Myocardial Metastasis of Tongue Cancer: A Rare Localization

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Patient: Male, 72-year-old
Final Diagnosis: Metastatic squamous cell carcinoma
Symptoms: None
Medication: —
Clinical Procedure: —
Specialty: Cardiology

Objective: Rare disease

Background: Cardiac metastases of head and neck tumors are extremely rare, and antemortem diagnosis is even rarer. In most cases, patients show symptoms or electrocardiographic abnormalities and expected survival is considerably low.

Case Report: A 72-year-old man was admitted to our cardiology ward with suspected endocarditis 2 months after a right hemiglossectomy for a squamous cell carcinoma. He was asymptomatic and showed no electrocardiogram abnormalities. Echocardiography showed an iso-echogenic round-shaped formation at the right ventricle apex that was suspected to be a metastasis owing to 2 cardiac magnetic resonance imaging examinations showing a significant increase in its volume over 14 days. The patient was evaluated by a multidisciplinary team and referred for first-line chemotherapy with carboplatin, 5-fluorouracil, and cetuximab. He died after almost 4 months of follow-up.

Conclusions: This case presents a very rare diagnosis as well as some unique features, including the intramyocardial localization, the absence of symptoms, and electrocardiogram abnormalities. Our report shows the relevance of imaging techniques in defining even the most unusual clinical findings, and it emphasizes the significance of early recognition of cardiac masses in order to identify the appropriate medical or surgical therapy for patients.

Keywords: Cardiology • Echocardiography • Head and Neck Neoplasms • Magnetic Resonance Imaging • Neoplasm Metastasis

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Background

Cardiac tumors represent a rare, yet challenging diagnosis and can be differentiated into primary tumors (rates between 0.001% and 0.3%) and secondary tumors (ranging from 2.3% and 18.3%) [1]. In theory, the heart can be metastasized by any malignant neoplasm, but antemortem diagnosis is very rare because clinical manifestations are subtle or even absent, especially at an early stage [1,2]. At more advanced stages, however, metastatic tumors gradually lead to heart failure, conduction disturbances, or valvular disease resembling mitral stenosis, anginal pain, or Stokes-Adams syndrome [3]. Cardiac metastasis is more frequently detected accidentally, usually at surgery or autopsy (in 2.3-7.1% of cases) [4].

In order of frequency, tumors metastasize to the pericardium, endocardium and valves, and the myocardium. The mechanisms of metastasis to the heart remain uncertain, but explanatory hypotheses include hematogenous spread via the coronary arteries, retrograde lymphatic extension, and intracavitary diffusion either by the inferior vena cava or by pulmonary veins [2,3]. Lymphatic spread tends to give rise to pericardial metastases, as seen in lung and breast cancer, owing to their topography; whereas, hematogenous spread leads to myocardial metastases [5]. The most common primary tumors that produce cardiac metastasis are melanoma, carcinoma of the bronchus, breast carcinoma, and lymphoma. Due to their propensity for generalized hematogenous spread, malignant melanomas frequently metastasize to the heart in this manner; whereas, esophageal and breast cancers metastasize to the heart predominantly by lymphatics [5].

Case Report

A 72-year-old man received a diagnosis of tongue carcinoma in December 2018. He subsequently underwent a right sub-total hemiglossectomy with right lateral neck dissection and a

Figure 1. Computed tomography-positron emission tomography scan. A hypermetabolic signal is visible near the apex of the right ventricle (arrow), showing a high standardized uptake value (maximum 15.1).

Video 1. First transthoracic echocardiography: subcostal view showing the intramyocardial mass near the right ventricle apex.

Figure 2. First transthoracic echocardiography. Two different (a and b) zoomed subcostal views of an intramyocardial mass (arrow) with a maximum diameter of approximately 1.6 cm.
reconstruction with a fasciocutaneous flap from the left thigh. Histological examination revealed poorly differentiated (G3) ulcerated squamocellular carcinoma infiltrating the muscular tissue and various metastatic lymph nodes (pT3, N2b, M0). About a month after surgery the patient was admitted to the Emergency Department for a surgical wound infection and dismissed with an oral antimicrobial therapy.

In January 2019, computed tomography-positron emission tomography (CT-PET) was performed and showed hypermetabolic activity in the patient’s tonsils and laterocervical lymph nodes. A hypermetabolic signal (standardized uptake value, maximum 15.1) was also shown in the heart at the ostium of the mitral valve and near the apex of the right ventricle (Figure 1). These images were thought to be suggestive of a septic thrombus. The patient was admitted to our cardiology ward with suspected endocarditis.

The patient presented no symptoms and, in particular, he did not report chest discomfort, palpitations, dyspnea, or syncope. Physical evaluation was normal, and an electrocardiogram (ECG) showed no significant abnormalities. The patient’s C-reactive protein and erythrocyte sedimentation rate were slightly increased. With few clinical data to confirm or exclude the suspected diagnosis of endocarditis, a better characterization of the hypermetabolic signal areas shown by CT-PET was essential.

A transthoracic echocardiography (TTE) (Figure 2A, 2B, Video 1) was done, and it showed an iso-ipoechogenic round-shaped formation at the right ventricle apex, with no signs of vascularization. The patient next underwent a cardiac magnetic resonance imaging (MRI) scan (Figure 3A, 3B, Video 2) for better tissue characterization. A round-shaped mass with a diameter of 17 mm, hardly distinguishable from the right ventricular wall, was found. The mass showed a hyperintense signal in T2-weighted images and in T1-weighted scans with late gadolinium enhancement and a hypointense signal in T1 scans with

**Figure 3.** First magnetic resonance imaging scan. (A) Four-chamber steady-state free precession and (B) short-axis T2 short tau inversion recovery, showing a 17-mm round-shaped mass (arrow) that is hardly distinguishable from the right ventricular wall.

**Video 2.** First magnetic resonance imaging scan: cine steady-state free precession sequences showing the intramyocardial mass and its movement, synchronous with the right ventricular wall.
These findings were suggestive of cardiac metastasis. The patient was re-evaluated by a multidisciplinary team of oncologists, radiologists, and maxillofacial surgeons. A second TTE (Figure 4, Video 3) and a follow-up MRI scan (Figure 5A, 5B, Video 4) were done 14 days later and showed significant augmentation in the sizes of the right ventricular lesion (34×25×34 mm) and infiltration of the right ventricular free wall and pericardial adipose tissue.

Video 3. Control transthoracic echocardiography: subcostal view showing a significative dimensional increase of right ventricular mass.

Figure 4. Control transthoracic echocardiography. After 14 days, the mass (arrow) showed a significant increase in linear dimensions with a maximum diameter of approximately 3 cm.

Figure 5. Control magnetic resonance imaging. Four-chamber (A) and short-axis (B) views confirming the lesion's (arrow) volumetric increase and showing infiltration of right ventricular free wall and pericardial adipose tissue.

Video 4. Control magnetic resonance imaging scan: cine steady-state free precession sequences, 4 chamber views confirming a significant augmentation in the sizes of the right ventricular lesion.

early gadolinium enhancement. These findings were suggestive of cardiac metastasis.

The patient was re-evaluated by a multidisciplinary team of oncologists, radiologists, and maxillofacial surgeons. A second TTE (Figure 4, Video 3) and a follow-up MRI scan (Figure 5A, 5B, Video 4) were done 14 days later and showed significant augmentation in the sizes of the right ventricular lesion (34×25×34 mm) and infiltration of the right ventricular
free wall and the pericardial adipose tissue. This formation also showed, in the patient's intracavitary portion, a nodular extension wedged between the right ventricular trabeculae with synchronous movement in cine sequences. Because of the lesion localization, it was impossible to obtain a sample for histological examination. During the first weeks of his hospital stay, the patient had also developed a subcutaneous lesion that appeared as a protuberance in the right frontal area that rapidly increased in volume. This lesion was not evident on admission, nor was it reported on the initial CT-PET. It was also investigated with an MRI scan of the head and neck district, and findings suggested that it was a metastasis. An incisional biopsy of the lesion was then performed, and the results were compatible with the localization of a squamous cell carcinoma.

From February to April 2019, the patient received 2 cycles of first-line chemotherapy according to the scheme carboplatin area under the curve (AUC) of 5 on day 1, 5-fluorouracil 800 mg/m²/d from day 1 to day 5 every 21 days, along with the anti-epidermal growth factor receptor drug cetuximab, at a loading dose of 400 mg/m² for the first cycle and then at the standard dose of 250 mg/m² weekly [6]. Treatment toxicity

### Table 1. Summary of published cases.

| Study                      | Year of publication | Primary site               | Location of metastases                                      | Symptoms                                      | ECG anomalies                                      | Treatment             | Follow-up period |
|----------------------------|---------------------|----------------------------|------------------------------------------------------------|------------------------------------------------|--------------------------------------------------|-----------------------|------------------|
| Werbel et al [9]           | 1985                | Tongue                     | Pericardium, right atrium                                  | Cardiac murmur, chest pain                    | ST depression, T waves inversion                  | Radiotherapy          | 7 wk             |
| Rivkin et al [10]          | 1999                | Tongue                     | Right ventricle                                            | Chest pain                                    | ST elevation, Q waves, AF                        | Chemotherapy          | “Several months” |
| Schwender et al [11]       | 2002                | Buccal mucosa              | Pericardium                                                | Weakness, respiratory distress                | AF                                               | None                  | N/A              |
| Hans et al [12]            | 2009                | Tongue                     | Right ventricle                                            | Dyspnea, lower extremity edema, hemoptyis     | Right bundle branch block                       | None                  | N/A              |
| Onwuchekwa et al [13]      | 2012                | Tongue                     | Right ventricle, interventricular septum                  | Syncope, dyspnea                              | None                                             | None                  | N/A              |
| Onwuchekwa et al [13]      | 2012                | Tongue                     | Left ventricle                                             | Palpitations, dyspnea                         | ST elevation, AF                                | Radiotherapy          | 2 mo             |
| Nagata et al [8]           | 2012                | Tongue                     | Pericardium, left lower pulmonary vein                    | Fever                                          | None                                             | Surgery               | 3 wk             |
| Nagata et al [8]           | 2012                | Soft palatal mucosa        | Right atrium, right ventricle, pericardium                 | None                                           | Q waves                                         | None                  | 3 wk             |
| Browning et al [14]        | 2015                | Tongue                     | Right ventricle                                            | N/A                                            | N/A                                             | None                  | N/A              |
| Malekzadeh et al [15]      | 2017                | Tongue                     | Right ventricle                                            | Acute chest pain                              | ST elevation                                     | Chemotherapy          | 2 mo             |
| Kumar et al [16]           | 2019                | Tongue                     | Interventricular septum                                    | Syncope                                       | Complete heart block                            | Pacemaker implant     | 1 wk             |
| Shafiq et al [17]          | 2019                | Tongue                     | Left ventricle                                             | None                                           | ST elevation                                     | Chemotherapy          | Months           |
| Tandon et al [18]          | 2019                | Tongue                     | Left and right ventricle                                   | Acute chest pain                              | ST elevation                                     | None                  | N/A              |
| Present case               | 2020                | Tongue                     | Right ventricle                                            | None                                           | None                                             | Chemotherapy          | 4 mo             |

AF – atrial fibrillation; ECG – electrocardiogram; N/A – not applicable.
was assessed every 3 weeks, according to the National Cancer Institute-Common Terminology Criteria for Adverse Events (CTCAE version 4.03, 2010).

Moderate renal toxicity with moderate anemia, associated with electrolyte imbalances, was recorded. The ECG started to show some changes, including repolarization abnormalities in the anterior leads that were not present in the first ECG. Moreover, a central venous catheter infection (that required removal of the catheter) was reported. Objective response was clinically evaluated before every cycle and by imaging after the 2 cycles. The patient was restaged by radiological imaging with a contrast-enhanced CT scan. The patient’s objective response rate was assessed according to Response Evaluation Criteria in Solid Tumors (RECIST) criteria [7]. The patient experienced progression of disease due to the appearance of a new cardiac metastasis. Because of disease progression, chemotherapy was definitively suspended, and the patient received supportive care during the rapid decline of his physical condition. The patient died in April 2019.

Discussion

Head and neck tumors represent an extremely rare source of cardiac metastasis, with fewer than 15 cases described in the medical literature [2,5,8] (Table 1). Differential diagnosis has to include a number of clinical conditions. In our case, the main imaging finding at admission was a hypermetabolic mass described in the apex of the right ventricle and at the ostium of the mitral valve. The suspected diagnosis on admission was endocarditis. This hypothesis was subsequently excluded based on clinical findings. The patient did not show any clinical characteristics of endocarditis. His levels of C-reactive protein and other inflammatory markers were just slightly altered, and his blood cultures were repeatedly negative. In addition, the TTE images were also highly atypical for endocarditis, with a round-shaped mass detected near the apex of the right ventricle. Other possible causes of cardiac hypermetabolic signals are primary tumors of the heart, but they are 40 times as rare as secondary tumors (with a reported prevalence in autopsy series of 0.001-0.3%). Furthermore, the most common tumor in this group is the myxoma, which generally shows very low or absent 2-[18F]fluoro-2-deoxy-d-glucose (FDG) uptake [19]. Blood thrombi can occasionally present as circular masses, but they tend to show no FDG uptake [20].

The main clinical findings that supported the diagnosis of a myocardial metastasis, even in the absence of a direct pathological confirmation, were the tissue characterization and demonstration of tissue infiltration based on MRI and the significant volumetric increase of the mass in a limited time frame. The diagnosis was also reinforced by the detection of a subcutaneous metastasis that suggested widespread disease.

We believe that our case presents a series of particularities that stand out even in the frame of a very rare diagnosis. In our literature search we observed that symptoms or electrocardiographic abnormalities were reported in most cases. In contrast, our patient remained completely asymptomatic for months and did not show any significant abnormality in his initial ECG. We also believe that our case can be considered unusual for the intramyocardial localization because most reported patients showed pericardial or intracavitary metastasis. Life expectancy is considered to be of few weeks in most patients, while our patient had a survival of about 4 months.

Conclusions

The paucity of antemortem diagnosis of cardiac metastasis emphasizes the importance of early recognition of cardiac masses in order to determine appropriate medical or surgical therapy for patients. Our case highlights the significance of echocardiography and MRI in the antemortem detection of a single metastasis, even in the most unusual localizations.
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