

Nodular Pulmonary Amyloidosis Mimicking Breast Carcinoma Metastasis

Meme Karsinom Metastazını Taklit Eden Nodüler Pulmoner Amiloidoz

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Abstract

Amyloidosis is a disease that is characterized by an extracellular accumulation of fibril proteins called amyloid in tissues, and organ dysfunction. There are various types of amyloidosis, with nodular pulmonary amyloidosis usually considered a subtype of AL amyloidosis. Surgical excision is usually curative and the prognosis is excellent. This case is presented to emphasize the rare occurrence of pulmonary amyloidosis and the need to keep malignancies in mind in differential diagnosis.

Key words: Amyloidosis, breast cancer, metastasis.

Özet

Amiloidoz, dokularda amiloid adı verilen fibril yapısındaki proteinlerin ekstrasellüler birikimi ve organlarda işlev bozukluğu ile seyreden hastalıktır. Çeşitli tipleri olup, nodüler pulmoner amiloidoz, genellikle AL amiloidozun bir subtipi olarak görülür. Cerrahi eksizyon genellikle küratiftir ve prognoz mükemmeldir. Bu olgu, pulmoner amiloidozun nadiren görülmesi, ayırıcı tanıda malignitelerin akılda tutulması gerekliliğini vurgulamak amacıyla sunulmuştur.

Anahtar Sözcükler: Amiloidoz, meme karsinomu, metastaz.

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Amyloidosis is characterized by an accumulation of Congo-Red positive amyloid fibrin deposits in the extracellular matrix of organs (1). The most common forms are systemic AL amyloidosis, systemic AA amyloidosis, systemic wild-type ATTR amyloidosis, systemic hereditary ATTR amyloidosis and localized AL amyloidosis (2). Nodular pulmonary amyloidosis is a usual subtype of localized AL amyloidosis (3) that may be associated with underlying inflammatory or malignant conditions. Patients are generally asymptomatic (4,5). Lesions are usually solitary, and may be mistaken for pulmonary malignancies. Biopsy material is stained with Congo-Red to give a green color under a polarized microscope, which is diagnostic (6). Excision of the nodules is usually curative and the prognosis is excellent (7). The case we present here is intended to emphasize that pulmonary amyloidosis is rare and may be mistaken for malignancies.

CASE

A 76-year-old female patient who underwent a right mastectomy 10 years previously for breast apocrine carcinoma followed by chemotherapy and radiotherapy, was diagnosed with pulmonary nodules in radiological follow-up and referred to us. No pathology was detected aside from the old operation scar on the right breast upon physical examination. Increased opacity was noted in the left paracardiac area on chest radiography. A thorax computed tomography (CT) showed a 13x15 mm solid nodule in the left upper lobe (Figure 1 and 2), a 15x20 mm nodule in the lingual (Figure 3 and 4), and an 6x8 mm nodule in the left lower lobe (Picture 5). A positron emission tomography/computed tomography (PET-CT) revealed a 1.5x1 cm nodule (SUVmax: 1.6) in the left upper lobe and a nodule of 2.2x1.4 cm (SUVmax: 1.4) in the lingula (Figures 6, 7 and 8). Since the patient could not be diagnosed through bronchoscopy and transthoracic, a fine needle aspiration biopsy (TTFNAB), operation was selected for diagnosis and treatment. In an exploration with a left thoracotomy, four nodules in the left upper lobe and one nodule in the lower lobe were palpated. A wedge resection was performed on the three peripheral nodules in the left upper lobe and in the lower lobe, and the frozen section was examined. The frozen result report read "The nodules may be breast carcinoma metastasis, but a definitive diagnosis should be confirmed with further investigations." A left upper lobectomy was performed, since the other central lesion in the upper lobe could not be completely removed due to its closeness to the superior pulmonary vein. The final pathology was reported as

nodular pulmonary amyloidosis (Figures 9, 10 and 11). The patient was discharged on the 7th postoperative day and is now in the 4th postoperative month.

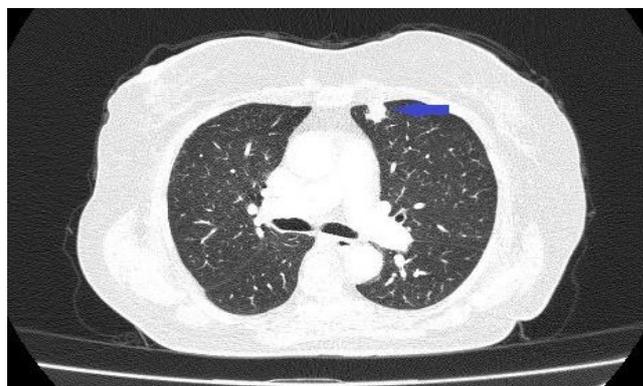


Figure 1: Thorax computed tomography showing a 13x15 mm solid nodule in left lung upper lobe

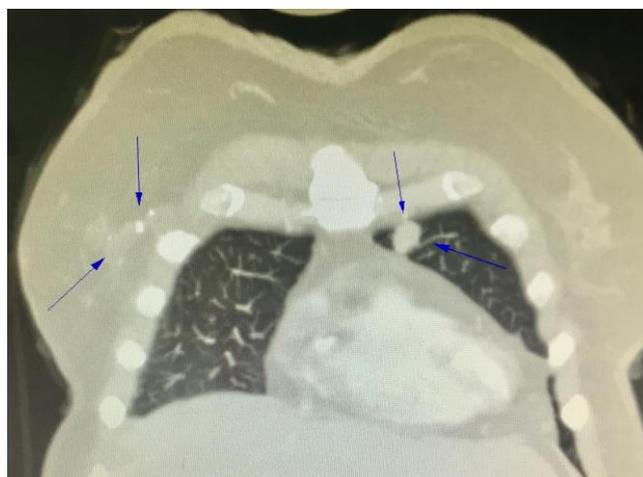


Figure 2: Thorax computed tomography showing a 13x15 mm solid nodule in left lung upper lobe (Coronal section)

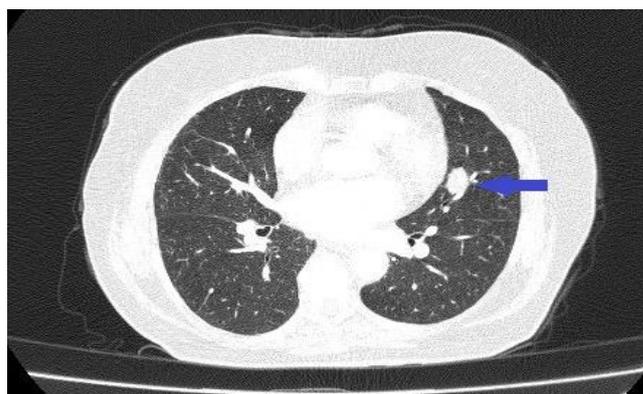


Figure 3: Thorax computed tomography showing a 15x20 mm nodule in left lung lingual lobe



Figure 4: Thorax computed tomography showing a 15x20 mm nodule in left lung lingual lobe (sagittal section)

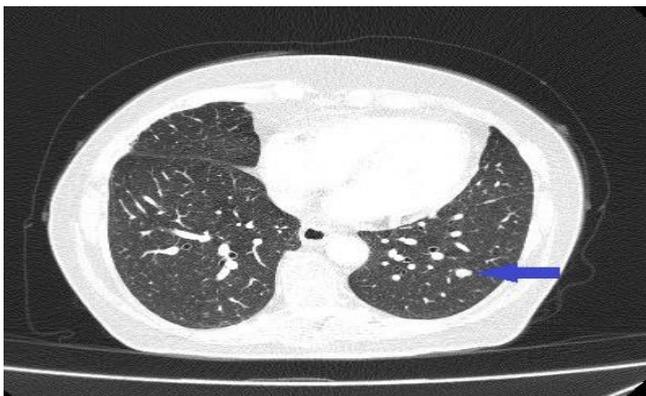


Figure 5: Thorax computed tomography showing a 6x8 mm nodule in left lung lower lobe

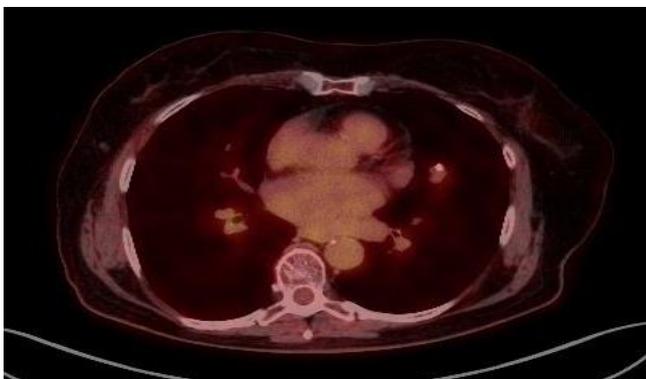


Figure 6: PET /CT showing a 1.5x1 cm nodule (SUVmax: 1.6) in left lung upper lobe

DISCUSSION

Nodular pulmonary amyloidosis, known also as nodular parenchymal amyloidosis or nodular amyloidoma, is a disease that is characterized by the presence of one or more tumor-like amyloid deposits in the lung (1). It was first described by Virchow in 1857 (7). It is rarely seen among the amyloid diseases (3,6), and is characterized primarily by a congophilic light chain amyloid (AL) deposition in the extracellular matrix, and may be solitary or, as with our patient, multiple (1,8). It is usually diagnosed in the sixth decade and is more common in men (1,9). Patients are generally asymptomatic, as in our patient (4,5), although it may cause cough, hemoptysis, dyspnea, pleural effusion and pulmonary arterial hypertension, depending on the location (10). It has no specific radiological findings and mimics malignancies radiologically. The present case emphasizes that pulmonary amyloidosis is rare and may be mistaken for malignancies. Our patient was operated on due to the suspicion of breast carcinoma metastasis.

18-Fluoro-deoxyglucose (18F-FDG) PET-CT has emerged as a tool for the diagnosis of pulmonary nodules to reduce invasive diagnostic examination. However, 18F-FDG shows a small amount of uptake in malignancies with low metabolic activity, such as bronchoalveolar cancer, carcinoid tumor and mucinous adenocarcinoma. As such, despite low involvement on PET-CT (SUVmax: 1,6), malignancies can be considered in a pre-diagnosis. Furthermore, it has high metabolic rates alongside such non-malignant conditions as tuberculosis, sarcoidosis and rheumatoid nodules. In our case, and as in other rarely reported cases, pulmonary nodular amyloidosis with low or moderate 18F-FDG involvement can be seen (11). As such, the results of an 18F-FDG PET-CT should be interpreted with caution in the differentiation of pulmonary amyloidosis from other malignant or benign lesions. A definitive diagnosis of localized pulmonary amyloidosis requires histological confirmation (12). In cases in which patient cannot be diagnosed via a CT-guided fine needle aspiration biopsy, an invasive surgical resection may be necessary. The patient in the present study could not be diagnosed via TFNAB, and so exploratory thoracotomy was decided upon. Furthermore, in the frozen examination, no differential diagnosis from malignancies could be made.



Figure 7: PET /CT showing a calcific nodul in left lung upper lobe



Figure 8: PET /CT showing a 2.2x1.4 cm nodule (SUVmax: 1.4) in left lung lingula lobe

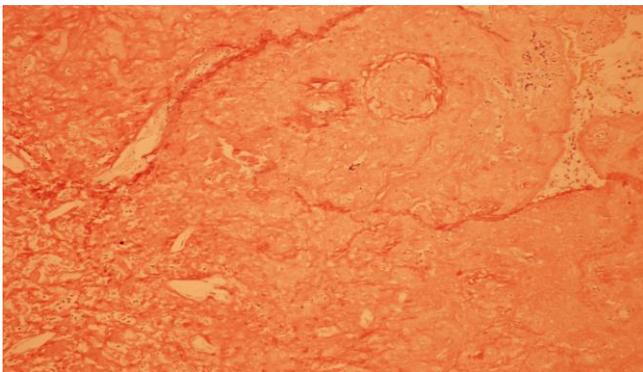


Figure 9: Congo red X 200

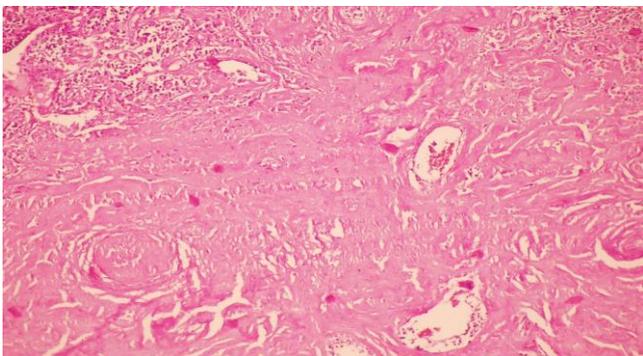


Figure 10: Acellular hyalinized material (H&E X200)

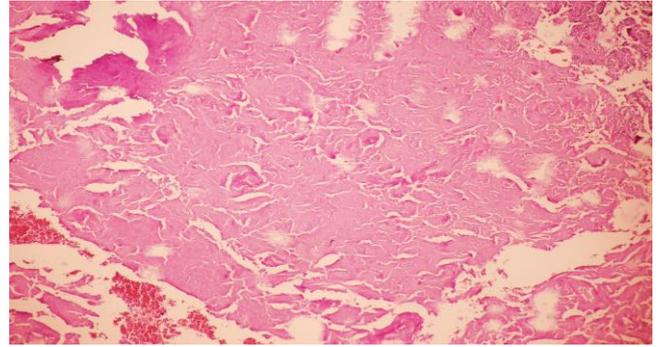


Figure 11: Acellular eosinophilic material (H&E X200)

Pulmonary amyloidosis may occur as a component of localized or systemic amyloidosis (12,13). Localized AL amyloidosis has a better prognosis than systemic amyloidosis. The 10-year survival rate after surgery is reported to be 97%. A surgical resection can be performed safely, and the prognosis is excellent (13). Our patient continues to be followed-up without problem in the postoperative 4th month.

In conclusion, local nodular pulmonary amyloidosis is a rare and unusual tumor of the lung, and surgical treatment is curative. Nodules may be solitary or multiple. It should be kept in mind that amyloidosis may mimic both benign and malignant pathologies, and should be considered in a differential diagnosis.

CONFLICTS OF INTEREST

None declared.

AUTHOR CONTRIBUTIONS

Concept - E.Ç.Ç., S.Y., S.G., A.U., Z.A.; Planning and Design - E.Ç.Ç., S.Y., S.G., A.U., Z.A.; Supervision - E.Ç.Ç., S.Y., S.G., A.U., Z.A.; Funding - E.Ç.Ç., Z.A.; Materials - S.Y., E.Ç.Ç., Z.A.; Data Collection and/or Processing - E.Ç.Ç., Z.A.; Analysis and/or Interpretation - S.Y., S.G., E.Ç.Ç.; Literature Review - E.Ç.Ç.; Writing - E.Ç.Ç.; Critical Review - S.Y., A.U., S.G.

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REFERENCES

1. Khor A, Colby TV. Amyloidosis of the Lung. *Arch Pathol Lab Med* 2017; 141:247-54. [\[CrossRef\]](#)
2. Sipe JD, Benson MD, Buxbaum JN, Ikeda S, Merlini G, Saraiva MJ, et al. Nomenclature 2014: amyloidfibril proteins and clinical classification of the amyloidosis. *Amyloid* 2014; 21:221-4. [\[CrossRef\]](#)
3. Kaplan B, Martin BM, Boykov O, Gal R, Pras M, Shechtman I, et al. Co-deposition of amyloidogenic immunoglobulin light and heavy chains in localized pulmonary amyloidosis. *Virchows Arch* 2005; 447:756-61. [\[CrossRef\]](#)
4. Beer TW, Edwards CW. Pulmonary nodules due to reactive systemic amyloidosis (AA) in Crohn's disease. *Thorax* 1993; 48:1287-8. [\[CrossRef\]](#)
5. Roden AC, Aubry MC, Zhang K, Brady JO, Levin D, Dogan A, et al. Nodular senile pulmonary amyloidosis: a unique case confirmed by immunohistochemistry, mass spectrometry, and genetic study. *Human Pathology* 2010; 41:1040-5. [\[CrossRef\]](#)
6. Howie AJ, Brewer DB. Optical properties of amyloid stained by Congo red: history and mechanisms. *Micron* 2009; 40:285-301. [\[CrossRef\]](#)
7. Utz JP, Swensen SJ, Gertz MA. Pulmonary amyloidosis The Mayo Clinic experience from 1980 to 1993. *Ann Intern Med* 1996; 124:4, 407-13. [\[CrossRef\]](#)
8. Milani P, Basset M, Russo F, Foli A, Palladini G, Merlini G. The lung in amyloidosis. *Eur Respir Rev* 2017; 26(145): pii: 170046. [\[CrossRef\]](#)
9. Yang MC, Blutreich A, Das K. Nodular pulmonary amyloidosis with an unusual protein composition diagnosed by fine-needle aspiration biopsy: a case report. *Diagn Cytopathol* 2009; 37:286-9. [\[CrossRef\]](#)
10. Scala R, Maccari U, Madioni C, Venezia D, La Magra LC. Amyloidosis involving the respiratory system: 5-year's experience of a multi-disciplinary group's activity. *Ann Thorac Med* 2015, 10:212-6. [\[CrossRef\]](#)
11. Standaert C, Herpels V, Seynaeve P. A solitary pulmonary nodule: pulmonary amyloidosis. *J Belg Soc Radiol* 2018; 102:20. [\[CrossRef\]](#)
12. Chen KT. Amyloidosis presenting in the respiratory tract. *Pathol Annu* 1989; 24:253-273.
13. Baumgart JV, Stuhlmann-Laeisz C, Hegenbart U, Nattenmüller J, Schönland S, Krüger S, et al. Local vs. systemic pulmonary amyloidosis-impact on diagnostics and clinical management. *Virchows Arch* 2018; 473:627-37. [\[CrossRef\]](#)