

Supplementary Materials

SUPPLEMENTARY TABLE S1 Phenotypes of chronic lung allograft dysfunction [1]

	BOS	RAS	Mixed^a	Undefined^b	
Obstruction (FEV ₁ /FVC <0.7)	✓	X	✓	✓	✓
Restriction (≥10% ↓ in TLC)	X	✓	✓	X	OR ✓
CT opacities ^c	X	✓	✓	✓	X

^aBy definition, all cases that transition from a BOS to a RAS phenotype and vice versa will meet these criteria.

^bUndefined means definite CLAD but with two possible combinations of variables making it difficult to categorise as one of the other phenotypes.

^cParenchymal opacities and/or pleural thickening consistent with a diagnosis of pulmonary and/or pleural fibrosis.

BOS: bronchiolitis obliterans syndrome; CLAD: chronic lung allograft dysfunction; CT: computed tomography; FEV₁: forced expiratory volume in 1 second; FVC: forced vital capacity; RAS: restrictive allograft syndrome; TLC: total lung capacity.

SUPPLEMENTARY TABLE S2 Other conditions/scenarios affecting pulmonary function in lung transplant or haematopoietic stem cell transplant (HSCT) recipients that need to be excluded during the diagnostic work-up of patients with suspected chronic lung allograft dysfunction or graft-versus-host-disease bronchiolitis obliterans syndrome

Lung transplantation [1]	HSCT [2]
<p>A. Conditions in which it may be valid to recalculate/rest FEV₁ reference value</p> <ul style="list-style-type: none"> • Decreasing lung due to normal aging process • Surgical factors (e.g. transplant lung resection, chest wall surgery, phrenic nerve damage) • Mechanical factors (e.g. persistent pleural effusion, persistent lung oedema due to significant kidney/heart/liver failure, airway stenosis, myopathy/neuropathy/Parkinson’s disease, weight gain, native lung hyperinflation after single-lung transplant) • Localised infection with chronic scarring (e.g. abscess/empyema/mycetoma) <p>B. Factors that cannot be differentiated easily from CLAD and do not ever allow</p>	<ul style="list-style-type: none"> • Infection • Idiopathic pneumonia syndrome • Cryptogenic-organising pneumonia • Pulmonary fibrosis • Late radiation effects • Asthma • Chronic obstructive pulmonary disease • Tracheomegaly • Tracheobronchomalacia • α_1-antitrypsin deficiency

recalculation/resetting of the FEV₁ reference value

- Any from (A) above where there is not stability for ≥6 months
 - Infiltration with tumour
 - Infiltration of the allograft with proven disease recurrence from the underlying transplant indication (e.g. LAM, sarcoidosis)
 - Drug or other induced pulmonary toxicity (e.g. sirolimus, methotrexate, amiodarone, radiotherapy)
 - Pulmonary arterial strictures or emboli
 - Acute/subacute generalised infection
 - Acute/subacute cellular or antibody-mediated rejection
 - Acute/subacute effects of aspiration
- C. Failing to reach normal predicted lung function (i.e. low FEV₁ reference value such that FEV₁ is ≤80% of the recipient predicted value)**
- May include an age difference between donor and recipient where older donor
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lungs are implanted or when an intra-operative allograft reduction surgery/lobectomy is performed	
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FEV₁: forced expiratory volume in 1 second; HSCT: haematopoietic stem cell transplant; LAM: lymphangioleiomyomatosis.

References

- 1 Verleden GM, Glanville AR, Lease ED, et al. Chronic lung allograft dysfunction: Definition, diagnostic criteria, and approaches to treatment - a consensus report from the Pulmonary Council of the ISHLT. *J Heart Lung Transplant* 2019; 38: 493-503.
- 2 Williams KM. How I treat bronchiolitis obliterans syndrome after hematopoietic stem cell transplantation. *Blood* 2017; 129: 448-455.