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**Radiotherapy for hyoid bone metastasis from lung adenocarcinoma: A case report**

Hsu J *et al*. Lung adenocarcinoma metastasis to hyoid bone

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**Abstract**

BACKGROUND

Metastasis to the hyoid bone is an exceptionally rare occurrence, with documented cases limited to breast, liver, colon, skin, lung, and prostate cancers. This report highlights an unusual case involving the metastasis of lung adenocarcinoma to the hyoid bone, accompanied by a distinctive headache. Previous documentation involved surgical resection of the hyoid mass. We present a case displaying the benefits of palliative radiotherapy.

CASE SUMMARY

A 72-year-old non-smoking, non-alcoholic woman, initially under investigation for a year-long elevation in absolute lymphocyte count, presented with a month-long history of intermittent throat pain. Despite negative findings in gastroenterological and otolaryngologic examinations, a contrast-enhanced chest computed tomography scan revealed a mediastinal mass and questionable soft tissue thickening in her left anterolateral neck. Subsequent imaging and biopsies confirmed the presence of lung adenocarcinoma metastasis to the hyoid bone. The patient was treated with platinum-based chemo-immunotherapy along with pembrolizumab. Ultimately, the lung cancer was unresponsive. Our patient opted for palliative radiation therapy instead of surgical resection to address her throat pain. As a result, her throat pain was alleviated, and it also incidentally resolved her chronic headaches. This is the second documented case of lung adenocarcinoma metastasizing to the hyoid bone.

CONCLUSION

Palliative radiotherapy may add to the quality of life in symptomatic patients with cancer metastatic to the hyoid bone.

**Key Words:** Metastasis; Radiotherapy; Adenocarcinoma; Hyoid; Throat; Headache; Case report

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**Core Tip:** Metastasis of lung adenocarcinoma to the hyoid bone is exceedingly rare. This case report spotlights the value of palliative radiation therapy for throat pain in patients with tumors metastatic to the hyoid bone.

**INTRODUCTION**

This case report accounts for the second reported instance of hyoid metastasis from lung adenocarcinoma. Metastatic tumors to the hyoid bone are rare, with sporadic instances reported in breast, liver, colon, skin, lung, and prostate cancers[1-6].We present the case of a 72-year-old female patient with stage IV terminal lung cancer presenting with throat pain attributed to a metastatic tumor on the hyoid. Choosing radiotherapy over surgery, she experienced significant relief from her throat pain.

**CASE PRESENTATION**

***Chief complaints***

A 72-year-old white female patient presents to an otolaryngology clinic complaining of intermittent throat pain for the past month.

***History of present illness***

Symptoms of throat pain persisted for a month with no response to antibiotics, antihistamines, or acid reflux medications. Alongside the throat pain, she was experiencing left, lateral, posterior headaches and intermittent episodes of vertigo.

***History of past illness***

Patient with no history of smoking or alcohol presented with a mildly elevated absolute lymphocyte count (ALC) of 4336 in February 2020. She denied systemic symptoms of fevers, night sweats, or unintentional weight loss.

***Personal and family history***

The patient reported only a paternal history of prostate cancer and denied melanoma, skin and squamous cell carcinomas in her family.

***Physical examination***

No masses were appreciated on palpation of her neck, flexible laryngoscopy, nor esophageal endoscopy.

***Laboratory examinations***

Mildly elevated ALC of 4336 in February 2020. A CBC in 2021 demonstrated an increase in lymphocyte count with an ALC of 5645. Given this increase in lymphocytosis, the patient was referred to a hematologist to be worked up for follicular lymphoma.

***Imaging examinations***

A computed tomography (CT) scan of the chest abdomen pelvis with contrast was performed, which incidentally showed a single para mediastinal 2.8 cm mass in the upper left lung lobe and questionable soft tissue thickening in her left anterolateral neck. Flow cytometry of her peripheral blood revealed an abnormal CD10 positive B-cell population, co-expressing CD20, CD19, and CD45 with low CD23 present, which is consistent with a follicular lymphoma. A CT neck with contrast, followed by magnetic resonance imaging (MRI) of the neck with and without contrast were ordered to further evaluate the neck tissue. The CT and MRI of the neck confirmed a focal soft tissue lesion with residual calcifications measuring 1.4 cm × 1.9 cm × 1.7 cm centered on the left limb of the hyoid bone (Figure 1). MRI of the neck also revealed a suspicious thyroid nodule which required an ultrasound to be further evaluated.

Due to the improbability of follicular lymphoma in the anterior mediastinum without associated lymphadenopathy, a lung biopsy was recommended. A CT-guided biopsy of the mass in the upper left lung lobe mass identified it as invasive mucinous adenocarcinoma. Figure 2 displays the biomarkers tested for the diagnosis. Positron emission tomography (PET)/CT revealed both the ipsilateral hilar node lesion and the hyoid bone lesion exhibited significantly higher metabolic activity compared to the primary lung cancer (Figure 3). Ultrasound-guided fine needle aspiration biopsy of the left hyoid mass and the right thyroid nodule resulted in a diagnosis of metastatic mucinous adenocarcinoma and papillary thyroid carcinoma respectively. Correlation of the biopsy with PET/CT confirmed that the hyoid bone tumor had metastasized from the primary lung adenocarcinoma.

**FINAL DIAGNOSIS**

Metastatic lung adenocarcinoma of invasive mucinous type to the ipsilateral hilar node and hyoid bone. The patient was diagnosed as papillary thyroid carcinoma and follicular lymphoma.

**TREATMENT**

The patient, now diagnosed with metastatic lung adenocarcinoma to the ipsilateral hilar node and hyoid bone, was presented to a tumor board. The tumor board recommended against a lobectomy due to the higher recurrence risk with stage III lung cancer. Dissection of the hilar lymph node was not recommended as the node was situated in the aortopulmonary zone. Since the lung adenocarcinoma lacked targetable mutations and did not express PDL1, the patient was treated with a platinum-based chemo-immunotherapy along with pembrolizumab based on the outcomes of a clinical trial in 2018[7]. The patient was also referred to head and neck surgery for consideration of resecting the metastatic cancer that engulfed the hyoid bone. However, she elected not to proceed with the procedure due to her concerns about the potential debilitating risks to swallowing and speech function.

Chemotherapy for metastatic lung cancer was initiated with Pemetrexed, Carboplatin, and Pembrolizumab for three cycles before a PET/CT scan was repeated. Despite undergoing chemotherapy, the patient continued to experience left-sided neck pain so 2 mg of dexamethasone OD was prescribed for managing her pain. PET/CT skull base to mid-thigh was repeated and after comparison to the previous PET/CT, the left upper lobe mass appeared to be stable to slightly increased in size, decreased SUV max in the ipsilateral left supra hilar node, and stable to minimally increased SUV max in the left hilar lesion which appeared grossly similar in extent to prior with some increased sclerosis of the engulfed left hyoid. These results indicated that the lung cancer was unresponsive to chemotherapy.

With disease persistent in the hyoid region and significant pain, the oncologist suggested the patient discontinue carboplatin and introduce palliative radiation for the hyoid metastatic disease. The patient chose to pursue radiotherapy. Prior to each treatment, a cone beam CT scan was obtained on the linear accelerator for targeting precision down to about 1 mm. Three hundred cGy was delivered in each treatment fraction and a total of 3600 cGy was given in 12 treatments over 2-1/2 wk.

**OUTCOME AND FOLLOW-UP**

The patient responded well to the treatment, completely remediating the patient’s complaint of throat pain and it also resolved the patient’s headaches. A repeated PET/CT showed a decrease of SUV max of 16.6 to 5.4 in the hyoid region.

Post palliative radiotherapy treatment, the patient remained asymptomatic for a year until succumbing to the advancing lung cancer.

**DISCUSSION**

Lung cancer is the leading cause of cancer death and the second most diagnosed cancer in the world[8]. Approximately 13.2% of non-small cell lung cancer metastasizes to bone[9], which are most commonly found on the spine, chest, and pelvis[10]. The hyoid bone is an unusual site for secondary tumors. This is the second reported case of lung adenocarcinoma metastasizing to the hyoid bone.

The previously documented patient elected for surgical resection of the hyoid bone[1]. Our patient opted out of surgery, due to its significant risks which could negatively impact her quality of life. The hyoid bone is positioned in the anterior midline of the neck and situated axially between the mandible and thyroid cartilage[11]. It plays a considerable role in phonation, conserving airway integrity, swallowing, and mastication; therefore, surgical resection of this bone potentially will result in the debilitation of these physiologic functions.

The benefits of palliative radiotherapy for bone cancer pain are well documented[12]. The radiotherapy of 300 cGy with each treatment, totaling 3600 cGy over 2-1/2 wk was well tolerated by the patient, and successfully mitigated the patient’s throat pain and headaches.

The relief of headaches was an unexpected positive outcome. One potential theory could be analogous to the Stylo-carotid syndrome. Patients with irregular stylohyoid complex either from an elongation of the styloid process, ossification of the stylohyoid ligament, or elongation of the hyoid bone sometimes experience neurological symptoms like headaches and vertigo. This clinical presentation of neck pain, headache, otalgia, and vertigo is due to ischemia, compression of the internal and/or external carotid arteries[11,13,14].

In conclusion, palliative radiotherapy may increase the quality of life for symptomatic patients with metastatic cancer to the hyoid bone.

**CONCLUSION**

Palliative radiotherapy may add to the quality of life in symptomatic patients with cancer metastatic to the hyoid bone.

**REFERENCES**

1 **Zhang HY**, Shan Li S, Guo X, Zhao N. Hyoid Bone Metastasis in Patients With Lung Adenocarcinoma: A Case Report. *Ear Nose Throat J* 2021; **100**: NP373-NP376 [PMID: 32283978 DOI: 10.1177/0145561320916985]

2 **Kusama H**, Okishiro M, Ishida T, Sato Y, Morimoto Y, Matsushita K, Hashimoto T, Kimura K, Katsura Y, Nitta K, Kagawa Y, Takeno A, Sakisaka H, Nakahira S, Taniguchi K, Egawa C, Takeda Y, Kato T, Tamura S, Takatsuka Y, Oku K, Goto T, Nagano T, Nakatsuka S. [A case of hyoid bone metastasis from breast cancer]. *Gan To Kagaku Ryoho* 2014; **41**: 1915-1917 [PMID: 25731374]

3 **Iguchi H**, Okabe Y, Takayama M, Wada T, Hachiya K, Matsushita N, Yamane H. [Solitary hyoid bone metastasis from hepatocellular carcinoma treated with surgery]. *Nihon Jibiinkoka Gakkai Kaiho* 2012; **115**: 783-786 [PMID: 23016270 DOI: 10.3950/jibiinkoka.115.783]

4 **Bracanovic D**, Vukovic V, Janovic A, Radosavljevic D, Rakocevic Z. Hyoid Bone and Thyroid Cartilage Metastases from Sigmoid Colon Adenocarcinoma: A Case Report. *Balkan Med J* 2017; **34**: 278-280 [PMID: 28443578 DOI: 10.4274/balkanmedj.2015.1817]

5 **Ryan JF**, Xie DX, Eytan DF, McCarthy EF, Mandal R, Gourin CG, Lipson EJ, Meyer CF, Vosler PS. Melanoma metastatic to the hyoid bone. *Clin Case Rep* 2021; **9**: 522-525 [PMID: 33489207 DOI: 10.1002/ccr3.3571]

6 **Wang SH**, Lee JC, Liu TC. Occult prostate cancer detected by hyoid bone metastasis after resection of thyroglossal duct cyst. *Int J Oral Maxillofac Surg* 2011; **40**: 1326-1328 [PMID: 21601425 DOI: 10.1016/j.ijom.2011.04.010]

7 **Gandhi L**, Rodríguez-Abreu D, Gadgeel S, Esteban E, Felip E, De Angelis F, Domine M, Clingan P, Hochmair MJ, Powell SF, Cheng SY, Bischoff HG, Peled N, Grossi F, Jennens RR, Reck M, Hui R, Garon EB, Boyer M, Rubio-Viqueira B, Novello S, Kurata T, Gray JE, Vida J, Wei Z, Yang J, Raftopoulos H, Pietanza MC, Garassino MC; KEYNOTE-189 Investigators. Pembrolizumab plus Chemotherapy in Metastatic Non-Small-Cell Lung Cancer. *N Engl J Med* 2018; **378**: 2078-2092 [PMID: 29658856 DOI: 10.1056/NEJMoa1801005]

8 **Sung H**, Ferlay J, Siegel RL, Laversanne M, Soerjomataram I, Jemal A, Bray F. Global Cancer Statistics 2020: GLOBOCAN Estimates of Incidence and Mortality Worldwide for 36 Cancers in 185 Countries. *CA Cancer J Clin* 2021; **71**: 209-249 [PMID: 33538338 DOI: 10.3322/caac.21660]

9 **da Silva GT**, Bergmann A, Thuler LCS. Incidence and Risk Factors for Bone Metastasis in Non-Small Cell Lung Cancer. *Asian Pac J Cancer Prev* 2019; **20**: 45-51 [PMID: 30678379 DOI: 10.31557/APJCP.2019.20.1.45]

10 **Zhang L**, Gong Z. Clinical Characteristics and Prognostic Factors in Bone Metastases from Lung Cancer. *Med Sci Monit* 2017; **23**: 4087-4094 [PMID: 28835603 DOI: 10.12659/msm.902971]

11 **AlJulaih GH**, Menezes RG. Anatomy, Head and Neck: Hyoid Bone. 2023 Aug 14. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2023 Jan- [PMID: 30969548]

12 **Zhu G**, Dong Y, He X, Zhao P, Yang A, Zhou R, Ma J, Xie Z, Song XJ. Radiotherapy Suppresses Bone Cancer Pain through Inhibiting Activation of cAMP Signaling in Rat Dorsal Root Ganglion and Spinal Cord. *Mediators Inflamm* 2016; **2016**: 5093095 [PMID: 26989332 DOI: 10.1155/2016/5093095]

13 **Jain S**, Bansal A, Paul S, Prashar DV. Styloid-stylohyoid syndrome. *Ann Maxillofac Surg* 2012; **2**: 66-69 [PMID: 23483633 DOI: 10.4103/2231-0746.95326]

14 **Colby CC**, Del Gaudio JM. Stylohyoid complex syndrome: a new diagnostic classification. *Arch Otolaryngol Head Neck Surg* 2011; **137**: 248-252 [PMID: 21422308 DOI: 10.1001/archoto.2011.25]

**Footnotes**

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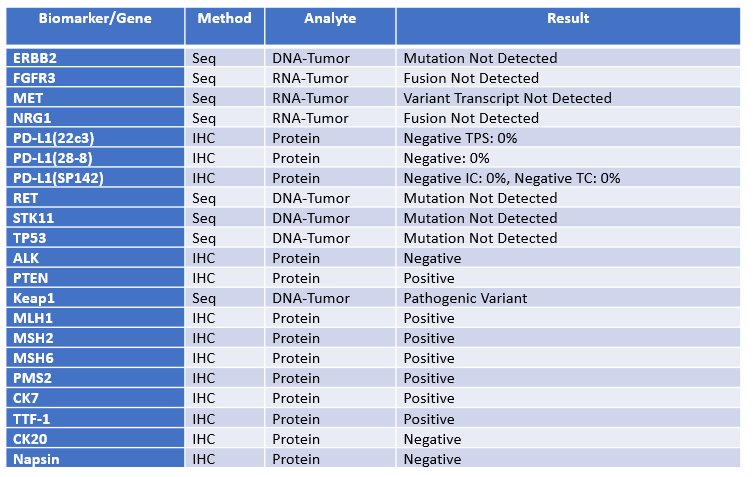
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**A B**

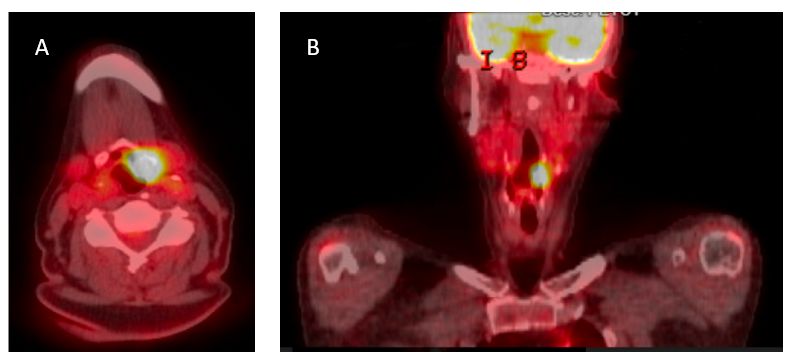
A close-up of an x-ray

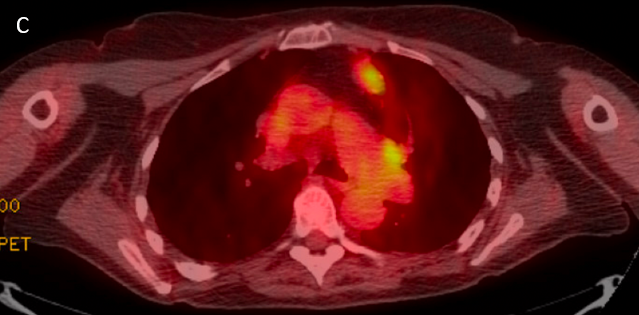
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**Figure 1 Computed tomography.** A: Computed tomography (CT) neck w/ contrast pre-radiotherapy; B: CT neck w/contrast post radiotherapy.



**Figure 2 Lung specimen biomarker/gene table.** Cancer-type relevant biomarkers and genes of the lung tumor.







**Figure 3 Positron emission tomography/**c**omputed tomography imaging.** A and B: Positron emission tomography/computed tomography (PET/CT) imaging indicating hyoid bone tumor; C: PET/CT imaging indicating lung tumors; D: Hyoid region post radiotherapy.