of these infections, and *Aspergillus* is the second most commonly isolated fungus in these patients⁴. Due to non-specific clinical symptoms, low test sensitivity, long wait for results, difficulty in obtaining positive blood cultures and lack of appropriate conditions for procedures such as tissue culture biopsy or bronchoscopy, the early diagnosis of aspergillosis in pediatric hematologic malignancies becomes difficult³. The case of laryngeal aspergillosis is extremely rare, and the symptoms of upper airway obstruction are the most common – hoarseness, stridor and sensation of pharyngeal globus. A high index of suspicion is necessary to indicate appropriate diagnostic procedures and to begin treatment properly. Antifungal treatment using Voriconazole is recommended in the literature⁵.

This report reinforces the importance of constant monitoring of patients with Acute Lymphoid Leukemia, the value of early diagnosis and treatment as the prevalent factor for a good clinical evolution of patients with invasive fungal infections.

Ethical matters

This case report was approved by the Research Ethics Committee of the Dr. Francisco das Chagas Lima e Silva Hospital - Santa Casa de Misericórdia of Belo Horizonte - SCMBH (CEP/CONEP nº 066601/2021).

***Correspondence**

Gabriela De Martin Silva
Santa Casa de Misericórdia
Av. Francisco Sales, 1111, Santa
Efigênia
CEP 30150-221, Belo Horizonte (MG),
Brasil
Tel.: +55 (31) 3238-8100
E-mail: gabrielamartin.gdms@gmail.com

Authors information

GMS - Specializing in Head and
Neck Surgery; B.Sc. in Medicine,
Universidade de Rio Verde/Goiás.
FCPL - Otorhinolaryngologist and
Head and Neck Surgeon. JBO -
Otorhinolaryngologist and Head
and Neck Surgeon. STO - Pediatric
Oncologist. JCAN - Coordinator of the
Pediatric Oncology Service.

References

1. Fujita TC, Sousa-Pereira N, Amarante MK, Watanabe MAE. Acute lymphoid leukemia etiopathogenesis. *Mol Biol Rep.* 2021;48(1):817-22. <http://dx.doi.org/10.1007/s11033-020-06073-3>. PMID:33438082.
2. Ceconello DK, Rechenmacher C, Werlang I, Zenatti PP, Yunes JA, Alegretti AP, Lanvers-Kaminsky C, Daudt LE, Michalowski MB. Implementation of the asparaginase activity assessment technique for clinical use: experience of a Brazilian Center. *Sci Rep.* 2020;10(1):21481. <http://dx.doi.org/10.1038/s41598-020-78549-y>. PMID:33293625.
3. Avcu G, Karapinar DY, Akinci AB, Sivas ZO, Sahin A, Bal ZS, Polat SH, Metin DY, Vardar F, Aydinok Y. Utility of the serum galactomannan assay for the diagnosis of invasive aspergillosis in children with acute lymphoblastic leukemia. *Int J Infect Dis.* 2017;54:8-12. <http://dx.doi.org/10.1016/j.ijid.2016.10.027>. PMID:27815226.
4. Vlaardingerbroek H, van der Flier M, Borgstein JA, Lequin MH, van der Sluis IM. Fatal Aspergillus rhinosinusitis during induction chemotherapy in a child with acute lymphoblastic leukemia. *J Pediatr Hematol Oncol.* 2009;31(5):367-9. <http://dx.doi.org/10.1097/MPH.0b013e3181983c71>. PMID:19415022.
5. Mariette C, Tavernier E, Hocquet D, Huynh A, Isnard F, Legrand F, Lhéritier V, Raffoux E, Dombret H, Ifrah N, Cahn JY, Thiébaud A. Epidemiology of invasive fungal infections during induction therapy in adults with acute lymphoblastic leukemia: a GRAALL-2005 study. *Leuk Lymphoma.* 2017;58(3):586-93. <http://dx.doi.org/10.1080/10428194.2016.1204652>. PMID:27397551.