Physiologic consequences of pneumonectomy

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Abstract: Although the incidence of pneumonectomies is decreasing all around the world, it is still required in centrally located tumors and in case of destroyed lung secondary to benign diseases. In a normal person, right lung contributes 55% of lung function due to the presence of three lobes. It is well known that remaining lung tissue expands its alveolar capacity following pneumonectomy. This compensatory growth has been attributed to stimulus by shear stress which leads to pulmonary fibroblast differentiation and neoalveolization. Pneumonectomy results in progressive deterioration of lung function (around 30%) over the years. This leads to an increase in pulmonary artery pressure and right ventricle workload. Elderly patients with preexisting pulmonary diseases are especially at risk. On the other hand, pediatric pneumonectomies under the age of five lead to a near normal pulmonary function. There are also effects on esophagus and main bronchi due to anatomic changes in the chest cavity. However, most of the pneumonectomized patients do not experience limitations in normal life. Pneumonectomy should be avoided whenever possible if an oncologically correct operation can be performed with a lesser resection.

Keywords: Pneumonectomy; pulmonary hypertension; pulmonary function

Introduction

Some of the pioneers of thoracic surgery, such as Harold Urschel and Jean Deslauriers have described pneumonectomy as a disease (1). This was especially true for right pneumonectomy as more lung tissue is removed and burden on the heart is much more profound.

Although the incidence of pneumonectomies is decreasing all around the world, it is still required in centrally located tumors and in case of destroyed lung secondary to benign diseases (2-4). In a recent multicenter French study on 4,498 patients who underwent pneumonectomy, in-hospital mortality rate was 7.8% (2).

Mortality and morbidity rate is similar in thoracoscopic pneumonectomies (7% and 28% respectively) and obviates the trauma and burden caused by removal of a single lung regardless of the approach (4).

In a normal person, right lung contributes 55% of lung function due to the presence of three lobes. However it is well known that remaining lung tissue expands its alveolar capacity following pneumonectomy (5). This compensatory growth has been attributed to stimulus by shear stress which leads to pulmonary fibroblast differentiation and neoalveolization.

Other than compensatory inflation of the remaining lung, there are significant changes involving other intrathoracic organs (heart, esophagus, hemidiaphragm, chest wall). These changes become more profound with time and there are significant differences between adults and children.

This article analyzes those physiologic changes following pneumonectomy.

Changes in pulmonary physiology

In adults

Removal of a single lung leads to a significant loss of pulmonary function. Deterioration in lung function is affected by several factors. Patient’s age, preoperative
pulmonary function, time after pneumonectomy and right/ left pneumonectomy are important (Figure 1) (6).

One of the early studies analyzed effect of thoracotomy on lung function 3–6 months after surgery (7). The patients underwent different magnitudes of lung resection. Only 13 out of 106 patients underwent pneumonectomy. While thoracotomy and wedge resection did not cause a significant decrease in exercise capacity, pneumonectomy caused a 28% decrease. In this study, forced expiratory volume in one-second (FEV\textsubscript{1}) decreased from 2.38±0.23 to 1.59±0.12 L without a significant change in oxygen saturation.

Another study was performed in Netherlands on 32 patients (surgery during 1993–1998) who were more than one year out from pneumonectomy (8). There was a decline in pulmonary function, however the results were heterogenous. Twenty-six patients were analyzed and 6 of them actually had improved pulmonary function postoperatively. FEV\textsubscript{1} declined at a rate of ≤50 mL/year in 10, 50–100 mL in 7 and ≥100 mL in 3 patients. Seventeen patients had analysis of exercise capacity and both workload in exercise capacity and maximal oxygen uptake (VO\textsubscript{2},max) decreased 27% and 30% respectively (8). In this study, at least 40% of the patients experienced impaired maximum exercise capacity due to ventilatory limitation following pneumonectomy.

The largest study was performed on 523 patients who underwent pneumonectomy over 20 years (1). Only 117 were alive at the time of analysis and 100 were included in the study. Decrease in FEV\textsubscript{1} were different for the side of pneumonectomy. For right side, FEV\textsubscript{1} decreased from 2.51±0.66 to 1.38±0.4L, whereas 2.44±0.65 to 1.53±0.37L for the left side. Change were -44% and -34% respectively, which shows the higher impact of removal of right lung. Although changes in carbon monoxide diffusion capacity (DLCO) showed similar changes with FEV\textsubscript{1}, arterial blood gases were completely normal and six minute walking distances (400±87 m) were over 80%. Hyperinflation was more profound after a left pneumonectomy (59% vs. 47%), which would be expected as right lung is bigger than the left one.

Another study analyzed the role of pulmonary rehabilitation of post-lung resection lung function in patients with chronic obstructive pulmonary disease (COPD) (9). Eighteen out of 103 patients underwent pneumonectomy. While loss in FEV\textsubscript{1} was much less profound in lobectomy patients who had preoperative physiotherapy (loss in FEV\textsubscript{1}: 23% vs. 7%, loss in vital capacity: 20% vs. 6%), the changes in pneumonectomy patients were minimal (38% vs. 29% for FEV\textsubscript{1}, 39% vs. 33% for vital capacity).

There are conflicting reports about the role of phrenic nerve function in the ipsilateral side of pneumonectomy. From the same series of 523 patients, 88 long term survivors had magnetic resonance imaging (MRI) to assess diaphragmatic function (10). It was normal in 44, whereas paralytic or paradoxic in 44. The authors concluded that patients with paralytic or paradoxic diaphragm movement has worse postoperative FEV\textsubscript{1}, forced vital capacity (FVC), DLCO and exercise capacity (6-minute walk test). These results were challenged in 12 patients who underwent ipsilateral phrenic nerve block following pneumonectomy which showed no difference in FEV\textsubscript{1} pre- and post-block (11). However these were only early analysis following block and prior study results were based on long-term follow-up (10).

In these studies, decrease in FEV\textsubscript{1} and DLCO is around 30% following pneumonectomy. Hyperinflation occurs and leads to recruitment of alveoli and better pulmonary function. Blood gas analysis shows a normal gas exchange, thus most of the patients enjoy a normal life style. Dyspnea is usually experienced upon exertion (12).

In children

Pneumonectomy is rarely indicated in children. A report of 18 children who underwent pneumonectomy for destroyed lung secondary to bronchiectasis, tuberculosis and aspergillosis showed no postoperative mortality and
complications in 3 patients (13). In a mean follow-up of 5 years, one patient developed scoliosis. Decrease in lung function was not as profound as adults and FVC was decreased by an average of 16.6%, whereas FEV$_1$ by an average of 18.2% as compared with preoperative values. All patients demonstrated a marked herniation of the remaining lung to the opposite side without any cardiopulmonary decompensation (13). We observed a similar case (17-year-old girl) who admitted with pneumothorax of an overly expanded right lung secondary to a hypoplastic left lung (14). The presentation was similar to bilateral pneumothorax as the right lung filled the whole thoracic cavity and there was scoliosis to the left side.

In another large study from South Africa, 59 children under the age of 14 underwent pneumonectomy (15). Other than one patient with lobar emphysema, all other cases were secondary to infectious etiologies. Follow-up was not adequate and authors did not report any long term effects of pneumonectomy in children.

In an earlier study with almost 30-year follow-up, patients who were younger than 5 years of age had similar lung capacity for two lungs which is explained with compensatory growth due to hyperplasia (16). Similar to the Turkish study, in the age group 6 to 20 years, there was a slight decrease in pulmonary function, but still much better than the deterioration experienced in adult patients (13,16).

Postpneumonectomy syndrome secondary to mediastinal shift occurs in 15% of the children and may lead to tracheal compression (17). It is managed by tissue expanders placed in the pneumonectomy space to prevent further shift or even aortic arch division and by-pass (17,18).

### Changes in cardiac physiology

Removal of a single lung leads to significant loss of pulmonary vascular bed and right side of the heart undergoes changes following pneumonectomy. The changes in pulmonary artery pressure, tricuspid valve problems and rate of pulmonary hypertension following pneumonectomy at various time periods postoperatively are shown in Table 1 (1,19,20).

Pneumonectomy leads to a change of cardiac position within the chest cavity (21). The study analyzed 15 patients and a lateral shift was observed following right pneumonectomy and a rotation of the heart following left pneumonectomy. Following right pneumonectomy there was an abnormally low right ventricular (RV) end-diastolic volume with a normal left ventricular (LV) function. Left pneumonectomy was associated with normal RV volumes and abnormal LV ejection fraction. Follow-up was over 5 years and patients were tachycardic with low stroke volume indicating impaired cardiac function (21).

Right pneumonectomy was associated with more impairment of cardiac functions, especially in elderly patients with low preoperative partial arterial oxygen measurements (19). Pulmonary hypertension was found to be common in elderly patients who underwent left pneumonectomy as well (1).

However in most of the patients RV index of myocardial performance was normal and late cardiac failure was uncommon (12).

#### Other physiologic effects

The empty space resulting from removal of the lung is usually filled up with serous fluid (Figure 2). In case of no postoperative complication, 70% of the chest cavity is filled within a mean of 3 days (22). Median time to complete opacification of the chest with fluid was slightly over two months. It has also been reported that a spontaneous emptying of pleural fluid from the pneumonectomy cavity can occur without any evidence of a bronchopleural fistula (23). This was thought to be the result of postoperative intrapleural pressure shifts.

Changes in esophageal emptying and dysphagia has been reported following right pneumonectomy (24). This

### Table 1 Changes in cardiac physiology following pneumonectomy

<table>
<thead>
<tr>
<th>Author and year</th>
<th>No. of patients</th>
<th>Follow-up time</th>
<th>Postop pulmonary artery pressure (mmHg)</th>
<th>Tricuspid valve problems (%)</th>
<th>Pulmonary hypertension (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Foroulis, 2004 (19)</td>
<td>35</td>
<td>6 months</td>
<td>Mean, 40.5±12.5</td>
<td>89</td>
<td>NS</td>
</tr>
<tr>
<td>Venuta, 2007 (20)</td>
<td>15</td>
<td>4 years</td>
<td>Systolic mean 34.3±7.6</td>
<td>100</td>
<td>NS</td>
</tr>
<tr>
<td>Deslauriers, 2011 (1)</td>
<td>100</td>
<td>NS</td>
<td>Mean, 36±9</td>
<td>88</td>
<td>36</td>
</tr>
</tbody>
</table>

LVEF, left ventricular ejection fraction; NS, not stated.
was related with mediastinal shift leading to angulation and compression of esophagus between vena cava and descending aorta.

We have also reported a gastric volvulus secondary to extreme elevation of left diaphragm 33 years following pneumonectomy (25). This leads to folding of the corpus of the stomach on itself towards the elevated left diaphragm dome. The patient was managed with laparoscopic Toupet fundoplication and gastropexy.

Conclusions

Pneumonectomy results in progressive deterioration of lung function over the years. This also leads to an increase in pulmonary artery pressure and right ventricle workload over the years. It is especially evident in elderly patients with preexisting pulmonary diseases. However, pediatric pneumonectomies under the age of five lead to a near normal pulmonary function, which proves that neoalveoalization is evident. Most of the patients do not experience limitations in normal life other than exercise or exertion. There are also effects on esophagus and main bronchi due to anatomic changes in the chest cavity.

In conclusion, it is advised to avoid pneumonectomy whenever possible due to long term physiologic burden, however it also well known that long term survival is possible after an oncollogically sound operation even though it is an extended pneumonectomy.

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Footnote

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