Abstract

Objective: To correlate forced expiratory volume in 1 second (V{EF}_1) and peak expiratory flow (PEF) with clinical parameters in children with moderate and severe asthma.

Methods: This was a non-concurrent cohort study, carried out at a pediatric pneumology clinic, in Belo Horizonte, MG, Brazil, between March and October 2002. The study enrolled children aged 5 to 16 years, with persistent asthma, being treated with a minimum of 500 µg/day beclomethasone, and with symptoms under control for at least 3 months. Seventy-five patients (96.1%) were selected by simple randomization and monitored for 3 months, via a clinical severity scale and pulmonary function tests (PEF and V{EF}_1). Results were analyzed using Pearson's coefficient.

Results: Correlations between absolute and percentage PEF figures and clinical severity score, were negative and very close to zero, signifying a weak correlation with no statistical significance. The same relationship was observed between V{EF}_1 and clinical severity score. The correlation between V{EF}_1 and PEF had a positive value with statistical significance (p = 0.000).

Conclusions: Since the best parameter for evaluating airway obstruction is V{EF}_1, the finding that there is a positive correlation between this measure and absolute PEF reinforces the importance of its use and allows for the recommendation that PEF be measured as part of the management of asthmatic children, particularly in severe cases.


Introduction

The prevalence of asthma among children and adolescents varies from 0 to 30% in different populations. In Brazil, it is the third greatest cause of hospital admissions among children and young adults, generating significant financial burden for the health service and elevated individual costs.

Furthermore, many patients are not capable of recognizing their own clinical deterioration. This reduced perception of their worsening state correlates with more severe asthma crises and increased risk of death. In Brazil, for example, it is estimated that in 1996 around 70% of deaths from asthma were of patients who did not recognize clinical decline and/or received inadequate treatment.

The Global Initiative for Asthma's management guide (GINA) recommends objective measurements of pulmonary function, such as spirometry or peak expiratory flow (PEF), for assessing the severity of asthma and response to treatment. Due to its simplicity and ease of assessment, daily monitoring of PEF at home has been recommended by GINA for patients with moderate and severe asthma, in order to help with management of symptoms and to alert of periods of exacerbation.

Peak expiratory flow represents the maximum flow generated during a forced expiration, at maximum intensity, starting from maximum lung inflation, i.e. total lung capacity. It is considered an indirect indicator of major airway obstruction and is affected by the degree of pulmonary inflation, thoracic elasticity, abdominal...
musculature and the patient’s muscular strength.\textsuperscript{5,6} It is effort-dependent and, because of this, requires the patient’s collaboration. The measurement of PEF can be performed using spirometers or portable measurement units that are available at an accessible cost and are relatively simple to operate.

International reference values have been published for PEF, according to age, stature and sex. However, the best way to evaluate them is by comparing patients with their previous best levels.\textsuperscript{7}

Some studies have questioned the true role of PEF measurements in the reduction of morbidity among asthmatic patients.\textsuperscript{8-10} Since asthma primarily affects the smaller airways, PEF will only be altered during later phases, after a significant increase in airway resistance. For this reason, it is argued that the best functional index for assessing small-airway function is the measurement of forced expiratory volume in 1 second (VEF\textsubscript{1}), which is measured using a spirometer.\textsuperscript{9} A spirometer is a relatively high-cost piece of equipment that must be operated by a trained professional.

Several authors have attempted to correlate PEF measurements with the alterations observed in asthmatic children’s symptoms. A majority of these studies analyzed children with moderate and severe asthma, controlled with inhaled corticoids. They demonstrated that there is a weak correlation (Pearson’s $r$ varying from -0.34 to 0.5) between alterations to PEF and symptoms suggestive of deteriorating pulmonary function.\textsuperscript{3,8,10,11} However, analogous studies are scarce in Brazil and Latin America.\textsuperscript{11}

The objective of the work reported here was to attempt to correlate VEF\textsubscript{1} and PEF measurements with clinical parameters in clinically stable children and adolescents with moderate and severe asthma on inhaled corticoids, in addition to verifying the contribution that serial PEF measurements have to offer for the outpatients follow-up of these patients.

**Methods**

This was a non-concurrent cohort study, carried out at the pediatric pneumology clinic at the Posto de Atendimento Médico Campos Sales. This is a secondary care clinic where around 400 asthmatic children are treated per year, referred by municipal health centers in Belo Horizonte. A sample of convenience was used, for which the participants were selected by simple randomization and followed-up for 3 consecutive months.

**Inclusion and exclusion criteria**

The study enrolled children and adolescents aged from 5 to 16 years old, with persistent asthma, on a minimum of 500 µg/day of beclomethasone and with symptoms under control for at least 3 months.

Exclusion criteria were exacerbations of symptoms, PEF and/or VEF\textsubscript{1} below 59\% of predicted, use of systemic corticoids during the 4 weeks prior to enrollment, infectious diseases of the upper airways or severe disease of any nature.

Children classified as having moderate or severe asthma, according to clinical parameters, were included in the study, as long as their pulmonary function was above 59\% of predicted.

**Operational definitions**

Asthma diagnosis was confirmed by a history of recurrent wheezing crises, coughing and dyspnea, clinically reversible with rapid-acting inhaled beta-2 agonist, combined or not with systemic corticoids.

Asthma was classified as persistent mild, moderate or severe, according to GINA.\textsuperscript{1} These criteria are based on the severity of exacerbations, intensity of symptoms, PEF and VEF\textsubscript{1} measurements and inhaled corticoid dosage.

The maximum dosages of beclomethasone or its equivalent that are recommended by GINA are 400 µg, 800 µg, or over 800 µg for the treatment of persistent mild, moderate and severe asthma, respectively. Since the presentation available in Brazil contains 250 µg of beclomethasone per spray, we considered doses of 500 µg, 750 µg, and over 750 µg/day for persistent mild, moderate and severe asthma respectively.

The occurrence and severity of exacerbations, nocturnal symptoms, limitations to physical activity and use of fast-acting beta 2-agonist were evaluated and classified according to the clinical severity score validated and published by Rosier et al.\textsuperscript{12} The scale ranges from 2 to 19 points depending upon the severity of the parameters assessed, as follows: scores from 2 to 8 correspond to mild asthma; 9 to 14 to moderate asthma; and 15 to 19 to severe asthma. Questionnaires were completed by parents, guardians or the children themselves.

**Monitoring of patients**

After enrollment, patients were monitored for 3 months, with clinical severity scores and pulmonary function test results (PEF and VEF\textsubscript{1}) produced by independent examiners who were unaware of the study objectives. Clinical functional assessments were undertaken every 2 weeks for the first 2 months and patients were reassessed once more at the end of he third month. Patients were kept on the same inhaled corticoid medication they had been using before enrollment, given via a 650 mL plastic spacer with a valve (Flumax\textsuperscript{®}, Flumax Equipamentos Médicos Ltda., Belo Horizonte).\textsuperscript{13-15}

A Mini-Wright Peak Expiratory Flow Meter (Clement Clarke, United Kingdom) was used at every consultation to...
measure PEF for the functional assessment, with the highest result chosen from three consecutive measurements for analysis, and adopting the reference values published by Polgar & Promadhat,\textsuperscript{16} these being accepted internationally.

Spirometry was performed with bronchodilation using 400 µg of salbutamol for all patients at enrollment, at week eight and at the end of the third month, in accordance with American Thoracic Society recommendations, with reference values calculated using equations published by Polgar & Promadhat.\textsuperscript{17}

At each visit, compliance with treatment and correct inhalation technique were assessed. Patients who had failed to use 25% or more of the inhaled corticoid dosage recommended for 2 weeks were excluded from the results.

### Statistical aspects

Descriptive statistics were used to characterize the study population, and linear regression employed to evaluate any correlation between the variables studied (VEF\textsubscript{1}, PEF and clinical severity score).

Data were correlated using Pearson’s coefficient, which varies from -1 to +1. Values close to -1 signify negative or inverse correlations, and values close to zero indicate an absence of correlation, while values close to +1 signify positive correlation. Results were considered significant when \( p < 0.05 \).

### Ethical considerations

The research protocol and informed consent form were approved by the Research Ethics Committee at the Universidade Federal de Minas Gerais.

### Results

The initial selection returned 87 children, of whom nine were excluded in accordance with the study’s exclusion criteria. Seventy-eight patients began the observation period, but three were excluded later: one patient had been incorrectly enrolled (VEF\textsubscript{1} < 59%), another reported clinical deterioration, and a third did not take sufficient medication, being excluded for partial compliance with treatment. Seventy-five patients therefore remained (96.1%) and were followed for 12 weeks.

Table 1 contains the characteristics of the 77 patients enrolled on the study.

There was a discrete predominance of the male sex, but without statistical significance (\( p = 0.58 \)), in addition to the asthma of the majority of the children being under good control, which can be confirmed with reference to their PEF and VEF\textsubscript{1} of over 80%.

![Scatterplot and regression line for the linear correlation between absolute PEF values and clinical severity scores](image)

**Figure 1** is a scatterplot of correlations between absolute PEF measurements and the clinical severity score, based on data obtained at all six assessments made during the study period.

The correlation observed between absolute PEF measurements and clinical severity score was negative and very close to zero, signifying a weak, almost non-existent correlation (\( r = -0.03 \)) and without statistical significance (\( p = 0.08 \)). The regression line helps visualization of the inverse correlation between the two variables.
Discussion

There were no correlations observed between the functional parameters PEF and VEF₁ and the clinical scale in the data published here. In contrast there was a positive correlation between absolute PEF values and VEF₁ reference values. The absence of any correlation may be because the children studied here were under control, from a clinical point of view, on medication (the majority scored between two and eight points on the clinical severity scale adopted), or due to the difficulty in recognizing deteriorated pulmonary function resulting from patients adapting to the limitations of their disease.

Correlations between VEF₁/PEF and clinical severity score did not have any statistical significance (p > 0.05). While, in the majority of cases, r was negative and close to zero, the distribution of points on the graphs demonstrated that there was no correlation between these variables. Patients classified with a score of 2 (mild asthma) exhibited PEF below 40% of predicted and VEF₁ of 50%. It is possible, as is observed with other chronic diseases, that these children have adapted to the limitations imposed by their asthma, leading to a reduced perception of the symptoms associated with airway obstruction.

There was a positive correlation between VEF₁ and PEF, demonstrating that if VEF₁ increases, PEF also increases (and vice-versa). This being so, PEF should be a useful indicator for assessing small airway obstruction, since VEF₁ is used as the gold standard in our country.

Clinical assessment is subjective and the results reveal their obvious limitations. This is why, before being relied on in isolation, it should be accompanied by objective measurements, such as PEF and spirometry.

This work was performed using a sample that is comparable to others in the literature, both in terms of number and age of participants and in terms of the follow-up period. The statistical analysis employed was also similar to other studies, in order to facilitate comparison and discussion. In contrast with the studies published to date, here all PEF measurements were taken by a trained professional who was unaware of the study objectives, which helped to control for possible measurement errors. In studies hitherto published, data were obtained, not just in the laboratory, but also from records of measurements taken by the patients themselves.

Brand et al. assessed 102 children aged 7 to 14 years over 2 weeks and found a weak correlation coefficient for PEF and a clinical severity score (r = -0.34, p < 0.01). The correlation observed for PEF and VEF₁ was r = 0.15, but p did not denote significance.¹⁰

Cabral et al., followed 92 children aged 6 to 16 years for 5 months, also observing a significant correlation
between PEF and symptoms, although a weak one and with a negative coefficient, (p < 0.05).11

Liam et al. monitored a group of 64 patients with stable asthma and found a weak correlation between a clinical severity score and PEF (r = -0.214, p = 0.104), and between the clinical severity score and VEF1 (r = -0.256, p = 0.041). The concluded that their results supported the need for objective measurements of airway obstruction.12

In their article, Goldberg et al. reported finding a strong correlation between PEF and o VEF1, with r values varying from 0.74 to 0.93 (p < 0.0001) and concluded that PEF measurements can be used as a useful follow-up parameter.13

In contrast, Paggiaro stated that the coefficient of the correlation between VEF1 and PEF for absolute figures varied from 0.78 to 0.95, and from 0.74 to 0.91 for percentages, which reinforces indications for the use of PEF measurements as an objective method for monitoring the pulmonary function of patients (r = 0.796, p = 0.0001).6

Newr et al. followed 244 children aged 4 to 18 years for 3 years and found that there was a good correlation between VEF1 and PEF (r = 0.73, p = 0.02). These values are comparable with figures found by Shingo et al., who observed a good correlation between VEF1 and PEF (r = 0.74) for measurements taken at home by patients and for PEF measurements taken at the laboratory (r = 0.85). Nevertheless, the correlation between VEF1 and a clinical severity score was weak (r = -0.13). All correlations had p < 0.005, however, the patients under investigation were more than 15 years old.20 These studies all help to corroborate the indication for using PEF when monitoring asthmatic children.

Among the studies published, it was Sly who found a positive, if weak, correlation between PEF and clinical severity score (r = 0.35, p < 0.01), and between PEF and variation in clinical severity score (r = 0.38, p < 0.01). He observed 80 children aged 6 to 16 years for 8 weeks.3

All of the published data commented on above, including data from the present study, found evidence of little or no correlation between clinical severity scores and VEF1 and PEF, and a positive correlation between VEF1 and PEF. Since the best parameter for assessing airway obstruction is VEF1, a positive correlation with PEF, as demonstrated by the research described here and by other studies in the literature, reinforces the importance of, and allows for the recommendation to use, PEF measurements for the management of asthmatic children, particularly in severe cases. However, even with periodic and sequential measures, they should not be relied on as the only objective parameter when monitoring asthmatics.

**References**

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