

# IMPACT OF AIR POLLUTION WITH PM<sub>10</sub> ON PRIMARY HEALTH CARE CONSULTATIONS FOR RESPIRATORY DISEASES IN CHILDREN IN ZASAVJE, SLOVENIA: A TIME-TREND STUDY

## VPLIV ONESNAŽENOSTI ZRAKA S PRAŠNIMI DELCI PM<sub>10</sub> NA ŠTEVILO OBISKOV V PRIMARNEM ZDRAVSTVENEM VARSTVU ZARADI BOLEZNI DIHAL PRI OTROCIH V ZASAVJU: ŠTUDIJA ČASOVNEGA TRENTA

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### Abstract

**Aim:** The aim of our study was to assess the temporal association between the number of consultations in the primary health care unit due to respiratory diseases in children and the level of particular matter of 10 micrometres in diameter (PM<sub>10</sub>) pollution in the Zasavje region.

**Methods:** A time-trend ecological study was carried out for the period between 1 January 2006 and 31 December 2011. The daily number of first consultations for respiratory diseases among children in the Zasavje region was observed as the outcome. Poisson regression analysis was used to investigate the association between the observed outcome and the daily PM<sub>10</sub> concentrations, adjusted to other covariates.

**Results:** The results showed that the daily number of first consultations were highly significantly associated with the daily concentrations of PM<sub>10</sub> in the Zagorje ( $p < 0.001$ ) and Trbovlje ( $p < 0.001$ ) municipalities. In the Hrastnik municipality, a significant association was not observed in all models.

**Conclusions:** It can be concluded that evidence of association between the daily PM<sub>10</sub> concentration and the daily number of first consultations for respiratory diseases among children exists, indicating that there is still a need for public health activities in the sense of reduction of harmful environmental factors in the region. Additionally, on the basis of these results, it can be assumed that with some improvements linkage of existing health and environmental data in Slovenia in general could be feasible in identifying a grounded need for future public health action.

**Key words:** outdoor air pollution, PM<sub>10</sub>, respiratory diseases, children, the Zasavje region, time-trend study

Izvirni znanstveni članek  
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### Izveček

**Namen:** Namen študije je bil oceniti časovno povezanost med številom obiskov v primarnem zdravstvenem varstvu zaradi boleznih dihal pri otrocih in prašnimi delci premera 10 mikrometrov (PM<sub>10</sub>) v Zasavju.

**Metode:** Ekološka študija časovnega trenda je bila izvedena za obdobje od 1. januarja 2006 do 31. decembra 2011. Opazovani izid je bil dnevno število prvih obiskov zaradi boleznih dihal pri otrocih v Zasavju. Za oceno povezanosti med opazovanim izidom in dnevnimi koncentracijami PM<sub>10</sub> standardizirano na preostale pojasnjevalne dejavnike, je bila uporabljena Poissonova regresijska analiza.

**Rezultati:** Rezultati so pokazali močno statistično povezanost med dnevnim številom prvih obiskov in dnevnimi koncentracijami PM<sub>10</sub> v občinah Zagorje ( $p < 0,001$ ) in Trbovlje ( $p < 0,001$ ). V občini Hrastnik nismo opazili značilne povezanosti pri vseh modelih.

**Zaključek:** Sklenemo lahko, da v Zasavju obstaja povezanost med boleznimi dihal pri otrocih in koncentracijo PM<sub>10</sub>, kar kaže na to, da je v tej slovenski regiji še vedno prisotna potreba po javnozdravstvenih ukrepih v smislu

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*zmanjševanja škodljivih okoljskih dejavnikov. Dodatno lahko na podlagi rezultatov študije sklepamo, da bi bilo lahko v prihodnje v Sloveniji povezovanje okoljskih in zdravstvenih podatkov z nekaterimi izboljšavami uporabno pri ocenjevanju utemeljenih potreb za javnozdravstveno ukrepanje.*

**Ključne besede:** onesnaženost zunanega zraka, PM<sub>10</sub>, bolezni dihal, otroci, Zasavje, študija časovnih trendov

## 1 INTRODUCTION

Given the scale and widespread distribution of outdoor air pollution, this negative health determinant is considered as one of the major public health concerns of today (1-8). It has been proven to be associated with a variety of adverse health outcomes, however most of the recent evidence focuses mainly on respiratory and cardiovascular effects (1, 6, 8-15). The most susceptible population group to respiratory effects of air pollution is children (16-18). One of the most important reasons is that children have a larger lung surface area per kilogram of body weight than adults and, under normal breathing, breathe in 50% more air per kilogram of body weight than adults. The other is that they have increased exposure to many air pollutants because of higher minute ventilation and higher levels of outdoor physical activity (16-20).

A variety of pollutants can be found in the outdoor air, however one of the most important is particulate matter (PM) of 10 micrometres in diameter (PM<sub>10</sub>). It can penetrate deep into the bronchial tree and trigger respiratory symptoms. Several studies have consistently demonstrated an association between emergency department visits or hospital admissions due to respiratory diseases and concentration of PM<sub>10</sub> (1, 6, 8). In the last decade, many studies have applied time-series methods to study the association between air pollution with PM<sub>10</sub> and its health effects (21-23). These studies mostly rely on routinely available outdoor air pollution and health registry data (24). Many of them have indicated a positive association between a short-term variation in outdoor levels of PM<sub>10</sub> and daily frequencies of events (e.g. primary health care visits, hospital admissions, deaths) for respiratory diseases (1, 4-6, 8).

In regards to air pollution, Slovenia is no exception. One of the most polluted areas in the country is the Zasavje region (or Zasavje) (Figure 1), which is located in the central part of the country and consists of municipalities Zagorje ob Savi (or Zagorje), Trbovlje and Hrastnik (Figure 2) (25, 26). In this region, there are three narrow

valleys located more or less perpendicular to the larger Sava river valley. The main characteristics of Zasavje are coal mines and various different kinds of heavy industry (cement, glass, chemical, etc.) (Figure 2) that were established in Zasavje in the past. Among others, one of the biggest steam power plants in Slovenia is located there, having the highest chimney in Europe (25). This is due to the proximity to the source of energy and the fact that the Sava river valley with the railway line for decades represented one of main transport pathways of Slovenia. Most industrial plants considered as the largest emitters in the region are situated in the Sava river valley or at the intersection of the Zagorje, Trbovlje and Hrastnik valleys and the Sava river valley (Figure 2). Ever since the establishment, this industry has had a huge impact on the outdoor air pollution. The most important pollution in the past has been due to sulphur dioxide (SO<sub>2</sub>) and PM<sub>10</sub>. However, according to the report of the Slovenian Environmental Agency (SEA), the situation in SO<sub>2</sub> has greatly improved, and the national legally defined maximum values are exceeded only occasionally, while measurements of PM<sub>10</sub> and ozone (O<sub>3</sub>) levels show that they constantly exceed the national legally defined maximum values at the existing monitoring stations in Zasavje (26, 27). On the bottoms of the valleys, the temperature inversions are also frequently present in winter and autumn. Unfortunately, only few studies have studied the association between outdoor air pollution and the health of the Zasavje population in the past (28, 29). In the last few years, there have been some new studies carried out. The first study was the study of Eržen et al. (30), which indicated the association between the level of outdoor air pollution and prevalence of chronic respiratory diseases in the Zagorje municipality by using a rough assessment of level of pollution in different parts of the municipality. In the study of Kukec et al. (31), researchers upgraded these research methods with multivariate statistical analyses in all three municipalities in the Zasavje. However, further research is needed in order to bring up evidence to prove the health impact due to the environmental factors.



Figure 1. Location of the Zasavje region in Slovenia.  
Slika 1. Lokacija regije Zasavje v Sloveniji.

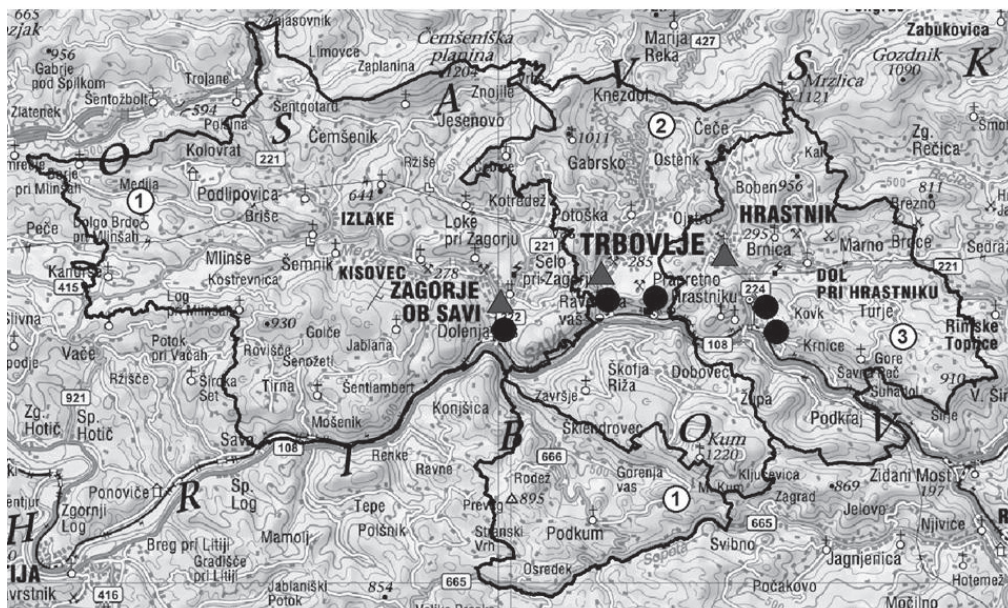


Figure 2. Zasavje region, Slovenia, map with locations of main point sources of outdoor air pollution in the region. Legend: ① = Zagorje municipality; ② = Trbovlje municipality; ③ = Hrastnik municipality; ● = location of cement, steam power, chemical, and glass plants; ▲ = location of environmental and meteorological data measuring stations; ■ = houses/settlements (font size of the settlement name indicate the rough size of the settlement).

Slika 2. Lokacija najpomembnejših točkovnih virov onesnaževanja zunanjega zraka v regiji Zasavje, Slovenija. Legenda: ① = občina Zagorje; ② = občina Trbovlje; ③ = občina Hrastnik; ● = lokacija kemične in cementne industrije, termoelektrarne in steklarne; ▲ = lokacija ekoloških in meteoroloških merilnih postaj; ■ = hiše/naselja (velikost črk naselja prikazuje okvirno velikost naselja)



The so-called linkage methods for environment and health analysis were proposed more than a decade ago by the World Health Organization (WHO) (24, 32). They belong to a wider group of epidemiological methods called ecological studies – a study design in which the relationships between environment and health are studied on population level, by analysing spatial and/or temporal variations in exposure and health outcome (33-35).

The aim of our study was to assess the feasibility of linkage of existing health and environmental data in Zasavje in identifying a grounded need for public health action. The specific goal of the study was to assess the temporal association between the number of consultations in primary health care units due to respiratory diseases in children and the level of PM<sub>10</sub> pollution in Zasavje. The hypothesis was that a positive temporal relationship between the daily number of first consultations for respiratory diseases and the daily concentration of PM<sub>10</sub> exists in the municipalities of Zasavje. The study was part of larger project that was performed at the Chair of Public Health, Faculty of Medicine, University of Ljubljana in collaboration with environmental and health experts (36).

## 2 METHODS

### 2.1 Study design and study population

The study design was an ecological time-trend study (34). The unit of observation was a single day of the observed period. The study population consisted of all children, aged 1-11 years, residing permanently in Zasavje, who visited the Community Health Centres in (CHC) Zagorje, Trbovlje or Hrastnik due to selected respiratory diseases between 1 January 2006 and 31 December 2011 (37). Altogether, 2,191 days were observed (365 in the years 2006, 2007, 2009, 2010 and 2011 and 366 in the year 2008).

### 2.2 Data acquisition

#### 2.2.1 Health data

Routinely collected health data were obtained from the health information systems of the CHCs of Zagorje, Trbovlje and Hrastnik. Daily numbers of first consultations due to the following diagnoses according to the WHO International Classification of Diseases, version 10 (ICD-10), were obtained: J00-J06 (acute upper respiratory tract infection), J10-J18 (influenza and pneumonia), J20-J22 (other acute lower respiratory tract infection), J30-J32 (other diseases of upper respiratory

tract) and J40-J46 (chronic lower respiratory tract disease).

#### 2.2.2 Environmental data

Immission data on daily PM<sub>10</sub> concentrations could be obtained at three fixed measuring stations in Zasavje, located in Zagorje, Trbovlje and Hrastnik (Figure 2), which are a part of the National automated network for monitoring air quality operated by SEA. In Zagorje and Trbovlje, the data could be obtained for the entire observation period, while in Hrastnik PM<sub>10</sub> measurements were only just started on 1 January 2010. Consequently, the observed period in Hrastnik only lasted between 1 January 2010 and 31 December 2011.

Immission data on other important outdoor air pollutants (co-pollutants) in Zasavje: SO<sub>2</sub>, O<sub>3</sub> and nitrogen dioxide (NO<sub>2</sub>), were obtained as well. Data on daily SO<sub>2</sub> and O<sub>3</sub> concentrations could be obtained at all the measuring stations for the entire observation period, while data on daily NO<sub>2</sub> concentration were only available at the Trbovlje measuring station.

Daily data on meteorological factors: air temperature and relative humidity, could be obtained at all the measuring stations for the entire observation period.

#### 2.2.3 Data on seasonal factors

In the analysis, the following seasonal factors were considered: season of the year (spring, summer, autumn, winter), work day (yes/no), holiday (yes/no) and influenza season (yes/no). Data on influenza season were obtained from annual reports (Epidemiological surveillance of communicable diseases in Slovenia) of the National Institute of Public Health of the Republic of Slovenia (38).

### 2.3 Statistical analysis

#### 2.3.1 Data description

The distributions of health and environmental data were statistically described by non-parametric typical statistical values (mean, standard deviation, minimum, maximum, 1<sup>st</sup>, 2<sup>nd</sup> and 3<sup>rd</sup> quartile). The temporal patterns of health and environmental data were presented by using sequence plots (33, 34).

#### 2.3.2 Relationship analysis

In the relationship analysis, the daily number of first consultations for all respiratory diseases was considered as the observed outcome, daily concentration of PM<sub>10</sub> (24-hr average PM<sub>10</sub> concentration) as the

explanatory factor and co-pollutants (24-hr average SO<sub>2</sub> concentration, 8-hr maximum average O<sub>3</sub> concentration), and 24-hr average NO<sub>2</sub> concentration), meteorological and seasonal factors were considered as covariates. In all pollutants, like in other similar studies (11, 39-44), lags from zero up to five days from exposure to the consultation day (lag 0, lag 1, lag 2, lag 3, lag 4 and lag 5 days respectively) were examined to determine the amount of time between exposure and effect. The association between the observed outcome, explanatory factor and covariates was analysed using Poisson regression models (45, 46). The modelling procedure was performed in three stages. In the first stage, univariate models for lags 0-5 days were built by relating the observed outcome to only the explanatory factor. In the second stage, single-pollutant models for lags 0-5 days were built by adding the explanatory factor to a core covariate model that included seasonal (season of the year, work day/weekend day, holiday/non-holiday day and influenza season) and meteorological factors (air temperature and relative humidity). In this stage, the best lags for the explanatory factor and co-pollutants were defined. In the third stage, multi-pollutant models were built by including best lags of the explanatory factor and co-pollutants along with the meteorological and seasonal factors. In order to achieve comparability of results for the Zagorje and Trbovlje municipalities, multi-pollutant models that only included SO<sub>2</sub> and O<sub>3</sub> as co-pollutants were built first. Afterwards, an additional model that

included SO<sub>2</sub>, O<sub>3</sub> and NO<sub>2</sub> was defined only for the Trbovlje municipality. The multi-pollutant model for the Hrastnik municipality was not defined due to the short observation time. The interpretable end result was the incidence rate ratio (IRR) (47). It was presented together with its 95% confidence interval (CI). P-value of 0.05 or less was considered as statistically significant in all the statistical tests.

All statistical analyses were carried out by using SPSS 18.0 software (SPSS Inc., Chicago, IL, USA).

The study protocol was approved by the National Medical Ethics Committee of the Republic of Slovenia.

### 3 RESULTS

#### 3.1 Data description

Complete health data were available at all three Zasavje CHCs for all 2,191 days of the study period. In the Zagorje municipality, there were 128/2,191 (5.8%) days with no first consultations for all respiratory diseases, and in the municipalities of Trbovlje and Hrastnik, there were 577/2,191 (26.3%) and 685/2,191 (31.3%) such days respectively. A statistical description of the distribution of the daily number of first consultations for all respiratory diseases is presented in Table 1. Temporal patterns of the daily number of first consultations for the observed outcome are presented in Figures 3a, 4a, and 5a. In all three municipalities, the observed outcome was the highest in winter months (from December to February).

Table 1. Descriptive statistics for the daily number of first consultations for respiratory diseases among children and environmental data in the Zasavje region, Slovenia, for 2,191 days between January 1, 2006 and December 31, 2011.

Tabela 1. Opisna statistika dnevnega števila prvih obiskov zaradi bolezni dihal pri otrocih in okoljskih podatkov v regiji Zasavje, Slovenija, za 2.191 dni med 1. januarjem 2006 in 31. decembrom 2011.

	Typical value/Tipična vrednost						
	Mean	SD	Min	Q <sub>1</sub>	Q <sub>2</sub>	Q <sub>3</sub>	Max
	<i>Povprečje</i>	<i>SD</i>	<i>Min</i>	<i>Q<sub>1</sub></i>	<i>Q<sub>2</sub></i>	<i>Q<sub>3</sub></i>	<i>Max</i>
<b>Zagorje municipality/Občina Zagorje</b>							
Number of consultations due to respiratory diseases Število obiskov zaradi bolezni dihal	4.99	4.0	0.0	2.0	4.0	7.0	29.0
PM <sub>10</sub> 24-hr average concentration (µg/m <sup>3</sup> ) PM <sub>10</sub> 24-urna povprečna koncentracija (µg/m <sup>3</sup> )	40.3	23.9	4.3	23.5	33.8	49.6	231.1
SO <sub>2</sub> 24-hr average concentration (µg/m <sup>3</sup> ) SO <sub>2</sub> 24-urna povprečna koncentracija (µg/m <sup>3</sup> )	5.8	4.6	0.0	2.7	5.1	7.8	47.0
O <sub>3</sub> maximum 8-hr average concentration (µg/m <sup>3</sup> ) O <sub>3</sub> maksimalna 8-urna povprečna koncentracija (µg/m <sup>3</sup> )	59.6	31.1	1.2	34.6	58.9	82.3	149.3
Temperature 24-hr average (°C) 24-urna povprečna temperatura (°C)	11.4	8.0	-10.2	4.9	11.9	18.1	28.3
Relative humidity 24-hr average (%) 24-urna povprečna relativna vlažnost (%)	73.4	14.6	3.0	65.7	75.1	83.9	98.4
<b>Trbovlje municipality/Občina Trbovlje</b>							
Number of consultations due to respiratory diseases Število obiskov zaradi bolezni dihal	4.91	4.9	0.0	0.0	4.0	8.0	33.0
PM <sub>10</sub> 24-hr average concentration (µg/m <sup>3</sup> ) PM <sub>10</sub> 24-urna povprečna koncentracija (µg/m <sup>3</sup> )	36.3	21.8	1.3	21.2	30.2	46.0	188.6
SO <sub>2</sub> 24-hr average concentration (µg/m <sup>3</sup> ) SO <sub>2</sub> 24-urna povprečna koncentracija (µg/m <sup>3</sup> )	4.0	4.5	0.0	1.0	3.0	5.6	43.0
O <sub>3</sub> maximum 8-hr average concentration (µg/m <sup>3</sup> ) O <sub>3</sub> maksimalna 8-urna povprečna koncentracija (µg/m <sup>3</sup> )	67.3	33.6	1.6	42.5	66.5	91.8	164.0
NO <sub>2</sub> 24-hr average concentration (µg/m <sup>3</sup> ) NO <sub>2</sub> 24-urna povprečna koncentracija (µg/m <sup>3</sup> )	20.0	9.1	1.7	19.6	18.9	25.0	60.5
Temperature 24-hr average (°C) 24-urna povprečna temperatura (°C)	11.3	8.0	-10.5	4.9	11.9	17.8	27.9
Relative humidity 24-hr average (%) 24-urna povprečna relativna vlažnost (%)	76.8	11.7	34.2	68.7	77.7	85.7	99.2
<b>Hrastnik municipality/Občina Hrastnik</b>							
Number of consultations due to respiratory diseases Število obiskov zaradi bolezni dihal	2.61	3.0	0.0	0.0	2.0	4.0	23.0
PM <sub>10</sub> 24-hr average concentration (µg/m <sup>3</sup> ) PM <sub>10</sub> 24-urna povprečna koncentracija (µg/m <sup>3</sup> )	28.0	16.2	3.1	17.3	23.9	34.7	123.4
SO <sub>2</sub> 24-hr average concentration (µg/m <sup>3</sup> ) SO <sub>2</sub> 24-urna povprečna koncentracija (µg/m <sup>3</sup> )	5.8	3.9	0.0	3.0	5.0	8.0	44.0
O <sub>3</sub> maximum 8-hr average concentration (µg/m <sup>3</sup> ) O <sub>3</sub> maksimalna 8-urna povprečna koncentracija (µg/m <sup>3</sup> )	73.5	33.1	3.8	48.3	72.8	97.0	178.1

Temperature 24-hr average (°C) <i>24-urna povprečna temperatura (°C)</i>	11.1	8.0	-10.4	4.8	11.7	17.6	28.1
Relative humidity 24-hr average (%) <i>24-urna povprečna relativna vlažnost (%)</i>	76.2	11.9	34.2	67.9	76.7	85.5	99.9

Legend/*Legenda*: SD -standard deviation/*standardni odklon*; Q<sub>1</sub> - the first quartile/*prvi kvartil*; Q<sub>2</sub> -the second quartile/*drugi kvartil*; Q<sub>3</sub> -the third quartile/*tretji kvartil*; Min/*Min* -minimum/*najnižja vrednost*; Max/*Max* -maximum/*najvišja vrednost*, \* = data available only for the period from January 1, 2010 to December 31, 2011/*podatki so dostopni le za obdobje od 1. januarja 2010 do 31. decembra 2011*

Complete data for the daily concentration of PM<sub>10</sub> were available for 2,135/2,191 (97.4%) days in the Zagorje municipality, 1,985/2,191 (90.6%) days in the Trbovlje municipality, and 730/730 (100%) days in the Hrastnik municipality. Statistical description of the distribution of daily PM<sub>10</sub> concentration is presented in Table 1. Temporal patterns of daily PM<sub>10</sub> concentration at all three measuring stations are presented in Figures 3b, 4b, and 5b. The highest daily PM<sub>10</sub> concentrations were observed in the months from November to February at all three measuring stations.

Complete data for the daily concentration of SO<sub>2</sub> were available for 2,080/2,191 (94.9%) days in the Zagorje municipality, 2,159/2,191 (98.5%) days in the Trbovlje municipality, and 2,132/2,191 (97.3%) days in the Hrastnik municipality. Complete data for the daily concentration of O<sub>3</sub> were available for 2,131/2,191 (97.3%) days in the Zagorje municipality, 2,101/2,191 (95.9%) days in the Trbovlje municipality, and 2,120/2,191 (96.7%) days in the Hrastnik municipality. Complete data for the daily concentration of NO<sub>2</sub> were available for 2,067/2,191 (94.3%) days in the Trbovlje

municipality. Statistical description of the distribution of daily co-pollutants (SO<sub>2</sub>, O<sub>3</sub> and NO<sub>2</sub>) concentration is presented in Table 1.

Complete data for the daily meteorological factors (air temperature and relative humidity) were available for 2,189/2,191 (99.9%) days in the Zagorje municipality, 2,184/2,191 (99.7%) days in the Trbovlje municipality, and 2,170/2,191 (99.0%) days in the Hrastnik municipality. Statistical description of the distribution of daily meteorological factors is presented in Table 1. In the observed period there were in total 541/2,191 (24.7%) winter days, 552/2,191 (25.2%) spring days, 552/2,191 (25.2%) summer days and 546/2,191 (24.9%) autumn days. There were also in total 626/2,191 (28.6%) weekend days and 1,565/2,191 (71.4%) workdays. In the observed period there were in total 1,588/2,191 (72.5%) non-holiday days and 603/2,191 (27.5%) holiday days (school holidays and work-free days). Also, there were in total 1,505/2,191 (68.7%) days without an influenza epidemic and 686/2,191 (31.3%) days with an influenza epidemic.

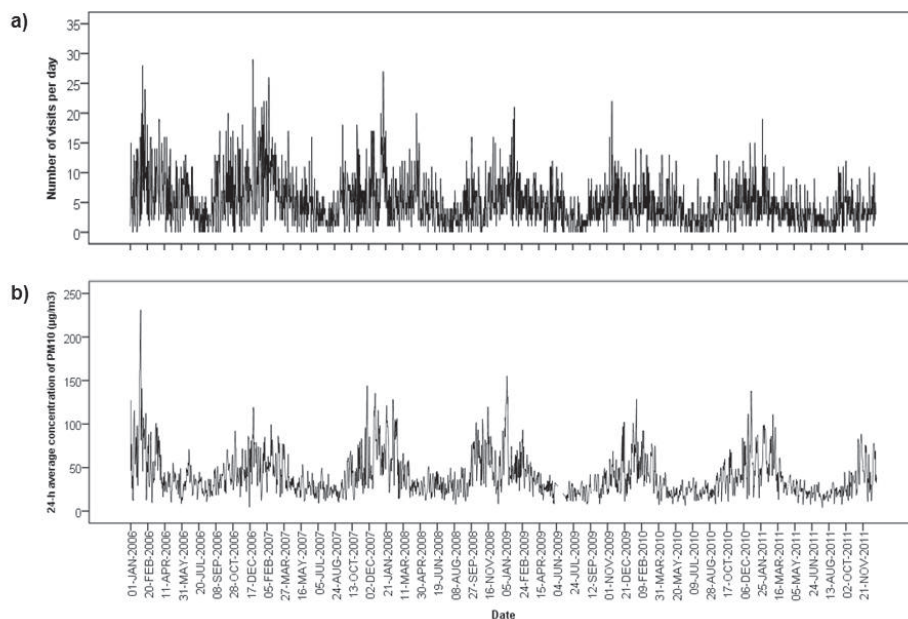


Figure 3. Temporal pattern of: a) daily number of consultations for respiratory diseases in children, and b) daily 24-hr average concentration of  $PM_{10}$  ( $\mu\text{g}/\text{m}^3$ ) in the Zagorje municipality, Slovenia, between January 1, 2006 and December 31, 2011.

Slika 3. Časovno spreminjanje: a) dnevne števila obiskov zaradi bolezni dihal pri otrocih in b) dnevne 24-urne povprečne koncentracije  $PM_{10}$  ( $\mu\text{g}/\text{m}^3$ ) v občini Zagorje, Slovenija, med 1. januarjem 2006 in 31. decembrom 2011.

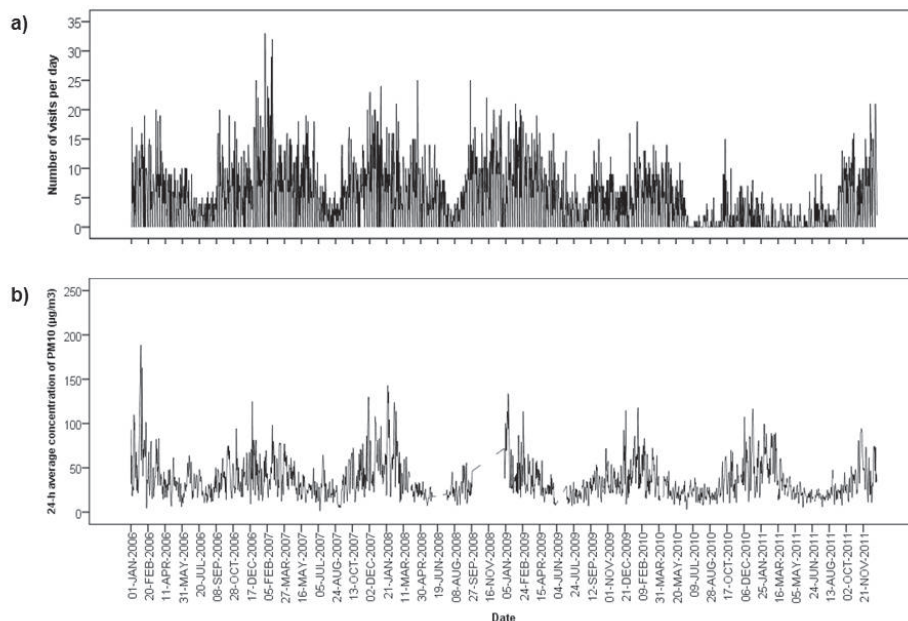


Figure 4. Temporal pattern of: a) daily number of consultations for respiratory diseases in children, and b) daily 24-hr average concentration of  $PM_{10}$  ( $\mu\text{g}/\text{m}^3$ ) in the Trbovlje municipality, Slovenia, between January 1, 2006 and December 31, 2011.

Slika 4. Časovno spreminjanje: a) dnevne števila obiskov zaradi bolezni dihal pri otrocih in b) dnevne 24-urne povprečne koncentracije  $PM_{10}$  ( $\mu\text{g}/\text{m}^3$ ) v občini Trbovlje, Slovenija, med 1. januarjem 2006 in 31. decembrom 2011.



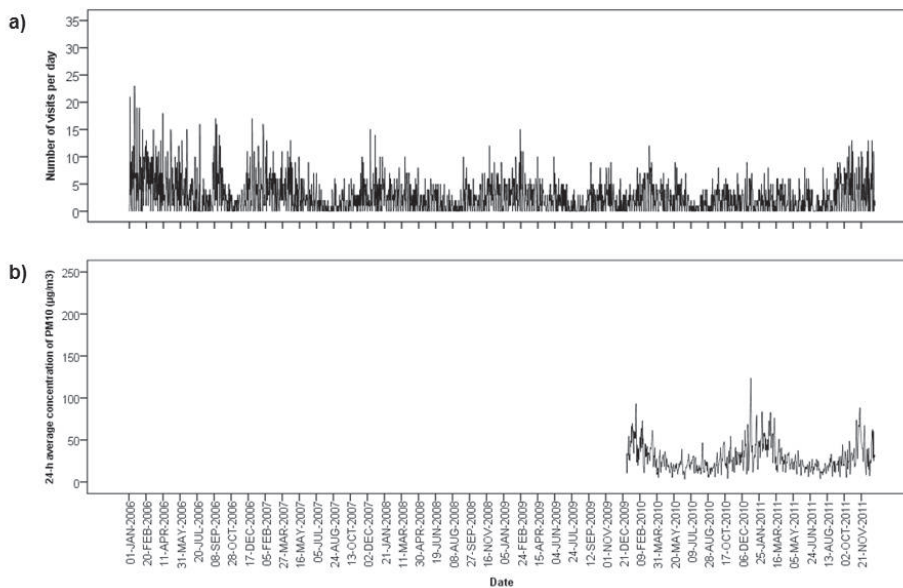


Figure 5. Temporal pattern of: a) daily number of consultations for respiratory diseases in children, and b) daily 24-hr average concentration of PM<sub>10</sub> ( $\mu\text{g}/\text{m}^3$ ) in the Hrastnik municipality, Slovenia, between January 1, 2006 and December 31, 2011.

Slika 5. Časovno spreminjanje: a) dnevne števila obiskov zaradi bolezni dihal pri otrocih in b) dnevne 24-urne povprečne koncentracije PM<sub>10</sub> ( $\mu\text{g}/\text{m}^3$ ) v občini Hrastnik, Slovenija, med 1. januarjem 2006 in 31. decembrom 2011.

### 3.2 Relationship analysis

The results of the univariate analysis showed that the daily number of first consultations for respiratory diseases among children was statistically significantly associated with PM<sub>10</sub> concentrations in all three municipalities. In all of them, the strongest association was observed in lag 0 (Zagorje municipality - IRR: 1.007, 95% CI: 1.007-1.008;  $p < 0.001$ ; Trbovlje municipality - IRR: 1.010, 95% CI: 1.009-1.011,  $p < 0.001$ ; Hrastnik municipality - IRR: 1.009, 95% CI: 1.006-1.011;  $p < 0.001$ ).

The results of single-pollutant multivariate models once again showed that the daily number of first consultations for respiratory diseases among children in the Zagorje and Trbovlje municipalities was statistically significantly associated with PM<sub>10</sub> concentrations. In

both municipalities, the strongest association was observed in lag 0 (Zagorje municipality - IRR: 1.003, 95% CI: 1.002-1.004;  $p < 0.001$ ; Trbovlje municipality - IRR: 1.003, 95% CI: 1.001-1.005;  $p < 0.001$ ). In the Hrastnik municipality, the association was no longer significant (IRR: 1.000, 95% CI: 0.995-1.005;  $p = 0.969$ ). The results of the multi-pollutant multivariate models with SO<sub>2</sub> and O<sub>3</sub> included as co-pollutants showed that daily number of first consultations for respiratory diseases among children was still significantly associated with PM<sub>10</sub> concentrations in both the Zagorje and Trbovlje municipalities. The detailed results are presented in Table 2. The results for co-pollutants are presented in the same table as well. In the Trbovlje municipality, the results didn't change much after the inclusion of NO<sub>2</sub> as an additional co-pollutant in the model (IRR: 1.004, 95% CI: 1.002-1.006,  $p = 0.001$ ).

Table 2. Results of the Poisson regression analysis of association between consultations for respiratory diseases and PM<sub>10</sub> concentration, controlling for selected covariates between January 1, 2006 and December 31, 2011 in the Zasavje region, Slovenia ( $N_{\text{days Zagorje}}=1.996$ ); ( $N_{\text{days Trbovlje}}=1.927$ ).

Tabela 2. Rezultati Poissonove regresijske analize povezanosti med obiski zaradi bolezni dihal in PM<sub>10</sub> standardizirano na izbrane pojasnjevalne dejavnike med 1. januarjem 2006 in 31. decembrom 2011 v regiji Zasavje, Slovenija ( $N_{\text{dni Zagorje}}=1.996$ ); ( $N_{\text{dni Trbovlje}}=1.927$ ).

Explanatory factor/Covariates Pojasnjevalni dejavnik/Sopojavi	IRR RIS	95% CI limits for IRR 95 % IZ za RIS			
		Lower Spodnji	Upper Zgornji	p p	
<b>Zagorje municipality/Občina Zagorje</b>					
PM <sub>10</sub> 24-hr average concentration (µg/m <sup>3</sup> ) PM <sub>10</sub> 24-urna povprečna koncentracija (µg/m <sup>3</sup> )	1.003	1.002	1.004	<0.001	
SO <sub>2</sub> 24-hr average concentration (µg/m <sup>3</sup> ) SO <sub>2</sub> 24-urna povprečna koncentracija (µg/m <sup>3</sup> )	0.983	0.976	0.989	<0.001	
O <sub>3</sub> maximum 8-hr average concentration (µg/m <sup>3</sup> ) O <sub>3</sub> maksimalna 8-urna povprečna koncentracija (µg/m <sup>3</sup> )	1.002	1.001	1.004	0.003	
Temperature 24-hr average (°C) 24-urna povprečna temperatura (°C)	0.998	0.991	1.005	0.633	
Relative humidity 24-hr average (%) 24-urna povprečna relativna vlažnost (%)	1.003	1.001	1.005	0.014	
Season of the year Letni čas					
	Summer/Poletje	1.000			
	Winter/Zima	1.732	1.478	2.030	<0.001
	Spring/Pomlad	1.111	1.039	1.188	0.002
	Autumn/Jesen	1.092	1.057	1.128	<0.001
Work day Delovni dan	No/Ne	1.000			
	Yes/Da	2.297	2.125	2.486	<0.001
Holiday Počitnice	No/Ne	1.000			
	Yes/Da	0.739	0.674	0.809	<0.001
Influenza season Sezona gripe	No/Ne	1.000			
	Yes/Da	1.260	1.160	1.369	<0.001
<b>Trbovlje municipality/Občina Trbovlje</b>					
PM <sub>10</sub> 24-hr average concentration (µg/m <sup>3</sup> ) PM <sub>10</sub> 24-urna povprečna koncentracija (µg/m <sup>3</sup> )	1.004	1.002	1.006	<0.001	
SO <sub>2</sub> 24-hr average concentration (µg/m <sup>3</sup> ) SO <sub>2</sub> 24-urna povprečna koncentracija (µg/m <sup>3</sup> )	0.986	0.977	0.995	0.002	
O <sub>3</sub> maximum 8-hr average concentration (µg/m <sup>3</sup> ) O <sub>3</sub> maksimalna 8-urna povprečna koncentracija (µg/m <sup>3</sup> )	0.998	0.996	1.001	0.180	
Temperature 24-hr average (°C) 24-urna povprečna temperatura (°C)	1.004	0.995	1.014	0.370	
Relative humidity 24-hr average (%) 24-urna povprečna relativna vlažnost (%)	0.996	0.992	1.001	0.132	
Season of the year Letni čas					
	Summer/Poletje	1.000			
	Winter/Zima	1.805	1.474	2.212	<0.001
	Spring/Pomlad	1.207	1.107	1.316	<0.001
	Autumn/Jesen	1.100	1.054	1.148	<0.001
Work day Delovni dan	No/Ne	1.000			
	Yes/Da	5.103	4.436	5.903	<0.001
Holiday Počitnice	No/Ne	1.000			
	Yes/Da	0.641	0.566	0.724	<0.001
Influenza season Sezona gripe	No/Ne	1.000			
	Yes/Da	1.148	1.032	1.278	0.012

Abbreviations/okrajšave: IRR/RIS – incident rate ratio/razmerje incidenčnih stopenj; CI/IZ – confidence interval/interval zaupanja; N – number of days/število dni

## 4 DISCUSSION

The main results of our study have consistently showed that in the Zagorje and Trbovlje municipalities the daily number of first consultations for respiratory diseases among children was significantly associated with the daily concentration of PM<sub>10</sub>. Only in the Hrastnik municipality was a significant association not observed in all models. This result is clearly in relation to the much shorter time series in this municipality compared to the other two municipalities in Zasavje. The hypothesis that a positive temporal relationship exists between the daily number of first consultations for respiratory diseases and the daily concentration of PM<sub>10</sub> was thus certainly confirmed in the Zagorje and Trbovlje municipalities. Moreover, these results are consistent with the results of many similar studies (23, 48-50) that also confirmed the positive association between respiratory diseases and PM<sub>10</sub> concentration.

Our study, in addition to the main results, also provides some additional findings. For example, in the Zagorje municipality, the daily number of first consultations for respiratory diseases among children was also significantly associated with the daily O<sub>3</sub> concentrations. Since the dynamics of this pollutant differ from the PM<sub>10</sub> dynamics, it would be useful to analyse this problem in depth in a separate study.

Our study has some potential limitations. First, when assessing the usefulness of environmental data for Zasavje as the input data for time-trend studies, we encountered some problems. The results of our study namely showed that all observed pollutants lacked a certain percentage of the measurement. The SEA provided an oral explanation that this was mainly due to the calibration of instruments or filter blockage of the measuring device but certainly not a deliberate shutdown of instrumentation. For the present study, the biggest problem was the lack of PM<sub>10</sub> data in the Hrastnik municipality. In this municipality, SEA only started with continuous measurements of 24-hr concentrations of PM<sub>10</sub> from 2010 onwards (26), however it is still not clear whether or not the measurements will continue to be implemented in the future (51-53). An additional problem related to PM<sub>10</sub> measurements, which must be stressed at this point, is that for now in Slovenia only concentrations of PM<sub>10</sub> are routinely measured, while concentration of PM<sub>2.5</sub> are only measured at three locations (Biotechnical Faculty, Ljubljana, Maribor and Maribor centre Vrbanski plateau) (26). Furthermore, the chemical composition of PM<sub>10</sub> is provided only in individual cases (27). However, these data are crucial for unbiased estimates of the health impact due to

the environmental factors. PM<sub>10</sub> can vary significantly and thus also have different effects on human health. Second, we have to take into account the potential problem with health data that was observed in the larger project that our study was a part of (36). In this project, the usefulness of health data for Zasavje as input data for time-trend studies was assessed, and some potential problems were encountered (36). On one side, there were no problems with the completeness of data collection - at all Zasavje CHCs, data for all days of the observed 6-year period were available. On the other side, certain ambiguities related to the outcome variable were encountered. In the Zagorje municipality, compared to the Trbovlje or Hrastnik municipalities, 1.3 to 1.4 times higher numbers of days were detected with at least one consultation due to respiratory diseases, among which acute respiratory diseases dominated (36). The project assumed that there may be a difference in the encoding of individual diagnoses. However, this problem could only be confirmed if time-trend studies of larger dimensions would be carried out in Slovenia. According to our knowledge, only two studies similar to ours were carried out in Slovenia so far. In both, the researchers observed the association between O<sub>3</sub> concentrations and the number of consultations due to respiratory diseases at CHCs. The first was carried out in the Nova Gorica (54, 55) and the second one in the Koper municipality (56). In both studies, only the data obtained at one CHC were analysed. As a result, the problems that surfaced in our study could not be observed. Third, a potential limitation could be that in our study the association between the observed outcome and explanatory factors was adjusted for covariates that were available in the frame of routinely collected data by SEA in Zasavje. However, we considered most of the generally recommended covariates in similar studies (50, 57, 58). Unfortunately, we could not take confounding factors such as the concentration of pollen into account, because the concentration of pollen is not monitored in the Zasavje region.

On the other hand, this study has several strengths. First, it is still one of the first and few such studies that actually need to become routine in monitoring the health of the population in relation to air pollution. Second, the results of the study indicate a positive association between PM<sub>10</sub> concentration and the daily number of first consultations for all respiratory diseases among children in Zasavje. Consequently, the study provides important information for further work in the field of public health activities, especially the implementation of environmental health promotion activities in the region. While there has been much done in the Zasavje region

in recent years in the sense of reduction of some air pollutants (for example installation of filter systems to reduce SO<sub>2</sub> emissions in the local cement and steam power plants), some problems still remain. The biggest problem at the moment is certainly the outdoor air pollution with PM<sub>10</sub>, but the results of our study also indicate a problem of the outdoor air pollution with O<sub>3</sub>. Here, new problems related to the chemical composition of PM<sub>10</sub>, in which a lot of invisible hazards could be hidden, are posing along the old problems related to PM<sub>10</sub> concentrations. Another important strength of our study is that it showed important deficiencies in the currently available input data for studies that integrate routinely collected health and environmental data in Slovenia. These deficiencies could be eliminated to a large extent and consequently make these kinds of studies in Slovenia more viable and useful in the field of health policy (59).

All issues arising during our study represent a new challenge for future research in the field of linkage of environmental and health data in Slovenia. Since this kind of research is in the beginning stages, there is a lot that has to be done. Although we now have some knowledge in the use of linkage methods for environment and health (24, 32), we first need to make the routinely collected data in both information systems - health and environment – more reliable. Although there will be a lot of difficulties in solving these problems, especially since changes in legislation should be addressed in this process, they are not unsolvable. Certainly, the multidisciplinary approach would be the most appropriate and successful.

## 5 CONCLUSIONS

In conclusion, we found positive correlation between concentrations of PM<sub>10</sub> and the daily number of first consultations for all respiratory diseases among children in the Zagorje and Trbovlje municipalities. On the basis of these results, it can be assumed that with some improvements (at least a uniform method to collect health-related data, more air pollution measuring sites in the more polluted areas and more detailed geographical studies), linkage of existing health and environmental data in Zasavje could be feasible in identifying a grounded need for public health action.

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## References

- Schwartz J, Slater D, Larson TV, Pierson WE, Koenig JQ. Particulate air pollution and hospital emergency room visits for asthma in Seattle. *Am Rev Respir Dis* 1993; 147: 826-31.
- Dockery DW, Pope CA. Acute respiratory effects of particulate air pollution. *Annu Rev Public Health* 1994; 15: 107-32.
- World Health Organization, Regional Office for Europe. Air quality guidelines for Europe. 3rd ed. Copenhagen: WHO Regional Office for Europe, 2005.
- Curtis L, Rea W, Smith-Willis P, Fenyves E, Pan Y. Adverse health effects of outdoor air pollutants. *Environ Int* 2006; 32: 815-30.
- Jin-Young M, Kyoung-Bok M, Sung-Il C, Domyung P. Lung effect of particulate air pollution on lung function in children. *Pediatr Pulmonol* 2008; 43: 476-80.
- Dockery DW. Health effects of particulate air pollution. *Ann Epidemiol* 2009; 19: 257-63.
- Künzli N, Perez L, Rapp R. Air quality and health. Lausanne: European Respiratory Society, 2010: 1-63.
- Nastos PT, Paliatsos AG, Anthracopoulos MB, Roma ES, Priftis KN. Outdoor particulate matter and childhood asthma admissions in Athens, Greece: a time-series study. *Environ Health* 2010; 9: 45.
- Albuquerque de Castro H, Hacon S, Argento R, Junger WL, de Mello C, Castiglioni JN et al. Air pollution and respiratory diseases in the Municipality of Vitória, Espírito Santo State, Brazil. *Cad Saúde Pública* 2007; 23: 5630-42.
- Dockery DW, Stone PH. Cardiovascular risks from fine particulate air pollution. *N Engl J Med* 2007; 356: 511-3.
- Middleton N, Yiallourous P, Kleanthous S, Kolokotroni O, Schwartz J, Dockery DW et al. A 10-year time-series analysis of respiratory and cardiovascular morbidity in Nicosia, Cyprus: the effect of short-term changes in air pollution and dust storms. *Environ Health* 2008; 7: 39.
- Farkaš-Lainščak J, Koprivnikar H, Kuček A, Košnik M. The most important risk factors for respiratory diseases (in Slovene). *Med Razgl* 2012; 51: 409-24.
- Wong CM, Thach TQ, Chau PY, Chan EK, Chung RY, Ou CQ et al. Part 4. Interaction between air pollution and respiratory viruses: time-series study of daily mortality and hospital admissions in Hong Kong. *Res Rep Health Eff Inst* 2010; 154: 283-362.
- Kan H, Chen B, Zhao N, London SJ, Song G, Chen G et al. Part 1. A time-series study of ambient air pollution and daily mortality in Shanghai, China. *Res Rep Health Eff Inst* 2010; 154: 17-78.
- Qiu H, Yu IT, Tian L, Wang X, Tse LA, Tam W et al. Effects of coarse particulate matter on emergency hospital admissions for respiratory diseases: a time-series analysis in Hong Kong. *Environ Health Perspect* 2012; 120: 572-6.
- Tamburini G, Ehrenstein OS, Bertolini R, editors. Children's health and environment: a review of evidence. Luxembourg: World Health Organisation, European Environmental Agency, 2002.



17. Kim JJ. American Academy of Pediatrics Committee on Environmental Health. Ambient air pollution: health hazards to children. *Pediatrics* 2004; 114: 1699-707.
18. World Health Organization. Effects of air pollution on children's health and development: a review of the evidence. Copenhagen: World Health Organization, 2005.
19. Buka I, Koranteng S, Osornio-Vargas AR. The effects of air pollution on the health of children. *Paediatr Child Health* 2006; 11: 513-6.
20. Suwanwaiphatthana W, Ruangdej K, Turner-Henson A. Outdoor air pollution and children's health. *Pediatr Nurs* 2010; 36: 25-32.
21. Pope CA, Dockery DW, Spengler JD, Raizenne ME. Respiratory health and PM<sub>10</sub> pollution—a daily time-series analysis. *Am Rev Respir Dis* 1991; 144: 668-74.
22. Katsouyanni K, Schwartz J, Spix C, Touloumi G, Zmirou D, Zanobetti A et al. Short term effects of air pollution on health: a European approach using epidemiologic time series data: the APHEA protocol. *J Epidemiol Community Health* 1996; 50: 18.
23. Zhang F, Li L, Krafft T, Lv J, Wang W, Pei D. Study on the association between ambient air pollution and daily cardiovascular and respiratory mortality in an urban district of Beijing. *Int J Environ Res Public Health* 2011; 8: 2109-23.
24. Briggs D, Corvalan C, Nurminen M, editors. Linkage methods for environment and health analysis: general guidelines. Geneva: World Health Organization, Office of Global and Integrated Environmental Health, 1996.
25. ERICo Velenje, Institute for ecological research. Pollution of the environment and natural resources as factors of the development in the Sava Basin region – a model approach: final report (in Slovene). Velenje: ERICo Velenje, Institute for Ecological Research, 2001.
26. Šegula A, Bolte T, Koleša T, Komar Z, Murovec M, Muri G et al. Quality of the air in the year 2011: annual report (in Slovene). Ljubljana: Slovenian Environment Agency, 2012.
27. Bolte T, Koleša T, Šegula A, Fašing J, Rode B, Planinšek A et al. Identifying sources of PM<sub>10</sub> in Zagorje ob Savi: final report (in Slovene). Ljubljana: Slovenian Environment Agency, 2010.
28. Lušič M. Allergic respiratory diseases in children in Zasavje. In: Healthy and safe environment to the children: proceedings (in Slovene). Ljubljana: Friends of Youth Association of Slovenia, 1990.
29. Lušič M. The impact of the air pollution on the frequency of hospitalization due to respiratory diseases in the paediatric ward of Trbovlje Hospital. In: Environmental pollution and a child: abstracts of discussions (in Slovene). Trbovlje, Ljubljana: Slovenian Medical Association, Paediatric Section, 1995.
30. Erzen I, Vertačnik G, Podkrajšek D, Juričič M, Uršič A, Zadnik V et al. Exploration of the impact of the environment on the occurrence of certain diseases and increased mortality rate of the population of the Zagorje ob Savi municipality: final report (in Slovene). Celje: Regional Institute of Public Health Celje, 2006.
31. Kukec A, Farkas J, Erzen I, Zaletel-Kragelj L. A prevalence study on outdoor air pollution and respiratory diseases in children in Zasavje, Slovenia, as a lever to trigger evidence-based environmental health activities. *Arh Hig Rada Toksikol* 2013; 64: 9-22.
32. Corvalan C, Nurminen M, Pastides H, editors. Linkage methods for environment and health analysis: technical guidelines. Geneva: World Health Organization, Office of Global and Integrated Environmental Health, 1997.
33. Morgenstern H, Thomas D. Principles of study design in environment epidemiology. *Environ Health Perspect* 1993; 101: 23-38.
34. Morgenstern H. Uses of ecologic analysis in epidemiologic research. *Am J Public Health* 1982; 72: 1336-44.
35. Pekkanen J, Pearce N. Environmental epidemiology: challenges and opportunities. *Environ Health Perspect* 2001; 109: 1-5.
36. Kukec A, Zaletel-Kragelj L, Bizjak M, Fink R, Jereb G, Košnik M et al. A study of linkage of environmental and health data in Zasavje as a model study to support the development and implementation of cross-sectoral policies on the environment and health. Final report (in Slovene). Ljubljana: Chair of Public Health, University of Ljubljana, Faculty of Medicine, 2012.
37. Statistical Office of the Republic of Slovenia, SI-STAT data portal. Data by municipalities. Population by age and sex, municipalities, Slovenia, semi-annual report (in Slovene). Available Sept 9, 2012 at: [http://pxweb.stat.si/pxweb/Dialog/varval.asp?ma=05C4002S&ti=&path=../Database/Dem\\_soc/05\\_prebivalstvo/10\\_stevilo\\_preb/20\\_05C40\\_prebivalstvo\\_obcine/&lang=2](http://pxweb.stat.si/pxweb/Dialog/varval.asp?ma=05C4002S&ti=&path=../Database/Dem_soc/05_prebivalstvo/10_stevilo_preb/20_05C40_prebivalstvo_obcine/&lang=2).
38. National Institute of Public Health of the Republic of Slovenia. Epidemiological surveillance of communicable diseases in Slovenia – annual reports (in Slovene). Available Sept 9, 2012 at: [http://www.ivz.si/Mp.aspx?ni=105&pi=5&\\_5\\_id=788&\\_5\\_PageIndex=0&\\_5\\_groupId=155&\\_5\\_newsCategory=&\\_5\\_action=ShowNewsFull&p=105-5.0](http://www.ivz.si/Mp.aspx?ni=105&pi=5&_5_id=788&_5_PageIndex=0&_5_groupId=155&_5_newsCategory=&_5_action=ShowNewsFull&p=105-5.0).
39. Lipsett M, Hurley S, Ostro B. Air pollution and emergency room visits for asthma in Santa Clara County, California. *Environ Health Perspect* 1997; 105: 216-22.
40. Hajat S, Anderson HR, Atkinson RW, Haines A. Effects of air pollution on general practitioner consultations for upper respiratory diseases in London. *Occup Environ Med* 2002; 59: 294-9.
41. Galan I, Tobias A, Banegas JR. Short-term effects of air pollution on daily asthma emergency room admissions. *Eur Respir J* 2003; 22: 802-8.
42. Anderson HR, Atkinson RW, Peacock JL, Sweeting MJ, Marston L. Ambient particulate matter and health effects: publication bias in studies of short-term associations. *Epidemiology* 2005; 16: 155-63.
43. Yi O, Hong YC, Kim H. Seasonal effect of PM<sub>10</sub> concentrations on mortality and morbidity in Seoul, Korea: a temperature-matched case-crossover analysis. *Environ Res* 2010; 110: 89-95.
44. Willocks LJ, Bhaskar A, Ramsay CN, Lee D, Brewster DH, Fischbacher CM et al. Cardiovascular disease and air pollution in Scotland: no association or insufficient data and study design? *BMC Public Health* 2012; 12: 227.
45. Cox S, West SG, Aiken LS. The analysis of count data: a gentle introduction to poisson regression and its alternatives. *J Pers Assess* 2009; 91: 121-36.
46. De Souza Tadano Y, Ugaya CML, Franco AT. Methodology to assess air pollution impact on human health using the generalized linear model with poisson regression. In: Khare M, editor. Air pollution - monitoring, modelling and health. Rijeka: InTech, 2012.
47. UCLA Academic Technology Services, Statistical Consulting Group. SPSS data analysis examples: poisson regression. Available June 20, 2012 at: <http://www.ats.ucla.edu/stat/spss/dae/poissonreg.htm>.
48. Moura M, Junger WL, Silva Mendonca GA, De Leon AP. Air quality and acute respiratory disorders in children. *Rev Saude Publica* 2008; 42: 1-8.
49. Strickland MJ, Darrow LA, Klein M, Flanders WD, Sarnat JA, Waller LA et al. Short-term associations between ambient air pollutants and pediatric asthma emergency department visits. *Am J Respir Crit Care Med* 2010; 182: 307-16.
50. Fraga J, Botelho A, Sá A, Costa M, Quaresma M. The lag structure and the general effect of ozone exposure on pediatric respiratory morbidity. *Int J Environ Res Public Health* 2011; 8: 4013-24.

51. European Parliament and the Council of the European Union. Directive 2008/50/EC of the European Parliament and of the Council of 21 May 2008 on ambient air quality and cleaner air for Europe.
52. Government of the Republic of Slovenia. Decree on ambient air quality (in Slovene). Ur List RS 2011; 21: 964-83.
53. Ministry of Environment and Spatial Planning of the Republic of Slovenia. Rules on assessment of ambient air quality and public information on ambient air quality (in Slovene). Ur List RS 2011; 21: 7771-800.
54. Šimac N. Air pollution by ozone in Nova Gorica region - assessment of the impact on human health: specialistic study thesis (in Slovene). Nova Gorica: Regional Institute of Public Health Nova Gorica, 2008.
55. Šimac N, Hladnik M, Zaletel-Kragelj L. The impact of temperature on tropospheric ozone in the Nova Gorica region. Zdrav Var 2011; 50: 121-30.
56. Rems-Novak MM. Effects of air pollution with ozone on primary health care consultations for respiratory tract diseases in children in Koper Municipality: master degree thesis. Nova Gorica: University of Nova Gorica, Graduate School of Environmental Sciences, 2013.
57. Gouveia N, Fletcher T. Respiratory diseases in children and outdoor air pollution in Sao Paulo, Brazil: a time series analysis. Occup Environ Med 2000; 57: 477-83.
58. Braga ALF, Saldiva PHN, Pereira LAA, Menezes JJC, Conceicao GMS, Lin CA et al. Health effects of air pollution exposure on children and adolescents in Sao Paulo, Brazil. Pediatr Pulmonol 2001; 31: 106-13.
59. Porta M, Greenland S, Last JM. A dictionary of epidemiology. 5th ed. New York: Oxford University Press, 2008.