# REVIEW ARTICLE

## Surgical approaches of endobronchial neoplasms

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#### **ABSTRACT**

Endobronchial tumors are a rare entity that presents with different pathological findings. The interventional pulmonologist, but also the thoracic surgeon have at their disposal the same techniques for diagnosis, however; the two modalities differentiate in the treatment approach. Diagnosis evaluation should include lymph node evaluation. Minimal invasive techniques under local or general anesthesia are usually preferred by the interventional pulmonologists, whereas in the surgical approach of the thoracic surgeons the general anesthesia is necessary. A more extensive surgical approach either lobotomy or pneumonectomy should be performed in cases with positive intrapulmonary lymph nodes. Carinal reconstruction should be performed skillfully to get a negative proximal margin whenever needed. In the current manuscript we will present the methods of patient evaluation and surgical techniques for the management of these lesions.

#### **KEY WORDS**

Endobronchial tumor; lung cancer; local treatment

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#### Introduction

Pure endobronchial neoplasm, defined as the tumor involving the bronchial lumen mainly, is rare and presents as diverse pathological distributions (1,2). Malignant diseases are more common than benign ones and mostly originate from the surface epithelium. Squamous cell carcinoma (SCC) has recently been replaced by adenocarcinoma as the most common lung cancer. However, SCC remains the most predominant malignant endobronchial histology in adults. In contrast, carcinoid cancer constitutes the majority of endobronchial tumors in children. Mucoepidermoid carcinoma (MEC) and adenoid cystic carcinoma (ACC) are two most common endobronchial cancers derived from submucosal gland. In our experience, MEC is more

common than ACC in the bronchus, and benign endobronchial neoplasm is extremely rare (Table 1).

Most patients with endobronchial tumors present with symptoms related to cough, chest pain, wheezing, hemoptysis, recurrent pneumonia and weight loss. Endobronchial neoplasms are often diagnosed earlier than tracheal lesions for their smaller diameter. The presence of hemoptysis and obstructive pneumonia indicates possible airway intra-luminal involvement by the tumor and the need for further bronchoscopic and imaging evaluation.

Parenchymal sparing surgery remains a clinical challenge, and is attempted mostly in endobronchial neoplasm treatment (3-6). However, simple segmental bronchial resection is not adequate to achieve curable resection in most malignant endobronchial tumors, especially in patients with lymph node metastasis (7). Carina reconstruction and sleeve lobe resection are often adopted in our patients with primary main bronchial malignancies (8-12). The lesions located in the lobe bronchus are usually treated as a lung tumor with lobectomy or sub lobectomy. More complex bronchial reconstruction can be tried in patients with benign and low-grade malignant tumors if expertise in management is available (13-15). Normal lung should be preserved in patients with damaged cardiopulmonary reserve. Interventional pulmonary therapy is helpful to release

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Table 1. Histology of endobronchial neoplasms.		
Benign	Malignancy	
Tumors of mesenchymal origin	Tumors of surface epithelial origin	
Hamartomas	Squamous cell carcinoma	
Lipomas	Adenocarcinoma	
Chondromas	Small cell carcinoma	
Primary pulmonary leiomyomas	Large cell neuroendocrine carcinoma	
Granular cell tumor	Carcinoid tumor	
Neurogenic tumors	Tumors of submucosal gland origin	
Tumors of submucosal gland origin	Mucoepidermoid Carcinoma (MEC)	
Mucous Gland Adenoma	Adenoid Cystic Carcinoma (ACC)	
Pleomorphic Adenoma	Pulmonary acinic cell carcinoma	
Tumors of surface epithelial origin	Tumors of mesenchymal origin	
Papillomas	Primary pulmonary sarcomas	
Solitary squamous papillomas	Metastases restricted to the bronchus	
Columnar cell papillomas	Biphasic malignant tumors	
Fibroepithelial polyps	Pleomorphic carcinoma	
	Carcinosarcoma	
	Pulmonary blastomas	
	Others	
	Lymphoma	
	Malignant melanoma	

airway obstruction in patients with advanced malignancy as a salvage procedure. Therapeutic bronchoscopy has also been employed in children with benign or low-grade malignancy to prevent late anastomotic stenosis from surgical sutures (16). Definitive radiation therapy, including brachytherapy, can be used for patients with localized malignancy and those who refuse to accept surgical treatment. However, any interventional or radiation therapy could increase the difficulty when delayed operation is performed and add more risk of surgery-related complications.

## Surgical approaches

#### Patient selection

It is possible to make parenchymal sparing resections rather than lobectomy or pneumonectomy for the treatment of endobronchial neoplasms with special anatomic locations. However, the selection of a specific surgical option should be determined by clinical factors.

The first is the histology status and pathologic staging. It is possible to preserve the normal lung in patients with benign tumors or low-grade malignancies, and a safe margin can be achieved in 2-4 mm. However, in patients with malignant lesions, especially in those with squamous cell lung cancer, segmental or limited bronchial resection should be carried out cautiously, and

a longer surgical margins more than 5mm is needed. On the other hand, in patients with bronchial lung cancer and positive lymph nodes adjacent to the bronchus, only segmental bronchial or lobar sleeve resection could predict early recurrence and poor long-term survival as compared with pneumonectomy (17).

The second is the pulmonary function. Predicted postoperative forced expiratory volume in 1 second (FEV $_1$ ) indicates a conservative surgical strategy, such as segmental or sleeve resection rather than simple pneumonectomy or lobectomy. However, more complex surgical plasty could easily lead to postoperative atelectasis, which is more likely to occur in cases of unmatched anastomosis. Although most strictures from edema could be released spontaneously, they should be managed carefully in patients with marginal respiratory reserve. Some patients with endobronchial tumors may suffer complete atelectasis distal to the diseased airway preoperatively. Lung perfusion of the impaired side can be restored after the airway is reopened with bronchial sleeve resection. For those patients, this "nascent" pulmonary function should be considered in calculating the postoperative predicted FEV $_1$  (18,19).

Meanwhile, preoperative assessment of the local lesion is the key point to tailor the surgical plan. The surgeon should be at hand or do it personally when the diagnostic bronchoscopy is carried out. Accurate and precise location should include each end of the margin, and the spatial relationships with the surrounding bronchial airways. In this aspect, fluorescence bronchoscopy could provide more information for potential submucosal invasion or minimal mucosal change. Bronchoscopy should be scheduled again at the time of operation to confirm the previous finding. Given the limited length of the bronchus, any preoperative endobronchal intervention could kiss goodbye to the opportunity for successful reconstruction. High resolution CT scan can reveal the detailed information of the bronchus and the lymph node status. PET/CT can show the involvement of regional and distal metastasis. However, any possible lymph node metastasis in mediastinum should be confirmed histologically by mediastinoscopy or EBUS. The 11<sup>th</sup>-13<sup>th</sup> lymph nodes can be accessed by mini-probe-ultrasound-guided needle aspiration.

The requirements on systemic conditions are not significantly different from other pulmonary operations. Long-term use of high-dose steroids is contraindicated to bronchial reconstruction. Pulmonary infection with fever and elevated white blood cell should be treated preoperatively. However, obstructive atelectasis is not a contraindication, because this condition can be converted immediately after the diseased airway is corrected with careful postoperative care. Active tuberculosis (TB) should be treated medically for at least 6 months before resection.

As most bronchial resections and reconstructions involve the conditions of open airway, selected single lung ventilation is mandatory. Double-lumen endobronchial intubation can satisfy most of the conditions. However, for patients with difficult airways and pediatric patients, a more complex approach is necessary.

#### **Evaluation**

Careful preoperative evaluation is very important for a tailored operation design. Bronchoscopy is a necessity in all cases to get pathologic results and define the range of tumor involvement. No indication is available for interventional resection if the curable surgical excision is planed shortly. The use of autofluorescence-reflectance bronchoscopy as an adjunct to white light bronchoscopy can improve the detection and localization of intraepitheal neoplasia and invasion (20,21). CT scan is a key tool to evaluate the lymph node status and extra-bronchial invasion. Any suspected metastasis should be evidenced by biopsy with EBUS or mediastinoscopy. Currently, PET can add more information to occult lymph node metastasis. Tri-dimensional reconstruction with CT scan is helpful to set up a more straight understanding of the disease.

#### Surgical technique

This section concentrates on the bronchial and tracheal reconstruction for bronchus-original neoplasms rather than regular lobectomy and pneumonectomy.

The standard surgical approach for bronchial disease is

through posterolateral thoracotomy. If the proximal end of the left main bronchus or carina is involved, sternotomy, bilateral thoracotomy, or clamshell incision could be selected for better exposure. With the availability of VATS pulmonary surgery, some centers have attempted to perform bronchial plasty via a minimally invasive approach. However, this new technique is limited for the right upper lobe sleeve resection in most cases.

If the bronchus/lobe sleeve resection is planned, extensive lymph node sampling should be performed to confirm the absence of local regional metastasis. The evaluated range should include any accessible area of the isolateral lung and mediastinum. A detailed procedure requires sampling and intraoperative frozen sections of abnormal and representative lymph nodes from the fissure and along the bronchus of the lobe to be preserved. Any metastasis along the bronchus or located at the fissure indicates a more extensive lung resection rather than bronchial sleeve resection along.

Isolation of the bronchus on the right side is easier than that on the left side. If a malignant lesion involves the right upper lobe bronchial orifice, exploration should be extended to the distal end of the intermittent bronchus and carina. Hilar release and pre-tracheal dissection is useful for the condition of carina reconstruction. Exposure of the left main bronchus, especially the proximal end, is usually difficult. Isolation of the aortic arch can increase more space for this area. Sometimes, we prefer to sacrifice the left superior segment to provide additional space for complex left main bronchial reconstruction, and the outcome is excellent.

When the bronchus is isolated and ventilation is controlled, the bronchus can be divided. In neoplastic disease, frozen section analysis is obtained by sending a thin ring of tissue from the two ends of airway to be anastomosed. A safe margin more than 5 mm is needed for the malignant tumor, and it can be compromised to 2 mm for the low-grade and benign lesions.

Currently, no attempt is made to tailor either end of the bronchus to correct the unmatched orifice in the case of end-toend anastomosis. A scuff anastomosis can provide a patent and complete connection for such a kind of airway reconstruction. Carinal involvement is common for the right main bronchial malignancy. Various reconstruction methods have been introduced for half carina resection. Miyamoto's technique is an excellent alternative to be selected in this condition (22). When the proximal end of the left main bronchus is invaded, left main bronchial sleeve resection and carinal reconstruction can be performed to secure a free margin. Several approaches for the left part of carinal operation have been reported, such as midline sternotomy, left thoracotomy, bilateral thoracotomy, midline sternotomy with left thoracotomy, and clamshell thoracotomy (23,24). However, left side carinal reconstruction is much more difficult than right side carinal reconstruction due to the narrow operative field and minimal mobilization of the airway. Excellent

exposure and reliable ventilation must be guaranteed no matter what approach is selected. Modified one-stoma-type carinoplasty could help avoid aggressive carinal resection for lung saving, and the outcome has proved to be good. To make more space and minimize tension under the aortic arch, sacrificing a lobe can be performed as an alternative to pneumonectomy.

For tumors involving the distal end of the main bronchus, upper bronchus, right intermediate and superior segment of the lower lobe, finely tailored bronchoplasy is demanded (25,26). When lobar and/or segmental sleeve resection with reconstruction is performed, sacrificing one adjacent segment could leave more space to do anastomosis.

Selection of the anastomotic technique depends on the surgeon's favor. Continuous suture with 4-0 or 3-0 PDS can save more time and achieve comparable results as compared with interrupted suture with 4-0 Vicryl (27,28). All anastomoses must be evaluated by bronchoscopy. The inflation test is also important to test the patency of the distal lung. Immediate collapse is more important to infer a patent anastomosis than enforced inflation. Any unsatisfactory anastomosis should be corrected immediately, even when conversion to lobectomy or pneumonectomy is necessary.

Autologous tissue wrapping is not routinely used for any kind of airway plasty. It is only indicated as a salvage procedure when the anastomosis is not satisfactory. Pericardial fat pad (PFP) sounds a good candidate for easy harvest and minimal invasion. However, in our experience, fat liquefaction is the inevitable fate of PFP with a risk of local infection. The intercostal muscle remains the optimal coverage tissue for airway anastomosis.

### Postoperative care

The postoperative course after endobronchial tumor resection is usually as even as regular lung surgery. Pain management is the key point to guarantee effective cough and physical therapy. Any sign of atelectasis distal to the anastomosis should be evaluated by bronchoscopy if the condition permits. Temporary anastomosis edema can be managed conservatively (29). Resurgery and interventional stent therapy should be considered when surgical-related stenosis occurs.

#### Perioperative outcome and long-term survival

There is no consistent result with respect to the operative mortality and morbidity in surgical treatment of endobronchial neoplasm because of the low incidence of the condition. If the carina is involved, especially at the left side, the incidence of surgical complications will be higher than other locations. Anastomotic stenosis is the most common complication after bronchial plasty including early and late strictures. Late stenosis is usually treated by interventional bronchoscopic therapy.

The airway should be evaluated with careful bronchoscopic examination. A personalized design is mandatory for any interventional therapy.

Long-term survival is different depending on the histology. For low-grade malignant and benign lesions, complete resection can achieve a very good survival outcome. The lymph node status and complete resection will generate different survival outcomes for malignant tumors, especially for SCC. Deliberate parenchyma-sparing surgery with misdiagnosis of metastasis could be harmful to the patients.

#### **Conclusions**

Endobronchial neoplasm is rare with a histological diversity. A personalized therapeutic strategy is important to maximize the efficacy of therapy. The lymph node status should be evaluated thoroughly before parenchyma-sparing surgery for malignant endobronchial tumors. Do not hesitate to do a lobectomy or pneumonectomy in cases with positive intrapulmonary lymph nodes. Carinal reconstruction should be performed skillfully to get a negative proximal margin whenever needed. Sacrificing a segment or lobe to obtain a wider surgical field may spare pneumonectomy in some cases.

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