

A case of myocardial metastasis from lung adenocarcinoma presenting as cerebral infarction

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ABSTRACT

Introduction: Myocardial metastasis from lung cancer is rarely found in antemortem settings. Herein we present a case of myocardial metastasis from lung adenocarcinoma presenting as cerebral infarction. **Case Report:** A 78-year-old Japanese female (non-smoker) who had been diagnosed with poorly differentiated lung adenocarcinoma suddenly complained of weakness in the left hand on a regular follow-up visit. Brain non-contrast-enhanced computed tomography (CT) scan showed no abnormality. Electrocardiography (ECG) showed abnormal Q waves, ST-segment elevation, and inverted T waves in leads II, III, and aVF that had not been demonstrated previously. The results of brain CT scan and symptoms were strongly suggestive of cerebral infarction. Brain magnetic resonance imaging (MRI) with diffusion-weighted imaging performed 5 days after admission showed a small, acute infarct on a knob on the precentral

gyrus. Transthoracic echocardiography performed to detect cardiac sources of embolism revealed an intracardiac mass. Cardiac 18F-fluoro-deoxy-glucose positron emission tomography (18F-FDG PET)/CT scan demonstrated increased uptake in the mass. After the patient died, autopsy revealed many metastatic lesions of various sizes accompanied by thrombi on the surface of the ventricular myocardium. **Conclusion:** Cardiac 18F-FDG PET/CT was useful for diagnosing myocardial metastasis in our patient. The possibility of cardiac metastases should be considered in patients with lung cancer in the event of sudden cerebral infarction.

Keywords: Myocardial metastasis, Lung cancer, Cerebral infarction, Cardiac PET/CT

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INTRODUCTION

Myocardial metastasis from lung cancer is often clinically invisible, and is thus very difficult to diagnose antemortem. Positron emission tomography (PET)/computed tomography (CT) scan offer additional metabolic information resulting in increased sensitivity for the detection of malignant tumor compared to purely morphological modalities [1]. We present a case

of myocardial metastasis from lung adenocarcinoma presenting as cerebral infarction.

CASE REPORT

Our patient was a 78-year-old Japanese female (non-smoker) who had been diagnosed with poorly differentiated lung adenocarcinoma staged as cT2aN2Mo, stage IIIA, in October 2008. She had been treated with chemoradiotherapy comprising cisplatin and docetaxel, followed by gefitinib. In February 2009, gefitinib treatment was stopped after the appearance of ground glass opacities in the lung. In June 2009, stereotactic radiotherapy was performed for local recurrence. Follow-up screening for metastasis with CT of the whole body, magnetic resonance imaging (MRI) of the brain, and bone scintigraphy in September 2009 showed negative results.

The patient suddenly complained of weakness in the left hand during a regular follow-up visit in October 2009. Neurological examination showed a positive Barré sign in the left upper extremity. No abnormalities were apparent on physical examination other than coarse crackles in the right lower lung field. Brain non-contrast-enhanced CT scan showed no abnormalities.

Blood testing showed normocytic anemia with a hemoglobin level of 9.8 g/dL. Serum creatinine level was 1.07 mg/dL and C-reactive protein concentration was 0.20 mg/L. Congestive heart failure was suspected based on the increased cardiac brain natriuretic peptide (BNP) level of 257.0 pg/mL. Levels of fibrin degradation products (FDP) and D-dimer were 43.6 µg/mL and 13.7 µg/mL, respectively. All other results of biochemical testing were normal.

Electrocardiography showed abnormal Q wave, ST-segment elevation, inverted T waves in leads II, III, and aVF that were not demonstrated in September 2009, but the patient reported no cardiac symptoms (Figure 1A).

Results of brain CT and patient symptoms were strongly suggestive of cerebral infarction. Treatment was started with edaravon and argatroban and neurological symptoms gradually improved. Brain MRI with diffusion-weighted imaging performed five days after admission showed a small, acute infarct on a knob on the precentral gyrus (Figure 1B–C). Magnetic resonance angiography of the circle of Willis yielded normal results.

Transthoracic echocardiography performed to detect cardiac sources of embolism revealed an intracardiac mass arising from the ventricular septum close to the apex (Figure 2A). Cardiac CT demonstrated a low-density mass located in the wall of the left ventricle (Figure 2B). Cardiac 18F-fluoro-deoxy-glucose (FDG) PET/CT demonstrated increased uptake in the mass in the inferior wall of the left ventricle close to the apex (Figure 2C–E). These findings were strongly suggestive of myocardial metastasis from lung adenocarcinoma.

On the other hand, brain MRI with gadolinium-enhanced T1-weighted imaging performed 32 days after

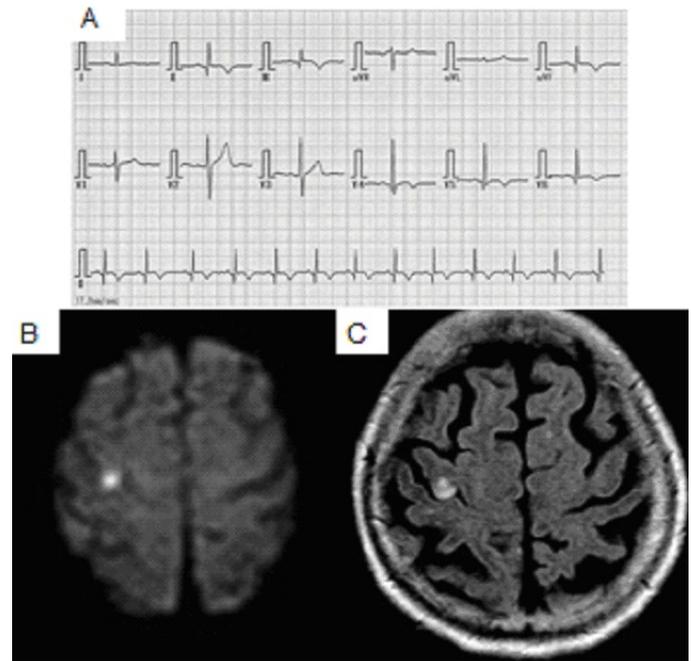


Figure 1: Electrocardiography showing abnormal Q waves, ST-segment elevation, and inverted T waves in leads II, III, and aVF (A). Axial diffusion-weighted imaging (DWI) (B) and axial T2-weighted magnetic resonance imaging (C) obtained 5 days after admission, showing hyperintensity on a knob on the precentral gyrus.

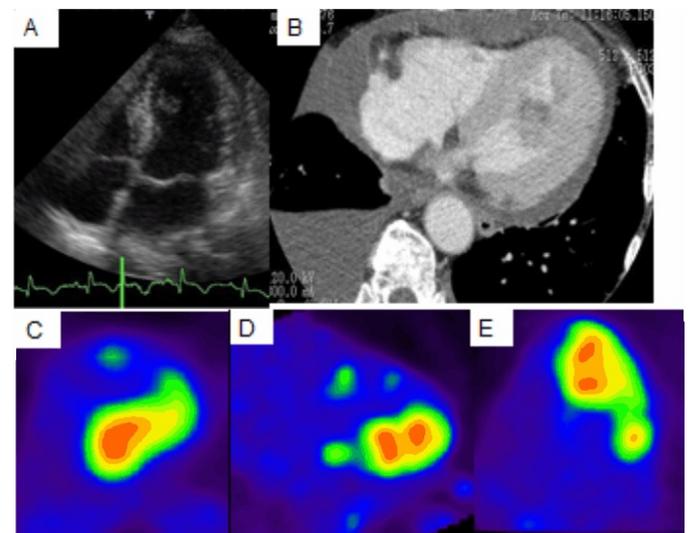


Figure 2: Transthoracic echocardiography obtained eight days after clinical ictus. (A) The apical four-chamber view on B-mode shows an intracardiac mass close to the apex, (B) Contrast-enhanced thoracic CT obtained 13 days after admission demonstrates a low-density mass located in the left ventricle, Cardiac 18F-FDG PET/CT in a fasting state in corresponding short-axis, (C) Vertical long-axis, (D) Horizontal long-axis, and (E) Slices showing abnormal increased uptake in the inferior wall close to the apex. Maximum standardized uptake value (SUVmax) was 10.14 g/mL. Imaging results are consistent with myocardial metastasis

admission showed enhancement at the same site demonstrated previously on MRI.

The patient was treated with erlotinib for recurrent lung adenocarcinoma, but died soon afterwards. Autopsy was performed, revealing many metastatic lesions of various sizes accompanied by thrombi on the surface of the ventricular myocardium (Figure 3).

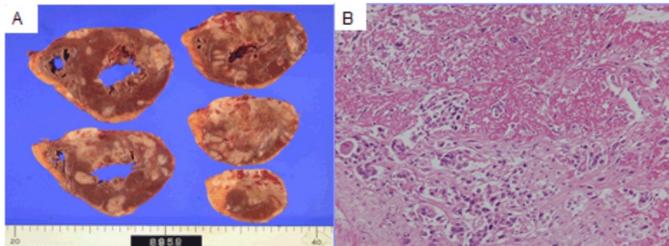


Figure 3: (A) Gross and (B) microscopic findings of cardiac metastases at autopsy. (A) Multiple metastases of various sizes in the myocardium were demonstrated in the resected heart. (B) Adenocarcinoma cells were demonstrated not only in myocardium but also in thrombus on the myocardial surface. (HE, x100 magnification).

DISCUSSION

Metastasis to the pericardium or heart is reportedly present in 30-45% of autopsies for lung cancer patients [2, 3]. Pericardium, including epicardium, is the most common location of cardiac involvement by secondary tumors [4], with myocardial metastasis reportedly comprising only about 10% of metastases to the heart [3]. Myocardial metastasis from malignant tumor is often clinically invisible, and can be very difficult to diagnose antemortem. Indications of metastases to the heart are often described as a rapid increase in heart size due to pericardial effusion, new signs of heart failure or valve disease, conduction defects, and atrial or ventricular heart rhythm disturbances [5]. In our patient, abnormalities were seen on ECG but the patient reported no cardiac symptoms.

Regarding the mechanisms of cardiac metastasis, it is reported that metastatic cells can reach the heart via the lymphatic or hematogenous route, or by direct or transvenous extension. In our patient, tumor cells seem to have attained to the heart mainly via hematogenous route considering the fact that hematogenous spread preferentially gives rise to myocardial metastasis [5].

Cerebral infarction has also been reported to result from obvious thromboembolism, probably from the surface of a metastatic tumor [6]. In this patient, cerebral infarction was the first presentation of cardiac metastasis from lung adenocarcinoma. Such embolic phenomena may be due to either tumor emboli or bland emboli that have dislodged from the tumor thrombus on the surface of the myocardial metastasis. The possibility of cardiac metastasis should be kept in mind for patients with lung cancer who suddenly present with cerebral infarction.

To explain the mechanisms of unexpected cerebral infarction in a patient with malignant tumor, the possible presence of Trousseau's syndrome should be considered. Trousseau's syndrome represents unexplained thrombotic events that precede the diagnosis of an occult visceral malignancy or appear concomitantly with the tumor. This syndrome is considered to be mediated by multiple mechanisms such as tissue factors, tumor-associated cysteine proteinase, tumor hypoxia, carcinoma mucins, or an overlap of these as a cause of prothrombotic condition [7]. Besides the presence of myocardial metastasis, the influence of Trousseau's syndrome might be one cause of cerebral infarction.

For the diagnosis of metastasis to the heart, echocardiography, enhanced CT, enhanced MRI, and the combination of these modalities may prove helpful [5]. The advantage of cardiac PET/CT scan for detecting myocardial metastasis has recently been reported [8]. Compared to purely morphological modalities, PET/CT scan offers additional metabolic information resulting in increased sensitivity for the detection of regional tumor growth and/or distant metastases, facilitating staging and follow-up of patients with malignancies [1]. In our patient, distinguishing metastatic cardiac tumor and thrombus was difficult using only echocardiography and contrast-enhanced CT scan. On the other hand, cardiac PET/CT scan demonstrated focally abnormal increases in the uptake of FDG in myocardium and correctly suggested metastatic tumor. Cardiac PET/CT scan is usually used for the detection of coronary artery disease (CAD), and is rapidly gaining popularity as a powerful and efficient alternative to conventional single-photon emission CT to evaluate regional myocardial perfusion and metabolism in patients with CAD [9]. With respect to cardiac tumors, cardiac PET/CT scan is useful not only for the detection of myocardial metastasis but also for specifying the exact location of the tumor compared with conventional PET/CT.

In contrast, a case report has described a patient with underlying malignancy who was misdiagnosed with cardiac metastasis due to an imaging pitfall in PET/CT scan. Efforts should therefore always be made to rule out false-positive interpretations [10].

CONCLUSION

We report here on a case of myocardial metastasis from lung adenocarcinoma that presented as cerebral infarction. Cardiac PET/CT was a useful modality for diagnosing myocardial metastasis in our patient. The possibility of cardiac metastasis should be kept in mind for patients with lung cancer who develop sudden cerebral infarction.

Author Contributions

Kentaro Suina – Substantial contributions to conception and design, Acquisition of data, Drafting the article,

revising it critically for important intellectual content, Final approval of the version to be published
Satoshi Hirano – Substantial contributions to conception and design, Acquisition of data, Drafting the article, Revising it critically for important intellectual content, Final approval of the version to be published
Junko Hirashima – Substantial contributions to conception and design, Acquisition of data, Drafting the article, Revising it critically for important intellectual content, Final approval of the version to be published
Kazutaka Shimizu – Substantial contributions to conception and design, Acquisition of data, Drafting the article, Revising it critically for important intellectual content, Final approval of the version to be published
Sosuke Takeuchi – Substantial contributions to conception and design, Acquisition of data, Drafting the article, Revising it critically for important intellectual content, Final approval of the version to be published
Miyako Morooka – Substantial contributions to conception and design, Acquisition of data, Drafting the article, Revising it critically for important intellectual content, Final approval of the version to be published
Kazuo Kubota – Substantial contributions to conception and design, Acquisition of data, Drafting the article, Revising it critically for important intellectual content, Final approval of the version to be published
Tateki Ito – Substantial contributions to conception and design, acquisition of data, Drafting the article, Revising it critically for important intellectual content, Final approval of the version to be published
Yuichiro Takeda – Substantial contributions to conception and design, Acquisition of data, Drafting the article, revising it critically for important intellectual content, Final approval of the version to be published
Haruhito Sugiyama – Substantial contributions to conception and design, Acquisition of data, Drafting the article, Revising it critically for important intellectual content, Final approval of the version to be published
Nobuyuki Kobayashi – Substantial contributions to conception and design, Acquisition of data, Drafting the article, Revising it critically for important intellectual content, Final approval of the version to be published
Koichiro Kudo – Substantial contributions to conception and design, Acquisition of data, Drafting the article, Revising it critically for important intellectual content, Final approval of the version to be published

Guarantor

The corresponding author is the guarantor of submission.

Conflict of Interest

Authors declare no conflict of interest.

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