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Title: Routine measurement of the LCI in CF with an ultrasonic device for multiple breath nitrogen washout

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Body: During the last decade multiple breath washout technique (MBW) for calculating the Lung Clearance Index (LCI) has become very popular for assessing ventilation inhomogeneity (VI) as an early manifestation of Cystic Fibrosis (CF) lung disease. However, routine use has been difficult not only due to limited availability of licensed equipment. Inert tracer gases (e.g. SF_6 , He) certified for medical purposes are not universally available. Switch to nitrogen washout (MBW $_{N2}$) using 100% oxygen may overcome this problem. The aim of this cross sectional study was to assess whether LCI derived from MBW $_{N2}$ discriminates as well as MBW $_{SF6}$ between patients with CF and healthy controls. 19 controls (7-51years) and 11 unselected patients with CF (7-25 years) performed 2-3 single MBW $_{N2}$ using the EasyOne Pro LABTM (ndd Switzerland) with 100% oxygen. Mean (SD) LCI was 6.5 (0.64) in controls and 9.3 (1.93) in CF with a mean difference (95% ci, p-value) of -2.83 (-4.14;-1.51, 0.001) between the groups. Within-test repeatability (CV%) was 5.3% in controls and 7.7% in CF. Assessment of LCI using licensed equipment for MBW $_{N2}$ was feasible and well tolerated in both, children and adults and patients and controls. LCI based on MBW $_{N2}$ differed significantly between patients with CF and controls and results were comparable to published data obtained with different equipment and with using SF $_6$ as tracer gas. We conclude that MBW $_{N2}$ reflects VI similar to MBW $_{SF6}$ and may thus be used for clinical application of MBW in patients with CF.