Primary congenital mediastinal cysts are relatively rare, accounting for approximately 12-30% of all mediastinal masses (1). The most common cyst is the bronchogenic/foregut cyst, which accounts for approximately half of these cystic lesions and typically occurs in the middle and posterior mediastinum. Recent case reports (2-6) have shown that several of these presumed foregut cysts exhibit Mullerian differentiation. Here we present two cases of posterior mediastinal ciliated cysts with Mullerian differentiation. In each case, the diagnosis was confirmed with immunostains for estrogen receptor, progesterone receptor, PAX8 and Wilm’s tumor 1, occurring in the posterior mediastinum and review the related literature.

**Case 1 presentation**

A 52 year-old female complaining of exertional shortness of breath was found to have a posterior mediastinal cyst that was originally identified on MRI scan for follow-up of a lipoma excision (Figure 1). A CT scan was subsequently performed that confirmed the presence of a 4.1 cm × 3.0 cm × 2.2 cm posterior mediastinal hypodense lesion consistent with a cyst, at the level of the carina, which was suspected to arise from either the right posterior lateral wall of the esophagus or the left main bronchus. The radiologic differential diagnosis included a bronchogenic cyst or esophageal duplication cyst. Surgical removal of the cyst was performed by robotic assisted thoracoscopy. The cyst was opened during the procedure. The removed tissue consisted of a collapsed unilocular cyst with a thin-wall and glistening, pink-tan inner surface. The specimen was submitted in its entirety for histologic examination.

Microscopically, the tissue consisted of a thin-walled cyst lined by cylindrical to cuboidal epithelium (Figure 2A, B, C, D, E, F). Ciliated, secretory, and intercalated cells, consistent with Mullerian epithelium were easily identified. Smooth muscle was found...
adjacent to the epithelial lining, and capillaries were easily identifiable in the remainder for the loose fibrous stroma. The epithelial cells had nearly 100% nuclear staining for estrogen receptor (ER), progesterone receptor (PR), PAX8, and Wilm’s Tumor 1 (WT1) by immunohistochemistry, supportive of Mullerian differentiation. Ultimately, the case was signed out as a cyst of the posterior mediastinum showing Mullerian differentiation (Hattori’s cyst).

**Case 2 presentation**

Case 2 was a 47 year-old female who presented with a left...
posterior mediastinal mass. Excision of the mass revealed a 5.0 cm × 2.0 cm × 1.0 cm cyst lined by a smooth, glistening pink-red surface. Microscopically, the cyst was lined by ciliated columnar-type cells (Figure 2G, H) and was originally diagnosed as an esophageal cyst. However, re-examination of the histology revealed focal areas consisting of Mullerian-type epithelium with secretory and intercalated-type cells. Immunohistochemical stains for ER and WT1 were performed and were diffusely positive, consistent with a Hattori cyst.

Discussion

Review of the literature reveals only five previous reports regarding mediastinal cysts in the posterior mediastinum with Mullerian differentiation (2-6). Hattori (2) was the first to describe such a cyst in 2005 in an 18-year-old female. Hattori later published a review of 19 mediastinal cysts, three of which contained Mullerian differentiation (3). Subsequently, a group in France re-reviewed all mediastinal cysts received in their laboratory, and found that 5.5% of the cysts (nine total cases) had histological features consistent with Mullerian differentiation (4). However, two of these nine cases were negative for estrogen and progesterone receptor by immunohistochemical study, bringing the diagnosis of these two cysts somewhat into question. In 2007, Businger et al. (5) published a case report of a single cyst, and Batt et al. (6) published the first case report of mediastinal cyst with Mullerian differentiation in the United States.

In total, including the present cases, fourteen definitive mediastinal cysts with Mullerian differentiation have been identified. All of the cases have occurred in females 40-60 years old, with the exception of Hattori’s original case report occurring in an 18-year-old female. The patients typically presented with cough or were asymptomatic. All cysts were paravertebral in location, occurring between T3-T8 vertebrae. All of the cysts were positive for either ER or PR. Our report is the first to show expression of PAX8 and WT1, both markers of Mullerian differentiation, in these cysts.

Mullerian cysts in locations outside of the female pelvis are rare. Some reported locations include the skin (7) and retroperitoneum (8). The etiology of Mullerian cysts arising in the mediastinum is unclear. Several authors have proposed various theories as to their origin. Batt (6) suggested that the lesion is derived directly from the primary Mullerian apparatus, similar to that of the postulated pathogenesis for the Mayer-Rokitansky-Kuster-Hauser syndrome. Hattori (2) suggested that the cyst could represent misplaced mesothelium with Mullerian characteristics. It should be noted that Mullerian cysts located in the posterior mediastinum are likely of a different etiology than those in the retroperitoneum, given that the latter often have endocervical differentiation, which typically is not seen in the mediastinal cysts (9).

While only 14 cases of mediastinal cysts with Mullerian differentiation have been described in the literature, it is likely that they are much more common than presumed in older females and often misdiagnosed as either bronchial or esophageal cysts. It is likely that a paravertebral mediastinal ciliated cyst in a middle-aged female should be considered a cyst with Mullerian differentiation until proven otherwise. However, given the limited number of described cases and lack of clinical follow-up, it is uncertain whether these cysts have different clinical outcomes than bronchial or esophageal cysts. It does appear that the cysts behave in a benign fashion.

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References
