MANIFESTATION OF BRONCHIAL REACTIVITY IN THE WORKERS EXPOSED TO VARIOUS GASES AT THE GASIFICATION DEPARTMENT OF THE POWER PLANTS OF KOSOVO

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INTRODUCTION

Air pollution has been a long time problem, but only in recent decades, it has been given relevant importance. Although numerous studies in this field have been reported, questions regarding the relationship between pathological changes and the deleterious effect of nitrogen oxide (NOx) in the workplace remain [1].

The effect of NOx causes hyperemia, edema, and hypertrophy of the bronchial mucosa, increased secretion of the glandular apparatus, and bronchial muscular tonus, which certainly causes damage to the ventilatory function of the lungs [2]. Chronic obstructive pulmonary disease (COPD) is one of the leading causes of death and disability worldwide. According to the WHO forecasts, COPD will become the fourth leading cause of mortality in the world by 2030 [3].

The main gaseous pollutants released during the gasification process by power plants are carbon monoxide and dioxide (CO and CO2), nitrogen dioxide (NO2), ozone (O3), and sulfur dioxide (SO2). Other important gaseous pollutants are ammonia and volatile organic compounds (e.g., methane, benzene, and chlorofluorocarbon) [4].

Air pollution has become the leading environmental cause of premature deaths [5]. Global estimates show that external pollution and indoor environments cause 1.15 million deaths worldwide (corresponding to nearly 2% of the total death toll) [6].

The harmful effects of O3, NO2, and particulate matter (PM), as well as the pollution of the traffic, in general, on respiratory symptoms and functions, are well documented. In addition to strong epidemiological evidence of a link between air pollution and asthma, recent studies, especially in urban areas, have shown the role of pollutants in the development of asthma and COPD [7,8].

Air pollution can also increase the risk of pneumonia by impairing the function of alveolar pulmonary macrophages and epithelial cells [9].

The interaction of air pollutants and their effect on bronchial hyperreactivity indicate that SO2 and O3 increase bronchial hyperreactivity. SO2 concentrations up to 0.5 ppm and O3 concentrations up to 0.2 ppm have shown this effect [10].

Studies have shown that respiratory obstruction and increased bronchial reactivity are present in the early lives of children who develop asthma [11].

Exposure to air pollution has been associated with increased cardiovascular disease (CVD), and the biological mechanism of oxidative stress and systemic inflammation plays an important role. Heart rate, systolic blood pressure, diastolic blood pressure, white blood cells, red blood cells, hematocrit, and hemoglobin are several biomarkers of CVD-related oxidative stress and systemic inflammation [12].

Air pollution poses a potential risk for increased lung cancer risk. The link between exposure to air pollution at work and lung cancer risk indicates that long-term exposure to workplace environmental pollution can cause lung cancer [13].

Oral, pharyngeal, and laryngeal cancers are linked to smoking and may also be linked to other sources of air pollution [14].

Keywords: Power plants of Kosovo, Workers during the process of gasification, Respiratory health.

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In Kosovo, the air quality has severely deteriorated in developed urban areas, especially in industrial power plants. The greatest impact on the environment is caused by Kosovo Energy Corporation power plants (TC-A and TC-B), industries (releasing gases at workplaces, surface mines, cement plants, etc.), and small ones. Pollution, however, is also caused by individual heating facilities [15].

On this basis, the essential purpose of our study is to compare the bronchial reactivity between healthy individuals and those exposed to high levels of gas-polluted air pollution in the workplace during the gasification process in Kosovo’s power plants. The most direct air pollution damage is, undoubtedly, reflected in the respiratory system; thus, all our study is focused on this system.

METHODS

Our study focused on detailed research to validate all cases verified with increased bronchial reactivity and COPD by examining the respiratory system as per the Medical Research Council (MRC) [16] questionnaire for COPD by recording the degree of pulmonary function impairment. Hence, we have ascertained the phenomenon of increased bronchial reactivity and COPD by measuring the objective parameters of the workers of the gasification department in Kosovo power plants.

Research at workplaces with exposure to different gases shows large differences in the prevalence of increased bronchial reactivity and obstructive pulmonary disease such as different habits, smoking working conditions, and workplaces.

Smoker and nonsmoker plethysmography values of the examined workers indicate an increasing tendency with increasing age and duration of exposure to harmful NOx in the workplace. It is important to note that there are statistically significant differences between smokers and nonsmokers. There are also statistically significant differences in these values compared to those of individuals suffering from chronic obstructive bronchitis and those without chronic obstructive bronchitis, which reduces the relevance of the MRC questionnaire [17].

The individual selection was randomized, and all research participants were informed about the procedures and purpose of the research.

All individuals who were part of the research were informed in advance to not administer any drugs at least 48 h before the examination to avoid interference with our test results. Thus, pulmonary functions are determined in tranquility. Specific airway resistance (SRaw) and intrathoracic gas volume (ITGV) measurements were then measured using body plethysmography. Based on the SRaw and ITGV values obtained, airway resistance (Raw) and specific airway conductance (SGaw) were calculated.

In the control group, we also applied an additional procedure to verify that the selected individuals were indeed healthy and did not possess bronchial reactivity in the respiratory tract airways. We demonstrated this by performing a bronchoconstriction provocation test with bronchoconstriction agents such as histamine. In our case, we have used histamine for airway constriction.

Relatively large numbers of patients are due to the interaction of many factors such as poor climatic conditions in the workplace, smoking, poor socioeconomic conditions, and inadequate medical care.

The research was carried out in two working groups: The control group and the experimental group. The control group included 32 healthy people, whereas the experimental group included 57 workers, with permanent work in the gasification process department at power plants in Kastriot, Kosovo.

Plethysmography is sophisticated equipment that measures the volume of air in the sternum. This equipment consists of a plastic booth dedicated to the individual entering to perform the measurements. Subject blows into a turbine, which, like spirometry, records the volumes and airflows that this subject performs. Using plethysmography, we can record the pressure-pressure curves as well as the pressure-flow curves. Calculate the ratios between cabin air pressures and airway pressures, and the pressure-flow curve is calculated as the ratio between the pressure in the booth and airflow in the airways. The main parameters we measure using a plethysmograph are the volumes of gases in the sternum, namely, ITGV. Hence, the volumes of air in the respiratory system can be calculated as well as the amount of air remaining in the alveolar cysts. Other very important parameters that are measured using a plethysmograph are the airway resistance (Raw). These are calculated by the pressure difference at the beginning and end of the airway when the flow passing through them is 1 l/s. These parameters indicate the level of obstacle faced by the air when breathing (inspirium-expirium). SRaw is calculated as the ratio between Raw and ITGV. Another parameter is SGaw, which represents the ratio of 1/SRaw.

If changes in the respiratory system are found, an appropriate treatment with drugs is initiated immediately. Drugs used to treat bronchial asthma and COPD were:

a. Bronchodilator
b. Anti-inflammatory drugs

Bronchodilator drugs: Beta-2 adrenergic agonists, methylxanthines, and anticholinergic drugs [18-20].

Anti-inflammatory drugs: Corticosteroids, antihistamines, and antileukotrienes [21-23].

As medical treatment of these diseases is limited, the only option is to take preventive measures [24-26].

This scientific study is about the difference between the two research groups. On this basis, we used the t-test as a differentiation mechanism regarding the level of significance between the two groups: The control group and the experimental group (gasification department workers). In the control group, we have two sets of plethysmography values, which were obtained as a result of pre- and post-provocation histamine measurements of individuals in the same group.

Statistical processing involves the determination of mean (X), standard deviation (SD), and standard error of the mean, as well as testing of statistical differences between workers and control group. Statistical method ANOVA was used for analysis. The ANOVA test was used to analyze the differences between the results obtained within the group.

The parameters are summarized in MS-EXCEL. The results obtained were tested using Student’s t-test to ascertain significant differences between the experimental and control groups. Statistical software INSTAT-3 and STATISTICA for Windows have been used.

RESULTS

We believe that the tracheobronchial system showed no sensitivity in healthy individuals; therefore, they can be used as a special control group to evaluate the condition of the experimental group. General data of the individuals of both groups are given in Tables 1 and 2 and Diagram 1.

Initially, we present the plethysmography results of the control group to avoid suspicions about the presence of bronchial reactivity in this group. These results are expressed in two series, before and after provocation with histamine bronchoconstriction test (Fig 1).

Statistical calculation with a t-test shows no significant difference (p>0.1) in healthy individuals before and after provocation with histamine. The SRaw and SGaw between the control and experimental groups were significantly different (p<0.05). The results show a pronounced presence of bronchial reactivity of workers who worked on...
Table 1: General characteristics of the subjects

| Group       | n  | Age (years)          | Height (cm) | Weight (kg) | VC (L)  | FEV<sub>1</sub> (L) |
|-------------|----|----------------------|-------------|-------------|---------|---------------------|
| Control     | 32 | 22.8±0.45            | 175±0.37    | 68.5±1.8    | 3.80±0.12| 3.82±0.12           |
| Experimental| 55 | 38.54±1.33           | 175.67±0.54 | 70.80±1.22  | 3.90±0.11| 3.68±0.12           |

Generalized mean values for: Control group n=32, X±standard error of the mean; Experimental group n=55, X±standard error of the mean. The general starting values for VC (L) and FEV<sub>1</sub> (L) are also given. VC: Vital capacity expressed in liters, FEV<sub>1</sub>: Enhanced expiratory volume in the 1<sup>st</sup> s, expressed in liters.

Table 2: Body-plethysmography features of individuals involved in the study

| Group       | n  | Raw (kPa×s/L) | ITGV (L) | SRaw (kPa×s) | SGaw (kPa×s) |
|-------------|----|---------------|----------|--------------|--------------|
| Control     | 32 | 0.17±0.5      | 3.05±0.3 | 0.51±0.3     | 1.96±0.13    |
| Experimental| 55 | 1.15±0.7      | 3.55±0.6 | 4.08±0.8     | 2.24±0.18    |

Raw (kPa×s/L): ITGV (L): SRaw: Raw×ITGV; SGaw: 1/SRaw; Raw: Airway resistance expressed in kilopascal×s/liter; ITGV: Volume of intrathoracic gas expressed in liters; SRaw: Specific airway resistance, which is the relationship between resistance and the volume of intrathoracic gas; SGaw: Expresses the relationship between 1/specific resistances. SGaw can also be referred to as specific airway passage or conductivity.

DISCUSSION

There is comprehensive literature on the health effects of gas pollutants in the workplace, including numerous epidemiological and toxicological studies. In each special study, there is a relationship between gaseous air pollution in the workplace and one or more other factors, depending on the level of exposure, the health status of the workers exposed, and their age.

Scientific evidence on the health effects of gaseous air pollution at workplaces has been increasing in recent years. Some questions remain open, but many epidemiological studies have shown the importance of air pollution in workplaces as a risk factor for their mortality and morbidity [27]. The impact of air pollutants on the respiratory tract has been widely and consistently reported in recent years [28].

Air pollution has broad systemic effects on the human body, affecting both the respiratory and cardiovascular systems through multiple mechanisms, including oxidative stress, inflammation, and endothelial dysfunction [29]. The correlation between air pollution and cardiorespiratory diseases is a global concern [30].

Statistical calculation of SRaw and SGaw was performed for the control and experimental group, indicating a significant difference between these groups (p<0.05). The presence of emphasized bronchial reactivity in the workers working in the gasification department at the Kosovo power plants has been registered.

The difference between the experimental subgroups confirmed the favorable impact of smoking on exacerbating the negative respiratory effects of air pollution at the gasification department of the power plant (p<0.01).

Due to indoor ventilation, employees who work indoors are subjected to the elimination of gases released by the ventilation system during the gasification process. The workers were less affected from bronchial reactivity than the residents who live near power plants as these workers leave the environment where they work after 8 h of work and travel farther away from the power plants where they live (up to 50 km away) with less air pollution or no pollution at all. This may also be the cause for lower susceptibility of bronchial reactivity in these workers (p<0.05) compared to residents who live near the power plants and are, therefore, constantly exposed to irritants for 24 h, which is manifested with emphasized bronchial reactivity (p<0.01), greatly endangering health [31].

In addition to cardiopulmonary morbidity and mortality, recent studies have found that air pollution is also associated with health effects, such as dementia, structural brain changes in children, cognitive impairments, and diabetes [32].

There is strong evidence that air pollution with different gases in the workplace exacerbates asthma morbidity and mortality in individuals with the disease and there is some evidence that air pollution with gases may affect asthma prevalence [33]. Children are at greater risk of developing asthma when exposed to higher concentrations of air pollutants [34].

A study carried out in continental France found that 9% of total mortality in France is due to anthropogenic PM<sub>2.5</sub> [35]. The calculated total mortality for PM<sub>2.5</sub> is higher than that due to PM<sub>10</sub> [36].

The American Heart Association published a "Scientific Statement" stating that "there is a correlation between PM<sub>2.5</sub> exposure and mortality - cardiovascular morbidity" [37].

In addition, a new health outcome in the United States showed that the incidence of lung cancer is related to exposure to PM<sub>2.5</sub> particles [38].
In the American Cancer Society study, the incidence of lung cancer increased by 8% for 10 μg·m⁻³ increases in PM₂.₅ levels measured between cities [39]. PM₂.₅ was also associated with decreased parathyroid hormone in exposed individuals [40].

In addition to CO and CO₂, other pollutants, including NO₂ and O₃, were also associated with workplace mortality [41]. SO₂ has also been associated with increased mortality and morbidity. Significant effects of SO₂ and cardiopulmonary mortality have been observed [42].

NO₂ concentrations are related to mortality rate and mortality is caused by cardiovascular and respiratory morbidity and lung cancer [43].

A study was conducted to ascertain the link between O₃ and mortality in 21 European cities and found that the 10 ppb increase in the 2-day 8-h O₃ average was 0.31% (95% confidence interval [CI]: 0.17%, 0.52%), 1.13% (95% CI: 0.74%, 1.51%), and 0.46% (95% CI: 0.22%, 0.73%) increase in total respiratory and cardiovascular mortality in summer [45].

Acute and inflammatory changes caused by sudden exposure to high concentrations of irritating gas or steam are associated with bronchoconstriction, manifested by dyspnea, and chest pain. Chronic inflammatory changes caused by constant or intermittent exposure to moderately increased concentrations of soluble (HCl, SO₂, NH₃, and the like) or less soluble irritants (O₃, NOₓ etc.) have been associated with variable airway reactivity. Different gases through their chlorides and carbon-containing compounds cause changes throughout the bronchial trunk and reversible bronchoconstriction that can be measured. The markedly increased trunk reactivity of the airways causes exposure to proteolytic enzymes. The inflammatory process associated with COPD is characterized by an increased number...
of activated alveolar macrophages, neutrophils, cytotoxic CD8+ T lymphocytes, B lymphocytes, CD4+ T lymphocytes, and the release of numerous inflammatory mediators (leukotrienes, cytokines, growth factors, chemokines, and oxidants). Chronic inflammation leads to a reconfiguration of small airways with lumen blockage due to increased mucus production and thickening of the airway walls due to edema and collagen formation, causing fibrosis and constriction [46].

Based on the research and results obtained, it can be concluded that continuous monitoring of the respiratory tract is necessary, especially because of the presence of an inflammatory reaction that occurs after prolonged exposure to the pollutant. These workers must undergo systemic visits and tracheobronchial provocation tests with histamine or methacholine, which is valuable for timely therapeutic intervention. Subjects with bronchial hyperreactivity show a special preference for the rapid onset of permanent changes in the respiratory tract, which is why they should be distinguished as subjects at higher risk.

CONCLUSION

These studies indicate undoubtedly that long-term exposure of workers at the gasification department of power plants in Kosovo to harmful gases such as carbon monoxide, carbon dioxide, sulfur dioxide, sulfur trioxide, NOx, soot, and other pollutants in the workplace cause the occurrence of respiratory tract inflammation with the release of chemical mediators manifested by bronchial hyper-reactivity.

Determining the value of the reaction against the inhalation of histamine hydrochloride resulted at a higher percentage of increase of bronchial reactivity, which is expressed with the increase in specific resistance (SRaw) and SGaw in groups exposed to toxic gases (p<0.05) in the airways compared to the control group (p>0.1).

Experimental group, as an entirety, and subgroup of smokers indicate about significant changes in the SRaw and SGaw (p<0.05) in the airways compared to the control group (p<0.1). The difference between the experimental subgroups confirmed the favorable impact of smoking in exacerbating the negative respiratory effects of air pollution (p<0.01).

Based on the research and the results obtained, it can be concluded that permanent monitoring of the respiratory tract is required, especially from the point of view of the presence of inflammatory reaction following prolonged exposure to environmental pollutants. People with increased bronchial hyperreactivity indicate a special predisposition for a quick manifestation of permanent changes in the respiratory tract, and this is just the reason of why they can be singled out as subjects with the highest riskiness.

AUTHORS’ CONTRIBUTIONS

PI conducted the literature search and determined studies for exclusion and inclusion. All extracted data from the included studies, performed the meta-analysis, and drafted the manuscript. HI conceived the idea for the study, designed the study, and critically revised the manuscript for important intellectual content. All authors contributed to data analysis, drafting and revising the article, gave final approval of the version to be published, and agree to be accountable for all aspects of the work.

CONFLICTS OF INTEREST

No conflicts of interest were declared by the authors.

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