Primary Hepatoid Adenocarcinoma of the Lung

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ABSTRACT

Hepatoid adenocarcinoma lung (HAL) is an extremely rare type of extrahepatic adenocarcinoma with morphological similarity to hepatocellular carcinoma. We report a case of 59 years male smoker who presented with anorexia, weight loss and cough. Chest X-ray revealed a homogenous opacity in the left upper and mid zone. Computed tomography (CT) chest showed a left upper lobe mass with enlarged hilar lymph nodes. Serum alpha fetoprotein (AFP) was normal. Guided fine needle aspiration cytology (FNAC) of the mass revealed a few atypical cells. Endobronchial biopsy showed sheets of polygonal cells with coarsely clumped chromatin, eosinophilic nucleoli and abundant eosinophilic to clear cytoplasm. The neoplastic cells were positive for hepatocyte-1, Carcinogenic embryonic antigen (CEA), Cytokeratin 7 (CK-7) and cytoplasmic positivity for thyroid transcription factor (TTF-1). After 3 months, the tumor had grown significantly. Due to its aggressive behavior, early and accurate diagnosis is necessary.

Keywords: Adenocarcinoma, Hepatoid, Pulmonary origin.


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INTRODUCTION

Hepatoid adenocarcinoma is an extremely rare type of tumor with its histological resemblance to hepatocellular carcinoma. Stomach is the most common site, whereas lung accounts for nearly 5% of its incidence. We describe a case of hepatoid carcinoma arising from the lung. It is generally associated with a poor prognosis.

CASE REPORT

A 59-year-old ex-smoker presented with complaints of anorexia and weight loss for 6 months, low grade fever for 3 weeks and cough for 10 days. On examination his pulse rate was 84/minute and blood pressure was 150/80 mm Hg. Respiratory examination revealed decreased breath sounds, impaired note on percussion and crepitations in right infraclavicular area. Rest of the systemic examinations was largely unremarkable. Lab investigations including hemoglobin, total count, platelet count, prothrombin, and activated prothrombin time were within the normal range. Serum alpha-fetoprotein (AFP) was normal (0.8 ng/ml). A homogenous opacity was noted in the left upper and mid zone in the chest X-ray. Computed tomography (CT) chest revealed a large solid heterogeneously enhancing mass lesion occupying the left upper lobe measuring 13 × 9.4 × 7.5 cm with multiple enlarged subcarinal lymph nodes largest measuring 2.7 cm (Fig. 1) and a solitary well-defined homogeneously enhancing benign cystic lesion without solid areas and necrosis in the liver measuring 1.2 cm. Clinical stage was stage IIIA (cT3N2M0). On bronchoscopy, left upper bronchus was completely occluded by intrabronchial growth. He underwent image-guided fine needle aspiration cytology (FNAC) of the mass which revealed a few clusters of atypical cells with abundant eosinophilic cytoplasm (Fig. 2). Endobronchial biopsy showed an infiltrating neoplasm composed of sheets of polygonal cells with distinct cell borders exhibiting round nuclei, coarsely clumped chromatin, prominent eosinophilic nucleoli, abundant eosinophilic to clear cytoplasm (Fig. 3). Both intra and extracellular periodic acid-Schiff (PAS) positive diastase resistant hyaline globules were noted in few of the cells. Bile plugs were absent. Amidst these cells, there were areas of necrosis. In view of hepatoid morphology, immunohistochemical stains were done. The neoplastic cells were positive for cytokeratin 7 (CK7), monoclonal carcinoembryonic antigen (CEA), hepatocyte-1, AFP, cytoplasmic positivity for thyroid transcription factor (TTF-1) and negative for CK20, P63, CK5/6 and CDX-2 (Figs 4 to 8). A final diagnosis of hepatoid adenocarcinoma of the lung was made.

DISCUSSION

In 1985, Ishikura et al first described hepatoid adenocarcinoma of gastric origin. In 1990, he described its origin from the lung. He adopted two criteria for diagnosing it:

- A mixture of tubular or papillary adenocarcinoma with sheets of neoplastic cells
- Neoplastic cells with centrally located nuclei, abundant eosinophilic cytoplasm morphologically resembling hepatocellular carcinoma cells. Our case fulfills the above criteria.
The most common location is the stomach (63%). The other sites of origin include ovary (10%), gallbladder (4%), pancreas (4%), and uterus (4%), whereas lung accounts for only 5% of its incidence.² Twenty-two cases were reported in the world literature.²⁻¹⁹ Among them, 21 were males and nine were heavy smokers. Majority

Fig. 1: Computed tomography chest showing a large solid heterogeneously enhancing lesion in the left upper lobe

Fig. 2: Fine needle aspiration cytology of the mass showing atypical cells with eosinophilic cytoplasm (hematoxylin and eosin stain. Original magnification 40×)

Fig. 3: Biopsy showing sheets of cells with hyaline globules resembling hepatocellular carcinoma cells (arrow mark (hematoxylin and eosin stain. Original magnification 40×)

Fig. 4: Hepatocyte-1 showing diffuse cytoplasmic positivity in the neoplastic cells (original magnification 40×)

Fig. 5: Monoclonal CEA showing cytoplasmic positivity in the neoplastic cells (original magnification 40×)

Fig. 6: Cytokeratin 7 showing diffuse cytoplasmic and membranous positivity in the neoplastic cells (original magnification 40×)
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Histogenetically, these tumors may arise from ectopic liver, germ cells in the lung or from the multipotential respiratory epithelial cells. The prognosis is usually worse as majority of these tumors present at an advanced stage. If the diagnosis is made at an early stage, it may be possible to resect the lesion which may improve the prognosis. Our case was staged as IIIA. The patient was offered chemotherapy/radiotherapy which he refused due to financial constraints. After 3 months a follow-up chest X-ray revealed marked increase in size of the homogenous opacity. Hence, the prognosis is expected to be worse in this patient.

CONCLUSION

Primary hepatoid carcinoma being rare in the lung should be diagnosed after ruling out metastasis from other organs. Recognition of the variant is important due to its aggressive behavior, hence warrants early diagnosis and treatment.

REFERENCES