

# Bronchogenic carcinoma in idiopathic pulmonary fibrosis patients after lung transplantation

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## Introduction

Lung transplantation (Ltx) is an accepted therapy for patients with end-stage lung disease. The most important indications are chronic obstructive pulmonary disease (COPD) and idiopathic pulmonary fibrosis (IPF).<sup>1</sup> The incidence of lung cancer after Ltx is 20-25 times higher than in the general population, but diagnosis is often difficult.<sup>2</sup> We describe three Ltx patients suffering primarily from IPF.

## Cases

Patient A presented 7 years after single Ltx with malaise. He was renovating his house. A HRCT of the chest showed an increasing opacity in the native lung replacing the fibrotic lesions (Figure 1). The differential diagnosis consisted of fungus infection, rejection and malignancy. Eventually he was diagnosed with a bronchogenic carcinoma, he died shortly after.

Patient B underwent a bilateral Ltx. In the explanted right lung a T2N2 carcinoma was found. Two years after radical chemo-radiotherapy progression appeared, shortly after he died.

Patient C complained of left pretibial pain before Ltx. <sup>18</sup>FDG-PET showed uptake in both lungs and the left tibia. The first was interpreted as compatible with her IPF, the latter was suggestive for Pierre Marie Bamberger. At the time of Ltx, however, she was diagnosed with an adenocarcinoma in both lungs. She died shortly after, see also Table 1.

Table 1: Summary of the clinical data of the presented cases.

|   | Patient A, male            | Patient B, male  | Patient C, female                      |
|---|----------------------------|--|--|
| Age diagnosis IPF / Ltx resp.                 | 46 / 48 years              | 53 / 57 years  | 47 / 53 years                          |
| Histology                                     | UIP                        | UIP  | NSIP, fibrotic type                    |
| Smoking status                                | ex, 30 packyears           | ex, 26 packyears                                       | never                                  |
| Treatment IPF (all acetylcysteine/prednisone) | cyclophosphamide           | azathioprine   | azathioprine                           |
| Single or bilateral Ltx                       | single (right)             | single (left), after rejection bilateral               | bilateral                              |
| Histology explanted lung                      | UIP, no malignancy         | left: UIP, right: squamous cell carcinoma              | massive bilateral adenocarcinoma       |
| Time Ltx - carcinoma                          | 8 years                    | in explanted lung                                      | in explanted lung                      |
| Stage and treatment                           | IV: none, poor performance | T2N2M0 (IIIA): chemo- radiotherapy, progression: chemo | CT4N2M1b (IV): palliative radiotherapy |
| Time carcinoma - death                        | 3 months                   | 22 months  | 3 months                               |

Figure 1: HRCT patient A.



HRCT of patient A showing an increasing opacity in the native lung replacing the fibrotic lesions, also ground glass areas in the transplant lung.

## Discussion

After Ltx the incidence of bronchogenic carcinoma is increased. Risk factors are IPF *per se*, immunosuppressive drugs, single versus bilateral Ltx, smoking, increasing age and male gender.<sup>3</sup>

In 6.9% of single Ltx a bronchogenic carcinoma arises in the native lung as we found in patient A.

This is rarely accounted when a bilateral Ltx is performed.<sup>3</sup> Moreover, nowadays most bilateral Ltx are done in Leuven. In 2% of patients a bronchogenic carcinoma is unexpectedly found in the explanted lung, as we found in patient B and C.<sup>1</sup>

Symptoms are usually aspecific or mimic an infection or rejection as in patient A.<sup>4</sup> Adenocarcinoma and squamous cell carcinoma represent the most frequent pathological types, followed by small cell carcinoma.<sup>4</sup> Although disease is often diagnosed in an early stage, the prognosis remains extremely poor.<sup>3</sup>

## Conclusions

Transplanted IPF patients are at risk for developing primary bronchogenic carcinoma. Symptoms are often aspecific, diagnosis is difficult and prognosis is extremely poor. These cases stress the importance of actively searching for bronchogenic carcinoma before as well after lung transplantation in patients with IPF.

## References

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