SLEEP STUDIES IN COPD PATIENTS

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ABSTRACT

BACKGROUND
Sleep is a highly organized, complex behaviour characterized by a relative disengagement from the outer world and variable, but specific brain activity. It is an endogenously generated, homeostatically regulated and reversible. During sleep, there are profound physiological changes. This is particularly true of breathing and in number of conditions this had important implications. Breathing alters according to the state of consciousness. In healthy subjects, ventilation falls with the onset of sleep and is reduced during all phases of sleep compared with waking levels, alterations in the pattern of breathing occur including periodic breathing, apnoea, hypopnoea with gradual progression from stage I, II (18% decreased) to REM sleep (35% of decrease ventilation to awake stage). REM sleep related alterations in ventilation is more severe in respiratory disordered patients who had already some Hypoxemia, COPD is of best example.

METHODS
Patients who attended outpatient Department of Pulmonary Medicine, Govt. General and Chest Hospital, OMC, Hyderabad, Telangana, between May 2012 and April 2014 with clinical features s/o COPD were admitted, thoroughly examined and evaluated with routine investigations and specific investigations like sputum for AFB, Gram’s stain, PFT, Chest X-ray to establish the diagnosis of COPD. (GOLD guidelines) All these patients (n=25) filled the COPD questionnaire along with Sleep symptoms questionnaire and Epworth Sleep Scoring (ESS). These patients are advised to undergo whole night PSG after informed consent.

RESULTS
The 24 patients with COPD were divided into two groups, Group 1: 6 patients out of 24 (25%) showed significant OSAHS (AHI >5/hr, i.e. overlap syndrome); Group 2: Rest of the 18 patients (75%) showed no significant OSAHS. Among these 18 patients, 12 are having asymptomatic hypopneas (Clinically insignificant SDB), 6 are not significant OSAHS. Among these 2 groups, mean values of 25 variables were tabulated, compared and analysed by using window stat version 8.5 from IndoStat services. Among the 25 variables, 11 were significantly different between 2 groups.

CONCLUSIONS
There is positive correlation between neck circumference, BMI and Epworth score with AHI (Apnoea-Hypopnoea Index). There is positive correlation between frequency of arousals, low sleep efficiency, REM sleep with O2 drop and snoring associated arousals with AHI. There is negative correlation between severity of COPD, duration of COPD and smoking with AHI. OSAHS in COPD patients (Overlap Syndrome) is incidental and does not share the pathology of two diseases (COPD and OSA). Sleep changes like reduced REM sleep, largest oxygen saturation drop during REM sleep, frequent arousals, frequent changing of sleep stages are observed in COPD patients, but more significantly seen in overlap syndrome. Independent risk factors for OSAHS like high BMI, more neck circumference, facial abnormalities and symptoms like excessive day time sleepiness should be looked for diagnosing overlap syndrome in COPD patients.

KEYWORDS
OLS (Overlap Syndrome); OSAHS (Obstructive Sleep Apnoea Hypopnoea Syndrome); PSG (Polysomnography); AHI (Apnoea Hypopnoea Index); COPD (Chronic Obstructive Pulmonary Disease); CPAP (Continuous Positive Airway Pressure).


INTRODUCTION
Sleep is a highly organized, complex behaviour characterized by a relative disengagement from the outer world and variable, but specific brain activity. It is an endogenously generated, homeostatically regulated and reversible. During sleep, there are profound physiological changes. This is particularly true of breathing and in number of conditions this had important implications.

Breathing alters according to the state of consciousness in healthy subjects, ventilation falls with the onset of sleep and is reduced during all phases of sleep compared with waking levels.1(1)

Alterations in the pattern of breathing occur including periodic breathing Apnoea, Hypopnoea with gradual progression from Stage I, II (18% decreased) to REM sleep.
REM sleep related alterations in ventilation is more severe in respiratory disordered patients, who had already some Hypoxemia, COPD is of best example.

**As patients with COPD have**

I) A higher physiological dead space than normal subjects and therefore the rapid shallow breathing that occurred during sleep will produce an even greater decrease in Alveolar ventilation.\(^{(2)}\)

II) Hyperinflation of lungs in COPD patients leads to more dependency upon Intercostal and Accessory muscles activity than normal individuals, but during REM sleep these Intercostal and Accessory Muscles are Flaccid, hence Diaphragm alone is unable to meet the demand.

III) Decrease FRC, V/Q Mismatch during sleep in COPD patients.\(^{(3,4,5)}\) OSAHS is SRBD of which shows oxygen desaturation during REM sleep.

If both OSAHS and COPD coexist in an individual, REM sleep related oxygen desaturation is more severe and as a result leads to many Cardiac, Non-Cardiac complications like Fluid Retention.\(^{(6)}\) polycythaemia.\(^{(7)}\) Cardiac arrhythmia.\(^{(8)}\) Pulmonary Hypertension.\(^{(9)}\) Hence, their early diagnosis and management is necessary.

All above changes should be well identified by doing Polysomnography (PSG) in COPD patients, where we know the (a) Impact of sleep on COPD, (b) Influence of COPD on sleep pattern.\(^{(10,11)}\)

Several studies demonstrate PaO\(_2\) decreased during sleep in COPD patients.\(^{(12,13)}\) mainly during REM sleep.\(^{(14,15)}\) being this decrease more pronounced during the Tonic phase of this sleep stage.\(^{(16)}\) It is natural to believe that Hypoxemia is responsible for sleep fragmentation in COPD and this Hypothesis is supported by experimental studies.\(^{(2)}\) and in diffuse parenchymal pulmonary disease patients.\(^{(17,18)}\)

Another Hypothesis to explain sleep fragmentation is the Hypocapnia, demonstrated in some studies.\(^{(19,20)}\) subjects wake up with PaCO\(_2\) close to 55 mmHg during NREM sleep and 60 mmHg during REM sleep. Paradoxically predominantly Pneumosoroma patients, usually Non-Hypocapnic and Non-Hypoxemic are the ones who present more sleep fragmentation suggesting that yet another factor such as increased ventilatory effort is responsible for awakening.\(^{(21)}\) In addition, more Hypoxemic patients individually present a better sleep pattern, although they can evolve to Cardiac arrhythmia.\(^{(22)}\) Right Ventricular Hypertrophy,\(^{(23)}\) polycythaemia.\(^{(24)}\) and Pulmonary Hypertension.\(^{(25)}\) the correlation between ventilatory functional test of COPD patients and PSG variables are still little studied.

This study as the objective of examining possible correlations among spirometric and polysomnographic variables in COPD patients.

**AIMS AND OBJECTIVES**

There is paucity of literature regarding sleep in COPD patients, particularly use of sleep studies in these patients. No direct study is seen in Indian population. As studies already done were giving variable prevalence and conclusions, we have conducted this study with the objective of,

a) Impact of sleep in COPD patients.

b) Influence of COPD on sleep pattern (Quality of sleep).

c) Overlap syndrome–prevalence.

**Study Design**

This is a prospective, analytical, transversal type of study.

**MATERIAL AND METHODS**

Patients who attended Outpatient Department of Pulmonary Medicine, Telangana Govt. General and Chest Hospital, OMC, Hyderabad, between May 2012 and April 2014 with clinical features s/o COPD were admitted, thoroughly examined and evaluated with routine investigations and specific investigations like sputum for AFB, Gram’s stain, PFT, Chest X-ray to establish the diagnosis of COPD. (GOLD guidelines).

All these patients (n=25), filled the COPD questionnaire along with Sleep Symptoms Questionnaire and Epworth Sleep Scoring (ESS).

These patients are advised to undergo whole night PSG after informed consent.

This study was considered as minimum risk in human research and was approved by the Ethics Committee of the University.

**PROCEDURE**

**Inclusion Criterion**

1. Patients having SOB, cough, history of smoking.
2. FEV\(_1\)/FVC<0.7 with irreversibility.
3. Snoring +ve and nocturnal sleep disturbance with or without EDS.

**Exclusion Criterion**

1. Asthma.
2. Other respiratory diseases.
3. Cor-pulmonale.
4. Patients who slept less than 4 hours.

Among these 25 selected patients of COPD with various degree of severity, one patient was excluded as he slept less than 4 hours.

These 24 patients underwent anthropometry in which weight and height were measured to the nearest 500 gms and 1 cm respectively and the BMI was calculated based on the formula. BMI = weight (kgs)/height\(^2\) (cm\(^2\)).

Neck circumference (cms) was measured at the level of cricothyroid membrane. Patients have reported to the sleep laboratory of the Department of Pulmonary Medicine at 8 PM on the day of testing.

All these patients underwent whole night Polysomnography. PSG is done by 54 multi-channel ALICE-5 system with whole night video recording. The entire testing process was done under the supervision of trained and capable technicians following the specifications and the criterion established by the R and K system.

**In PSG the following Montages were used**

1. EEG–7 sites–for staging of sleep (Gold coated cup electrodes are used).
2. EOG–1 cm upwards and outwards from the right outer canthus.
3. 1 cm downwards and outwards from the left outer canthus.
4. CEMG–2 sites—one above and one below to the chin for muscle tone measurement and reorganization of Bruxism.
4. LEMG–4 sites–two on each side–over shin and lower tibia–for the measurement of muscle tone and identification of PLMI.
5. Abdominal and thoracic belts – for identification of efforts.
6. Nasal Cannula – For the measurement of air flow.
7. Positional sensor – For posture of patient.
8. Pulsoximeter – for SpO₂ measurement.
9. EKG – For cardiac monitoring.
10. Video monitoring.

Patient was requested to sleep around 10 PM. The recording of sleep study was started after ensuring the impedance of the electrodes was set to zero. Raw data is analysed manually. Sleep stage is scores as per R and K rules. Respiratory events are scored according to revised AASM task force recommendations.

RESULTS
The 24 patients with COPD were divided into two groups
Group 1: 6 patients out of 24 (25%) showed significant OSAHS (AHI >5/hr, i.e. overlap syndrome); Group 2: Rest of the 18 patients (75%) showed no significant OSAHS. Among these 18 patients, 12 are having asymptomatic hypopneas (Clinically insignificant SDB), 6 are not significant OSAHS. Among these 2 groups, mean values of 25 variables were tabulated, compared and analysed by using Window Stat Version 8.5 from IndoStat services.

Among the 25 variables, 11 were significantly different between 2 groups.
1) Epworth score, which is indicative of excessive daytime sleepiness, well correlating with AHI, i.e. high (>10) in Group 1, less in Group 2.
2) Neck circumference and body mass index were high among Group 1, low in Group 2 patients according to student T test (p= 0.009 and p= 0.002) and Mann Whitney (p= 0.002 and p= 0.001).

It is suggesting that clinical features (e.g., EDS and smoking) and anthropometric findings (e.g., NC and BMI) are useful to screen OSAHS.
3) Duration of NREM sleep is low in Group 1 patients according to T test (p=0.001) and Mann Whitney (p=0.000) and TAI more in Group 1 according to T test (p=0.014) and Mann Whitney (p=0.000).
4) TAI (Total Arousal Index) with oxygen drop/AHI (Apnoea Hypopnoea Index) and OA are high in Group 1 patients. According to T test (p=0.000) and Mann Whitney (p=0.000), the inference is that obstructive apnoea is causing more number of arousals with significant oxygen drop among this group, which results in complications earlier than Group 2.
5) Desaturation with NREM sleep: TAI is more pronounced in Group 1 patients according to T test (P=0.005 and 0.014) and Mann Whitney (p=0.0052 and 0.020).

DISCUSSION
Though there is no common pathological link between OSAHS and COPD, their coexistence is not uncommon. The patients with clinical features suggesting OSAHS should be evaluated with Epworth score and overnight PSG for early detection of OSAHS.

Hence, sleep study in COPD patients even though not indicated in all COPD patients but should be done at least in high risk patients to avoid premature cardiac and non-cardiac complications secondary to OSAHS.

Limitations of Study
Arterial blood gas analysis not done, oesophageal pressure monitoring is not done. Sample size too small to know the prevalence of overlap syndrome among COPD patients.

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Table 1
CONCLUSIONS
There is positive correlation between neck circumference, BMI and Epworth score with AH1 (Apnoea-Hypopnoea Index).

There is positive correlation between frequency of arousals, low sleep efficiency, REM sleep with O2 drop and nor snoring associated arousals with AH1.

There is negative correlation between severity of COPD, duration of COPD and smoking with AH1.

OSAHS in COPD patients (Overlap Syndrome) is incidental and does not share the pathology of two diseases (COPD and OSA).

Sleep changes like reduced REM sleep, larger oxygen saturation drop during REM sleep, frequent arousals, frequent changing of sleep stages are observed in COPD patients, but more significantly seen in overlap syndrome.

Independent risk factors for OSAHS like high BMI, more neck circumference, facial abnormalities and symptoms like excessive day time sleepiness should be looked for diagnosing overlap syndrome in COPD patients.

Clinical Implication
Though there is no common pathological link between OSAHS and COPD, their core existence will not be uncommon. With various clinical features, these suspicious clinical features should be evaluated with Epworth score and thorough investigation with overnight PSG for early detection of OSAHS.

Hence, sleep studies in COPD patients even though not indicated in all COPD patients, but should be done at least in high risk group patients to avoid cardiac and non-cardiac premature complications.

REFERENCES