Squamous Cell Carcinoma Arising from the Pleural Cavity After Pneumonectomy for Chronic Empyema
Yeong Jeong Jeon, M.D., Sumin Shin, M.D., Young Mog Shim, M.D., Ph.D.

Department of Thoracic and Cardiovascular Surgery, Samsung Medical Center, Sungkyunkwan University School of Medicine

Malignant tumors associated with chronic empyema have been reported in the literature, and a majority of these tumors are lymphomas. Epithelial tumors originating from the post-pneumonectomy space in patients with chronic empyema are extremely rare. Here, we present the cases of 2 patients with squamous cell carcinoma arising from the pleural cavity after pneumonectomy for chronic empyema.

**Key words:**
1. Squamous cell carcinoma
2. Empyema
3. Pneumonectomy

**Case reports**

1) **Case 1**
A 74-year-old man underwent left pleuropneumonectomy for tuberculous empyema in 1997. He developed a bronchopleural fistula (BPF) and underwent an Eloesser operation following BPF closure with an omental flap in 1998. The patient was then doing well. However, in 2015, he was referred to Samsung Medical Center because of increased wound discharge and pain. Chest computed tomography (CT) demonstrated pleural thickening and irregularity, along with bony destruction in the left 10th rib, at the Eloesser aperture (Fig. 1A). A F-18 fluorodeoxyglucose positron emission tomography (FDG-PET) revealed hypermetabolic pleural thickening with invasion into the left 10th rib (maximum standardized uptake value [SUV]=21.0) (Fig. 1B). He underwent incisional biopsy for tissue confirmation. The biopsied specimen was examined microscopically; the results led to a diagnosis of squamous cell carcinoma. The patient then received definitive radiotherapy with a dose of 60 Gy in 20 fractions, and the tumor regressed markedly. He is currently in a good general condition and is being followed up as an outpatient.

2) **Case 2**
A 53-year-old woman underwent right pneumonectomy because of a destroyed tuberculous lung with pleurisy in 1990. She was diagnosed with post-pneumonectomy empyema without BPF in 2006 and underwent repeated irrigation and debridement, followed by the Clagett procedure. She visited Samsung Medical Center complaining of right shoulder and chest pain in October 2010. Chest CT revealed extrathoracic extension of enhancing soft-tissue lesions originating from the right pneumonectomy space. Percutaneous needle biopsy was performed, and the pathologic findings included a fibrinous exudate and inflammatory cells. The patient was treated with antibiotics; however, her pain was progressive, and she received a follow-up chest CT scan in March 2011. The chest CT revealed progression of the extrathoracic extension of the enhancing soft-tissue lesion,
Fig. 1. (A) Chest computed tomography demonstrated pleural thickening and irregularity, along with bony destruction in the left 10th rib, at the Eloesser aperture. (B) An F-18 fluorodeoxyglucose positron emission tomography scan revealed hypermetabolic pleural thickening with invasion into the left 10th rib (maximum standardized uptake value=21.0).

Fig. 2. (A) Chest computed tomography revealed progression of the extrathoracic extension of the enhancing soft-tissue lesion, including rib invasion (black arrow) and particularly involving the neural foramen and the epidural space at the level of the thoracic spine (T7 vertebral body) (white arrow), indicating an empyema-associated malignancy. (B) An F-18 fluorodeoxyglucose positron emission tomography scan showed a hypermetabolic malignant mass in the right pneumonectomy space (maximum SUV=35.3), right chest wall (maximum SUV=9.9), right seventh rib, and T7 (maximum SUV=26.1). SUV, standardized uptake value.

including rib invasion and particularly involving the neural foramen and the epidural space at the level of the thoracic spine (T7 vertebral body), indicating an empyema-associated malignancy (Fig. 2A). An F-18 FDG-PET scan showed a hypermetabolic malignant mass in the right pneumonectomy space, right chest wall, right seventh rib, and T7 spine (SUV=35.3) (Fig. 2B). Pathologic diagnosis of the squamous cell carcinoma was confirmed by the percutaneous needle biopsy of the soft-tissue lesions. The patient received palliative radiotherapy with a total dose of 42 Gy in 12 fractions. She developed bladder dysfunction in addition to back and leg pain. A spine CT showed an increase in the size of the malignant mass involving the T6, T7, and T8 vertebral bodies. The patient's status declined, and she eventually died.

Discussion

Chronic empyema, due to either tuberculosis or bacterial causes, can result in the development of a malignancy [1]. The most common tumor is non-Hodgkin lymphoma; however, thus far, only a few cases of epithelial tumors have been reported [2-4]. In the literature, only 6 cases of squamous cell carcinoma arising from the pleura have been reported [5].
At our medical center, 72 patients experienced post-pneumonectomy empyema between 1998 and 2015. Nineteen of these 72 patients underwent a pneumonectomy because of an infective disease resulting from tuberculosis, aspergillosis, or chronic empyema due to BPF. Two of these patients developed malignancies in the pleural cavity.

The pathogenesis of a malignant tumor arising in the post-pneumonectomy cavity associated with chronic empyema remains unknown. According to Deaton [6], squamous cell carcinoma may originate from the metaplasia of the pleural endothelium, metaplastic bronchial epithelium that has extended into the cavity, or metaplastic skin epithelium growing into the cavity.

In the cases discussed here, 2 patients experienced tuberculous empyema for several years before undergoing pneumonectomy. Chronic tuberculous empyema is known as a predisposing factor for developing a thoracic malignancy [1]. Both patients developed post-pneumonectomy empyema with or without BPF, and underwent a thoracotomy (Eloesser or Clagett). The tumors in these 2 cases were diagnosed after 17 years and 4 years, respectively. The chronic inflammatory state after post-pneumonectomy empyema is believed to have led to the development of the malignancy.

Radiographic findings in patients with chronic empyema, such as increased opacity in the thoracic cavity, soft-tissue bulging, nodular pleural thickening, and destruction of the bone near the empyema, suggest the possibility of a chronic empyema-associated malignancy [7]. Osteolytic changes with pleural thickening in the imaging findings were observed in the 2 cases reported, and were suggestive of a malignancy. Squamous cell carcinoma was confirmed by biopsy.

The treatment of tumors arising from the post-pneumonectomy space should be determined on the basis of the patient's condition. Most of the patients who underwent pneumonectomy had poor pulmonary function; therefore, radiation therapy was recommended. The abovementioned 2 patients received radiotherapy because of poor pulmonary function.

These are rare case reports of squamous cell carcinoma arising from the pleural cavity after pneumonectomy for chronic empyema. We suggest that the possibility of a malignancy originating from the pleural cavity be considered in patients who experience post-pneumonectomy empyema.

**Conflict of interest**

No potential conflict of interest relevant to this article was reported.

**References**