CASE REPORT



Devastating fungal endocarditis involving ascending aorta in a patient with a history of aortic valve replacement: a case report

Parisa Koohsari^{1†}, Akram Nakhaee^{2†}, Mehrzad Rahmanian³, Faeze Salahshour⁴, Roya Parkhideh² and Farnoosh Larti^{2*}

Abstract

Background Fungal endocarditis is a rare but serious condition associated with high mortality rates. Various predisposing factors contribute to its occurrence, such as underlying cardiac abnormalities, cardiac surgeries, prosthetic cardiac devices, and central venous catheters. Diagnosing fungal endocarditis, particularly Aspergillus, poses challenges, often complicated by negative blood cultures.

Case presentation This report details a case of extensive ascending aorta involvement in Aspergillus endocarditis (AE) in a 24-year-old man with a history of bioprosthesis aortic valve replacement (AVR). Three months post-AVR, he presented with pericardial effusion and aortic rupture, leading to a redo biological valved conduit aortic root replacement (Bentall surgery). Despite the intervention, the tubular graft exhibited extensive Aspergillus involvement, resulting in graft disruption and significant peri-aortic infection. A second redo procedure involving aortic homograft root replacement was performed. Unfortunately, the patient succumbed two days after the surgery.

Conclusion A combined approach of medical and surgical therapies is recommended to manage fungal endocarditis. Despite efforts, the mortality rate associated with Aspergillus endocarditis remains unacceptably high, with no significant difference observed between combination therapy and antifungal treatment alone. Further research is essential to explore novel therapeutic strategies and improve outcomes for patients with this challenging condition.

Keywords Aspergillus, Fungal endocarditis, Aortic pseudoaneurysm, Aortic CT angiography, Echocardiography

[†]Parisa Koohsari and Akram Nakhaee contributed equally to this work.

*Correspondence:

⁴ Department of Radiology, Advanced Diagnostic and Interventional Radiology Research Center (ADIR), Imam-Khomeini Hospital Complex, Tehran University of Medical Sciences, Tehran, Iran

Background

Fungal endocarditis (FE) is a rare condition with an overall mortality rate of over 50%. However, the incidence is expected to increase due to advances in medical and surgical therapies, like reconstructive cardiovascular surgery, implantation of intracardiac prosthetic devices, prolonged use of IV catheters, exposure to multiple broad-spectrum antibiotics, and immunosuppression. *Candida* and *Aspergillus* are two common species causing FE. It is shown that *Aspergillus* endocarditis (AE) is related to higher mortality rates than other microorganisms [1]. Herein, we report a rare and fatal AE affecting the ascending aorta in a 24-year-old man, causing multiple complications.



© The Author(s) 2024. **Open Access** This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit http://creativecommons.gr/licenses/by/4.0/. The Creative Commons Public Domain Dedication waiver (http://creativecommons.gr/licenses/by/4.0/. The Creative Commons Public Domain and redit line to the data.

Farnoosh Larti

Farnooshlarti@gmail.com

¹ School of Medicine, Tehran University of Medical Sciences, Tehran, Iran ² Cardiology Department, Imam Khomeini Hospital Complex, Tehran

Liniversity of Medical Sciences Tehran Ira

University of Medical Sciences, Tehran, Iran

³ Cardiothoracic Surgery Department, Imam Khomeini Hospital Complex, Tehran University of Medical Sciences, Tehran, Iran

Case presentation

A 24-year-old man with a history of aortic surgery was admitted to our hospital with malaise. Three months before admission to our center, he had undergone AVR in another hospital for congenital aortic stenosis. Ten days after discharge, dyspnea urged him to seek medical contact, and he was re-hospitalized in the same hospital. Upon arrival, severe hypotension, massive pericardial effusion, and cardiac tamponade were detected, and he was transferred to the operating room emergently. The diagnosis of aortic rupture due to extensive endocarditis was made in that hospital. Based on the surgical report, the surgeon observed multiple vegetation attached to the aortic valve and ascending aorta. The bioprosthesis was excised, and he underwent a redo biological valved conduit aortic root replacement (Bentall surgery), and antibiotics therapy was started. The exact antimicrobial initiation dosage and timeline were unclear as we gathered these data based on limited discharge notes. Still, the positive results of the vegetation culture changed the treatment regimen to intravenous (IV) anti-fungal. Based on the discharge sheet, he was treated with the proper intravenous (IV) anti-fungal regimen for one month. Then, he was discharged with oral antifungal treatment. Despite two months of therapy with Voriconazole, he continued to suffer extreme fatigue and was referred to our team for further assessment. Figure 1 shows the brief timeline of the disease presentation. After admission to our hospital, transthoracic echocardiography (TTE) followed by transesophageal echocardiography (TEE) was performed. Diffuse multiple masses in the entire tubular graft with significant flow restriction in ascending aorta were detected. A disrupted aortic tubular graft and a large pseudoaneurysm ($1.5 \text{ cm} \times 1.8 \text{ cm}$) in its posterior part were seen. Extending this bulky vegetation from the pseudoaneurysm neck to the peri-aortic space created an extraordinary echocardiographic view (Fig. 2, Videos 1, 2, 3 and 4). Following TEE, the patient underwent a contrast-enhanced Computed tomography (CT) scan, which demonstrated polypoid soft tissue densities in the previous surgery site of the ascending aorta in favor of infective vegetations along with contrast media out pouching adjacent to the aortic root indicating pseudoaneurysm (Fig. 3).

Upon admission, three blood culture sets were obtained from separate venipuncture sites before initiating anti-fungal therapy. The intravenous (IV) anti-fungal treatment with Voriconazole and Caspofungin) was restarted. Despite treatment, his symptoms began to worsen during the following three days after admission. He had a fever, malaise, vomiting, and right abdominal, back, and flank pain with right upper quadrant and right costovertebral angle tenderness. The results of three blood cultures were negative. An abdominal ultrasound showed a narrow free fluid rim in Morison's pouch and splenorenal recess. There was no evidence of appendicitis or hydronephrosis. Increased bladder thickness with debris was seen in favor of pyelonephritis. A spiral



Fig. 1 The timeline of the clinical presentation and surgical management



Fig. 2 Transesophageal echocardiography: diffuse multiple masses (red arrow) in the tubular aortic graft with extension to the peri-aortic space (yellow arrow), A large disruption (cyan arrow) and pseudoaneurysm in the posterior part of the tubular graft. Ao, Ascending aorta; LV, Left ventricle



Fig. 3 A, Oblique coronal reformatted; **B**, axial oblique reformatted at the aortic valve level; **C**, axial oblique reformatted; and **D**, Volume rendering reconstruction of Aortic CT angiography depicts disruption (blue arrow) of aortic root just above the valve (red arrow) along with a large pseudoaneurysm (red dots). An Apple core-like appearance (yellow four-point stars) due to vegetation is seen in the mid-ascending aorta. Most of the pseudoaneurysm is not opacified as it is full of vegetation

abdominopelvic CT scan with oral and intravenous contrast was recommended for evaluating mesenteric ischemia and septic embolism as differential diagnoses. Still, a Doppler ultrasound was performed instead due to acute kidney injury (rise in creatinine level from 1.09 to 1.7 mg/dl). In this situation, a high-risk 2nd redo homograft aortic root replacement surgery was planned. He and his family were informed about the risks and finally underwent surgery. The excised biologic valve was sent for culture, and *Aspergillus flavus* was isolated in the specimen. After surgery, he had acute renal failure and septicemia and received antibiotic and antifungal treatment. Acute tubular necrosis occurred, and he, unfortunately, expired in the intensive care unit (ICU) two days later despite hemodialysis.

Discussion and Conclusions

Fungal endocarditis (FE) is a rare disorder with increased incidence over the past decades and a more than 50% mortality rate. The most common signs of FE are fever and peripheral embolism, commonly in large brain vessels, mesenteric organs, kidneys, coronary arteries, and the limbs [2]. Several predisposing conditions are identified for FE, including underlying cardiac abnormality and surgery, prosthetic cardiac devices, and central venous catheters as the most common situations [1].

Diagnosing FE is made by following after clinical suspicion: transthoracic or transesophageal echocardiography demonstrating large, bulky vegetations, peripheral blood cultures, histological examination or culture of accessible emboli, and fungal antigen and antibody detection tests. Histologic examination of removed valve vegetations had the highest sensitivity (95%), whereas blood cultures had the least one (54%) [3]. Diagnosing mold-related endocarditis, especially *Aspergillus*, is even more challenging due to more negative blood cultures (81.2% in yeast vs. 30.8% in mold) [1].

Aortic pseudoaneurysm is a rare complication after cardiac surgeries, especially aortic procedures. It is shown that most patients have a history of aortic root replacement or repair (60%) and ascending aorta replacement (32%). Proximal and distal suture lines of implanted valves and aorta were indicated as pseudoaneurysm formation sites in half of the patients [4]. Meshaal et al., in a retrospective cohort study, showed that healthcare-associated endocarditis (HAE), prosthetic valve endocarditis (PVE), and aortic abscess/pseudoaneurysm as strong predictors of *Aspergillus* endocarditis (AE) [5].

Echocardiography is an essential tool in establishing etiological diagnosis. According to European Society of Cardiology guidelines, TTE is recommended as the first-line imaging modality of infective endocarditis (IE). TEE is recommended in patients with prosthetic valves or the presence of a negative or non-diagnostic TTE. The sensitivity of TTE for diagnosing vegetation in IE is approximately 70% in native and 50% in prosthetic valves, compared to 96% and 92% in TEE. Specificity is reported at around 90% for both modalities [6]. Large vegetations are characteristic features of fungal endocarditis, but unfortunately, the rate of false negative echocardiographic results has not been negligible [2]. The role of echocardiography in diagnosing complications after surgeries involving the ascending aorta is vital as the first imaging modality [7].

Multislice CT has a similar accuracy with TEE in detecting vegetations, abscesses, and pseudoaneurysms and is possibly superior in providing information toward the extent and consequences of any perivalvular extension, including the anatomy of pseudoaneurysms, abscesses, and fistulae [8]. Another study demonstrated the usefulness of CT in defining the size, anatomy, and calcification of the aortic valve, root, and ascending aorta in aortic IE, which can be helpful in surgical planning [9].

The aortic valve is the most commonly affected location in FE (62%), followed by the mitral valve (25%) and tricuspid valve (10%). It is shown that the mitral valve's FE is associated with the highest mortality rate either alone or with concomitant aortic valve involvement. On the other hand, tricuspid valve FE had the best survival rate (89%) [3].

Conclusions

Aspergillus is an essential species causing FE. It is shown that the mortality rate in patients with mold endocarditis is significantly higher than in yeast one (82.1% vs. 40.3%) [1].

A combination of medical and surgical therapy is still suggested as the best treatment for FE. Nevertheless, the mortality rate of *Aspergillus* endocarditis is still unacceptably high without a significant difference between combination therapy and antifungal therapy alone [3].

Abbreviations

- FE Fungal endocarditis
- HAE Healthcare-associated endocarditis
- PVE Prosthetic valve endocarditis
- AE Aspergillus Endocarditis
- TEE TTE (Trans thoracic echocardiography), Transesophageal echocardiography
- AVR Aortic valve replacement

Supplementary Information

The online version contains supplementary material available at https://doi.org/10.1186/s13019-024-02733-8.

Additional file 1: Video 1. Transthoracic long-axis view showed diffuse masses in the aortic tubular graft.

Additional file 2: Video 2. Transesophageal view showed a large pseudoaneurysm (1.5 cm × 1.8 cm) in the posterior part of the tubular graft.

Additional file 3: Video 3. Transesophageal view showed a large pseudoaneurysm (1.5 cm × 1.8 cm) in the posterior part of the tubular graft with significant flow restriction.

Additional file 4: Video 4. Transesophageal view (short axis of the aortic tubular graft) showed diffuse multiple masses in the tubular graft. Extending these bulky vegetations from the pseudoaneurysm neck to the peri-aortic space created an astonishing echocardiographic view.

Authors' contributions

FL. designed and conducted the study. FL. collaborated in the conceptualization and critically edited the first and final drafts. R.P. F.S. and M.R. were involved in data gathering and data processing. P.K. and A.N. wrote the first and final draft. All authors read and approved the final manuscript.

Funding

None.

Availability of data and materials

Data is available on reasonable request from the authors.

Declarations

Ethics approval and consent to participate

The Medical Ethics Committee of Tehran University of Medical Sciences approved the study.

Consent for publication

Informed consent was obtained from the patient and his legal executors. They were reassured that all information remains confidential.

Competing interests

The authors declare no competing interests.

Received: 12 November 2023 Accepted: 29 March 2024 Published online: 08 April 2024

References

- Pierrotti LC, Baddour LM. Fungal endocarditis, 1995–2000. Chest. 2002;122(1):302–10. Available from: https://pubmed.ncbi.nlm.nih.gov/ 12114375/. [Cited 2023 Aug 2].
- Rubinstein E, Lang R. Fungal endocarditis. Eur Heart J. 1995;16 Suppl B(SUPPL. B):84–9. Available from: https://pubmed.ncbi.nlm.nih.gov/ 7671932/. [Cited 2023 Aug 2].
- Ellis ME, Al-Abdely H, Sandridge A, Greer W, Ventura W. Fungal endocarditis: evidence in the world literature, 1965–1995. Clin Infect Dis. 2001;32(1):50–62. Available from: https://pubmed.ncbi.nlm.nih.gov/ 11118386/. [Cited 2023 Aug 2].
- Malvindi PG, Van Putte BP, Heijmen RH, Schepens MAAM, Morshuis WJ. Reoperations for aortic false aneurysms after cardiac surgery. Ann Thorac Surg. 2010;90(5):1437–43. Available from: https://pubmed.ncbi.nlm.nih. gov/20971235/. [Cited 2023 Aug 2].
- Meshaal MS, Labib D, Said K, Hosny M, Hassan M, Abd Al Aziz S, et al. Aspergillus endocarditis: Diagnostic criteria and predictors of outcome, A retrospective cohort study. PLoS One. 2018;13(8). Available from: https:// pubmed.ncbi.nlm.nih.gov/30092074/. [Cited 2023 Aug 2].
- Habib G, Lancellotti P, Antunes MJ, Bongiorni MG, Casalta JP, Del Zotti F, et al. 2015 ESC Guidelines for the management of infective endocarditis: The Task Force for the Management of Infective Endocarditis of the European Society of Cardiology (ESC). Endorsed by: European Association for Cardio-Thoracic Surgery (EACTS), the European Association of Nuclear Medicine (EANM). Eur Heart J. 2015;36(44):3075–123. Available from: https://pubmed.ncbi.nlm.nih.gov/26320109/. [Cited 2023 Aug 2].

- Nakhaee A, Parkhideh R, Rahmanian M, Pouraliakbar H, Sardari A, Larti F, et al. Re-emphasis on the role of echocardiography in the diagnosis of aortic root surgery complications. Echocardiography. 2022;39(10):1344– 7. Available from: https://pubmed.ncbi.nlm.nih.gov/36198100/. [Cited 2023 Aug 2].
- Feuchtner GM, Stolzmann P, Dichtl W, Schertler T, Bonatti J, Scheffel H, et al. Multislice computed tomography in infective endocarditis: comparison with transesophageal echocardiography and intraoperative findings. J Am Coll Cardiol. 2009;53(5):436–44. Available from: https://pubmed. ncbi.nlm.nih.gov/19179202/. [Cited 2023 Aug 2].
- Fagman E, Perrotta S, Bech-Hanssen O, Flinck A, Lamm C, Olaison L, et al. ECG-gated computed tomography: a new role for patients with suspected aortic prosthetic valve endocarditis. Eur Radiol. 2012;22(11):2407– 14. Available from: https://pubmed.ncbi.nlm.nih.gov/22622348/. [Cited 2023 Aug 2].

Publisher's Note

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.