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Spatial Associations of Long-term Exposure to Diesel Particulate Matter with Seasonal and Annual Mortality Due to COVID-19 in the Contiguous United States

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Research Article

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Abstract

Background

People with certain underlying respiratory and cardiovascular conditions might be at an increased risk for severe illness from COVID-19. Diesel Particulate Matter (DPM) exposure may affect the pulmonary and cardiovascular systems. The study aims to assess if DPM was spatially associated with COVID-19 mortality across three waves of the disease and throughout 2020.

Methods

We tested an ordinary least square (OLS) model, then two global models, spatial lag model (SLM) and spatial error model (SEM), designed to explore spatial dependence, and a geographically weighted regression (GWR) model designed to explore local associations.

Results

The GWR model found that associations between COVID-19 deaths and DPM concentrations may increase up to 57, 36, 43, and 58 deaths per 100,000 people in some US counties for every 1 μ g/m³ increase in DPM concentration. Relative significant positive association are observed in New York, New Jersey, eastern Pennsylvania, and western Connecticut for the wave from January to May, and in southern Florida and southern Texas for June to September. The period from October to December exhibit a negative association in most parts of the US, which seems to have influenced the year-long relationship due to the large number of deaths during that wave of the disease.

Conclusions

Our models provided a picture in which long-term DPM exposure may have influenced COVID-19 mortality during the early stages of the disease, but that influence appears to have waned over time as transmission patterns evolved.

Background

In 2020, more than 20 million cases of coronavirus disease 2019 (COVID-19) were identified in the United States (U.S.), and more than 350,000 people died (Dong et al. 2020, John Hopkins University 2021). Many studies established the relationship between the prevalence of COVID-19 and underlying health conditions, social determinants of health, being of Black race, and environmental exposures. The most common underlying conditions reported to increase risk of severe illness from COVID-19 included obesity, hypertension, cardiovascular disease, type 2 diabetes, and chronic respiratory diseases, including asthma and chronic obstructive pulmonary disease (Clark et al. 2020, Nieman 2020). Incident COVID-19 deaths among people ages 65 years and older comprised at least 85% of all incident COVID-19 deaths in the U.S. on any day in 2020 (CDC 2021). In addition to age, socioeconomic status, access to healthcare, physical environment, and education have been identified as social determinants of COVID-19 hospitalization and mortality (Phillips et al. 2020, Mollalo et al. 2020). Several studies have observed a disproportionate share of COVID-19 incidence and mortality among predominantly Black U.S. communities, which may be partly attributable to social and economic inequalities and preexisting comorbidities (Yancy 2020, Reyes 2020, Phillips et al. 2020, Gayam et al. 2021, Peek et al. 2021). Yancy (2020) reported that the COVID-19 death rate in predominantly Black U.S. counties (6.3/100,000) was more than three times higher than in predominantly White counties in April 2020. Through July 21, 2020, Reyes (2020) reported a death rate twice as high among African Americans in the U.S. (97.9/100,000) compared with the White population, based on data from 45 states and the District of Columbia. Phillips et al. (2020) found that the increased risk of complications is related to the high prevalence of pre-existing comorbid conditions in part due to hereditary genetic predisposition.

The impact of particulate matter exposures on COVID-19 outcomes have also been evaluated, with some studies centered around diesel particulate matter (DPM). In an investigation of the role of long-term exposure (2000-2016) to air pollution during the first months of COVID-19, Wu et al. (2020) found that an increase of 1 µg/m³ in particulate matter with a nominal diameter of 2.5 µm (PM_{2.5}) was associated with an 11% increase in the COVID-19 death rate for January 1-June 18, 2020. Bozack et al. (2022) performed a similar analysis to test associations of COVID-19 intensive care unit (ICU) admission and mortality with long-term concentrations of PM2.5, nitrogen dioxide, and black carbon for the period March 8-August 30, 2020 in New York City. They noted an association of ICU admission and mortality with long-term PM_{2.5} concentrations (collected December, 2018-December, 2019). Petroni et al. (2020) investigated the association of COVID-19 mortality with respiratory hazard index calculated across 3223 U.S. counties using emissions data for 2014 and COVID-19 data through May 13, 2020. They observed a 9% increase in COVID-19 mortality per unit increase in respiratory hazard index, which includes DPM. Their analyses with only DPM demonstrated an increased effect of 182% in the mortality rate ratio with a 0.5 µg/m³ increase in DPM concentration. Hendryx and Luo (2020) studied the association of long-term exposure to ozone (obtained from 2016), PM2.5 (obtained from 2016), and DPM (obtained from 2014) with COVID-19 prevalence and mortality through May 31, 2020. They showed an increase of 14.3 deaths per 100,000 U.S. residents for each DPM concentration increase of 1 µg/m³ in a single-pollutant model adjusted for demographic, health, smoking, and COVID-19 testing covariates. These findings collectively suggest that longterm PM exposure may predispose an individual to COVID-19 mortality. However, association of COVID-19 mortality with long-term DPM may change over time with the evolution of the coronavirus and changes in policies and personal behaviors. Our understanding of the effect of long-term DPM exposure on COVID-19 mortality during different waves of the disease and over the locations impacted by those waves remain unknown, hampering anticipation of disease hotspots.

DPM is composed of a complex mixture of black carbon and organic carbon. Studies have shown that 80–90% of particles emitted by diesel engines are less than 2.0 µm (Lee et al. 2015, Douglas et al. 2019), small enough to penetrate the alveoli (Ristovski et al. 2012). Long-term DPM exposure has been associated with adverse respiratory and cardiovascular effects (Pronk et al. 2009, Ristovski 2012, Douglas et al. 2019). Diesel engines power school buses, heavy-duty trucks, a variety of off-road heavy equipment, shipping, and commercial boating (Lee et al. 2015, Douglas et al. 2019). DPM emissions are higher in urban areas, where most of the global population lives (Ristovski 2012, Douglas et al. 2019). Likewise, greater DPM concentrations have been observed in socioeconomically disadvantaged communities (Douglas et al. 2019, Clougherty et al. 2014).

Methods

Our study explores spatial associations between long term average concentrations of DPM, as a metric for past air pollution exposure, and COVID-19 mortality across each pandemic wave and throughout 2020 in the U.S. The objectives of the study are 1) to assess if living near DPM sources increased the risk of death from COVID-19, 2) to estimate how associations between mortality and long-term exposure to DPM may have changed over time with changes in the Coronavirus and in the population's behavior, and 3) to test if models accounting for spatial autocorrelation improve model estimates. Data for air pollution, health, demographic, and social determinants of health were merged for this analysis, and global and local models were both applied to examine these relationships.

Population Data

County-level number of COVID-19 deaths were obtained from the publicly-available Johns Hopkins Coronavirus Resource Center (Johns Hopkins University 2020) for the period January 1- December 31, 2020. Data for potential confounders associated with COVID-19 deaths, including access to health care, education, poverty, demographics, transportation, and occupation were obtained from the American Community Survey (ACS; U.S. Census Bureau 2020) and the County Health Rankings (CHR; Robert Wood Johnson Foundation 2020) (Table 1).

Table 1 Potential confounders tested in the models.							
Race/ethnicity	Poverty and Wealth						
Fraction Black	Fraction homelessness						
Fraction White	Fraction with a severe housing burden						
Fraction Hispanic	Food-environment index						
Fraction American Indian	Fraction Income inequality						
Fraction Asian	Fraction Unemployment						
Fraction Pacific Islander	Fraction Median income						
Transportation	Fraction in poverty						
Fraction who walks to work	Median home value						
Fraction who takes public transportation to work	Demographics						
Fraction who takes a bicycle or motorcycle to work	Population density						
Fraction who drives a car to work							
Fraction average time to work							
Traffic volume							
Age							
Fraction over 65							
Median age							
Occupation							
Fraction in a service occupation							
Fraction in a manual occupation							
Fraction working in a mining or agricultural occupation							
Fraction working in a construction occupation							
Fraction working in a utilities occupation							
Health							
Fraction in poor health							
Fraction obese							
Fraction with diabetes							
Fraction reporting inactivity							
Fraction smoking							
Access to Healthcare							
Fraction uninsured							
Fraction population receiving health care							
Fraction hospitals per county							
Fraction hospital beds							
Education							
Fraction Incomplete school							

Exposure Data

Long-term average DPM concentrations were obtained from the 2014 National Air Toxics Assessment (NATA) database, the most recently modeled concentrations of hazardous air pollutants and select other pollutants (U.S. Environmental Protection Agency (EPA) 2018). EPA used a hybrid model that coupled a Community Multiscale Air Quality (CMAQ) chemical transport model to the American Meterological Society/Environmental Protection Agency Regulatory Model (AERMOD) dispersion model to estimate NATA air pollutant concentrations at the census tract level through a multi-step process. CMAQ v5.2 was first run over a 12 km x 12 km grid based on DPM emissions inputs from the National Emissions Inventory (U.S. EPA 2018). Next, the AERMOD

dispersion model was run for each source using the same inputs but with receptors distributed over census tract centroids. Finally, concentrations estimated by AERMOD along the census tract centroids were scaled by the ratio of the CMAQ concentration to the average of the AERMOD concentrations over that same grid cell. This formulation allows for more accurate representation of the chemistry and physics of the DPM than the AERMOD dispersion model can provide alone, while maintaining the finer census tract level spatial resolution of the dispersion model.

Model Runs

We tested the association between COVID-19 mortality and long-term DPM concentrations across the contiguous United States for time periods coinciding with each COVID-19 wave in 2020: January 1-May 31, 2020, June 1-September 30, 2020, and October 1-December 31, 2020. We also ran the model for the entire year: January 1-December 31, 2020.

We used regression analysis to examine spatial non-stationarity in the relationship between COVID-19 and DPM while accounting for potentially confounding effects. This work is similar to spatial modeling approaches used by Sun et al. (2020) and Rahman et al. (2020). Sun et al. (2020) investigated different spatial regression models and compared them with an ordinary least squares (OLS) regression model to explain the transmission pattern of COVID-19. County-level race/ethnicity and socio-economic covariates were included in their models. We adapted their approach by focusing on associations of COVID-19 mortality with DPM and by investigating different time periods. Three global models, OLS, spatial lag model (SLM), and spatial error model (SEM), were run to produce a nationwide effect estimate. One local model, geographically weighted regression (GWR), produced effect estimates at the county scale. The R Statistical Software version 3.6.3 was used to run all code. We performed spatial regression modeling with the following libraries: *spdep*, *spgwr*, and *spatialreg*.

OLS models are designed to minimize the sum of squared differences between the true data and the prediction across the dataset (Goldberger, 1964). Mollalo et al. (2020) studied county-level variations of COVID-19 incidence in the U.S. From a list of 35 demographic, socio-economic, topographic and environmental variables, they used a stepwise forward selection procedure and then checked for multicollinearity to determine the most significant predictors of COVID-19. Then, using the same selected explanatory variables, they tested their model using OLS and several spatial models including SEM, SLM, and GWR (described below). Accounting for spatial autocorrelation in their model improved performance over OLS. Karaye and Horney (2020) also compared OLS to spatial regression models to analyze the impact of social vulnerability on COVID-19 cases. Spatial autocorrelation of the residuals may compromise the validity of the OLS model and produce biased estimators (LeSage and Pace 2009, Loonis and De Bellefon 2018). The model assumptions of zero mean, independence, heteroscedasticity, and normal distribution are met for the case where OLS is a complete and correct model in which the variables capture all of the spatial variation without specifying spatial positions (DeAngelis and Yurek 2017, Schabenberger and Gotway 2017). Spatial autocorrelation in residuals may occur due to an omitted variable.

SLMs estimate an autocorrelation parameter ("spatial lag") using a weighted average of the response variable across neighboring areas, testing if neighboring observations affect one another (LeSage and Fischer 2008, Sun et al. 2020). As the autocorrelation parameter approaches zero, the SLM approaches the OLS (LeSage and Fischer 2008). In SEMs, errors across neighboring areas are autocorrelated ("spatial error") (Le Gallo et al. 2005). SEMs estimate the relationship between the residuals in a spatial region and those in adjacent regions (Sun et al. 2020). The spatial structure is in the residuals, meaning that some important predictors are omitted in the model (Chi and Zhu 2020).

SLM and SEM have only one spatial dependence parameter. The single-valued characteristic makes it impossible for global spatial models to reveal local spatial patterns (Chi and Zhu 2020, Fotheringham et al. 2003). Another limitation of global spatial models is that the model is dependent on the spatial weighting matrix (Chi and Zhu 2020). In contrast, GWR allows for local models to be fit to each observation using spatial distance as a weighting factor for the influence of all other points (Fotheringham et al. 2003). To determine local associations between COVID-19 cases in the U.S. and demographic, socio-economic, topographic and environmental parameters, Mollalo et al. (2020) examined two local models including GWR. The variables incorporated in the model are the same set used for OLS, SLM, and SEM. Similarly, Karaye and Horney (2020) compared GWR to OLS to understand the spatially varying effect in the relationship between social vulnerability and COVID-19 case counts. The main advantage of GWR as a local model is the ability to test for spatial variability among the effects of different variables in the model (Chi and Zhu 2020, LeSage and Pace 2009, Fotheringham et al. 2003). Another strength is that GWR has the same model structure as the OLS, which facilitates comparison between the two models (Fotheringham et al. 2003).

For our spatial autoregressive models, we estimated spatial relationships between regions based on contiguous boundaries shared between 2 or more counties, assuming that COVID-19 spread in a county is influenced by adjacent counties. For GWR, a cross validation function minimizes the root mean square prediction error that defines the weight matrix. We evaluated spatial autocorrelation among contiguous cells in the model residuals using Moran's I (Moran 1950). Statistically significant Moran's I indicates either correlation or anticorrelation among neighboring units. Additionally, we used Lagrange multiplier test statistics to understand whether the spatial lag or spatial error pattern is more important for interpreting the local results.

The level of urgency of the COVID-19 outbreak contributed to uncertain policy decisions and interventions in health in compressed timeframes coupled with the complex social, economic and political events of 2020 (Lancaster et al. 2020). Therefore, effects related to pandemic waves could have influenced the importance of specific variables during these different times of the year. Therefore, a set of different covariates have been integrated into the model for each time period. To determine which covariates to include in the regression models of COVID-19 mortality, we applied a stepwise selection algorithm for each season (Table 1). Then, the same covariates were incorporated in the best model for OLS, SLM, SEM, and GWR for each specific wave (Table 2), based on the following framework:

COVID-19 deaths = DPM concentration + Confounder variables + error term (1)

Table 2 Model framework for each wave modeled.

Wave Dates	Models
Jan 1- May 31, 2020	COVID-19 deaths = DPM concentration + Fraction Black + Fraction American Indian + Fraction who take public transportation to work + Fraction average time to work + Fraction uninsured + Fraction smoking + Fraction Income inequality + Population density (2)
Jun 1- Sep 30, 2020	COVID-19 deaths = DPM concentration + Fraction Black + Fraction Hispanic + Fraction American Indian + Fraction who take public transportation to work + Fraction reporting inactivity + Fraction Incomplete school + Population density (3)
Oct 1- Dec 31, 2020	COVID-19 deaths = DPM concentration + Fraction Black + Fraction American Indian + Fraction working in a mining or agricultural occupation + Fraction average time to work + Fraction reporting inactivity + Fraction obese + Fraction over 65 + Fraction homelessness (4)
Jan 1- Dec 31, 2020	COVID-19 deaths = DPM concentration + Fraction Black + Fraction Hispanic + Fraction American Indian + Fraction Pacific Islander + Fraction working in a mining or agricultural occupation + Fraction reporting inactivity + Fraction with a severe housing burden + Fraction Income inequality (5)

The confounder selection procedure was based on minimizing the Akaike information criterion (AIC) after controlling for multicollinearity. We used this same process for each of the three waves and throughout 2020 to find the most significant models for determining the nationwide and local associations between COVID-19 mortality and DPM concentration.

Results

County-level annual average DPM concentration varied from 0.0003 to 1.13 μ g/m³ with a nationwide median of 0.095 μ g/m³. Median regional concentrations for the Northeast, Southeast, West, and Mountain states were 0.117, 0.111, 0.079, and 0.037 μ g/m³, respectively (Fig. 1). Elevated DPM concentration could be observed at specific points corresponding to cities.

During the January May wave, the highest cumulative numbers of COVID-19 deaths were found in roughly the same regions as elevated DPM (Fig. 2a). As 2020 progressed, most counties experienced a higher mortality rate. The New York region exhibited lower cumulative deaths during the October-December wave of our study (Fig. 2c), with a mean of 98 deaths per 100,000 compared with the January-May wave, which had a mean of 280 deaths per 100,000 (Fig. 2a). As shown in Fig. 2a and 2b, cumulative deaths increased substantially from the first wave to the second wave in the Southeast region. In the West region, New Mexico, Arizona and California displayed the same pattern as the Southeast, with a significant increase during the second wave. For the September-December wave, COVID-19 deaths expanded across almost all of the US, exhibiting nearly the same pattern as for the all-year distribution (Fig. 2c and 2d).

At a global level, all models demonstrated a statistically significant association between long-term average DPM concentration and COVID-19 mortality for the first nine months of 2020, as represented by the January-to-May and June-to-September waves (Table 3). SLM and SEM produced slightly higher associations for the June-September wave. For the wave from October to December, none of the global models were found to produce positive associations or to be statistically significant. For the entire year, both the OLS and SLM produced positive associations, while the SEM produced a negative association.

Table 3

Independent variables value per 100,000 people. Where cells are left blank, the forward stepwise variable selection process did not identify those variables

							potential							
Variabl	es	January – May, 2020				June – September, 2020			October- December, 2020				Januar	
		OLS	SLM	SEM	GWR	OLS	SLM	SEM	GWR	OLS	SLM	SEM	GWR	OLS
DPM		3.05	1.9	2.05	-14.45- 57.19	4.59	4.24	4.56	-30.63- 36.49	-0.73	-1.12	-3.32	-157.98- 43.21	5.56
Black		52.45	37.83	43.13	-906.15- 540.33	109.9	86.73	105.28	-1148.6- 414.02	-31.41	-18.35	-27.95	-3523.5- 1267.7	124.3
Hispan	ic	-	-	-	-	63.1	49.68	59.26	-246.24- 280.38	-	-	-	-	77.91
Americ Indian	an	35.4	30.31	27.8	-4001.9- 639.59	59.69	57.98	60.75	-1349.6- 8239.5	140.2	85.79	89.96	-2130.6- 3225.9	167.5
Pacific Islande	er	-	-	-	-	-	-	-	-	-	-	-	-	-4839
Mining Agricul	or ture	-	-	-	-	-	-	-	-	403.4	192.43	177.57	-2076.3- 2390.5	364.4
Public Transp	ortation	-539.9	-738.58	-853.25	-2742- 3836.1	626.8	736.42	676.29	-10054- 4309.5	-	-	-	-	-
Time to	o work	194.3	87.3	23.06	-1630.1- 2133	-	-	-	-	-776	-347.01	-475.26	-5455.5- 9276.3	-
Inactivi	ity	-	-	-	-	113.3	96.5	90.52	-1447.5- 4008.4	109.6	88.59	114.85	-677.69- 816.65	178.9
Uninsu	red	-54.69	-46.12	-37.53	-528.7- 921.85	-	-	-	-	-	-	-	-	-
Smokir	ng	-92.44	-82.97	-85.9	-1555.5- 346.31	-	-	-	-	-	-	-	-	-
Obese		-	-	-	-	-	-	-	-	45.47	31.96	41.67	-536.1- 742.75	-
Over 65	5	-	-	-	-	-	-	-	-	59.62	58.2	58.82	-810.75- 891.23	-
Income inequa	e lity	3.81	3.55	2.85	-13.41- 80.69	-	-	-	-	-	-	-	-	12.87
Homele	essness	-	-	-	-	-	-	-	-	-330.8	-184.44	-182.37	-2234.8- 1384.1	-
Housin burden	g	-	-	-	-	-	-	-	-	-	-	-	-	-405
Incomp school	olete	-	-	-	-	1039	1009.8	1054.5	-3356.4- 6255.9	-	-	-	-	-
Popula density	tion	0.013	0.013	0.016	-0.72 - -0.55	-0.014	-0.014	-0.015	-0.062- 0.65	-	-	-	-	-
R ²		0.37	0.42	0.45	0.65	0.41	0.44	0.44	0.59	0.17	0.33	0.32	0.48	0.15

OLS did not seem to be the most appropriate model to study spatial association between COVID-19 mortality and DPM. Smaller associations for the spatial autoregression models compared with OLS suggested that the OLS covariates were positively biased due to spatial autocorrelation. Both Moran's I and visual inspection of the residuals maps (Figure S1) indicated spatial clusters of high values and of low values. The SLM and SEM models provided modest improvements in model fit, as indicated by slightly higher values of coefficient of determination (R²). Model fit testing indicates that the SLM provided a better fit than the SEM for the year-long data, based on the Lagrange multiplier test.

The local spatial differences estimated using the GWR model are presented as a range of values (Table 3). The mean COVID-19 mortality – DPM association for GWR is identical to that of the OLS, but overall R^2 for GWR indicates improved performance over all global models. Spatial distribution of the DPM coefficients indicates changing conditions across the country during the three parts of the year (Fig. 3). During the January-May wave, associations were mostly positive across the U.S. (Fig. 3a), ranging from an increase of 57.19 deaths per 100,000 for every 1 µg/m³ increase in DPM concentration. During the June-September wave, about half of the contiguous US presented a positive association (Fig. 3b), while associations were more negative for the October-December wave (Fig. 3c). Year-round COVID-19 associations were similar to those for the October-December wave, likely due to the large number of cases during that timeframe. Local variations in R^2 across the waves showed high (>70%) values in the Northeast and Southwest during the January-May and June-September waves and in the year-long model. High R^2 persisted into the October-December wave for the Southwest, albeit with a smaller area (Fig. 4). Low R^2 (<40%) were observed in the areas with greatest decrease in mortality with increasing DPM concentration, suggesting much greater uncertainty in those associations than in the positive ones seen in the New York area during the first wave. Moreover, COVID-19 mortality was only statistically significantly associated DPM concentration during the January-May wave.

Among all confounding covariates incorporated in the models, fraction Black race and fraction American Indian ethnicity were statistically significantly positive in all global models. In addition to these two covariates, Inactivity is significant in the June-September and October-December waves and in the year-long model, and the confounders Hispanic, Mining or Agriculture, Public Transportation, Time to Work, Income Inequality, and Population Density were significant at different time periods of the model.

Discussion

Our study analyzed the spatial correlation between COVID-19 mortality and long-term DPM concentration as a surrogate for exposure across the continental United States during three waves of the COVID-19 pandemic during 2020. Our results suggested that long-term exposure to DPM may have been an important factor in COVID-19 mortality during the first two waves of the disease and that long-term DPM exposure may have been more highly influential during the January-May wave. Sidell et al. (2022) examined associations between air pollution exposure and COVID-19 incidence for monthly and annual averages of PM_{2.5}, nitrogen dioxide (NO₂), and ozone (O₃) over four waves corresponding to those in our study plus January-February, 2021 for a Southern California cohort. They similarly observed that PM_{2.5} had a larger effect during the first wave and that the effect diminished over time. A spatial autocorrelation term was controlled for in these models, but Sidell et al. (2022) did not incorporate local methods. Differences in the outcome variable and the specific exposure also necessitates further examination of spatial and temporal patterns.

Our results indicate that the OLS does not account for the spatial associations of COVID-19 mortality with DPM concentrations. These results are similar to those of Sidell et al. (2022) and Mollalo et al. (2020), although their studies considered COVID-19 incidence rate rather than mortality. Mollalo et al. (2020) used OLS, SLM, SEM, and two versions of the GWR to model COVID-19 incidence and mortality for the time period of January 22-April 9, 2020 and found notable spatial associations of both COVID-19 incidence and mortality with several predictors. The study of Hendryx and Luo (2020), covering the January-May wave, revealed strong associations of COVID-19 prevalence and mortality with long-term DPM and PM_{2.5} concentrations. Their study estimated a coefficient of 14.3-18.7 deaths per 100,000 U.S. residents for each increase of $1 \ \mu g/m^3$ in DPM concentration. Inflation of the DPM effect shown in their results is possibly due to correlation between covariates and their mixed linear multiple regression model that does not account for spatial correlation. Stakhovych and Bijmolt (2007) emphasized that correlated spatial errors lead to bias and uncertainty in the OLS results. Moreover, LeSage and Fischer (2008) noted that spatial correlation in the OLS error terms is a sufficient motivation to employ spatial autoregression models for discerning spatial relationships between dependent and independent variables.

The spatial global models outperformed the OLS model in terms of model fit for all models except June-September. This improved performance may be related to spatial autocorrelation. A difference in coefficients and R² among the OLS, SLM, and SEM models was not observed during the June-September wave, when the modeled relationships between COVID-19 mortality and long-term DPM concentrations lost statistical significance. Kim (2021) reported an inflated effect of spatial autocorrelation on OLS predictor coefficients, suggesting less spatial autocorrelation during the June-September wave consistent with Bini et al. (2009) and Smith and Lee (2011).

Among the modeling techniques analyzed for our study, GWR provided the best model fit, based on estimated global R^2 . Our results revealed where and when local long-term exposure to DPM may have been associated with COVID-19 mortality, consistent with results from both Karaye and Horney (2020) and Mollalo et al. (2020) regarding patterns of local prevalence and local mortality of the disease based on local R^2 . Some areas in the Northeast and West regions presenting a high R^2 in our study align with Mollalo et al. (2020) for incidence rate.

As noted by Fotheringham (1998), our GWR results illustrates the need to account for local phenomena.

Socio-economic disparity could explain the non-stationary effect of DPM exposure on COVID-19 mortality, due to drastic differences between contiguous areas. Socially vulnerable communities, including minoritized racial groups, have seen spatially associated COVID-19 incidence (Karaye and Horney, 2020). This is consistent with the strong association we observed for the fraction Black confounder (Table 3). Moreover, Paolella et al. (2018) pointed out spatial associations among fine particulate matter concentration, health effects, and minoritized groups and found out that finer spatial resolution reveals substantially higher fine particulate matter concentrations in Black and Hispanic communities.

The differences among associations of COVID-19 mortality and DPM concentrations found by the SLM and SEM for the year-long time period, when SLM was demonstrated to be more significant by a Lagrange test, helped to illustrate that neighboring effects were more relevant in modeling the spatial relationship with COVID-19 deaths than unobserved latent variables contained in the error term. Counties near other counties with high COVID-19 incidence are likely to have higher incidence. Nonetheless, since the weighting matrix chosen for our study is based on spatial adjacency, the county size differences between the Eastern and Western U.S. may have affected the parameter estimates creating more uncertainty in the larger counties (Chi and Zhu 2020). Some variability in the association between COVID-19 and DPM exposure within those might have not been captured although DPM sources are more likely to be found in urban areas. However, since the SLM for the year-long time period was not statistically significant, other models should be considered when data are combined across multiple waves.

Several limitations of this study need to be acknowledged with respect to the input data. It is possible that, with more data and/or more time, the association would disappear. Exposure measurement error could bias the results (Villeneuve and Goldberg, 2020). Our spatial modeling approach is intended to account for spatial exposure measurement errors. However, errors from applying cross-sectional analyses persist. Although we studied different waves of the disease, our models were not truly longitudinal. Long-term exposure to DPM was estimated using concentrations from the 2014 NATA. This is the most recent

nationwide prediction of DPM concentrations produced by the U.S. EPA and was also used in Hendryx and Luo (2020) and Petroni et al. (2020). The dataset likely includes higher DPM concentrations than for 2020 given fleet turnover, suggesting that the magnitude of the effects of DPM calculated by our study and these other studies were underestimated. Widely reported undercounting during the January-May wave would further contribute to this underestimation (Dubrow, 2021). The set of potential confounders employed in our models was chosen to evaluate the influence of factors other than DPM potentially associated with COVID-19 outcomes (Wu et al. 2020). However, it was impossible to represent all influential factors in the relationship between each wave of COVID-19 mortality and long-term DPM concentrations, so uncertainty in the potential for confounding existed (Wu et al. 2020, Hendryx and Luo 2020). Furthermore, the study was designed at county level. Spatial variation within counties was not captured and, the difference in county size could have caused uncertainty since the weighting matrix defined for our analyses on which SLM, SEM and GWR relied, was spatial adjacency. Therefore, associations at scales finer than county-level, including individual- and neighborhood-level associations, could not be inferred (Wu et al. 2020). Despite these limitations, our study included a rigorous analysis of spatial relationships for different time periods and tested a variety of potential confounders to minimize these limitations.

Conclusions

Our study used spatial econometric models alongside a local GWR to assess spatial relationships between COVID-19 mortality waves and long-term DPM exposure. Our findings are consistent with prior studies showing a positive association between air pollution and COVID-19 mortality (Wu et al. 2020, Petroni et al. 2020), specifically for DPM (Hendryx and Luo 2020). Our study built on these previous findings by exploring associations of COVID-19 mortality with long-term DPM concentrations across waves of the pandemic. In doing so, our models provided a picture in which long-term DPM exposure may have influenced COVID-19 mortality during the early stages of the disease, as observed specifically for the periods of January-May and June-September 2020. Waning influence of DPM during October to December suggested that person-to-person disease transmission regardless of past exposures may have become more influential in the spread of COVID-19 and in mortality rates once the disease became widespread throughout the U.S. Further investigation might focus on factors associated with COVID-19 mortality during the October-December wave. Although COVID-19 data were available beyond this period, the introduction of vaccines during 2021 were likely to have been so influential that combination of the two years of data may produce misleading conclusions.

Abbreviations

ACS: American Community Survey AERMOD: American Meterological Society/Environmental Protection Agency Regulatory Model AIC: Akaike information criterion CHR: County Health Rankings COVID-19: Coronavirus disease 2019 DPM: Diesel particulate matter EPA: Environmental Protection Agency GWR: Geographically weighted regression ICU: Intensive care unit NATA: National Air Toxics Assessment NO2: Nitrogen dioxide OLS: Ordinary least squares 0₃: Ozone PM2 5: Particulate matter with a nominal diameter of 2.5 µm R²: Coefficient of determination SEM: Spatial error model SLM: Spatial lag model U.S.: United States Declarations Ethics approval and consent

Not applicable

Consent for publication

Not applicable

Availability of data and materials

The datasets supporting the conclusions of this article are in the following repositories:

COVID-19 mortality data can be found in the Johns Hopkins University Global Coronavirus (COVID-19) Database, https://data.world/covid-19-data-resourcehub/covid-19-case-counts/workspace/file?filename=COVID-19+Cases.csv.

Diesel particulate matter data can be found in the U.S. Environmental Protection Agency 2014 National Air Toxics Assessment, https://www.epa.gov/nationalair-toxics-assessment/2014-nata-assessment-results#self.

Community characteristics data can be found in the U.S. Census Bureau 2014-2018 American Community Survey, https://data.census.gov/cedsci/table? d=ACS%205-

Year% 20 Estimates% 20 Data% 20 Profiles&g=0400000 US22.050000&tid=ACSDP5Y2018.DP05&hidePreview=false&tp=true&moe=true&vintage=2018.DP05&hidePreview=false&tp=true&vintage=2018.DP05&hidePreview=4005BidePreview=4005BidePreview=4005BidePreview=4005BidePreview=4005BidePreview=4005BidePreview=4005BidePreview=4005BidePreview=4005Bid

Community characteristics data can also be found in the Robert Wood Johnson Foundation 2020 County Health Rankings, http://www.countyhealthrankings.org.

Competing interest

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

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Authors' contributions

MEM designed the modeling process, wrote the R code, and wrote the manuscript. JG advised MEM on the study design and modeling processes and edited the manuscript. JRB advised MEM on the study design and modeling processes, assisted in writing the manuscript, and edited the manuscript. All authors have read and approved the manuscript.

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References

- Bini LM, Diniz-Filho JAF, Rangel TF, Akre TS, Albaladejo RG, Albuquerque FS, Aparicio A, Araújo MB, Baselga A, Beck J, Bellocq MI, Böhning-Gaese K, Borges PAV, Castro-Parga I, Chey VK, Chown SL, De Marco Jr. P, Dobkin DS, Ferrer-Castán D, Field R, Filloy J, Fleishman E, Gómez JF, Hortal J, Iverson JB, Kerr JT, Dissling WD, Kitching IJ, León-Cortés JL, Lobo JM, Montoya D, Morales-Castilla I, Moreno JC, Oberdorff T, Olalla-Tárraga MÁ, Pausas JG, Chi G, Zhu J. (2020) Spatial Regression Models for the Social Sciences. SAGE Publications, Inc., Newbury Park, CA.
- Bozack A, Pierre S, DeFelice N, Colicino E, Jack D, Chillrud SN, Rundle A, Astua A, Quinn JW, McGuinn L, Yang Q, Johnson K, Masci J, Lukban L, Maru D, Lee AG. (2022) Long-term air pollution exposure and COVID-19 mortality. Am J Respir Crit Care Med. 205(6):651-662. Doi://10.1164/rccm.202104-08450C
- 3. Centers for Disease Control (CDC). (2020) COVID-19. https://www.cdc.gov/coronavirus/2019-ncov/index.html. Accessed August 30, 2021.
- 4. Chumney ECG, Simpson KN. (2006) Methods and Designs for Outcomes Research. American Society of Health-System Pharmacists, Bethesda, MD.
- 5. Clark A, Jit M, Warren-Gash C, Guthrie B, Wang HH, Mercer SW, Sanderson C, McKee M, Troeger C, Ong KL, Checchi F, Perel P, Joseph S, Gibbs HP, Banerjee Am Eggo RM. (2020) Global, regional, and national estimates of the population at increased risk of severe COVID-19 due to underlying health conditions in 2020: A modelling study. The Lancet Global Health. 8(8): e1003-e1017; Doi://10.1016/S2214-109X(20)30264-3
- 6. Clougherty JE, Shmool JL, Kubzansky LD. (2014) The role of non-chemical stressors in mediating socioeconomic susceptibility to environmental chemicals. Current Environmental Health Reports. 1: 302-313; Doi://10.1007/s40572-014-0031-y
- 7. Davis JA, Meng Q, Sacks JD, Dutton SJ, Wilson WE, Pinto JP. (2011) Regional variations in particulate matter composition and the ability of monitoring data to represent population exposures. Sci Total Environ. 409: 5129-5135. Doi://10.1016/j.scitotenv.2011.08.013.
- 8. DeAngelis DL, Yurek S. (2017) Spatially explicit modeling in ecology: A review. Ecosystems. 20: 284-300; Doi://10.1007/s10021-016-0066-z
- 9. Dong E, Du H, Gardner L. (2020) An interactive web-based dashboard to track COVID-19 in real time. Lancet. 20: 533-534; Doi://10.1016/S1473-3099(20)30120-1

- Douglas JA, Archer RS, Alexander SE. (2019) Ecological determinants of respiratory health: Examining associations between asthma emergency department visits, diesel particulate matter, and public parks and open space in Los Angeles, California. Preventive Medicine Reports. 14: 100855; Doi://10.1016/j.pmedr.2019.100855
- 11. Dubrow JK. (2021) Local data and upstream reporting as sources of error in the administrative data undercount of Covid 19. Int J Soc Res Method. Doi://10.1080/13645579.2021.1909337.
- 12. Fotheringham AS, Brunsdon C, Charlton M. (2003) Geographically Weighted Regression: The Analysis of Spatially Varying Relationships. John Wiley & Sons, New York.
- 13. Fotheringham AS, Charlton ME, Brunsdon C. (1998) Geographically weighted regression: a natural evolution of the expansion method for spatial data analysis. Environ Plan A. 30: 1905-1927; Doi://10.1068/a301905
- 14. Gao C, Feng Y, Tong X, Lei Z, Chen S, Zhai S. (2020) Modeling urban growth using spatially heterogeneous cellular automata models: Comparison of spatial lag, spatial error and GWR. Computers, Environment and Urban Systems. 81: 101459; Doi://10.1016/j.compenvurbsys.2020.101459
- 15. Goldberger AS. (1964) Classical Linear Regression. In: Econometric Theory. John Wiley & Sons, New York.
- 16. Hendryx M, Luo J. (2020) COVID-19 prevalence and fatality rates in association with air pollution emission concentrations and emission sources. Environ Pollut. 265: 115126; Doi://10.1016/j.envpol.2020.115126.
- 17. Johns Hopkins University. (2020) Global Coronavirus (COVID-19) Data. Accessed April 26, 2020. https://data.world/covid-19-data-resource-hub/covid-19case-counts/workspace/file?filename=COVID-19+Cases.csv.
- 18. Karaye IM, Horney JA. (2020) The impact of social vulnerability on COVID-19 in the US: an analysis of spatially varying relationships. American Journal of Preventive Medicine. 59: 317-325; Doi://10.1016/j.amepre.2020.06.006
- 19. Kim D. (2021) Predicting the magnitude of residual spatial autocorrelation in geographical ecology. Ecography. 44: 1121-1130; Doi://10.1111/ecog.05403
- Lancaster K, Rhodes T, Rosengarten M. (2020) Making evidence and policy in public health emergencies: lessons from COVID-19 for adaptive evidencemaking and intervention. Evidence Policy Doi: https://doi.org/10.1332/174426420X15913559981103.
- 21. Lee K-H, Jung H-J, Park D-U, Ryu S-H, Kim B, Ha K-C, Kim S, Yi G, Yoon C. (2015)Occupational Exposure to Diesel Particulate Matter in Municipal Household Waste Workers. PLoS ONE. 10(8): e0135229; Doi://10.1371/journal.pone.0135229
- 22. Le Gallo J, Baumont C, Dall'Erba S, Ertur C. (2005) On the property of diffusion in the spatial error model. Applied Economics Letters. 12: 533–536; Doi://10.1080/13504850500120722
- 23. LeSage JP, Fischer MM. (2008) Spatial growth regressions: model specification, estimation and interpretation. Spatial Economic Analysis. 3: 275-304; Doi://10.1080/17421770802353758
- 24. LeSage J, Pace RK. (2009) Introduction to Spatial Econometrics. Chapman and Hall/CRC, London.
- 25. Loonis V, de Bellefon MP. (2018) Handbook of Spatial Analysis: Theory and practical application with R. Insee Méthodes, 131.
- 26. Miyara M, Tubach F, Pourcher V, Morélot-Panzini C, Pernet J, Haroche J, Lebbah S, Morawiec E, Gorochov G, Caumes E, Hausfater P, Combes A, Similowski T, Amoura Z. (2020) Low rate of daily smokers in patients with symptomatic COVID-19. medRxiv. Doi://10.1101/2020.06.10.20127514.
- 27. Mollalo A, Vahedi B, Rivera KM. (2020) GIS-based spatial modeling of COVID-19 incidence rate in the continental United States. Science of The Total Environment. 138884; Doi://10.1016/j.scitotenv.2020.138884
- 28. Moran PAP. (1950) Notes on Continuous Stochastic Phenomena. Biometrika. 37: 17-23; Doi://10.2307/2332142
- 29. Nieman DC. (2020) Coronavirus Disease-2019: A tocsin to our aging, unfit, corpulent, and immunodeficient society. J Sport Health Sci. Doi://10.1016/j.jshs.2020.05.001
- 30. Paolella DA, Tessum CW, Adams PJ, Apte JS, Chambliss S, Hill J, Muller NZ, Marshall JD. (2018) Effect of model spatial resolution on estimates of fine particulate matter exposure and exposure disparities in the United States. Environ Sci Technol. 5: 436-441. Doi://10.1021/acs.estlett.8b00279.
- 31. Peek ME, Simons RA, Parker WF, Ansell DA, Rogers SO, Edmonds BT. (2021) COVID-19 among African Americans: an action plan for mitigating disparities. American Journal of Public Health. 111: 286-292; Doi://10.2105/AJPH.2020.305990
- 32. Petrilli CM, Jones SA, Yang J, Rajagopalan H, O'Donnell L, Chernyak Y, Tobin KA, Cerfolio RJ, Francois F, Horwitz LI. (2020) Factors associated with hospital admission and critical illness among 5279 people with coronavirus disease 2019 in New York City: prospective cohort study. BMJ. 369; Doi://10.1136/bmj.m1966.
- 33. Petroni M, Hill D, Younes L, Barkman L, Howard S, Howell IB, Mirowsky J, Collins MB. (2020) Hazardous air pollutant exposure as a contributing factor to COVID-19 mortality in the United States. Environmental Research Letters. 15: 0940a9; Doi://10.1088/1748-9326/abaf86/meta
- 34. Phillips N, Park IW, Robinson JR, Jones HP. (2020) The Perfect Storm: COVID-19 Health Disparities in US Blacks. Journal of Racial and Ethnic Health Disparities. 1-8; Doi://10.1007/s40615-020-00871-y
- 35. Pronk A, Coble J, Stewart PA. (2009) Occupational exposure to diesel engine exhaust: a literature review. Journal of Exposure Science and Environmental Epidemiology. 19: 443-457.
- 36. Qian H, Rahbek C, Rodríguez MÁ, Rueda M, Ruggiero A, Sackmann P, Sanders NJ, Terribile LC, Vertaas OR, Hawkins B. (2009) Coefficient shifts in geographical ecology: an empirical evaluation of spatial and non-spatial regression. Ecography. 32: 193-204; Doi://10.1111/j.1600-0587.2009.05717.x
- Rahman MH, Zafri NM, Ashik FR, Waliullah M. (2020) GIS-based spatial modeling to identify factors affecting COVID-19 incidence rates in Bangladesh. medRxiv. Doi://10.1101/2020.08.16.20175976.
- 38. Reyes MV. (2020) The disproportional impact of COVID-19 on African Americans. Health and Human Rights. 22: 299.

- Ristovski ZD, Miljevic B, Surawski NC, Morawska L, Fong KM, Goh F, Yang IA. (2012) Respiratory health effects of diesel particulate matter. Respirology. 17(2), 201-212; Doi://10.1111/j.1440-1843.2011.02109.x
- 40. Robert Wood Johnson Foundation. (2020) County Health Rankings. Accessed August 25, 2020. http://www.countyhealthrankings.org.
- 41. Schabenberger O, Gotway CA. (2017) Statistical Methods for Spatial Data Analysis. CRC press, London.
- 42. Sidell MA, Chen Z, Huang BZ, Chow T, Eckel SP, Martinez MP, Lurmann F, Thomas DC, Gilliland FD, Xiang AH. (2022) Ambient air pollution and COVID-19 incidence during four 2020-2021 case surges. Environ Res. 208: 112758. Doi://10.1016/j.envres.2022.112758
- 43. Smith TE, Lee KL. (2012) The effects of spatial autoregressive dependencies on inference in ordinary least squares: a geometric approach. Journal of Geographical Systems. 14: 91-124; Doi://10.1007/s10109-011-0152-x
- 44. Stakhovych S, Bijmolt TH. (2009) Specification of spatial models: A simulation study on weights matrices. Papers in Regional Science. 88: 389-408; Doi://10.1111/j.1435-5957.2008.00213.x
- 45. Sun F, Matthews SA, Yang TC, Hu MH. (2020) A spatial analysis of the COVID-19 period prevalence in US counties through June 28, 2020: Where geography matters? Annals of Epidemiology. Doi://10.1016/j.annepidem.2020.07.014
- 46. U.S. Census Bureau. (2020) 2014-2018 American Community Survey. Accessed April 24, 2020. https://data.census.gov/cedsci/table?d=ACS%205-Year%20Estimates%20Data%20Profiles&g=0400000US22.050000&tid=ACSDP5Y2018.DP05&hidePreview=false&tp=true&moe=true&vintage=2018.
- 47. U.S. Environmental Protection Agency. Technical Support Document. (2018) 2014 National Air Toxics Assessment. U.S. Environmental Protection Agency, Research Triangle Park, NC. https://www.epa.gov/national-air-toxics-assessment/2014-nata-assessment-results#self
- 48. Villeneuve PJ, Goldberg MS. (2020) Methodological considerations For epidemiological studies of air pollution and the SARS and COVID-19 coronavirus outbreaks. Environ Health Perspect. 128(9): 095001. Doi://10.1289/EHP7411.
- 49. Williamson EJ, Walker AJ, Bhasharan K, Bacon S, Bates C, Morton CE, Curtis HJ, Mehrkar A, Evans D, Inglesby P, Cockburn J, McDonald HI, MacKenna B, Tomlinson L, Douglas IJ, Rentsch CT, Mathur R, Wong AYS, Grieve R, Harrison D, Forbes H, Schultze A, Croker R, Parry J, Hester F, Harper S, Perera R, Evans SJW, Smeeth L, Goldacre B. (2020) Factors associated with COVID-19-related death using OpenSAFELY. Nature. 584: 430-436; Doi://10.1038/s41586-020-2521-4.
- 50. Wu X, Nethery RC, Sabath MB, Braun D, Dominici F. (2020) Air pollution and COVID-19 mortality in the United States: Strengths and limitations of an ecological regression analysis. Science Advances. 6: Doi://10.1126/sciadv.abd4049
- 51. Yancy CW. (2020) COVID-19 and African Americans. JAMA. 323(19): 1891-1892; Doi://10.1001/jama.2020.6548

Figures

Figure 1

Spatial distribution of DPM concentration across contiguous U.S. counties (µg/m³).

a) January – May

b) June - September



Figure 2

Spatial distribution of COVID-19 deaths for (a, top left) January-May, (b, top right) June-September, (c, bottom left) October-December, and (d, bottom right) all of 2020.

a) January - May





Figure 3

Map of associations between COVID-19 mortality and long-term DPM concentration for U.S. counties.



Figure 4

Spatial distribution of local R² for the GWR model.

Supplementary Files

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