

In contrast, SpO<sub>2</sub>% indicated significant respiratory depression in only 4/10 patients, with small absolute changes in SpO<sub>2</sub>% from 96.5 (95.1 to 99.2)% at baseline to 96.2 (95.2 to 97.0%) at 30 min. A non-significant decline in NRD from baseline (109.5 (69.5 to 185.1) a.u.) to 30 min post IOT 84.3 (59.2 to 118.1) a.u.,  $p = 0.12$  was also observed. Baseline NRD and opioid-induced drop in SpO<sub>2</sub>% were inversely related ( $r = -0.67$ ,  $p = 0.04$ ).

**Conclusion** Significant acute respiratory depression is commonly induced by opioid drugs prescribed to treat opioid addiction. Hypoventilation is reliably detected by capnography, but not by SpO<sub>2</sub>% alone. Chronic suppression of NRD in the presence of underlying lung disease may be a risk factor for acute opioid-induced respiratory depression.

### S51 ARTERIAL OXYGEN CONTENT REFLECTS HAEMOGLOBIN MORE THAN OXYGENATION INDICES IN 440 PATIENTS WITH PULMONARY ARTERIOVENOUS MALFORMATIONS

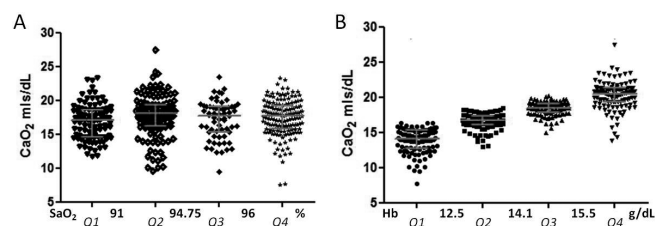
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**Introduction and objectives** Our goal was to use a long term model of human hypoxaemia to evaluate factors that reduce arterial oxygen content (CaO<sub>2</sub>) and therefore demand higher cardiac outputs to maintain tissue oxygen delivery. This is important for clinical practice; for clinical trials that use cardiac index as a primary outcome measure; and particularly relevant for patients with pulmonary and systemic arteriovenous malformations (AVMs) due to hereditary haemorrhagic telangiectasia (HHT).

**Methods** Presentation data were evaluated on 497 consecutive patients with pulmonary AVMs due to HHT, reviewed between 1999 and 2013. SaO<sub>2</sub> was measured by pulse oximetry in the supine and erect postures, and the mean SaO<sub>2</sub> calculated after 7, 8, 9 and 10 min standing. Same-day haemoglobin was measured in venous blood samples in 440 patients. Presentation CaO<sub>2</sub> was calculated by the equation  $\text{oxygen saturation (SaO}_2, \%) \times \text{haemoglobin (gram/dL)} \times 1.34/100$ .

**Results** There was a four-fold difference in CaO<sub>2</sub> across the 440 patients (range 7.6–27.5, median 17.6) mls of oxygen per decilitre (dL) of arterial blood. SaO<sub>2</sub> ranged from 59–100% (median 94.8%), but CaO<sub>2</sub> did not change appreciably across the SaO<sub>2</sub> quartiles (median CaO<sub>2</sub> 17.1; 18.1; 17.7; 17.8 mls/dL;  $p = 0.34$ , Figure 1A). In contrast, CaO<sub>2</sub> was primarily determined by haemoglobin which ranged from 5.9–21.8 g/dL (median 14.1 g/dL). The median CaO<sub>2</sub> across quartiles of haemoglobin were 14.1; 16.7, 18.5; and 20.5 mls/dL ( $p < 0.0001$ , Figure 1B). For each 1 g/dL rise in haemoglobin, there was a 10% increase in mls of oxygen per unit blood volume.



**Abstract S51 Figure 1** Distribution of arterial oxygen content (CaO<sub>2</sub>) across the quartiles of A) oxygen saturation (SaO<sub>2</sub>), and B) haemoglobin (Hb) in 440 patients with pulmonary AVMs

**Conclusions** Currently, in long term conditions, more attention is paid to modest differences in SaO<sub>2</sub> than to haemoglobin.<sup>1</sup> It has been shown that patients with PAVMs maintain CaO<sub>2</sub>, and deliver the same amount of oxygen per heart beat (oxygen pulse) before and after correction of hypoxaemia by PAVM embolisation.<sup>2,3</sup> For patients where higher cardiac outputs may be detrimental, further attention should be given to minor incremental falls in haemoglobin that substantially reduce arterial oxygen content.

### REFERENCES

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### S52 THE EFFECT OF AGE ON ARTERIAL OXYGEN CONTENT IN PATIENTS WITH PULMONARY ARTERIOVENOUS MALFORMATIONS (PAVMs)

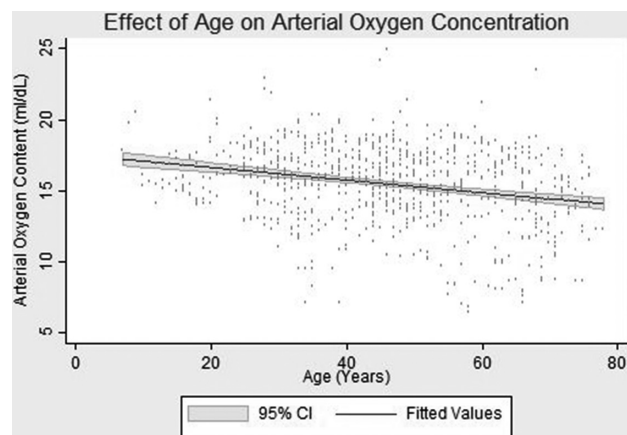
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**Introduction and objectives** It is recognised that age-associated changes in the chest wall and lung parenchyma lead to decreased efficiency of ventilation and gas exchange, resulting in reduced arterial partial pressure of oxygen (PaO<sub>2</sub>) and haemoglobin saturation (SaO<sub>2</sub>). The total oxygen content of arterial blood (CaO<sub>2</sub>) depends upon SaO<sub>2</sub>, as well as haemoglobin concentration. Our goal was to examine serial changes in arterial oxygen content with age in a cohort with hypoxaemia due to pulmonary arteriovenous malformations (PAVMs).

**Methods** Retrospective longitudinal follow-up data was collected for 100 consecutive PAVM patients presenting to a tertiary care institutional clinic between 1984 and 2001, and reviewed until 2015. Subjects provided up to 30 (median 9) separate annual datasets. SaO<sub>2</sub> was measured by pulse oximetry in the supine and erect postures, and the mean SaO<sub>2</sub> was calculated after 7, 8, 9 and 10 min standing. Haematological and biochemical blood indices evaluated haemoglobin, haematinics, and iron indices. CaO<sub>2</sub> in mls of oxygen per dL (ml/dL) of blood was calculated using the equation:  $[\text{SaO}_2 (\%) \times \text{haemoglobin (g/dL)} \times 1.34]/100$ . Data were analysed using STATA IC v13.1.

**Results** Age and PAVM-treatment associated changes in SaO<sub>2</sub> were mostly accompanied by opposing changes in haemoglobin levels that maintained the CaO<sub>2</sub>. Two major patterns were observed. The first was the expected increase in haemoglobin with lower SaO<sub>2</sub>, due to secondary erythrocytosis and polycythemia. The second, less well recognised, was an increase in SaO<sub>2</sub> when haemoglobin fell, most commonly when subjects developed iron deficiency and anaemia. Nevertheless, excluding participants with iron deficiency, CaO<sub>2</sub> decreased with age (Figure 1,  $r^2 = -0.0654$ ;  $p < 0.001$ , Figure 1).



Abstract S52 Figure 1

**Conclusions** The body maintains arterial oxygen content within a normal range using well-known erythropoietic mechanisms in response to hypoxaemia. Despite this feedback mechanism, in patients with pulmonary arteriovenous malformations, overall arterial oxygen content still decreases with age.

## Advances in cystic fibrosis

### S53 OUTCOMES FOLLOWING BRONCHIAL ARTERY EMBOLISATION FOR HAEMOPTYSIS IN ADULTS WITH CYSTIC FIBROSIS

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**Introduction** Bronchial artery embolisation (BAE) is recommended as the therapy of choice for massive haemoptysis in cystic fibrosis (CF) but there are no randomised controlled trials of BAE in this setting. Outcomes from BAE are uncertain and the efficacy of BAE in sub-massive haemoptysis is unclear. We performed a single-centre observational study to investigate the role of BAE in CF-related haemoptysis.

**Methods** All patients with CF undergoing BAE from March 2011 to January 2015 were identified at the time of the procedure. Patient records were reviewed following hospital discharge or death. Severity of haemoptysis was classified as: massive (>240 ml/24 h or >100 ml/day for  $\geq 2$  days), severe (>20 ml/24 h) or mild (<20 ml/24 h). Data were collected on adjuvant therapies, time to recurrence, complications and survival.

**Results** Twenty-seven patients underwent 49 BAE procedures. Median age was 30 years (range 18–72) and 16 (59%) were male. Mean baseline FEV<sub>1</sub>-predicted was 51.0% (SD 19.3). BAE was indicated for massive haemoptysis in 18 episodes (37%), severe in 27 (55%) and mild in 4 (8%). Adjuvant therapies included tranexamic acid in 48 (98%), intravenous antibiotics in 47 (96%), intravenous vitamin K in 31 (63%), ethamsylate in 8 (16%), terlipressin in 7 (14%) and propranolol in 5 (10%) of episodes.

Twenty-nine (59%) BAEs were complicated by adverse events including chest pain (29%), headache (12%), paraesthesia (10%), groin pain (6%), limb weakness (4%) and limb ischaemia (2%).

Eight patients (30%) required  $\geq 2$  BAEs during the study (range 2–7). Median time to first repeat BAE was 213 (range 18–682) days. Overall, haemoptysis recurred after 31/49 (63%) procedures with no significant difference between massive and sub-massive haemoptysis (61.1% vs 64.5%).

Five patients (18.5%) died during the study and this group had a median FEV<sub>1</sub>-predicted of 32% (range 28–82%). Mortality was 3.7% at 30 days following first BAE and 11.1% at six months. Four out of 8 (50%) patients requiring repeat BAE died compared with 1/19 (5%) who needed a single BAE only ( $p = 0.006$ ).

**Conclusion** BAE may be life-saving but is associated with considerable morbidity in CF. Need for repeat BAE is associated with increased mortality.

### S54 COGNITIVE FUNCTION IN ADULTS WITH AND WITHOUT CYSTIC FIBROSIS RELATED DIABETES (CFRD) ATTENDING A LARGE UK CYSTIC FIBROSIS UNIT

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**Introduction and objectives** On reaching adulthood many cystic fibrosis (CF) sufferers develop cystic fibrosis related diabetes (CFRD). CFRD shares clinical characteristics with type 1 (T1DM) and type 2 diabetes mellitus (T2DM). Impaired glucose tolerance (IGT), T1DM and T2DM have deleterious effects on cognitive performance. Hence, patients with CFRD are hypothesised to show similar impairment. This study aimed to elucidate the nature and severity of any cognitive impairment in patients with CFRD compared to non-diabetic patients with CF and healthy controls matched as closely as possible for age, gender and education level. Patients with CF were also matched as closely as possible on CFTR genotype.

**Methods** Adult (>16 years old), pancreatic insufficient patients registered to a large UK CF unit who had adequate verbal and written English were eligible. 49 patients with insulin-treated CFRD and 49 CF non-diabetics who had received a normal oral glucose tolerance test (OGTT) within the past 12 months were recruited. 46 healthy matched controls were recruited from relatives of patients and the general population. Cognitive performance was assessed using the Cambridge Neuropsychological Test Automated Battery (CANTAB). Subjective measures of sleep, stress, mood and cognitive functioning were also collected.

**Results** Matched controls performed better than both groups of patients with CF on tests of visual memory and learning, verbal memory, visual sustained attention, processing speed and executive function. Patients with CFRD performed significantly worse than controls on tests of mental flexibility and processing speed, which is consistent with the pattern of impairment shown in T1DM, and on verbal memory and learning, which is consistent with the pattern of impairment shown in T2DM. Compared to non-diabetic patients with CF, those with CFRD performed worse on tests of visual sustained attention, verbal memory, working memory, and processing speed.

**Conclusion** CFRD has a negative impact on cognitive performance akin to T1DM and T2DM. Non-diabetic patients with CF also show impaired cognition but to a lesser degree than CFRD. Even modest cognitive impairment in adults with CF may impact upon their self-management skills, health and quality of life.