THE ASSOCIATION OF EXPOSURE TO PM$_{10}$ WITH THE QUALITY OF LIFE IN ADULT ASTHMA PATIENTS

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Abstract
Objectives: Air pollution has become a critical environmental issue, which severely threatens the well-being of asthma patients. The quality of life of these patients, when exposed to air pollutants such as particulate matter 10 (PM$_{10}$), has been poorly studied. The current research examined the association between the concentration of PM$_{10}$ in the air and the quality of life of patients with asthma. Material and Methods: The study group consisted of 300 adult asthma patients treated in 2 allergy outpatient clinics in Kraków, who declared they would not leave the city in the 14-day study period. Daily concentrations of PM$_{10}$ from air monitoring stations were recorded over a period of 2 weeks, following which the patients filled out the standardized Asthma Quality of Life Questionnaire (AQLQ) regarding the monitored period to assess the total quality of life and its 4 domains (symptoms, limitation of activity, emotional functioning and environmental stimuli). Results: The average PM$_{10}$ exposure was 65.2 μg/m$^3$ and only 30% of the patients were exposed to values of ≤50 μg/m$^3$, i.e., the highest 24-h threshold value considered acceptable by the World Health Organization. The observed effect of an increased level of exposure to airborne PM$_{10}$ was associated with reduced scores in AQLQ from 0.40 at the medium level to 0.46 at the high level, in comparison to the low level. The total score of the asthma-related quality of life and its domains showed poorer outcomes as the concentration of PM$_{10}$ was increasing (every 0.08 pt per a 10 μg/m$^3$ increase). Conclusions: The increase in the concentration of PM$_{10}$ in the air impacts on the overall quality of life and its particular domains in people with exceptional predispositions, such as patients with bronchial asthma. Physicians taking care of asthma patients should pay special attention to the quality of patient’s life in response to the course and control of that illness, in relation to air pollution.

Key words: quality of life, asthma, air pollution, environmental health, PM$_{10}$, AQLQ

INTRODUCTION
Asthma is a common chronic non-communicable disease that affects as many as 339 million people of all ages in all parts of the world, and is increasing as people adopt modern lifestyles and live in a more urbanized environment. With a predicted increase in the proportion of the world’s population living in urban areas, there is likely to be a significant increase in the number of people with asthma worldwide over the next 2 decades. It is estimated that there may be an additional 100 million people with asthma by 2025 [1,2].

Asthma causes a substantial health burden to societies, reducing the quality of life, not only due to physical and psychological health, but also due to socio-economic out-
tic perspective, air pollutants cause oxidative injury to the airways, leading to inflammation, remodeling, and an increased risk of sensitization. Ambient levels of PM exacerbate existing asthma, especially by contributing to oxidative stress and allergic inflammation. There is also some evidence in support of PM as a cause of new cases of asthma in all populations, both children and adults [10].

The association between air pollution and asthma morbidity is well proven. Patients with asthma living in urban areas with high particulate pollution levels are more likely to have frequent asthma symptoms, asthma-related emergency department visits and hospitalizations than those living in areas with low pollution [11]. A systematic review and meta-analysis of cohort studies have proven the association between long-term exposure to air pollution and the incidence of asthma. Epidemiological studies have shown the impact of air pollutants on the increasing incidence of asthmatic symptoms and other adverse health effects [33]. These results suggest a deleterious effect of ambient air pollution on the incidence of asthma in adults [12–14].

There is epidemiological evidence that asthmatic symptoms can be worsened by an increase in PM$_{10}$. Exposure to air pollution contributes to asthma development, aggravations and exacerbations. Typical outdoor air pollutants that can trigger asthmatic symptoms include PM, with the severity of symptoms varying with the level of exposure. An increase in air pollutants (i.e., ozone and PM) raises the risk of both asthma development and its symptoms. Exposure to PM$_{10}$ can trigger or exacerbate asthma attacks, and control is essential for proper risk management and risk communication for the exposed asthma patients, especially those living in urban environments [15,16].

Asthma as a chronic disease could cause problems and difficulties in everyday functioning, in psychological and mental health related to the prevalence of anxiety and depression symptoms, which impact on subjective health. A lot of asthma patients have screened posi-
tive for depressive symptoms, and they report a worse quality of life during control visits in the outpatient allergy clinic, but it is not well confirmed by epidemiological studies. Therefore, the asthma-related quality of life (ARQL) measured by disease-specific scales should also be considered in the assessment of the social and health status of these patients [17,18]. Measurements of health among asthma patients, and also the effects of health care, must include not only an indication of changes in the frequency and severity of asthma or its symptoms, but also an estimation of self-assessed well-being. This should be assessed by measuring the changes in the patient’s quality of life. The health-related quality of life (HRQL) is a broad ranging concept affected in a complex way by the person’s own physical health, psychological state and mental health, the level of independence, social relationships, personal health beliefs, and the relationship to the salient features of their physical environment, including air pollutants [19].

It seems that verification of the quality of life of patients with bronchial asthma may be as useful as routine diagnostic methods that assess the disease course control.

In the asthma management and treatment process, HRQL is an important cause and outcome. In clinical practice, the assessment of the quality of life gives valuable information that can indicate areas in which the patient is most affected and needs support. It helps the physician in advising and making the best choices in patient care. Surveys have shown that differences in assessing the quality of life depend on the patient’s gender and place of residence, which may indicate the groups to which therapeutic and educational efforts should be directed [20,21]. It seems that in assessing the impact of air pollution on asthma control, an analysis of the patient’s quality of life should be prognostic in the treatment process.

The Asthma Quality of Life Questionnaire (AQLQ) is one of the most popular tools used to measure changes in HRQL over the course of treatment of this chronic disease. The use of this questionnaire has proven that a 0.5-pt decrease in the total score of AQLQ is a harbinger of an unfavorable course of the disease [22,23].

The HRQL in patients with asthma exposed for a long time to air pollutants, such as PM$_{10}$, in the urban environment has been poorly studied. There is little evidence that PM$_{10}$ reduces the quality of life in asthma patients, and could be related to the likelihood of the initial sensitization and induction of both the disease and syndromes. Only limited evidence exists on the impact of PM$_{10}$ on the HRQL of patients with bronchial diseases. There is no conclusive confirmation of the unambiguously detrimental connection between the quality of life and the concentration of PM$_{10}$ in the outdoor air. A pilot study conducted in 2013 in Kraków, Poland, did not significantly prove the association between PM$_{10}$ concentrations in the outdoor air and the quality of life of asthma patients [24]. Since the assessment of the impact of exposure to air pollutants, such as PM$_{10}$, on the quality of life of patients with bronchial asthma has not yet been the subject of a sufficient number of studies, this study gives an additional opportunity to assess and discuss this problem, in order to improve the well-being of adult asthma patients.

The objective of the study was to assess the impact of PM$_{10}$ exposure on the quality of life of patients with asthma living in Kraków, and to determine whether this exposure was related to changes in their subjective health. Results of this study should be important in planning preventive programs, risk communication and patients’ education to improve their well-being.

**MATERIAL AND METHODS**

A group of 349 adults patients (>18 years old) with moderate asthma, treated in 2 allergy outpatient clinics located in Kraków, were recruited for a 2-year survey conducted in 2013–2015. These patients were with partially controlled asthma, living in Kraków, and they declared not to leave the city during 14 days of the survey. Each pa-
tient was subjected to observation for 2 weeks. The surveyed patients with bronchial asthma were diagnosed according to the applicable standards, and they were under permanent control of a physician specializing in allergic diseases. In the study group, the recruited patients were under regular treatment in an outpatient allergy clinic. The group was fairly homogenous, i.e., it included patients with partially controlled asthma, well educated in asthma, recognizing and controlling symptoms of the disease, with a good knowledge of the types and use of medication, and regularly taking their medications.

The patient’s diary was used as a next tool. Its first part was completed by a physician specializing in allergic diseases during the first visit, including the date of birth, the place of residence, education, employment, workplace, way to work, smoking, keeping pets at home, and the type of treatment applied. The second part was used for everyday completion by the patient over 2 weeks of observation. Information for each patient was obtained that pertained, inter alia, to the number of hours and places they spent outside.

For each patient, a daily concentration of PM$_{10}$ on each day of observation, from all available air pollution monitoring stations of the Voivodeship Environmental Protection Inspectorate in Kraków, was reported and introduced to the database. Then, the patients were assigned to the station which was the closest to their declared outdoor place of stay in order to determine what concentration of the PM$_{10}$ fraction the patients were exposed to in the survey period. A selected address for each patient was verified and then entered into Google maps. The point measured on the map was determined with the command “measure distance.” Then, the locations for 3 stations of the Voivodeship Environmental Protection Inspectorate in Kraków were searched for and, at each, a point was designated by the command “distance to that place.” Data about PM$_{10}$ concentrations recorded by particular stations were obtained from the webpage of the Voivodeship Environmental Protection Inspectorate [25].

Overall, 349 patients were recruited, but 49 of them were eventually excluded due to leaving the city in the observation period, or not filling out/losing the patient’s diary. Finally, data of 300 patients aged 20–80 years (average 53±15.3 year) were used for the analysis, including 145 women (48.3%) and 155 men (51.7%).
The study was conducted with the approval of the Bioethics Committee of the Jagiellonian University (No. KBET/167/B/2012).

Statistical analyses
All characteristics were presented as means with standard deviations (SD) or frequency with percentage distribution, respectively to the measurement scale. Results regarding the quality of life of asthma patients (the AQLQ total score and its domains) were presented using the median with quartile distribution. The 2-week average PM$_{10}$ concentrations were additionally categorized with cut-offs at 30th and 70th percentiles (values of 50 and 80 $\mu$g/m$^3$, respectively). The first cut-off at 30th percentile corresponded to the 24-h threshold guided by the WHO, while 70th percentile of the highest PM$_{10}$ concentration was selected as the second cut-off. In addition, PM$_{10}$ concentrations were analyzed both as continuous and categorical variables. Potential confounders, such as gender, education, smoking, employment and keeping pets at home, were controlled in the analyses.

Associations between nominal variables were verified using the $\chi^2$ test or the exact Fisher test, as appropriate. Differences in the age of the patients exposed to different concentrations of PM$_{10}$ categories were tested using one-way ANOVA, whereby the equality of variances was verified with Levene’s test and normal distribution in subgroups with the Kolmogorov-Smirnov test. Outdoor temperature differences across the PM$_{10}$ categories were tested with the Kruskal-Wallis test as the assumption of variance homogeneity was not met in ANOVA. The non-parametric trend in AQLQ scores across the PM$_{10}$ categories was also checked [26,27].

The potential impact of airborne PM$_{10}$ on the quality of life of asthma patients (the AQLQ total score and its domains) was estimated in linear regression models with 2-week average PM$_{10}$ concentrations, both as categorical and continuous variables, with a unit of 10 $\mu$g/m$^3$, adjusted for age (in years), gender, university education (yes vs. no), smoking (yes vs. no) and the mean outdoor temperature in the measurement period. Among the variables considered as potential confounders, age, university education and temperature in the measurement period were associated with AQLQ and its domains, in a univariate analysis. Gender and smoking were a priori added to the models. The potential collinearity between PM$_{10}$ concentrations and temperature was checked using the variance inflation factor (VIF = 1.86), whose values allowed for putting those variables into 1 model.

The AQLQ itself was classified into 3 categories, i.e., a good quality of life (a score of 6–7), a reduced quality of life (a score of 4–5) and a poor quality of life (a score of 1–3), where “1” meant good and “3” meant poor. The PM$_{10}$ exposure level (both as a categorical and continuous variable) was used to assess its impact on the poor quality of life using ordered logistic regression models adjusted for age, age$^2$, gender, university education, active smoking and temperature in the measurement period. The same analyses were conducted for all 4 domains, as 4 separate tests. The proportional odds assumption was verified using the Brant test. The term “age$^2$” was added as the age variable did not comply with the proportional odds assumption.

All the tests were 2-tailed, and significance was set at $p < 0.05$. All the analyses were performed using STATATA/IC 13.1 software.

RESULTS
The average PM$_{10}$ exposure (a mean in the 2-week period) was 65.2 $\mu$g/m$^3$ (a range of 18.0–128.2 $\mu$g/m$^3$) and only 75 patients (30%) were exposed to values of $\leq$50 $\mu$g/m$^3$, i.e., the 24-h threshold recommended by WHO. The PM$_{10}$ measurements were equally distributed across seasons. There were no significant differences between the patients in the number of hours spent outdoors. In general, all the patients declared that they spent 2–3 h/day outdoors, on average.
The groups of patients with various average PM<sub>10</sub> exposure levels did not differ significantly in terms of gender, education (university vs. lower), employment or keeping pets at home. Nevertheless, the patients exposed to higher levels of PM<sub>10</sub> were older, and the studied subgroups were significantly different in terms of smoking (Table 1). Declining trends in the total and specific domains of AQLQ were observed with the increasing levels of airborne PM<sub>10</sub> exposure (Figure 1).

Table 2 shows the effects of categorized PM<sub>10</sub> exposure on the ARQL adjusted for potential confounders. The observed effect of an increased level of exposure to airborne PM<sub>10</sub> was associated with reduced scores in AQLQ from 0.40 at the medium level, as compared to the low level, to 0.46 at the high level, as compared to the low level. A similar decrease was observed in the symptoms and activity limitation domains with differences

<table>
<thead>
<tr>
<th>Variable</th>
<th>Respondents exposed to PM&lt;sub&gt;10&lt;/sub&gt; (N = 300)</th>
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<tbody>
<tr>
<td></td>
<td>total (≤50 μg/m&lt;sup&gt;3&lt;/sup&gt;) (N = 90)</td>
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<tr>
<td>Gender [n (%)]</td>
<td></td>
</tr>
<tr>
<td>women</td>
<td>145 (48.3)</td>
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<tr>
<td>men</td>
<td>155 (51.7)</td>
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<tr>
<td>Age [years] (M±SD)</td>
<td>53±15.3</td>
</tr>
<tr>
<td>Education [n (%)]</td>
<td>127 (42.3)</td>
</tr>
<tr>
<td>Active smoker [n (%)]</td>
<td>31 (10.3)</td>
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<tr>
<td>Current employment [n (%)]</td>
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</tr>
<tr>
<td>student</td>
<td>11 (3.7)</td>
</tr>
<tr>
<td>employed</td>
<td>172 (57.5)</td>
</tr>
<tr>
<td>unemployed</td>
<td>27 (9.0)</td>
</tr>
<tr>
<td>retired</td>
<td>89 (29.8)</td>
</tr>
<tr>
<td>missing</td>
<td>1 (0.3)</td>
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<tr>
<td>Keeping pets at home [n (%)]</td>
<td>133 (43.3)</td>
</tr>
<tr>
<td>Temperature [°C] (M±SD)</td>
<td>7.8±5.88</td>
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<tr>
<td>Asthma Quality of Life Questionnaire (Me (Q1–Q3))</td>
<td></td>
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<tr>
<td>total score</td>
<td>5 (4–6)</td>
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<tr>
<td>symptoms</td>
<td>6 (4–6)</td>
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<tr>
<td>limitation of activity</td>
<td>5 (4–6)</td>
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<td>emotional functioning</td>
<td>5 (4–6)</td>
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<tr>
<td>environmental stimuli</td>
<td>5 (4–5)</td>
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<sup>a</sup> Chi<sup>2</sup> test.
<sup>b</sup> ANOVA.
<sup>c</sup> Fisher exact test.
<sup>d</sup> Kruskal-Wallis test.
<sup>e</sup> Non-parametric test for trend.
Figure 1. Distribution of a) Asthma Quality of Life Questionnaire (AQLQ) total scores and its domains: b) symptoms, c) limitation of activity, d) emotional functioning, e) environmental stimuli according to PM$_{10}$ levels in adult asthma patients, Kraków, Poland, 2013–2015
of 0.51 and 0.46 at the high level of exposure, compared to the low level, respectively. In terms of emotional functioning and environmental stimuli, the quality of life did not differ significantly between the medium and high levels, compared to the low level of PM$_{10}$.

In order to assess the effects of PM$_{10}$ exposure on the risk of poorer outcomes in AQLQ (both total and domains scores), multivariable ordinal logistic regression models were applied. It was observed that the medium levels of PM$_{10}$ exposure were associated with an over 4-fold higher risk of poorer outcomes in AQLQ (OR = 4.29, 95% CI: 1.94–9.52), while the high exposure with a 6-fold higher risk (OR = 6.03, 95% CI: 2.45–14.88), compared to the low levels of PM$_{10}$ exposure. Similar results were seen in the symptoms domain. The risk of a poorer quality of life in terms of symptoms reached a value that was 3.76 (95% CI: 1.60–8.90) times higher for the medium level of PM$_{10}$ and 6.77 (95% CI: 2.56–17.92) for the high level, compared to the low one. The risk of poor outcomes in terms of limitation of activity was 2.81 times higher, in emotional functioning 2.37 times higher, and in environmental stimuli 2.45 times higher (95% CI: 1.67–6.26), but only in the high vs. low level of the PM$_{10}$ concentration (Table 2).

The effects of the continuously expressed mean of the 14-day level of PM$_{10}$ exposure on ARQL, adjusted for potential confounders, are shown in Table 3. All the domains of AQLQ, except environmental stimuli, present...
due to primary particles from combustion sources, mainly vehicles, with a contribution from secondary particles. So, the effects of coarse fractions, such as PM$_{10}$, cannot be excluded, and should be analyzed [30]. There is considerable evidence linking ambient particles measured as PM with an aerodynamic diameter of <10 μm (PM$_{10}$) to daily mortality and hospital admissions. The adverse health impact of PM$_{10}$ on the human respiratory system is well documented by studies showing associations between the concentrations of PM$_{10}$ and the quantified health effects, all cause and cause-specific mortality, morbidity and life expectancy, the incidence of respiratory diseases, and exacerbation of such diseases [31,32]. The overall epidemiological evidence suggests that these adverse health effects are dependent on both exposure concentrations and the length of exposure, and that long-term exposure has stronger and more persistent cumulative effects than short-term exposure [33]. In the study by Lu et al. [34] for a standardized increment in PM with aerodynamic diameters of <10 μm (PM$_{10}$), there is an excessive risk of mortality and hospital admissions. The adverse health impact of PM$_{10}$ on the human respiratory system is well documented by studies showing associations between the concentrations of PM$_{10}$ and the quantified health effects, all cause and cause-specific mortality, morbidity and life expectancy, the incidence of respiratory diseases, and exacerbation of such diseases [31,32]. The overall epidemiological evidence suggests that these adverse health effects are dependent on both exposure concentrations and the length of exposure, and that long-term exposure has stronger and more persistent cumulative effects than short-term exposure [33]. In the study by Lu et al. [34] for a standardized increment in PM with aerodynamic diameters of <10 μm (PM$_{10}$), there is an excessive risk of mortality and hospital admissions. Short-term exposure to PM$_{10}$ was found to be positively associated with increases in mortality for non-accidental causes, cardiovascular diseases and respiratory diseases [34]. The study by Zhang et al. [35] suggests that short-term exposure to outdoor air pollution may induce or exacerbate respiratory

<table>
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<th>AQLQ and its domains</th>
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<td>original scale*</td>
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<td></td>
<td>B</td>
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<tr>
<td>Total score</td>
<td>-0.08</td>
</tr>
<tr>
<td>Symptoms</td>
<td>-0.08</td>
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<tr>
<td>Limitation of activity</td>
<td>-0.08</td>
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<tr>
<td>Emotional functioning</td>
<td>-0.08</td>
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<tr>
<td>Environmental stimuli</td>
<td>-0.07</td>
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Explanations as in Table 2.

DISCUSSION

A lower quality of life is an important predictor of hospitalization and mortality among patients with chronic obstructive pulmonary disease (COPD) and asthma [28]. Coarse PM fractions can lead to adverse responses in the lungs, triggering processes leading to hospital admissions [29]. This suggests that special consideration should be given to studying and regulating coarse particles separately from fine particles. Results of the study by Anderson et al. [30] suggest that the active component of PM$_{10}$ resides mostly in the fine fraction, and this is mostly

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Table 3. Impact of the average 14-day concentrations of PM$_{10}$ (unit: 10 μg/m$^3$) on the asthma-related quality of life and the risk of a poorer quality of life in the study group of adult asthma patients, Kraków, Poland, 2013–2015

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Explanations as in Table 2.
Figure 2. Predicted probabilities (with 95% CI) of a “poor” asthma-related quality of life: a) Asthma Quality of Life Questionnaire (AQLQ) total scores and its domains; b) symptoms, c) limitation of activity, d) emotional functioning, e) environmental stimuli, according to PM$_{10}$ levels in adult asthma patients, Kraków, Poland, 2013–2015
diseases, upper respiratory tract infections, and chronic obstructive pulmonary diseases, leading to considerable medical expenditure of the affected patients. Despite the above-mentioned epidemiological studies, the body of research investigating the influence of air pollution, including various concentrations of PM$_{10}$, on the HRQL in asthma patients living in urban areas still appears insufficient. This study included an observation of the subjective health of patients with asthma, living in the urban environment of Kraków, when exposed to various concentrations of PM$_{10}$. The results revealed that increased levels of PM$_{10}$ were related to a lower quality of life in the surveyed patients. The total score of AQLQ and the scores for its 4 domains, such as symptoms, limitation of activity, emotional functioning, and environmental stimuli, as assessed in the study, were significantly reduced by increasing concentrations of this air pollutant. A decrease in HRQL was not only caused by exceeding the highest acceptable value of PM$_{10}$ exposure that was significant, but also by the degree of exceeding the norm as well. In situations where the norm was exceeded multiple times, all the domains of the patient’s quality of life showed significantly lower scores. Symptoms and limitation of activity were the most affected domains of the quality of life, which decreased with every single case of exceeding the norm. An assessment of subjective health in patients suffering from asthma has a great value in controlling the course of the disease although it is not always appreciated by physicians. The assessment of the patient’s well-being consists not only of the observation of symptoms but also of the information obtained from the patient concerning his/her quality of sleep, limitation of daily activities, and experiencing negative emotions related to health threats. Based on the results of this research, one can also talk about personal exposure to environmental factors, such as dust pollution by PM$_{10}$ fractions and their impact on the patient’s quality of life. Patients with asthma often report a considerable deterioration of their well-being and an intensification of symptoms during increased air pollution periods. Physicians may have problems with recognizing the reasons, i.e., whether these result from media coverage or exposure to air pollution. Popular diagnostic methods that assess the severity of bronchial asthma and its control, such as spirometry and the peak flow meter, do not ensure a full assessment of the patients’ health and well-being. There are certain health-related tools to measure the quality of life in asthma patients, including sleeping disorders, limitation of activity, emotional and mental health problems, anxiety, mood swings, and depression symptoms [19,36,37]. Once applied in this study, AQLQ proved to be a good standardized tool, sensitive in detecting changes in subjective health, and in measuring asthma-specific problems related to the patient’s quality of life [22,23]. The quality of life is an important issue for the contemporary understanding and assessment of the health status of patients with chronic diseases. Thus, it is important to identify and evaluate various factors which interact with and impact on their quality of life. The environmental hazards, such as air pollutants, belong to these factors. One of the goals in asthma management is to ensure an individual patient that the burden of his/her disease is controlled and limited, and that the patient is still able to have the best possible quality of life. Therefore, taking into consideration the outdoor air quality, including the PM$_{10}$ concentration levels, in planning preventive and educational measures addressed to these patients, in order to improve their well-being, seems both valuable and useful [19,38]. This study had some methodological limitations regarding the assessment of individual exposure to PM$_{10}$. The individual exposure to PM$_{10}$ was assessed based on records for the district of the city which was the closest to the patient’s place of residence. Mobile dust monitors seem to be a more precise method of evaluating PM$_{10}$ exposure. However, due to their technical limitations and difficulties in patients’ compliance, the use of mobile dust monitors was not possible.
CONCLUSIONS
The results suggest that in order to attain the management goals in asthma treatment, the quality of life should be measured next to lung function, especially in the period of air pollution and exposure to PM_{10} in the patient’s place of residence and in the surrounding urban environment. Brief and short self-administered instruments such as AQLQ provide a good opportunity to identify the patients at risk, who could benefit more from preventive interventions and patient education.

The study has shown that any exposure to increased values of PM_{10} impacts on both the overall quality of life and its specific domains among patients with bronchial asthma. Being aware that there is no minimal level of safe exposure to PM_{10}, physicians should pay special attention to their patients’ quality of life in relation to air pollution, especially in places where high or medium concentrations of PM_{10} occur during many days over long periods. This should be remembered by both physicians and patients.

It seems important to educate patients about the sources of air pollutants, such as PM_{10} factors influencing concentration values, and the possible strategies of reducing their individual exposure and susceptibility.

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