

**Methodological weaknesses regarding the use of PEF-meters when assessing physiological effects of air pollution risk factors**

Metodološke pomanjkljivosti uporabe PEF-metrov za oceno fizioloških učinkov zračnih polutantov

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**Abstract:** The manuscript discusses the applicability of self-administered monitoring of respiratory function in asthma patients in order to discern between negative health effects of different air pollutants on respiratory function. Thirteen asthma patients measured their respiratory function twice daily over a one month winter period. They used PEF-meters to monitor peak expiratory flow (PEF; L/min) and forced expiratory volume in the first second of expiration (FEV<sub>1</sub>; L). Subjects' vital capacity (VC; L) was measured in a laboratory setting at the end of the measuring period. Respiratory data were evaluated in respect to the ambient concentrations of NO<sub>2</sub>, NO<sub>x</sub>, SO<sub>2</sub>, O<sub>3</sub>, and PM<sub>10</sub>, which were monitored during the same period. The concentrations of some air pollutants (PM<sub>10</sub> and NO<sub>x</sub>) exceeded the critical levels on several days during the study. PEF-meter data (PEF, FEV<sub>1</sub> and FEV<sub>1</sub>/VC), however, did not respond significantly to the ambient conditions (P>0.05). The results speak against the use of self-administered PEF-meter monitoring for the recognition and investigation of air pollutant related negative health effects. Limitations and delimitations of the method are presented.

**Keywords:** respiratory function; asthma; self-administered respiratory monitoring; particulate matter; PM<sub>10</sub>; PEF; FEV<sub>1</sub>

**Izvleček:** Prispevek analizira uporabnost samo-apliciranega spremljanja dihalne funkcije pri astmatikih, uporabljenega z namenom razločiti med negativnimi zdravstvenimi učinki različnih zračnih onesnaževalcev na dihalno funkcijo. Trinajst astmatikov je dvakrat dnevno v obdobju enega meseca v zimskem času spremljalo svojo dihalno funkcijo. Maksimalni pretoka zraka ob izdihu (PEF; L/min) in prisiljen ekspiratorni volumen v prvi sekundi izdiha (FEV<sub>1</sub>; L) so spremljali s PEF-metri. Vitalna kapaciteta preiskovancev (VC; L) je bila izmerjena v laboratoriju ob koncu raziskovalnega obdobja. Respiratorni podatki so bili analizirani glede na okoljske koncentracije onesnaževalcev NO<sub>2</sub>, NO<sub>x</sub>, SO<sub>2</sub>, O<sub>3</sub> in PM<sub>10</sub>, ki so bili spremljani v istem časovnem obdobju. Koncentracije nekaterih zračnih onesnaževalcev (PM<sub>10</sub> in NO<sub>x</sub>) so presegle kritično mejo v več dnevih raziskovalnega obdobja. Kljub temu, s podatki s PEF-metrov (PEF, FEV<sub>1</sub> in FEV<sub>1</sub>/VC) ni bilo mogoče pokazati statistično značilnih odzivov dihalne funkcije na okoljske razmere (P>0.05). Rezultati raziskave izpostavljajo problematičnost uporabe samo-apliciranega spremljanja dihalne funkcije s PEF-metri za prepoznavo in raziskave negativnih zdravstvenih učinkov, povezanih z zračnimi onesnaževalci. V besedilu so podrobneje prikazane prednosti in slabosti predstavljene metodologije.

**Ključne besede:** respiratorna funkcija; astma; samo-aplicirano spremljanje dihalne funkcije; PM<sub>10</sub>; PEF; FEV<sub>1</sub>

## Introduction

Respiratory tract is an organic system that is maximally exposed to air pollution. People with respiratory diseases are thus one of the most susceptible population groups to different air pollutants. Indeed, studies suggest that the quality of air in the environment can affect the respiratory function in humans (Rios et al. 2004), and that respiratory conditions, such as asthma, can be aggravated in a polluted environment (Heinrich et al. 1999).

Asthma is a chronic disease that severely affects the respiratory function. Asthma associated inflammation of the airways constricts the bronchia, which increases resistance and decreases the airflow through the respiratory system. It is generally accepted that the prevalence of asthma has increased significantly over the last years. It has been suggested that an increase in the number of asthmatic conditions is most likely related to environmental factors, as the period in which the changes have been observed is too short for genetic mutations to occur (Joseph et al. 1996). Epidemiological and controlled exposure studies of human volunteers have indeed demonstrated that exposure to a variety of pollutants induces asthma exacerbations (Peden 1996).

The intensity of asthma symptoms and signs varies in time and the aggravation of health status has been related to several factors, such as pollen, ambient temperature and viral infections (Gent et al. 2003). Furthermore, studies have demonstrated that the aggravation of disease is related to several air pollutants, although, the relation between asthma outbreaks and any particular air substance has not yet been determined unequivocally (Gent et al. 2003). Different studies suggest that particulate matter (PM) and NO<sub>2</sub> can be pointed out as air pollutants that cause the most negative health effects in individuals with already impaired respiratory function, namely, in COPD and asthma patients (Lagorio et al. 2006).

As air pollutants cause negative health effects worldwide, this topic has gained an increasing interest. Epidemiological approach has most often been used when the effects of air pollutants on health

have been studied. In contrast, physiological data on this matter are rather scarce or even non-existent. Thus rather surprisingly, data on the effects of different air pollutants on a day to day respiratory function of both, healthy people and people with respiratory diseases, are limited, although they can provide vital information about the detection and consequent prevention against the environmental risk factors. In one of the few studies, for example, Linn and co-workers (Linn et al. 1996) reported that in healthy schoolchildren forced vital capacity measured in the morning decreases significantly with increase in particulate air pollution or NO<sub>2</sub> measured over the preceding 24 hours, and that morning-to-afternoon change in FEV<sub>1</sub> becomes significantly affected with increase in particulate matter, NO<sub>2</sub>, or O<sub>3</sub> on the same day. Daily assessment of respiratory function can therefore serve as a fast and non-subjective tool, which is able to discern critical environmental factors.

It is likely that one of the reasons for the lack of data about the effects of different air pollutants on a day to day respiratory function is logistical, as daily ambulatory measurements of respiratory function are expensive and rather difficult to be organized. The availability and development of new techniques, which allow self-administered monitoring of respiratory function at home, thus provide promising new research opportunities and the use of peak expiratory flow meters (PEF-meters) has been suggested as a potentially useful methodological approach to address this issue (Bellia et al. 2003).

To monitor the respiratory state of asthma patients, pneumotachs or spirometers are usually used in ambulatory monitoring, and portable peak expiratory flow-meters (PEF-meters) by patients themselves. Peak expiratory flow (PEF, L/min) is the maximal flow of air during a forced expiration. PEF measurements are used in outpatient monitoring of asthma and in emergencies (Cross and Nelson 1991). PEF measurements, however, have been criticized for use with chronic obstructive pulmonary disease patients (COPD) due to unreliability, poor reproducibility, and unavailable reference values (Pauwels et al. 2001). To improve

the monitoring, the measurements of forced expiratory volume in the first second of expiration (FEV<sub>1</sub>, L) have therefore become regularly used and have also become available with newer portable PEF-meters. It has to be noted, that PEF and FEV<sub>1</sub> measurements provide different information, since PEF reflects flows in the large airways, and FEV<sub>1</sub> reflects obstruction in different parts of the airways (Paggiaro et al. 1997).

As data about the application of portable PEF-meters for a daily respiratory monitoring for the assessment of the effects of air pollutants on respiratory function are limited (Bellia et al. 2003), the present study aimed to assess the feasibility, sensitivity and applicability of self-administered monitoring of respiratory function by PEF-meters for the detection of environmental conditions that have previously been demonstrated to negatively affect respiratory function in humans. Should the negative effects of air pollutants be reflected in diminished respiratory function as self-assessed by PEF meters, asthma patients would gain a simple tool to recognize the environmental risk and consequently decrease it by modifying their behavior, i.e. by modifying the time or location of their outdoor exposure. We hypothesized that self-administered PEF-meter monitoring will be able to detect changes in respiratory function of asthma patients in relation to the concentrations of specific air pollutants in an urban area.

## Methods

Asthma patients were selected for the study, because they, being respiratory disease patients, are one of the most sensitive groups to air pollution. Daily monitoring of the respiratory function of 13 asthma patients was performed over a period of one month. The study was performed in wintertime, as this is the period in which the likelihood for temperature inversion and thus the high concentrations of air pollutants in the selected urban area is the highest. Furthermore, the concentration of pollen, which can considerably worsen the respiratory function of asthmatic patients, and can thus interfere with the respiratory measurements, is smallest in winter. If the study was, for example, performed during seasons change, when the temperature fluctuation are

highest and the pollen concentration increases, this two influential factors could severely interfere with the study results (Gent et al. 2003).

### *Ethical clearance*

The study was conducted in accordance with the Declaration of Helsinki and the protocol of the study gained approval by the Ethics Committee of the Republic of Slovenia (approval number 110/12/04). Each subject provided written informed consent before participating.

### *Subjects*

Asthma patients were asked to voluntarily participate in the study and were recruited by their attending pulmonologist. The subjects with emphysema, chronic bronchitis, and heart failure could not participate in the study. Also, the subjects with acute upper respiratory infections could not be recruited. The main inclusion criteria for the subjects was a settled medical asthmatic condition, for which large alterations in regular medications were not expected to occur, as a substantial change in medication during the course of the study would likely interfere with the study results. Furthermore, as air pollution is affected by geographical features, the study aimed to minimize this effect by recruiting only the subjects living in an area within approximately 1.5 km of air radius around the measuring station of Environmental Agency of the Republic of Slovenia. Thus, only the subjects permanently living in the strict town area of Ljubljana were included in the study.

### *Protocol and instrumentation*

The information about the daily concentrations of pollutants in the air of the town of Ljubljana was obtained from the Environmental Agency of the Republic of Slovenia, an official body that monitors air pollution in Slovenia. The data about the following pollutants were obtained: particulate matter of less than 10 micrometers in diameter (PM<sub>10</sub>; µg/m<sup>3</sup>); ozone (O<sub>3</sub>; µg/m<sup>3</sup>); sulphur dioxide (SO<sub>2</sub>; µg/m<sup>3</sup>); nitrogen dioxide (NO<sub>2</sub>; µg/m<sup>3</sup>), and nitrogen oxides (NO<sub>x</sub>; µg/m<sup>3</sup>).

Peak expiratory flow-meters (Piko-1 Ferraris Respiratory Europe Ltd., Great Britain) were

used in the study to monitor the respiratory state of asthma patients. These PEF meters determine both, peak expiratory flow (PEF, L/min), thus the maximal flow of air during forced expiration, as well as the forced expiratory volume in the first second of expiration (FEV<sub>1</sub>, L). According to manufacturers' recommendations, the PEF and FEV<sub>1</sub> values measured by PiKo-1 PEF meter correlate well at all flows with those measured by a pneumotach. The devices used correspond both, to the American Thoracic Society standards and to standards of the European Union.

Prior to the study, all subjects were given thorough instructions about the functioning of the PEF meters and aims of the study. A correct procedure of the measurement was demonstrated to them. To perform the measurement, the subjects were asked to stand up, hold a PEF-meter horizontally and be careful not to cover the expiratory outlets. They were instructed to blow as much air as they could through the mouthpiece of the PEF-meter after an audible signal. They were asked to perform the measurement three times at each occasion and to write down the obtained values of PEF and FEV<sub>1</sub>

into the suitable tables after each measurement. The subjects performed the measurements in the morning, immediately when they got up, and in the evening, just prior they went to sleep. Apart from providing the values of PEF and FEV<sub>1</sub>, all subjects were asked to daily fulfill the questionnaire with any remarks they thought might be relevant to the measurements, such as: their state of health, fever, cold, a change in measurement place (if they would be absent from home), and a change in medication. For each subject, the data acquisition tables were prepared in advance and were collected at the end of a one-month period.

Vital capacity (VC, L), i.e. the maximum volume of air that can be exhaled after a maximum inhalation, was also measured in a laboratory setting with a spirometer (Cosmed K4b2, Rome, Italy) at the end of the study period. The subjects' age was recorded and their height and weight were determined on a certified scale (Vita Libela Elsi, Celje, Slovenia). The spirometer was first calibrated with a 3-litre calibrating syringe (Cosmed, Rome, Italy), then the subjects were fitted with a nose clip. The standing subjects first breathed normally

Table 1: Subjects characteristics, including gender, with individual and average age, height, mass and smoking status presented

Tabela 1: Značilnost preiskovancev. Predstavljeni so spol ter posamezni in povprečni podatki za starost, višino, maso in kadilski status.

| <b>Subjects' characteristics</b>           |                                                |                                  |                               |                           |                                                |
|--------------------------------------------|------------------------------------------------|----------------------------------|-------------------------------|---------------------------|------------------------------------------------|
| <b>Lastnosti preiskovancev</b>             |                                                |                                  |                               |                           |                                                |
| Subject number<br>Številka<br>preiskovanca | Gender<br>(male/female)<br>Spol (moški/ženska) | Age<br>(years)<br>Starost (leta) | Height<br>(cm)<br>Višina (cm) | Mass<br>(kg)<br>Masa (kg) | Smoker<br>(yes/ex/no)<br>Kadilec (da/bivši/ne) |
| 1                                          | male / moški                                   | 56                               | 177                           | 84                        | Yes / Da                                       |
| 2                                          | male / moški                                   | 58                               | 189                           | 97                        | Yes / Da                                       |
| 3                                          | male / moški                                   | 31                               | 186                           | 81                        | Yes / Da                                       |
| 4                                          | male / moški                                   | 70                               | 164                           | 52                        | No / Ne                                        |
| 5                                          | male / moški                                   | 39                               | 183                           | 80                        | Ex / Bivši                                     |
| 6                                          | female / ženska                                | 26                               | 166                           | 54                        | Yes / Da                                       |
| 7                                          | female / ženska                                | 85                               | 159                           | 67                        | Ex / Bivši                                     |
| 8                                          | female / ženska                                | 41                               | 175                           | 60                        | No / Ne                                        |
| 9                                          | female / ženska                                | 40                               | 166                           | 61                        | No / Ne                                        |
| 10                                         | female / ženska                                | 64                               | 163                           | 76                        | Ex / Bivši                                     |
| 11                                         | female / ženska                                | 59                               | 166                           | 94                        | Yes / Da                                       |
| 12                                         | female / ženska                                | 59                               | 155                           | 92                        | Ex / Bivši                                     |
| 13                                         | female / ženska                                | 45                               | 173                           | 83                        | Ex / Bivši                                     |
| <b>AVG±SD</b>                              | /                                              | <b>52±17</b>                     | <b>171±11</b>                 | <b>75±15</b>              | /                                              |

through a spirometer and then took a maximally deep breath, after which they blew through the mouthpiece as strongly and as for as long as they could. The whole procedure was repeated three times and the maximal measured volume of expired air was taken as representative for the subjects' vital capacity. No broncho-dilatators were used for the vital capacity determination.

#### Statistical analysis

For each pollutant, a day with its minimum and maximum measured concentration was selected. Then, PEF and FEV<sub>1</sub> values of all subjects measured for these two days were compared with a two tailed paired Student T-test. Both, the absolute and relative values of PEF and FEV<sub>1</sub> were analysed.

Table 2: The maximum hourly concentrations of fine particles (PM<sub>10</sub>; µg/m<sup>3</sup>) and nitrogen oxides (NO<sub>x</sub>; µg/m<sup>3</sup>) in the air in the town of Ljubljana during the experimental period. The two experimental days with the maximum and minimum concentrations of the two air pollutants are presented in bold. The shaded areas denote days in which the legislative boundary concentrations of PM<sub>10</sub> or NO<sub>x</sub> were exceeded

Tabela 2: Največji urni koncentraciji finih delcev (PM<sub>10</sub>; µg/m<sup>3</sup>) in dušikovih oksidov (NO<sub>x</sub>; µg/m<sup>3</sup>) v zraku v mestu Ljubljana med potekom raziskave. Dneva z največjima in najmanjšima koncentracijama obeh onesnaževalcev sta prikazana odebeljeno. Osenčena polja označujejo dneve, v katerih so bile zakonsko določene največje koncentracije PM<sub>10</sub> ali NO<sub>x</sub> presežene

| The maximum hourly concentrations of pollutants in the experimental days |                                          |                                      |                                         |
|--------------------------------------------------------------------------|------------------------------------------|--------------------------------------|-----------------------------------------|
| Največja urna koncentracija onesnaževalcev v raziskovalnih dneh          |                                          |                                      |                                         |
| Experimental Day<br>Raziskovalni dan                                     | PM <sub>10</sub><br>(µg/m <sup>3</sup> ) | Experimental Day<br>Raziskovalni dan | NO <sub>x</sub><br>(µg/m <sup>3</sup> ) |
| <b>4</b>                                                                 | <b>207.45</b>                            | <b>4</b>                             | <b>208.55</b>                           |
| 3                                                                        | 181.47                                   | 5                                    | 200.55                                  |
| 5                                                                        | 155.19                                   | 6                                    | 170.30                                  |
| 25                                                                       | 133.18                                   | 25                                   | 162.35                                  |
| 6                                                                        | 126.29                                   | 12                                   | 138.90                                  |
| 2                                                                        | 121.15                                   | 3                                    | 136.80                                  |
| 27                                                                       | 115.01                                   | 11                                   | 132.35                                  |
| 28                                                                       | 102.92                                   | 18                                   | 119.35                                  |
| 1                                                                        | 95.42                                    | 27                                   | 112.75                                  |
| 12                                                                       | 90.02                                    | 1                                    | 102.85                                  |
| 24                                                                       | 88.35                                    | 17                                   | 94.10                                   |
| 26                                                                       | 84.01                                    | 13                                   | 93.50                                   |
| 11                                                                       | 75.33                                    | 8                                    | 92.30                                   |
| 17                                                                       | 70.00                                    | 26                                   | 91.25                                   |
| 13                                                                       | 69.75                                    | 28                                   | 91.15                                   |
| 18                                                                       | 67.60                                    | 2                                    | 90.70                                   |
| 19                                                                       | 59.27                                    | 22                                   | 87.05                                   |
| 16                                                                       | 55.49                                    | 24                                   | 86.70                                   |
| 14                                                                       | 52.45                                    | 23                                   | 78.65                                   |
| 23                                                                       | 52.14                                    | 16                                   | 75.15                                   |
| 20                                                                       | 50.90                                    | 19                                   | 75.10                                   |
| 21                                                                       | 46.38                                    | 9                                    | 57.70                                   |
| 8                                                                        | 44.08                                    | 20                                   | 48.75                                   |
| 15                                                                       | 42.22                                    | 10                                   | 40.80                                   |
| 10                                                                       | 39.70                                    | 15                                   | 31.65                                   |
| 22                                                                       | 38.32                                    | 7                                    | 30.10                                   |
| 9                                                                        | 37.32                                    | 14                                   | 22.70                                   |
| 7                                                                        | <b>29.20</b>                             | <b>21</b>                            | <b>15.30</b>                            |

To enable inter-individual comparisons, FEV<sub>1</sub>/VC ratio was calculated for each subject and compared between the days with minimal and maximal concentrations of air pollutants. A P-value of less than 0.05 was adopted as statistically significant. All the following results are presented as average±SD.

## Results

### *Subjects characteristics*

The average age of the subjects was 52±17 years, height 171±11 cm and mass 75±15 kg. Eight of the subjects were females (two smokers, four ex-smokers, two non-smokers) and five of them males (three smokers, one ex-smoker, one non-smoker) (Table 1). During the study the subjects pursued their regular daily activities and reported no changes in their day-to-day medication. During the period of the study, none of the asthmatics required any emergency medical treatment, medicine alterations, or hospitalization.

### *Concentrations of pollutants*

During the course of the study, the concentrations of PM<sub>10</sub> and NO<sub>x</sub> exceeded the legislatively determined critical values, i.e. the boundary daily value of 50 µg/m<sup>3</sup> for PM<sub>10</sub> and the hourly boundary value of 200 µg/m<sup>3</sup> for NO<sub>x</sub>, on several occasions (Table 2). The concentrations of ozone, SO<sub>2</sub> and NO<sub>2</sub> remained below the legislatively determined critical values for a particular substance throughout the course of the present study. The concentrations of air pollutants at the measuring location are presented on Figure 1.

### *Respiratory function measurements*

Maximal evening values of PEF and FEV<sub>1</sub>, both absolute and relative data, were used for the analysis.

On a day with maximal (207.5 µg/m<sup>3</sup>) PM<sub>10</sub> concentration, the PEF of the subjects was 342.5±146.7 L/min, and on a day with a minimal (29.2 µg/m<sup>3</sup>) PM<sub>10</sub> concentration 352.3±124.9 L/min (P=0.55). PEF, relative to the first day of

measurements was 90±23 % on the day of the maximal, and 97±12 % on the day of the minimal PM<sub>10</sub> concentration (P=0.26). The differences were thus not statistically significant (P>0.05).

FEV<sub>1</sub> of the subjects (Figure 2) was similar at 2.5±0.9 L on a day with maximal and minimal PM<sub>10</sub> concentration (P=0.67). FEV<sub>1</sub>, relative to the first day of measurements was 90±22 % on the day of the maximal, and 94±21 % on the day of the minimal PM<sub>10</sub> concentration (P=0.40). The differences were thus not statistically significant (P>0.05).

The ratio of FEV<sub>1</sub>/VC of the subjects was 69±23 % on a day with maximal, and 71±19 % on a day with minimal PM<sub>10</sub> concentration, with no significant differences (P=0.58>0.05) between the two days.

On a day with maximal (208.6 µg/m<sup>3</sup>) NO<sub>x</sub> concentration, the PEF of the subjects was 342.5±146.7 L/min, and on a day with minimal (15.3 µg/m<sup>3</sup>) NO<sub>x</sub> concentration 355.0±148.2 L/min (P=0.64). PEF, relative to the first day of measurements was 90±23 % on the day of the maximal, and 96±28 % on the day of the minimal NO<sub>x</sub> concentration (P=0.41). The differences were thus not statistically significant (P>0.05).

FEV<sub>1</sub> of the subjects (Figure 2) was 2.5±0.9 L on a day with maximal NO<sub>x</sub> concentration, and 2.9±1.0 L on a day with minimal NO<sub>x</sub> concentration (P=0.18). FEV<sub>1</sub>, relative to the first day of measurements was 90±22 % on the day of the maximal, and 105±28 % on the day of the minimal NO<sub>x</sub> concentration (P=0.12). The differences were thus not statistically significant (P>0.05).

The ratio of FEV<sub>1</sub>/VC of the subjects was 69±23 % on a day with maximal, and 78±26 % on a day with minimal NO<sub>x</sub> concentration, with no significant differences (P=0.15>0.05) between the two days.

The concentrations of other air pollutants (ozone, NO<sub>2</sub>, SO<sub>2</sub>) did not exceed the legislatively determined critical levels during the course of the study. PEF, FEV<sub>1</sub> and PEF/FEV<sub>1</sub> were nevertheless compared with respect to the maximal and minimal concentrations of these pollutants and no significant differences between the conditions were observed (P>0.05).

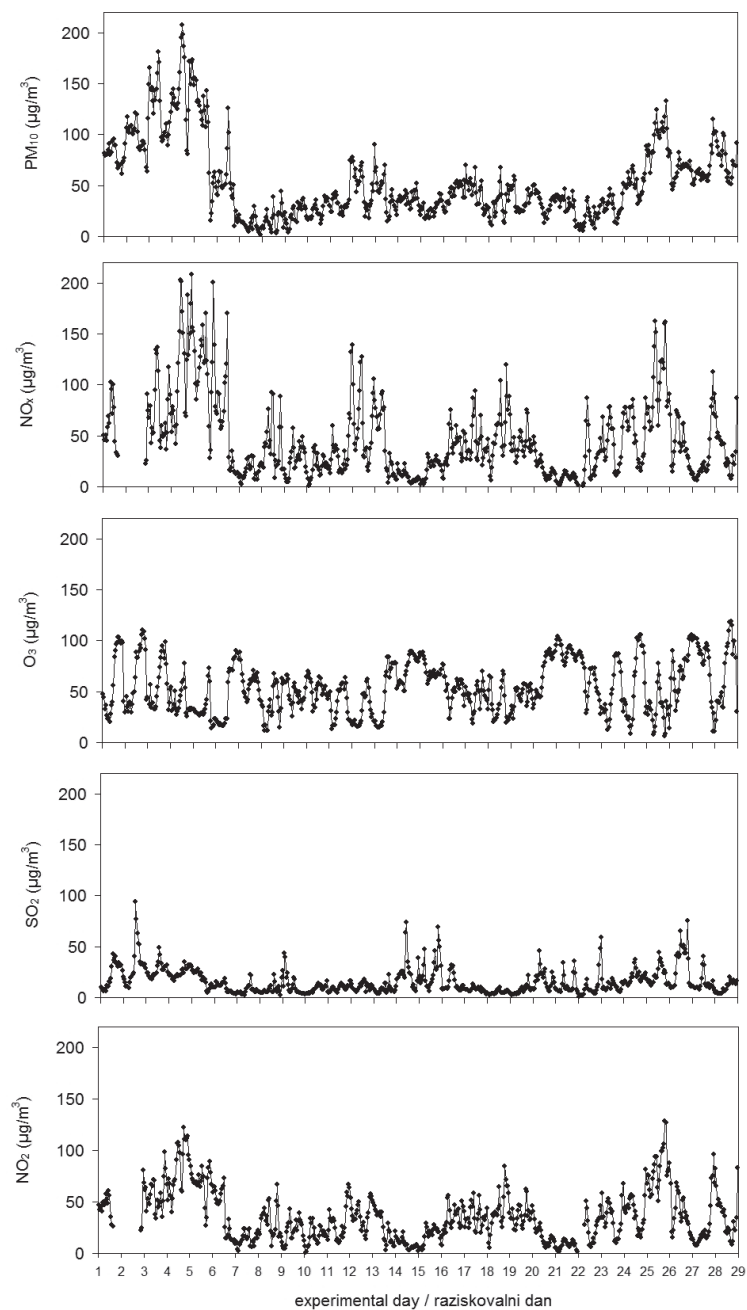


Figure 1: Measured concentrations of PM<sub>10</sub>, NO<sub>x</sub>, O<sub>3</sub>, SO<sub>2</sub> and NO<sub>2</sub>, all measured in  $\mu\text{g}/\text{m}^3$  in ambient air, during the course of the study

Slika 1: Izmerjene koncentracije PM<sub>10</sub>, NO<sub>x</sub>, O<sub>3</sub>, SO<sub>2</sub> in NO<sub>2</sub> v okoljskem zraku med potekom raziskave, vse izražene v  $\mu\text{g}/\text{m}^3$



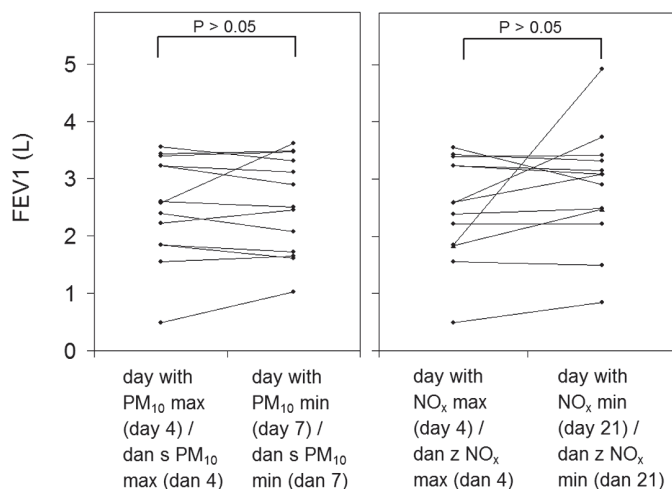


Figure 2: Individual values of forced expiratory volume in the first second of expiration ( $FEV_1$ , L) measured in the days with maximal and minimal ambient concentrations of  $PM_{10}$  ( $\mu\text{g}/\text{m}^3$ ) and  $NO_x$  ( $\mu\text{g}/\text{m}^3$ ). For any of the pollutants, no significant differences in  $FEV_1$ , as assessed by self-administered PEF-meter monitoring, were found between the two days ( $P > 0.05$ ). The individual data also demonstrate that no consistent trend or pattern in the  $FEV_1$  responses was evident

Slika 2: Posamezne vrednosti prisiljenega ekspiratornega volumna zraka v prvi sekundi izdiha ( $FEV_1$ , L), izmerjene v dneh z največjima in najmanjšima okoljskima koncentracijama  $PM_{10}$  ( $\mu\text{g}/\text{m}^3$ ) in  $NO_x$  ( $\mu\text{g}/\text{m}^3$ ). Za nobenega od onesnaževalcev se  $FEV_1$  vrednosti, izmerjene s samo-apliciranim spremljanjem dihalne funkcije s PEF-metrom, med obema dnevnoma niso značilno ( $P > 0.05$ ) razlikovale. Posamezni podatki kažejo tudi na odsotnost kakršnegakoli trenda ali vzorca v izmerjenih  $FEV_1$  vrednostih

## Discussion

The present study investigated whether self-administered monitoring of respiratory function by PEF-meters can be used for the recognition of critical environmental conditions that have been demonstrated to aggravate the severity of asthma symptoms. Although the exact levels of air pollutants, as well as time of outdoor exposure cannot be standardised for a field study, both, the concentrations of  $PM_{10}$  and  $NO_x$  were elevated above the legislatively determined critical levels in several days during the selected study period, and were above the concentrations that have been proven to affect the respiratory function in asthmatics (Pope, III and Kanner 1993).

The respiratory values of PEF,  $FEV_1$ , and  $FEV_1/VC$  were investigated in the view of minimal and maximal concentrations of air pollutants detected in a one-month winter period. In contrast to the expectations, the elevated concentrations of

the selected air pollutants were not significantly reflected in the monitored respiratory parameters of the subjects as self-assessed by PEF-meters. The lack of the observed effect might result either from the actual lack of a physiological effect, or from the inappropriateness of the used method to detect the effect – the two options are discussed below.

To a certain extent, the absence of effect can be explained by the fact that during the course of the present study, the concentrations of the majority of monitored air pollutants did not exceed the legislatively determined critical values. This might suggest that the actual respiratory function of asthma patients was indeed not affected by air pollutants. However, both,  $PM_{10}$  and nitrogen oxides did exceed the critical values for several days and therefore the absence of any significant effects of  $PM_{10}$  on the respiratory function as determined by PEF-meters should be seriously questioned.



The fact that exposure to particulate matter affects respiratory function in asthmatics has been demonstrated several times. In asthmatic children, an increase in particulate matter (PM<sub>10</sub>) elevates the incidence of lower respiratory symptoms and cough (Aekplakorn et al. 2003). Similarly, small airway function significantly decreases, and oxidative stress significantly increases in relation to PM<sub>2.5</sub> (as well as to SO<sub>2</sub> and NO<sub>2</sub>) concentrations in ambient air (Liu et al. 2009). Studies have also demonstrated that acute increases in PM<sub>10</sub> result in a greater use of asthma medications and increased hospital admissions due to asthma (Lipsett et al. 1997). Even more, Pope and Dockery (Pope and Dockery 1999) noticed an average of 2 % increase in hospitalizations and related health care visits, and an approximate 3 % increase in asthma symptoms for each 10 µg/m<sup>3</sup> rise in PM<sub>10</sub>.

Furthermore, Pope and Kanner (1993) reported that in smokers with mild to moderate chronic obstructive pulmonary disease PM<sub>10</sub> of approximately 100 µg/m<sup>3</sup> was associated with a significant decline in FEV<sub>1</sub>. The PM<sub>10</sub> level of 100 µg/m<sup>3</sup> was exceeded in several days during the course of the present study, but nevertheless no significant diminishment in PEF or FEV<sub>1</sub> as self-assessed by PEF-meters has been detected. It therefore seems more reasonably to believe that the self-administered respiratory monitoring by PEF-meters failed to detect the actual physiological effect of air pollutants, rather than to believe in the absence of the actual physiological effect. The reasons for such conclusions are several and are presented in details below.

A potential difference between personal exposures and ambient pollution may be a source of variation. However, Janssen and co-workers (Janssen et al. 1998) demonstrated that personal PM<sub>10</sub> concentrations are well correlated with ambient PM<sub>10</sub> concentrations over time, which provided support for using fixed site outdoor measurements of air pollutants for studies aiming to link day-to-day variations in respiratory function with ambient conditions. In the present study, as PM<sub>10</sub> concentrations were one of the crucial pollutants, the failure of self-administered PEF-meter monitoring to detect negative effects of air pollutants on respiratory function as being due to the difference between indoor and outdoor conditions, is therefore excluded.

One may also argue that the failure of self-administered PEF-meter monitoring to detect negative effects of air pollutants on respiratory function might be due to the number of subjects included in the study. However, as evident from the results, in the present study the calculated P values were far from statistical significance, although the inter-individual variability was accounted for by a repeated-measures design. Furthermore, no consistent changes were observed in the individual responses, thus no pattern or trend in the PEF or FEV<sub>1</sub> as related to the air pollutant concentrations was detected, rather the responses were chaotic (Figure 2). It is therefore rather unlikely that simply increasing the sample size would yield any significant response. Furthermore, the present study was designed on the basis of published studies, which used similar methodological approach, and found significant results with similar sample sizes (Chalmers et al. 2002; Kim et al. 2007; Fujimura et al. 2003). Kim et al. (2007), for example, investigated the effects of on-site ozone concentrations on peak expiratory flow in 17 asthmatics by the use of pocket PEF-meters. Kim et al. (2007) found that although the degree of asthma symptoms was influenced significantly by the ozone concentration, no significant correlation between PEF as self determined by pocket PEF-meters and ozone concentration was found. The results of Kim et al. (2007) provide further support to the present study, suggesting that it is highly unlikely to expect different performance assessment of PEF-meters, should indoor instead of outdoor measuring sites be used in the present study.

Furthermore, although the PEF-meters used to determine the respiratory function of the subjects in the present study have been assessed as reliable (Rodriguez-Pascual et al. 2006), self-administered PEF-meter monitoring still largely depends on the compliance of the subjects and should be therefore closely scrutinized. As usual with any self-administered monitoring, a certain percentage of the subjects will not follow the suggested protocol precisely, therefore, individual data have to be carefully examined. In the present study, for example, 20 subjects were enrolled, but data from 13 subjects satisfied the criteria for the final analysis, namely, three subjects did not perform the measurements regularly, as evident from the

post-hoc memory recall of the PEF meters, one of the subjects experienced some problems with handling the PEF meter and therefore the respiratory data were not recorded, one of the subjects did not consent to vital capacity monitoring, and two of the PEF meters were broken due to misuse during the measuring period, which the subjects failed to report. It is therefore suggested that the subjects should be very thoroughly instructed about the proper use of PEF-meters. Ideally, a short test period of several days prior to the actual beginning of the study with repeated self-administered PEF-meter monitoring may prove beneficial in improving the reliability of measurements. During such a test period, subjects should follow a similar protocol as planned for the study, and data from the test period should be screened by the investigators. Subjects, who would fail to produce reliable self-administered PEF-meter measurements during the test period, should be further instructed, and, if no progress made, they should be excluded from the actual study. Finally, an insufficient compliance with long-term monitoring and falsified measurements will diminish the reliability of data; the option of automatic saving of the results provided by some PEF-meters, such as the one used in the present study, help to identify the falsified reports. In longer longitudinal studies, loss of PEF-meter accuracy over time (Irvin et al. 1997) should also be considered, as this will play a crucial role in the final success or failure of a study.

## Povzetek

Kakovost zraka vpliva na dihalno funkcijo ljudi in zdravstveno stanje respiratornih bolnikov, denimo astmatikov, se v onesnaženem ozračju poslabša. Kljub temu obstaja znatno pomanjkanje podatkov o učinkih različnih onesnaževalcev na vsakodnevno dihalno funkcijo ljudi, saj, tudi zaradi različnih metodoloških pristopov, ustreznih rezultatov še nismo pridobili. Namen raziskave je bil zato oceniti uporabnost samo-apliciranega spremljanja dihalne funkcije pri astmatikih, uporabljenega z namenom razločiti med negativnimi zdravstvenimi učinki različnih zračnih onesnaževalcev na dihalno funkcijo.

Trinajst astmatikov je dvakrat dnevno v obdobju enega meseca v zimskem času spremljalo svojo dihalno funkcijo. Maksimalni pretoka zraka ob izdihu (PEF; L/min) in prisiljeni ekspiratorni volumen v prvi sekundi izdiha (FEV<sub>1</sub>; L) smo spremljali s PEF-metri. Vitalna kapaciteta preiskovancev (VC; L) je bila izmerjena v laboratoriju ob koncu raziskovalnega obdobja. Respiratorni podatki so bili analizirani glede na okoljske koncentracije NO<sub>2</sub>, NO<sub>x</sub>, SO<sub>2</sub>, O<sub>3</sub>, in PM<sub>10</sub>, ki so bili spremljani v istem časovnem obdobju. Koncentracije nekaterih zračnih onesnaževalcev (PM<sub>10</sub> in NO<sub>x</sub>) so presegle kritično mejo v več dnevih raziskovalnega obdobja. Kljub temu s podatki s PEF-metrov (PEF, FEV<sub>1</sub> in FEV<sub>1</sub>/VC), ni bilo mogoče pokazati statistično značilnih odzivov dihalne funkcije na okoljske razmere ( $P > 0.05$ ).

Rezultati pričujoče raziskave kažejo, da samo-aplicirana uporaba PEF-metrov ne omogoča razločevanja med negativnimi učinki različnih zračnih onesnaževalcev na dihalno funkcijo niti pri najbolj občutljivih skupinah ljudi, niti v okolju, v katerem so koncentracije zračnih onesnaževalcev dovolj velike, da po obstoječih podatkih povečajo število hospitalizacij, in niti ob tako povečanih koncentracijah zračnih onesnaževalcev (trdnih delcev), da dokazano negativno učinkujejo na dihalno funkcijo človeka. Nadalje, rezultati raziskave kažejo, da ostaja ambulantno spremljanje dihalne funkcije, čeprav je logistično zahtevno, eden redkih, če ne edini zanesljivi vir podatkov o negativnih učinkih zračnih onesnaževalcev na dihalno funkcijo, tako pri zdravih posameznikih, kot pri respiratornih bolnikih.

## Conclusions

Quality of air affects respiratory function in humans, and respiratory conditions such as asthma are aggravated in a polluted environment. Nevertheless, data on the effects of different air pollutants on a day-to-day respiratory function are lacking, as due to different methodological approaches the issue in question has not been adequately addressed yet. The aim of this study was therefore to assess the applicability of self-administered monitoring of respiratory function in asthma patients in order to discern between negative health effects of different air pollutants

on respiratory function. Thirteen asthma patients measured their respiratory function twice daily over a one month winter period. They used PEF-meters to monitor peak expiratory flow (PEF; L/min) and forced expiratory volume in the first second of expiration (FEV<sub>1</sub>; L). Subjects' vital capacity (VC; L) was measured in a laboratory setting at the end of the measuring period. Respiratory data were evaluated in respect to the ambient concentrations of NO<sub>2</sub>, NO<sub>x</sub>, SO<sub>2</sub>, O<sub>3</sub>, and PM<sub>10</sub>, which were monitored during the same period. The concentrations of some air pollutants (PM<sub>10</sub> and NO<sub>x</sub>) exceeded the critical levels on several days during the study. PEF-meter data (PEF, FEV<sub>1</sub> and FEV<sub>1</sub>/VC), however, did not respond significantly to the ambient conditions (P>0.05).

The novel findings of the present study are that the self-administered PEF-meter monitoring can not discern between the negative health effects of different air pollutants even in the most susceptible groups of population, even in the environments where the concentrations of air pollutants are high enough to induce increased level of hospitalizations, and even with increased levels of the air

pollutant (particulate matter) that seems to cause the largest negative effects on human respiratory health. Furthermore, the results of the present study suggest that, although logistically demanding, ambulatory monitoring of respiratory function may be one of the few, if not the only reliable non-subjective source of information on the negative effects of air pollutants on respiratory function in both healthy individuals and respiratory patients.

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