

controlled asthma with a mean predicted FEV<sub>1</sub> of 54%. 13/24 (61%) were on a maintenance dose of oral corticosteroids (mean dose 19 mg/day). The mean inhaled corticosteroids was 1006 µg. The cause of death was not known in 6 cases, with majority of cases died from non-asthma causes (78%) with cardiovascular disease being the most common. Asthma was the cause of death in 4/18 (22%) cases. Co-morbid diseases were prevalent particularly those that form the metabolic syndrome. Non-concordance with asthma medications and smoking history (current and ex-smokers) were also common (50% and 60% respectively).

**Conclusion** Although death is a rare event in our severe asthma service (0.7%), patients did die prematurely (mean age 51 yrs) usually from non-asthma causes, but asthma still accounted for death in the fifth of this group. Larger multicentre study with control data will be needed to confirm these findings and look for drivers and predictors of mortality in severe asthma.

## New markers of lung physiology

### P79 COMPARISON OF CF AND NON CF LCI RESULTS USING THE EXHALYZER D AND INNOCORTM DEVICES

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**Background** Multiple breath washout (MBW) is a pulmonary function test that allows measurement of lung clearance index (LCI), a marker of ventilation heterogeneity. LCI is increasingly being considered as a clinical trial outcome measure. The Exhalyzer D N<sub>2</sub> washout (N<sub>2</sub>/ExD, Ecomedics AG, Switzerland) is the recommended technique by the ECFS CTN. Mostly our group's experience with LCI uses the Innocor<sup>TM</sup> gas analyser (Innovision, Denmark), SF<sub>6</sub> as a tracer gas (SF<sub>6</sub>/Inn). To understand the N<sub>2</sub>/ExD technique, the aims of this study were to compare a) intra-test variability; b) LCI values c) LCI and FEV<sub>1</sub> for the two techniques.

**Methods** 21 CF (14F, mean 20 ± 12 yrs) and 10 non-CF (8F, mean 28 ± 10 yrs) have completed MBW trials with the 2 techniques in random order on the same visit; methods previously described (Horsley *et al.* 2009, Jensen *et al.* 2013).

**Results** Intra-test CoV (2–3 repeats) was similar for SF<sub>6</sub>/Inn and N<sub>2</sub>/ExD (3.57 vs 3.78 CF, ns). LCI was significantly higher with N<sub>2</sub>/ExD than with SF<sub>6</sub>/Inn in both CF (14.05 vs 9.33; p = <0.0001) and non-CF (7.08 vs 6.64, p = 0.04). Both techniques show a significant correlation for LCI and FEV<sub>1</sub>%, although the regression slopes were significantly different (p = 0.005); much steeper for N<sub>2</sub>/ExD.

**Conclusions** LCI values cannot be used interchangeably between the Innocor<sup>TM</sup> and Exhalyzer D. In non CF, there was a statistically significant difference, with higher values in the N<sub>2</sub>/ExD. The difference was much greater in CF and increased as LCI

did. In the context of similar intra-test variability, the hypothesis that this may translate into improved power in future clinical trials needs to be tested. Sponsored by a non-restricted grant from Novartis.

### REFERENCES

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- Jensen R, Stanojevic S, Gibney K, *et al.* Multiple breath nitrogen washout: a feasible alternative to mass spectrometry. *PLoS One*. 2013;**8**:e56868

### P80 EXTRAPOLATING LUNG CLEARANCE INDEX (LCI) FROM SHORTENED MEASUREMENTS

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**Introduction and objectives** Shortened multiple breath washouts (MBW) are attractive in young children, where a long test time may not be tolerable (Thorax 2013;68:586–7). Lung clearance index (LCI) is usually calculated at 1/40<sup>th</sup> of the starting concentration (LCI<sub>std</sub>), but in shorter MBWs the LCI is calculated at 1/20<sup>th</sup> of the starting concentration (LCI<sub>0.5</sub>). LCI<sub>0.5</sub> and LCI<sub>std</sub> are closely correlated but not interchangeable, and so we calculated an extrapolated full LCI (LCI<sub>ex</sub>) from LCI<sub>0.5</sub>. Our hypothesis was that extrapolated LCI (LCI<sub>ex</sub>) will better reflect LCI<sub>std</sub> than LCI<sub>0.5</sub> and be more sensitive to intervention.

**Methods** Condition-specific equations for cystic fibrosis (CF), primary ciliary dyskinesia (PCD) and asthma, and an overall equation for all 3 groups, were developed to convert LCI<sub>0.5</sub> to LCI<sub>ex</sub> from data previously analysed (n = 90, Thorax 2014;69 [Suppl<sup>2</sup>]A166). LCI<sub>std</sub>, LCI<sub>0.5</sub> and LCI<sub>ex</sub> were then calculated for new cohorts (n = 70, 20 asthma, 30 CF, 20 PCD), and LCI<sub>ex</sub> was compared to LCI<sub>std</sub>. CF patients receiving IV antibiotics (n = 17) and asthma patients receiving triamcinolone (n = 32) also had LCI<sub>std</sub>, LCI<sub>0.5</sub> and LCI<sub>ex</sub> calculated and compared. The upper limit of normal for LCI<sub>std</sub> was calculated from healthy controls. LCI<sub>ex</sub> was compared to LCI<sub>std</sub> with a Bland-Altman plot.

**Results** In CF, at higher LCIs, the spread between LCI<sub>ex</sub> and LCI<sub>std</sub> grew but there was no bias. For PCD and asthma agreement remained very good. There was no significant difference between LCI<sub>std</sub> and LCI<sub>ex</sub>. Results for positive and negative prediction of LCI<sub>std</sub> for LCI<sub>0.5</sub> and LCI<sub>ex</sub> are shown in the Table 1. LCI<sub>ex</sub> and LCI<sub>0.5</sub> were also sensitive to interventions in asthmatic and CF patients, although in general LCI<sub>std</sub> had better p values.

**Conclusions** LCI<sub>ex</sub> performed better than LCI<sub>0.5</sub> in predicting results of LCI<sub>std</sub>. However, LCI<sub>ex</sub> did not always reflect LCI<sub>std</sub>, particularly in CF, so the two cannot be used interchangeably in lung disease.

**Abstract P80 Table 1** LCI<sub>ex</sub> had an improved false negative rate compared with LCI<sub>0.5</sub>

LCI	Correctly categorised (%)				Sensitivity to intervention	
	True positive	True negative	False positive	False negative	Asthma	CF
LCI <sub>std</sub>	-	-	-	-	p = 0.001	p = 0.03
LCI <sub>ex</sub>	37	26	5	2	p = 0.02	p = 0.02
LCI <sub>0.5</sub>	38	26	0	6	p = 0.02	p = 0.04