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## Should pulse oximetry be included in GPs' assessment of patients with obstructive lung disease?

Lene G. Dalbak<sup>a,b</sup>, Jørund Straand<sup>b</sup>, and Hasse Melbye<sup>a</sup>

<sup>a</sup>General Practice Research Unit, Department of Community Medicine, University of Tromsø, Norway; <sup>b</sup>Department of General Practice, Institute of Health and Society, University of Oslo, Norway

#### **ABSTRACT**

Objective: To explore the associations between decreased pulse oximetry values ( $SpO_2$ ) and clinical, laboratory, and demographic variables in general practice patients diagnosed with asthma or chronic obstructive pulmonary disease (COPD), including those with both COPD and asthma in combination.

Design/setting: A cross-sectional study in seven Norwegian general practices of patients aged 40 years or over who were diagnosed by their general practitioner (GP) with asthma and/or COPD. The patients were examined during a stable phase of their disease. Patients diagnosed with COPD (including those with combined COPD/asthma) and those diagnosed with asthma only were analysed separately.

Main outcome measures: Decreased SpO<sub>2</sub> values ( $\leq$  95% and  $\leq$  92%).

Results: Of 372 patients included (mean age 61.5 years, 62% women), 82 (22.0%) had SpO $_2 \le 95\%$ , of which 11 had SpO $_2 \le 92\%$ . In both asthma and COPD patients, SpO $_2 \le 95\%$  was significantly associated with reduced lung function (spirometry), a diagnosis of coronary heart disease and older age ( $\ge 65$  years). In the COPD group, haemoglobin above normal was associated with SpO $_2 \le 95\%$ . These associations were confirmed by multivariable logistic regression, where FEV $_1\%$  predicted < 50 was the strongest predictor of SpO $_2 \le 95\%$  (odds ratio 6.8, 95% confidence interval 2.8–16.4). Conclusion. Pulse oximetry represents a useful diagnostic adjunct for assessing the severity of obstructive pulmonary disease. Decreased pulse oximetry values in stable-phase patients with asthma and/or COPD should prompt the GP to consider revising the diagnosis and treatment and to look for co-morbidities.

#### **KEY POINTS**

- Despite its common use in general practice, the diagnostic benefits of pulse oximetry remain to be established.
- Decreased pulse oximetry values are associated with both reduced lung function (spirometry) and with a diagnosis of coronary heart disease.
- Decreased pulse oximetry values may reflect suboptimal treatment and/or undiagnosed comorbidity.
- Pulse oximetry may therefore be a useful measure in the follow-up of asthma and COPD patients in general practice.

#### ARTICLE HISTORY

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#### **KEYWORDS**

Asthma, chronic obstructive pulmonary disease, comorbidity, general practice, Norway, oximetry

#### Introduction

Pulse oximetry is a non-invasive, simple, inexpensive, and rapid test to estimate haemoglobin oxygen saturation. Hand-held pulse oximeters have become available in general practice and have been reported to be useful diagnostic tools for the assessment of chronic obstructive pulmonary disease (COPD) during both stable phase [1,2] and exacerbations [1–3], and in particular for confirming the need for oxygen therapy [1,4,5]. Pulse oximetry may also be helpful in assessing the severity of asthma exacerbations [1,6]. An oxygen

saturation  $(SpO_2) \le 92\%$  indicates hypoxaemia, but values between 93% and 95% are lower than normal [1,7,8]. In COPD patients,  $SpO_2$  values  $\le 95\%$  predict hypoxia during exercise and air travel [8,9]. Pulse oximetry is regarded as a valid screening test for systemic hypoxia [10], but current guidelines do not inform general practitioners (GPs) how to deal with  $SpO_2$  values between 93% and 95%. More evidence of the usefulness of pulse oximetry is urgently needed, because its use is rapidly increasing in general practice [1,11,12].

GPs implementing pulse oximetry should be able to explain to patients the implications of a lower-thannormal SpO<sub>2</sub> value. The aim of this study was to describe conditions and patient characteristics associated with decreased pulse oximetry values in primary care patients with stable obstructive lung disease.

#### Material and methods

The study was carried out at seven Norwegian GP group practices. The practices were not randomly selected, but were chosen based on the availability of spirometry results from the previous five years and the type of electronic medical record system used. Of 43 241 patients listed at the seven practices, 18931 were adults aged 40 or over, among whom 1784 were identified from the medical records as being diagnosed by the GP with asthma and/or COPD within the previous five years. For reasons of feasibility (e.g. extra workload for the participating GPs), each group practice decided the proportion of registered patients with asthma or COPD that they would invite to participate in the study. A total of 1111 patients were invited to participate, and in all the practices these were randomly selected in alphabetical order among the eligible patients. Invitations were sent by surface mail without additional reminders. Participation required completion of a guestionnaire, a meeting during a stable phase of their disease for a clinical examination including spirometry (i.e. baseline assessment), plus examinations during any exacerbations in the subsequent year. This report is based on the baseline examinations, which took place between March 2009 and March 2010. The participants were instructed not to take their regular respiratory medication on the day of the examination.

#### **Measurements and instruments**

The GPs recorded comorbidities including cardiovascular diseases on a computerized questionnaire linked to the patient's medical record. On a separate questionnaire, patients recorded their smoking habits. The patient's height and weight were recorded to calculate their body mass index (BMI).

Oxygen saturation was measured with a digital handheld pulse oximeter, Onyx II model 0550 (Nonin Medical, Inc., Plymouth, MN, USA). The highest value obtained from three measurements was recorded.

The HemoCue Haemoglobin system (Quest Diagnostics, Madison NJ, USA) was used for haemoglobin measurements. The thresholds for raised values were based on the reference values used at the University Hospital of North Norway. The upper normal limit was

16.0 g/dL for women and 17.0 g/dL for men. C-reactive protein (CRP) was analysed using an Afinion AS100 Analyser (Axis-Shield, Oslo, Norway), Orion Quickread CRP (Orion Diagnostica, Espoo, Finland) or ABX Micros CRP (Horbia ABX SAS, Montpellier, France), all of which could display values down to 8 mg/L.

Spirometry was carried out after the pulse oximetry test, following European Respiratory Society/American Thoracic Society guidelines [13], using a Spirare SPS310 spirometer (Diagnostica AS, Oslo, Norway). During spirometry, the patients were seated and a nose clip was not used. Post-bronchodilator spirometry was carried out 20 min after inhalation of 0.4 mg salbutamol. The post-bronchodilator forced expiratory volume in one second (FEV<sub>1</sub>) and forced vital capacity (FVC) were used in the analyses. Norwegian reference values for spirometry were applied [14].

#### Statistical analyses

Two thresholds of abnormal SpO<sub>2</sub> values were used as outcome measures,  $\leq$  95% and  $\leq$  92%. The frequencies of reduced SpO<sub>2</sub> associated with each patient characteristic (Table I) were analysed separately in asthma and COPD patients. The division into these two groups was based on the GPs' diagnosis coded as R96 (COPD) and R95 (asthma) according to the International Classification for Primary Care [15]. Patients who were given both diagnoses were allocated to the COPD group. Continuous variables (age, BMI, FEV<sub>1</sub>, and CRP) were categorized. BMI was categorized as underweight  $(< 20 \text{ kg/m}^2)$ , normal weight/overweight  $(20-30 \text{ kg/m}^2)$ , and obese (> 30 kg/m<sup>2</sup>) [16]. The FEV<sub>1</sub>% predicted was categorized as severely reduced (<50%), moderately reduced (50–80%), and normal ( $\geq$  80%), in line with the Global Initiative for Chronic Obstructive Lung Disease grading of COPD [5]. CRP values were dichotomized into  $\geq$ 8 mg/mL and <8 mg/L. The significance of associations between reduced SpO2 and patient characteristics was analysed using the chi-square test. Age, sex, and variables significantly associated with a decreased SpO<sub>2</sub> (p < 0.05) in univariable analysis were entered into a multivariable logistic regression  $SpO_2 \le 95\%$  as the outcome variable. The multivariable analysis was also performed without categorizing the continuous variables. SPSS version 18.0 (IBM, Armonk, NY, USA) was used in all analyses.

#### **Results**

Of the 1111 patients invited to participate, 380 (34.2%) accepted and attended the baseline examination. Eight patients were excluded from analysis, two because they



Table 1. Frequency of  $SpO_2$  values  $\leq 95\%$  in 372 patients aged 40 years or more diagnosed with asthma or COPD.

	COPD			Asthma		
		$SpO_2 \le 95\%$			$SpO_2 \le 95\%$	
Patient characteristics	Patients, n	n (%)	p-value	Patients, n	n (%)	p-value
All	166	51 (31.7)		206	31 (15.1)	
Age (years)						
≥ 65	96	37 (38.5)	0.01	55	14	0.01
<65	70	14 (20.0)		151	(25.5) 17 (11.3)	
Sex						
Men	67	21 (31.3)	0.9	75	13	0.5
Women	99	30 (30.3)		131	(17.3) 18 (13.7)	
Smoking habits						
Never	22	5 (22.7)	0.7	72	13	0.7
Previous	90	28 (31.1)		83	(18.1)	
Current	54	18 (33.3)		51	11 (13.3) 7 (13.7)	
Coronary heart disease						
Yes	42	18 (42.9)	0.05	22	7 (31.8)	0.02
No	124	33 (26.6)		184	24 (13.1)	
Other cardiovascular disease						
Yes	49	18 (36.7)	0.3	28	4 (14.3)	0.9
No	117	33 (28.2)		178	27 (15.2)	
FEV₁% predicted						
>80	37	6 (16.2)	0.01	122	10 (8.2)	< 0.01
50-80	84	24 (28.6)		78		
< 50	45	21 (46.7)		6	17 (21.8) 4 (66.7)	
BMI (kg/m <sup>2</sup> ) <sup>a</sup>						
< 20	17	9 (52.9)	0.06	4	0 (0.0)	0.3
20-30	103	27 (6.2)		142	24	
≥ 30	34	13 (38.2)		51	(16.9) 5 (9.8)	
CRP (mg/L)						
≥8	29	12 (41.4)	0.2	29	6 (20.7)	0.4
<8	137	39 (28.5)		177	25 (14.1)	
Haemoglobin above normal <sup>b</sup>						
Yes	16	10 (62.5)	< 0.01	15	2 (13.3)	0.9
No	148	40 (27.0)		190	29 (15.3)	

Notes: p-values denote the significance of differences between categories of patient characteristics assessed by chi-square test. <sup>a</sup>Value missing for 21 patients. <sup>b</sup>Hb above 16.0 g/dL in women and above 17.0 g/dL in men; Hb values missing in three patients.

were undergoing an acute exacerbation, two because they did not complete post-bronchodilator spirometry, and four because they did not undergo pulse oximetry. For the 372 included patients, the mean age was 61.5 years and 62% were women. A diagnosis of COPD only was registered in 74 patients, asthma only in 206 patients, whereas 92 patients were registered with both diagnoses. Median SpO<sub>2</sub> was 97%. In total, 82 patients (22.0%) had  $SpO_2$  values  $\leq 95\%$ ; 11 had an  $SpO_2 \le 92\%$ ; of these, 10 were COPD patients and two were on long-term oxygen therapy.

The frequency of both  $SpO_2 \le 92\%$  and  $SpO_2 \le 95\%$ increased with decreasing levels of FEV<sub>1</sub>% predicted (Figure 1). When patients with COPD and asthma were analysed separately, the frequency of  $SpO_2 \le 95\%$ increased significantly with increasing patient age, decreasing FEV<sub>1</sub>% predicted, and the presence of coronary heart disease (Table 1). The association of coronary heart disease with decreasing  $SpO_2$  values (p < 0.001) is illustrated in Figure 2. Haemoglobin above the threshold value was associated with  $SpO_2 \le 95\%$  only in the COPD group (Table 1). Predictors of SpO<sub>2</sub> < 92% were evaluated only in the COPD group, for which  $SpO_2 \le 92\%$  was associated with decreasing levels of FEV<sub>1</sub>% predicted, BMI < 20, and raised haemoglobin values (Table 2).

Multivariable logistic regression revealed that age, comorbidity with coronary heart disease, decreased FEV<sub>1</sub>% predicted, and haemoglobin above normal were all independent predictors of  $SpO_2 \le 95\%$  (Table 3). The strongest predictor of SpO<sub>2</sub> ≤ 95% was FEV<sub>1</sub>% predicted < 50%, recording an odds ratio of 6.8 (Table 3). Using FEV<sub>1</sub>% predicted as a continuous variable gave similar results and did not improve the model, with the Nagelkerke R<sup>2</sup> increasing only from 0.21 to 0.22. In the logistic regression analyses, there was no significant interaction effect between age and the other predictors. Multivariable analysis with  $SpO_2 < 92\%$  as outcome was not performed because of the low number of patients with such values (n = 11).

#### **Discussion**

#### **Principal findings**

SpO<sub>2</sub> ≤ 95% was strongly associated with reduced lung function, coronary heart disease, greater age, and abovenormal haemoglobin. Among the COPD patients,

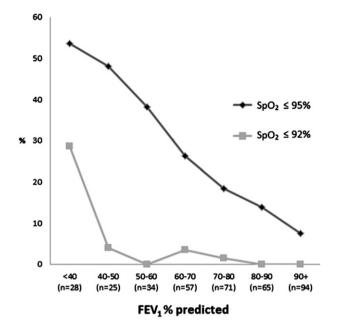


Figure 1. Frequency of reduced SpO<sub>2</sub> by FEV<sub>1</sub>% predicted in 372 adult patients in general practice with a diagnosis of asthma and/or COPD.

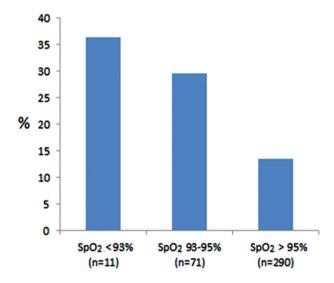


Figure 2. Frequency of coronary heart disease by pulse oximetry values in 372 adult patients in general practice with a diagnosis of asthma and/or COPD.

 $SpO_2 \le 92\%$  was also more frequently found in patients with BMI < 20.

#### Strengths and weaknesses

In clinical practice, asthma and COPD may sometimes be hard to differentiate [17]. The inclusion of patients diagnosed with either or both of these conditions may make our results relevant for the population of patients in primary care who have obstructive lung disease. More

Table 2. Frequency of  ${\rm SpO_2} \le 92\%$  by patient characteristics in 166 patients aged 40 years or over diagnosed with COPD by a GP.

		$SpO_2 < 92\%$	
Patient characteristics	Patients, n	n (%)	p-value
All	166	10 (6.0)	
Age (years)		` ,	
> 65	96	7 (7.3)	0.4
_ <65	70	3 (4.3)	
Sex		` '	
Men	67	3 (4.5)	0.5
Women	99	7 (7.1)	
Smoking habits			
Never	22	0	0.4
Previous	90	7 (7.8)	
Current	54	3 (5.6)	
Coronary heart disease			
Yes	42	4 (9.5)	0.3
No	124	6 (4.8)	
Other cardiovascular disease			
Yes	74	5(6.8)	0.7
No	92	5(5.4)	
FEV <sub>1</sub> % predicted			
> 80	37	0 (0)	
50–80	84	3 (3.6)	
< 50	45	7 (15.6)	< 0.01
BMI (kg/m²) <sup>a</sup>			
< 20	17	5 (29.4)	< 0.01
20–30	103	4 (3.9)	
≥ 30	34	0 (0.0)	
CRP (mg/L)			
≥8	29	3 (10.3)	0.3
<8	137	7 (5.1)	
Haemoglobin above normal <sup>b</sup>			
Yes	16	4 (25.0)	< 0.01
No	148	6 (4.1)	

Notes: aValue missing for 12 patients. bValue missing for two patients.

Table 3. Predictors of  $SpO_2 \le 95\%$  in 369 patients aged 40 years or over diagnosed in general practice with asthma and/or COPD.

Patient characteristics	Odds ratio (95% CI)	p-value
FEV <sub>1</sub> % predicted		
>80	Reference	0.002
50-80	2.9 (1.5-5.8)	< 0.001
< 50	6.8 (2.8–16.4)	
Coronary heart disease	2.0 (1.1–3.8)	0.03
Age	1.0 (1.0–1.1)	0.004
Sex male	0.8 (0.4–1.4)	0.43
Asthma-only diagnosis	1.1 (0.6–2.1)	0.75
Haemoglobin above normal	2.5 (1.0-6.0)	0.04

Notes: CI = confidence interval;  $SpO_2 = arterial$  oxygen saturation measured by pulse oximetry;  $FEV_1 = forced$  expiratory volume in one second.

than half the patients in the COPD group had also been diagnosed with asthma within the previous five years. COPD was most frequently the later diagnosis [17], and this may represent the trend to a change in diagnosis from asthma to COPD [18]. Asthma can develop into COPD [19], but the tendency to choose an asthma diagnosis even if COPD would be more appropriate may partly reflect the changed reimbursement regulations for respiratory medication introduced in Norway in 2006. At the time that this study was performed, the costs of inhaled corticosteroids and of inhaled corticosteroids

combined with long-acting  $\beta_2$ -agonists could generally only be reimbursed in patients with a diagnosis of asthma. Mixed diagnoses may also reflect consultations with more than one doctor during the five-year period.

In this study, current smoking was not associated with decreased pulse oximetry readings. The oxygen saturation in current smokers may have been overestimated, because the presence of carboxyhaemoglobin may falsely increase SpO<sub>2</sub> readings [20]. Pulse oximeters also have other limitations [7]. Oxygen saturation may be overestimated in patients with darkly pigmented skin and anaemia [7]. Nail polish, dirt, and artificial nails may cause falsely low readings [21]. Poor perfusion (cold digits) may also result in uncertain results [7]. Analysis of arterial blood obtained by arterial puncture remains the gold standard for measurement of oxygen saturation [7].

The low participation rate (34%) increases the risk of including a study sample that is not representative of the patients in primary care with asthma or COPD. It is likely that the most severely impaired COPD patients, who had been followed up closely in secondary care, were less interested in taking part in the study. The same would apply to patients at the other end of the spectrum, namely those who felt healthy. This is discussed in more detail in a previous report from the study [17]. Although the prevalence of low SpO<sub>2</sub> values in our population might have been affected by a somewhat skewed selection of participants, it is not likely that the associations between decreased SpO<sub>2</sub> values and lung function were substantially affected.

#### Findings in relation to other studies

The prevalence of  $SpO_2 \le 95\%$  (22%) in the participants in our study was considerably higher than that found in the general population in a Norwegian study of 6317 adults of similar age [22]. In that study, the overall prevalence of  $SpO_2 \le 95\%$  was 6.3%, and 15.5% among individuals with self-reported COPD [22]. In a retrospective analysis of 81 COPD patients attending a US university medical centre and tested for desaturation during exercise, the prevalence of SpO<sub>2</sub> < 95% was 46% [8]. In our study, 32% of the COPD patients had an  $SpO_2 \le 95\%$  (see Table 1).

We found greater age to be an independent predictor of  $SpO_2 \le 95\%$ . This was also the case in the Norwegian population-based study [22], and we know from previous studies that lower oxygen saturation can be found in the healthy elderly compared with that in younger healthy adults [23].

In a Dutch study of primary care patients, Schermer et al. [1] showed a positive association between FEV<sub>1</sub>% predicted < 50 and SpO<sub>2</sub> in COPD patients with acute exacerbation or worsening dyspnoea, whereas the association observed in patients with stable COPD did not reach significance. In the Norwegian populationbased study referred to above [22], both age  $\geq$  65 years and FEV<sub>1</sub>% predicted < 50 were strong predictors of  $SpO_2 \le 95\%$ , as in our study, but increased BMI was also an independent predictor. This latter association was not found in our study. A reason for this discrepancy may be that obesity is proportionally a more common reason for breathing problems in the general population than among patients diagnosed with asthma or COPD. In our study, BMI < 20 was strongly associated in COPD patients with  $SpO_2 \le 92\%$ . The increased occurrence of decreased oxygen saturation in underweight COPD patients fits with the increased COPD mortality that has been found in patients with very low BMI [16]. In a COPD population from primary care in the Netherlands [24], 11.7% had a BMI < 18.5, similar to the 11.5% in our COPD patients with BMI < 20.

In both univariable and multivariable analyses, coronary heart disease was a strong predictor of  $SpO_2 \le 95\%$  in both asthma and COPD patients. An association between coronary heart disease and decreased oxygen saturation was also found in a study from Spain, which concluded that pulse oximetry is useful in establishing the diagnosis and severity of heart failure in acute myocardial infarction [25]. Cardiovascular disease (including ischaemic heart disease, heart failure, atrial fibrillation, and hypertension) is probably both the most frequent and the most important comorbidity coexisting with COPD [18, -26], because they share similar aetiology (i.e. smoking) [27]. Cardiovascular comorbidity is probably underdiagnosed in COPD patients [28].

#### Implications for primary care practice

We could not find any particular recommendations in the guidelines for the follow-up of stable COPD and asthma patients with  $SpO_2$  values  $\leq 95\%$  [2,5,7]. Our study shows that patients with moderately decreased oxygen saturation have an increased risk of severely reduced lung function and comorbid coronary heart disease. Some may have received suboptimal pharmacological treatment. Although old age may sometimes be the only explanation, the increased risk of comorbidity and of suboptimal treatment in patients with SpO2 ≤ 95% indicate that these patients should be given special attention and followed up more closely than patients with normal oxygen saturation. The pulse oximeter may be a particularly helpful tool in primary care, because it is tolerable for the patients and is easy to use, and is thus acceptable within the time constraints of a busy practice.

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#### **Ethical approval**

Participating patients signed a written consent form and the study was approved by the Regional Committee for Medical and Health Research Ethics in North Norway.

#### **Declaration of interest**

The authors report no conflicts of interest. The authors alone are responsible for the content and writing of the paper.

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