Short burst oxygen therapy after activities of daily living in the home in chronic obstructive pulmonary disease

S J Quantrill, R White, A Crawford, J S Barry, S Batra, P Whyte, C M Roberts

Background: Short burst oxygen therapy (SBOT) is widely prescribed in the UK with little evidence of benefit. A study was performed to examine whether SBOT benefits patients when undertaking normal activities at home among those who already use it.

Methods: Twenty-two patients with chronic obstructive pulmonary disease (COPD) were included in the study. All regularly used SBOT at home and claimed that it helps them. Each patient chose two daily living activities for which they used SBOT for relief of breathlessness. Patients were then randomised to use either an air or oxygen gas cylinder. At least 1.5 min later the same activity was performed using the other gas cylinder. The same process was then repeated for the second chosen activity. The main endpoints were subjective and objective times to recovery, analysed for each activity separately or taking the average over the two activities. A paired statistical analysis was performed.

Results: All patients used SBOT with nasal prongs after exercise. Using the average recovery time over two activities for each patient, the mean objective recovery time was 38 s lower (95% CI – 81 to +5) using oxygen and the mean subjective recovery time was 34 s lower (95% CI – 69 to +2). Five patients were correctly able to distinguish oxygen from air after both activities and there was a suggestion that their recovery times were shorter than those who did not correctly identify the gases (91 s vs 20 s using objective recovery times, and 80 s vs 22 s using subjective recovery times), although this was a subgroup analysis based on only five patients with non-significant results.

Conclusions: There is some evidence that SBOT shortens recovery time after activities of daily living in a selected group of patients with COPD, but the effect is small. There appears to be a subgroup of patients who may benefit to a much greater degree.

METHODS

Subjects
Thirty-nine patients with COPD defined by NICE criteria who regularly used SBOT at home were identified from the Waltham Forest general practice prescribing database and screened using a telephone questionnaire to ascertain suitability for the study.
B’ and were blinded to both patients and researchers. Patients performed each activity until they felt the need to stop and use oxygen. Subsequent tests were only performed when the patient had completely recovered from the previous activity, both subjectively and objectively. Pulse oximetry (Minolta Pulsox 3iA, Sunrise Medical, UK) was monitored throughout.

Endpoint measures were subjective and objective recovery times. Subjective recovery was defined as the point at which each patient stated their breathing had returned to normal. Objective recovery was defined as occurring when the oxygen saturation had returned to within 2% and heart rate to within 5 beats/min of pre-activity values. Breathlessness was measured using a 10 cm visual analogue score (VAS) with the end points of “not breathless at all” and “the most breathless I have ever been” at the start and end of each activity and at the point of subjective recovery. After each activity had been repeated, patients were asked if one cylinder enabled them to recover quicker than the other. If they did notice a difference, they were asked if the less effective cylinder was “better than nothing at all”.

Statistical analysis
The recovery time for each patient was taken as the average time for the two activities undertaken. Recovery times tended to have a skewed distribution (long tail to the right), so both the mean and median values are presented. However, the differences in recovery times (oxygen minus air) tended to be normally distributed (as determined by a probability plot) so paired t tests were used. Data were analysed using SPSS. The sample size was originally estimated to be 20 patients based on differences in VAS measurements for breathlessness to detect a difference of 8 mm with 90% power. However, VAS measurements would only have been useful for patients using SBOT after exercise, and during the course of the study we discovered that all patients used it after exercise. We therefore considered recovery times to be more appropriate, but it was difficult to perform a power calculation at the outset when the range of recovery times and expected differences between the groups was not known and could not be anticipated from existing data.

RESULTS
Patient characteristics
Twenty-two patients (14 men, mean age 72 years, range 56–86) were deemed suitable and agreed to take part. Reasons for exclusion among the other 17 patients included comorbidity (n = 6), SBOT not used in relation to exertion (n = 6), current exacerbation (n = 2), too unwell (n = 1), away from home (n = 1), and SBOT not yet prescribed (n = 1). The characteristics of patients included in the study are shown in table 1.

Seventeen patients had moderate or severe COPD as measured by spirometry but, although forced expiratory volume in 1 s (FEV1) in the remaining five patients was >50% predicted, carbon monoxide transfer factor (TlCO) in four of these five patients was <42% predicted indicating significant emphysema (unmeasured in the other). Most desaturated on exercise. Eleven patients (50%) used LTOT via a concentrator. All patients stated that they breathed oxygen via nasal prongs and none used masks. Flow rates were 2 l/min in 18 patients and 4 l/min in 4. In addition, all used their oxygen after activity and none before activity. Activities selected by patients for use of oxygen included walking upstairs (n = 20), walking around the house or garden (n = 13), vacuum cleaning (n = 8), walking uphill (n = 1), kitchen work (n = 1) and sweeping up (n = 1). Activity time and change in VAS after activity was not significantly different between oxygen and air.

Duration of activity
Because the same activity was performed twice (once with oxygen and once with air), we checked that the length of activity was similar when using oxygen or air in case this could explain the differences in recovery times. There was no evidence that activity length differed. During the first activity the mean difference in the length of activity (oxygen minus air) was –8 s (p = 0.64, paired t test), and during the second chosen activity the mean difference was 11 s (p = 0.60).

Recovery time
Subjective and objective times to recovery with oxygen and air are shown in table 2. The table shows the effect of oxygen therapy for each activity separately and the mean of the two activities. There was no difference in recovery times with oxygen compared with air. There was, however, a tendency for the effect on objective recovery times to be greater during the first activity (table 2). This may not be surprising given that, by the time the patient performed the second activity, he/she would already have performed the first activity twice.

Ability to distinguish oxygen from air
Five patients (22.7%) were correctly able to distinguish the oxygen from air on both occasions, five were able to identify the oxygen after one activity only, and 10 could not identify the oxygen on either occasion. Seven patients thought that, even though they had identified one gas as not being oxygen, it was better than nothing at all. Data were not available for two patients. A subgroup analysis was performed on the five patients who correctly distinguished oxygen from air on both occasions and the results were compared against those who could not. This subgroup of five correct identifiers had shorter subjective and objective recovery times using oxygen compared with air (80 s and 91 s, respectively, vs 22 s and 20 s for incorrect identifiers), although these results did not reach statistical significance (fig 1).

There were no significant differences in age, % predicted FEV1, baseline oxygen saturation or mean lowest desaturation between the two subgroups. There was no significant difference in either objective recovery time (p = 0.77) or subjective recovery time (p = 0.13) for those patients on LTOT compared with those who were not.

DISCUSSION
The data in this study suggest that SBOT shortens recovery time after activities of daily living in a highly selected group of patients with COPD. However, the beneficial effect is small overall. The shortening of recovery times did not reach statistical significance, but this is likely to be due to the relatively small number of subjects. One of the limitations of the study was that sample size was powered on a different outcome measure (VAS score) which was subsequently found to be inappropriate for this particular study. To show a difference in recovery times of about 35 s would require at least 50 patients (paired analysis), more than twice as many as the current study.

Table 1 Characteristics of subjects (n = 22)

<table>
<thead>
<tr>
<th>Characteristic</th>
<th>Mean (SD)</th>
<th>Range</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age (years)</td>
<td>72.0 (7.3)</td>
<td>56–86</td>
</tr>
<tr>
<td>FEV1 (litres)</td>
<td>0.87 (0.38)</td>
<td>0.40–1.69</td>
</tr>
<tr>
<td>FEV1 (% predicted)</td>
<td>38.0 (16.1)</td>
<td>17–74</td>
</tr>
<tr>
<td>SaO2 (%) resting, on air</td>
<td>93.1 (3.8)</td>
<td>82–98</td>
</tr>
<tr>
<td>Desaturation with activity (%)</td>
<td>–7.5 –25 to –0.5</td>
<td></td>
</tr>
<tr>
<td>FEV1, forced expiratory volume in 1 s; SaO2, oxygen saturation.</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>
Patients who correctly distinguished oxygen from air both after both sets of activities, a finding which is even more striking considering that all our patients had claimed that SBOT alleviated their breathlessness. Interestingly, all our patients used oxygen after the activity and none before, suggesting that they do not perceive benefit from pre-oxygenating. This concurs with the results of studies in which patients were administered oxygen before exercise.\textsuperscript{13,15,16} The difficult question of when a change becomes clinically significant in terms of relief of breathlessness, shortening of recovery time and improvement of exercise capacity can be illustrated by the wide variation in cut-off points used in studies of SBOT and ambulatory oxygen.\textsuperscript{12,15,17–22} Although the subjective recovery time was 34 s or 15\% shorter with oxygen than with compressed air, the clinical importance of this must be debatable. McDonald et al.\textsuperscript{23} in a study of ambulatory oxygen in COPD, also make the point that the increase in 6 minute walking distance of <20 m in their laboratory tests, while of “statistical significance”, did not translate into any useful improvement in day-to-day functioning and the prescription of oxygen on the basis of these tests could not therefore be justified. Perhaps we should move away from a statistical model of what is significant to a more patient-centred view, reflecting the change in philosophy of the UK healthcare system as a whole over recent years.

The limitations of our study are the relatively small numbers included which means that a statistically significant reduction in overall objective recovery time could have been missed, but there is still the question of whether this would be of any clinical value. Our study also included a mixture of patients, half of whom were receiving LTOT. However, this simply reflects actual clinical practice where SBOT is prescribed. Some patients with a concentrator used this at times instead of a cylinder (although not in the study); however, they were all known to use their oxygen cylinder regularly as this is how they were originally identified from prescribing records. All patients used nasal prongs, which is a possible confounder as some who are predominantly mouth breathers might benefit more from oxygen via a mask. We did not study the effect of SBOT during exacerbations which is another important area for research, particularly as oxygen usage may increase at these exacerbations which is another important area for research.

Prior studies have shown similar small improvements with SBOT or no benefit overall for the alleviation of breathlessness on exertion in COPD.\textsuperscript{11,13,15–17} There may, however, be concerns that these trials have missed a subgroup of patients for whom SBOT is beneficial. In our study only 5 of 20 patients questioned were correctly able to distinguish oxygen from air after both sets of activities, a finding which is even more striking considering that all our patients had claimed that SBOT helped them in undertaking these activities. However, there was a much greater difference in recovery time for the 5 patients who correctly distinguished oxygen from air both times (80 s for subjective recovery and 91 s for objective recovery) and, although not statistically significant, this could represent a subgroup for which oxygen is beneficial.

Previous studies have all been conducted in the artificial setting of the lung function laboratory or hospital environment and have studied a mixture of patients who may or may not be regular oxygen users.\textsuperscript{12,15,17–22} Our study is unique as we aimed to overcome these problems by specifically targeting the subgroup that regularly use SBOT, testing patients at home undertaking activities of daily living with which they are familiar. Furthermore, it was a specific requirement that patients claimed SBOT alleviated their breathlessness. Interestingly, all our patients used oxygen after the activity and none before, suggesting that they do not perceive benefit from pre-oxygenating. This concurs with the results of studies in which patients were administered oxygen before exercise.\textsuperscript{13,15,16}

\textbf{Table 2} Comparison of the effect of oxygen therapy and air on recovery times

<table>
<thead>
<tr>
<th>Activity 1</th>
<th>Recovery time with oxygen [s]</th>
<th>Recovery time with air [s]</th>
<th>Mean difference (oxygen minus air) (95% CI)</th>
<th>p Value</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Median</td>
<td>IQR</td>
<td>Mean</td>
<td>Median</td>
</tr>
<tr>
<td>Objective</td>
<td>65</td>
<td>55</td>
<td>90</td>
<td>120</td>
</tr>
<tr>
<td>Subjective</td>
<td>170</td>
<td>98</td>
<td>169</td>
<td>165</td>
</tr>
<tr>
<td>Objective</td>
<td>58</td>
<td>78</td>
<td>104</td>
<td>110</td>
</tr>
<tr>
<td>Subjective</td>
<td>180</td>
<td>120</td>
<td>207</td>
<td>230</td>
</tr>
</tbody>
</table>

Mean of activities 1 and 2

<table>
<thead>
<tr>
<th>Activity 2</th>
<th>Recovery time with oxygen [s]</th>
<th>Recovery time with air [s]</th>
<th>Mean difference (oxygen minus air) (95% CI)</th>
<th>p Value</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Median</td>
<td>IQR</td>
<td>Mean</td>
<td>Median</td>
</tr>
<tr>
<td>Objective</td>
<td>75</td>
<td>82</td>
<td>97</td>
<td>110</td>
</tr>
<tr>
<td>Subjective</td>
<td>186</td>
<td>110</td>
<td>186</td>
<td>240</td>
</tr>
</tbody>
</table>

IQR, interquartile range.

\textbf{Figure 1} (A) Subjective time to recovery and (B) objective time to recovery in subgroup who correctly identified oxygen after both activities (n = 5) and those who did not (n = 17).
times and also be prescribed for patients on discharge from hospital.

In conclusion, we have shown that SBOT does appear to shorten recovery time overall after activities of daily living in this highly selected patient group, but it is debatable whether this is of clinical significance. However, there appears to be a small subgroup of patients who are able to perceive this benefit, and a larger study is needed to verify this and determine how they may be identified.

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REFERENCES
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