

## SHORT REPORT

# Use of peak expiratory flow for assessing bronchodilator responsiveness

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### Abstract

Lung function in 1686 adult patients was measured before, and 15 minutes after, salbutamol inhalation. Bronchodilator responsiveness (BDR) was defined as a 12% improvement over baseline in either FEV<sub>1</sub> or FVC, along with an absolute volume increment of 200ml. Peak expiratory flow (PEF) change, both absolute and relative to baseline, was also calculated ( $\Delta$ PEF and  $\Delta$ PEF%, respectively). Change in PEF significantly correlated with change in FEV<sub>1</sub>. However,  $\Delta$ PEF and  $\Delta$ PEF% had poor discrimination in identifying BDR, with all specific cut-off values for  $\Delta$ PEF and  $\Delta$ PEF% having low or moderate sensitivity, specificity and predictive values.

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### Introduction

Spirometry is the recommended investigation both for diagnosis and categorisation of the severity of airflow limitation.<sup>1,2</sup> Objective measurement of bronchodilator responsiveness (BDR) is useful in the clinical evaluation of these patients. The BDR test is carried out by performing baseline spirometric evaluation, with repeat spirometry after administration of a short-acting bronchodilator, and noting absolute and relative increments in observed vital capacity (VC) and/or forced expiratory volume in the first second (FEV<sub>1</sub>).

However, spirometry is not widely available in primary care settings, and is often not performed routinely due to technical and logistic constraints. Peak expiratory flow (PEF) measurement is a simpler test in the assessment of obstructive airway disorders. The PEF instrument is cheap, portable and easy to operate and maintain. Guidelines on asthma suggest that either FEV<sub>1</sub> or PEF can be used to categorise patients into various grades of disease severity and control.<sup>3,4</sup> Many clinicians also assume a general parity between these two measurements, although we have recently shown that this may not be correct.<sup>5</sup>

Nevertheless, it would be helpful if PEF could be used as a surrogate for spirometry in BDR assessment in primary care settings. A few investigators have studied this previously in

both adults and children.<sup>6-9</sup> However, these studies were conducted on small numbers of selected patients, and variable criteria were used to define BDR, making interpretation and comparison difficult. Some data suggest that, although lack of responsiveness in PEF can be used to exclude BDR, it is of much less value as a diagnostic test.<sup>8,9</sup>

We therefore studied adult patients undergoing BDR testing to evaluate the performance of PEF measurement as an alternative to spirometry in diagnosing BDR.

### Patients and methods

Records of all consecutive adult patients (aged more than 15 years) undergoing BDR testing during a six-year period (1999 to 2004) were retrieved and analysed retrospectively. Sources of referral, reasons for performing spirometry, and other clinical details were not analysed. All subjects had performed spirometry on a dry rolling seal spirometer (Spiroflow; P K Morgan Ltd.; Kent, U.K.) followed by PEF estimation using Wright's peak flow meter. VC, FEV<sub>1</sub> and PEF were measured by experienced technicians using American Thoracic Society guidelines.<sup>10</sup> The recorded PEF was corrected for the non-linearity of the PEF meter using the equation: corrected PEF = 0.00075 PEF<sup>2</sup> + 0.585 PEF + 53.2. This value was used for all subsequent analyses.<sup>11</sup>

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A diagnosis of BDR based on standard spirometric criteria was considered the gold standard. A patient was considered to have BDR if there was a 12% improvement over baseline in either FEV<sub>1</sub> or FVC, along with an absolute volume increment of 200ml, 15 minutes after inhalation of 400 micrograms of salbutamol via a metered dose inhaler.<sup>1,2</sup> The change in PEF, both absolute and relative to baseline, was also calculated ( $\Delta$ PEF and  $\Delta$ PEF% respectively). The correlation between change in pre-bronchodilator and post-bronchodilator observations in PEF, and a corresponding change in FEV<sub>1</sub> or FVC was studied. Overall discrimination of both  $\Delta$ PEF and  $\Delta$ PEF% in identifying BDR was quantified as area under receiver operating characteristic (AUROC) curves. Performance of specific cut-off values of both  $\Delta$ PEF and  $\Delta$ PEF%, as well as various combinations of both together in correctly identifying subjects with BDR, was also assessed.

## Results

A total of 1686 records were analysed. There were 910 men (age 15-83 years) and 776 women (age 15-80 years). BDR was documented by standard spirometric criteria in 565 (33.5%) of these patients.

$\Delta$ PEF values showed statistically significant correlation with corresponding changes in FEV<sub>1</sub> (Pearson's correlation coefficient 0.365,  $p < 0.001$ ) and FVC (Pearson's correlation

coefficient 0.253,  $p < 0.001$ ).  $\Delta$ PEF% values correlated marginally better with corresponding changes in FEV<sub>1</sub> (Pearson's correlation coefficient 0.421,  $p < 0.001$ ) and FVC (Pearson's correlation coefficient 0.363,  $p < 0.001$ ) values relative to their respective baselines. However, the absolute values of the correlation coefficients were low. In addition, both  $\Delta$ PEF and  $\Delta$ PEF% were poor discriminators at predicting BDR, as AUROC values for both parameters were low (0.673, 95% confidence limits 0.646-0.701, and 0.720, 95% confidence limits 0.694-0.747, respectively).

Individual performance characteristics of specific  $\Delta$ PEF values (an increment of 20, 40, 60 or 80 L/min over baseline) and  $\Delta$ PEF% values (an increment of 10%, 15% or 20% over baseline) were also studied. All these cut-off values had low or moderate sensitivity, specificity and predictive values in identifying BDR (Table 1). None of these values appeared to possess clinically useful sensitivity, while a moderately high specificity was achieved only for  $\Delta$ PEF exceeding 80 L/min.

## Discussion

There are few data on the use of PEF to assess BDR. The present retrospective analysis is the largest study attempted on this subject. It was designed so that the results reflect an average clinician's day-to-day concerns at a primary health care level.

**Table 1. Performance characteristics of various cut-offs of post-bronchodilator increment in PEF when compared to standard spirometric criteria to define post-bronchodilator responsiveness.**

Change in PEF over baseline	Sensitivity	Specificity	Positive predictive value	Negative predictive value	Likelihood ratio (positive)
>10%	0.710	0.601	0.473	0.804	1.780
>15%	0.609	0.723	0.525	0.786	2.195
>20%	0.501	0.815	0.578	0.764	2.712
>20 L/min	0.727	0.491	0.419	0.782	1.431
>40 L/min	0.556	0.677	0.464	0.751	1.721
>60 L/min	0.342	0.806	0.470	0.708	1.756
>80 L/min	0.203	0.880	0.462	0.687	1.703
>10% and >20 L/min	0.690	0.616	0.476	0.798	1.799
>10% and >40 L/min	0.549	0.698	0.478	0.754	1.820
>10% and >60 L/min	0.342	0.806	0.470	0.708	1.757
>10% and >80 L/min	0.203	0.880	0.462	0.687	1.703
>15% and >20 L/min	0.603	0.726	0.526	0.784	2.204
>15% and >40 L/min	0.522	0.769	0.532	0.761	2.260
>15% and >60 L/min	0.329	0.829	0.492	0.710	1.922
>15% and >80 L/min	0.203	0.883	0.467	0.687	1.742
>20% and >20 L/min	0.501	0.815	0.578	0.764	2.712
>20% and >40 L/min	0.455	0.830	0.574	0.751	2.670
>20% and >60 L/min	0.304	0.856	0.516	0.709	2.120
>20% and >80 L/min	0.195	0.896	0.485	0.688	1.865

See reference 2 for details on spirometric criteria for bronchodilator responsiveness

Our study population was not limited to any particular disease subset, but instead constituted a large number of patients routinely referred to our Pulmonary Function Laboratory for BDR assessment. Consequently, our observations are relevant to the routine assessment of patients seen in primary care settings. Although we understand the limitations and bias of such a retrospective analysis, and we are hampered by the lack of complete clinical details for all patients, the large number of subjects studied still provides much more precise results compared to those achieved by previous investigators.<sup>6-9</sup> In addition, we based our analysis on PEF values corrected for non-linearity of the PEF meter. Therefore the results have a wider applicability in light of the new European standards introduced in 2004.

Our results show that PEF changes after bronchodilator inhalation correlate well with corresponding FEV<sub>1</sub> and FVC changes. However, it would appear that, irrespective of the criteria used to describe BDR based on PEF measurements, BDR defined this way is a poor discriminator in identifying true BDR based on traditional spirometry. The poor sensitivity and specificity obtained from most cut-off values studied would suggest a good deal of overlap in values between those patients who have BDR (as defined on spirometric criteria) and those who do not. Although it is difficult to compare our results with previous work due to differences in the subjects studied and the definition of BDR, the specificity and positive predictive value of PEF-based BDR in our study appear slightly lower than that reported by others.

In a study on 73 patients with asthma or COPD, the sensitivity and specificity of a 60 L/min increase in PEF in detecting a 9% or more increase in FEV<sub>1</sub> as a percentage of predicted value were 68% and 93%, respectively, with a positive predictive value of 87%.<sup>6</sup> In another study on 44 patients with suspected asthma, a >18% increase in the PEF showed a sensitivity of 85%, specificity of 79%, positive predictive value of 77% and negative predictive value of 86%, with respect to an increase in FEV<sub>1</sub> >15%.<sup>7</sup> BDR using PEF and FEV<sub>1</sub> were compared using various definitions in another study on 48 patients with cough. In general, results showed high specificities, but low sensitivities and positive predictive values.<sup>8</sup> The highest positive predictive value of 83% was found with  $\Delta$ PEF% increment of 20% against an absolute FEV<sub>1</sub> increase of 200 mL. In another study on 176 children with asthma,  $\Delta$ PEF% of 20% or 25% had a high specificity (96% and 96%, respectively), but moderate sensitivity (51% and 53%, respectively), in identifying BDR (defined as a 9% increase in FEV<sub>1</sub> predicted after inhalation of 800mcg salbutamol).<sup>9</sup>

The sensitivity and specificity values are important in the actual clinical scenario. A test with high sensitivity is needed if PEF were to be used as a screening test to identify BDR. This

is the more common situation in primary care, and clearly PEF measurements yield low sensitivities in this regard at various cut-offs. On the other hand, high specificity is preferred for a diagnostic test. Although some cut-off values do have specificity values above 0.8, PEF still cannot be used clinically for this purpose in view of the low negative predictive values.

The timing of spirometry and PEF measurement may have introduced some bias in the results since PEF was always performed after spirometry, both for baseline and post-bronchodilator assessments. Even then, post-bronchodilator PEF reading was obtained within a time-frame consistent with the nearly peak bronchodilator action of salbutamol. Although PEF is an effort-dependent test and patients may have had minor fatigue after spirometry, the same sequence was followed for all patients, and the impact (if any) on comparison between PEF and spirometry data is likely to be small in the vast majority of patients.

In conclusion, our data do not support the use of  $\Delta$ PEF or  $\Delta$ PEF% as a surrogate for standard spirometric criteria for BDR assessment.

#### Conflict of interest declaration

None to declare.

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