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The effects of increased temperatures (Heat waves) on Cardiovascular health
among adults in Atlanta

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An abstract of
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Abstract

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By Wandave Tizhe

Background: In the coming years, average temperatures will increase as climate change continues. These temperature changes can have a direct effect on human health, including in people with cardiovascular diseases; numerous studies in the United States have reported a link between high ambient temperatures and increased emergency department (ED) visits due to cardiovascular diseases. In this study, we assessed the association between summer temperature (May and September, 2016-2018) and hypertension using ED data for adults in Atlanta Georgia.

Methods: We examined the association between ambient temperature and daily ED visits for hypertensive diagnoses among adults who visited four of Emory University's hospitals. We also explored the impact of having a history of hypertension. We analyzed the data using a time-stratified case-crossover study design, with conditional logistic regression, at up to three days of lag. The case-crossover study design controls for time trends and individual-level confounders, as each individual serves as their own control. We estimated odds ratios (ORs) and 95% confidence intervals (CI) for maximum temperature changes for each degree celsius. We also investigated potential non-linearity in the temperature-morbidity relationship, by evaluating the association within each temperature quartile.

Results: During the 3 year study period from 2016 – 2018, there were 220,210 adult ED visits. Of these, 33,565 (15.2%) were patients with a previous history of hypertension. The reason for the visit was hypertension in twelve percent of cases. Those who presented with hypertension and had a hypertensive history were 15,159 (6.8%) of total visits. About a quarter of all visits were in the elderly. We observed significant ($p < 0.05$) associations between heat and all ED visits at lags 0, 1 and 2 days. We observed no significant associations with visits for hypertension or in visits where the cases had a history of hypertension. There was no evidence of non-linearity.

Conclusion: We found that that increased temperatures increased the rate of visits to the ED but not for hypertensive visits in particular or visits in patients with a history of hypertension.

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INTRODUCTION

The Earth is warming due to elevated concentrations of greenhouse gases, and will continue to warm in the future (Holli Riebeek et al, 2010). The decadal global land and ocean surface average temperature for 2011–2020 was the warmest decade on record, with a surface temperature of +0.82°C (+1.48°F) above the 20th-century average (NOAA., State of climate 2021). In the USA, like many other places, heatwaves have become more frequent and intense, and cold waves have become less frequent (Kristiane Huber et al, 2017). Annual average U.S temperatures are projected to increase by 2.5°F and 8°F by the end of this century, depending primarily on the level of future emissions of greenhouse gases, under the RCP2.6 and RCP 8.5 scenarios respectively. (John Walsh, et al., 2014). These temperature changes will have direct effects on human health and the environment (Balbus et al).

Climate change has increased the frequency and the intensity of high temperatures, which are currently the leading cause of weather-related deaths in the United States (Greg Fischer et al 2021). For example, one study reported that extreme heat accounted for about 31% of all weather-related deaths from 2006 to 2010 (Balbus, J et al., 2016). Numerous study results in the United States have shown a link between high ambient temperatures and increased mortality, including for cardiovascular diseases (Medina-Ramón, M., & Schwartz, J. 2017). For example, a large study of 43 cities in the United States estimated that the daily mortality rate during heatwave days was 3.7% higher on average than non-heatwave days during 1987–2005 (Anderson GB, Bell ML 2011). A 2017 meta-analysis reported that the risk of cardiovascular mortality,

specifically, increased by 1.3% per degree rise in temperature (RR, 1.013; 95% CI [1.011–1.015]) (Moghadamnia MT, et al 2017).

In addition to mortality, exposure to hot outdoor temperatures is also associated with severe morbidities, including emergency department (ED) visits and/or hospital admissions (Harikrishna, H et al 2020). This has been reported for many specific causes, including cardiovascular disease (Harikrishna .H., et al 2020, Astrom DO et al 2011,). Another study showed that heatwaves appeared to be associated with cardiovascular and respiratory morbidities (RE: 0.999, 95%CI: 0.996, 1.002, p-value = 0.61 for cardiovascular morbidity; RE: 1.043, 95%CI: 0.995, 1.093; p-value = 0.08 for respiratory morbidity) (Cheng J et al, 2019). Studies also indicate that the strengths of associations between temperature and ED visits varies by age; the elderly and children have been identified as two particularly susceptible populations (Schifano P, et al 2009, Astrom et al 2011).

Associations between heat and cardiovascular disease are also supported by plausible biological mechanisms. For example, one theory is that high temperatures can increase the occurrence of heart attacks and strokes in susceptible patients because of increased blood viscosity (Baaghideh M, Mayvaneh F, et al 2017). Other proposed mechanisms between heat and cardiovascular outcomes include increased surface blood circulation and sweating, which may lead to increased cardiac workload, dehydration and salt depletion, hemoconcentration, elevated blood viscosity, and the risk of thrombosis (Bouchama, A. and J.P. Knochel, 2012). Moreover, heat stress was suggested to induce the release of interleukins modulating local and systemic

acute inflammatory responses (Bouchama, A. and J.P. Knochel, 2012). These inflammatory responses can result in heart failure by increasing damage to heart tissue and inflammation (Wilker EH, et al, 2012).

The elderly population may be at a higher risk for physiological and behavioral reasons, such as existing cardiovascular diseases, impaired kidney function, and living alone with limited social support (Kovats RS, Kristie LE. 2017). Individuals who are confined to bed and unable to care for themselves may be at high risk of death during heat waves, possibly due to their limited access to emergency care (Hajat S, Armstrong, et al, 2006, Knowlton K, et al, 2009). Children may be vulnerable in part because their renal systems are stressed by a series of thermoregulatory adjustments under excessive heat (Xu Z et al 2012) as well as their activity patterns (Vanos JK, 2015).

When put together, these results suggest that more heat-related cardiovascular mortality may be expected in the future with climate change (Chen, B., et al., 2021). This is a particular concern because of the high disease burden from cardiovascular disease worldwide, including in the United States (Ponjoan, A. et al 2017).

To prevent temperature-related morbidity, it is important to protect vulnerable populations. In addition to children and the elderly, another potentially vulnerable population is people with a history of cardiovascular disease. In this study, our objective was to estimate warm-season associations between

increased temperatures (heat) and daily ED visits for hypertension in Atlanta, Georgia. Atlanta is located in the southeastern United States, which tends to experience more intense heat and humidity than rest of the United States (Bonan, G.B. 2017). Atlanta has also experienced rates of increase in heatwave frequency and duration that are higher than the national averages from 1961 to 2010 (Habeeb, D., et al, 2015)

This study aimed to answer the following two research questions (1) Does increased temperature prompt ED visits for hypertension in Atlanta?, and (2) does a history of hypertension make people more vulnerable to ambient temperature? To answer these questions, we used a time-stratified case-crossover study design, a statistical technique well suited to examining short-term exposures with the acute outcome.

Our findings can play an important role in supporting local emergency preparedness, performing detailed risk assessment, and protecting public health (Ebi KL, et al 2015, Frumkin H et al 2008). For example, identification of heat metrics most associated with adverse health outcomes may result in more effective local warning systems.

METHODS

As mentioned above, we conducted a case-crossover study on the association between ambient temperature and ED visits for hypertension in Atlanta, GA, USA.

Study area

Atlanta is the capital of Georgia and its most populous city. With an estimated 6 million people in the metropolitan area in 2020, it is also the 37th most populous city in the United States, with 51% black, 38% white, 4% Asian, and other groups comprising 2.3%. The median age of 33.5 (Census.gov). The climate of Atlanta and its metropolitan area is humid subtropical according to the Köppen classification, with four seasons including hot, humid summers and cool winters that are occasionally cold by the standards of the southern United States. (climaterealityproject.org) Summers are long and consistently hot and humid.

Health dataset

Data on ED visits comes from the Emory Clinical Data Warehouse dataset from the Emory University School of Medicine Emergency Department, and includes individual-level patient records from four of Emory's hospitals. We collected CDW data on all ED visits for three consecutive summers between 2016 and 2018. We restricted our analysis to the warm season of May 1st to September 30th to focus on the effects of high temperatures. We included any ED visit if the patient was an adult (18 or older) and lived in Atlanta. We

looked at all ED visits and those diagnosed with the ICD 10 code for hypertension (I10).

We defined each ED visit as a case; thus, a patient could be in the dataset more than once if the person experienced multiple visits to the ED over the study period. Relevant data elements included the date of the visit, patient age, prior medical history and diagnosis (reason for visit). We were also able to extract data on whether the patient had a self-reported history of hypertension.

Temperature dataset

We obtained daily observations of minimum and maximum temperature during the summer season May to September 2016-2018 from the Atlanta Hartsfield international airport station for the 3-year study period of 2016 through 2018. The daily temperatures were merged with the ED data by date.

Statistical Analysis

We analyzed the data using a time-stratified case-crossover study design, with conditional logistic regression. The approach is a modification of the matched case-control study, where each case serves as his or her own control so that known and unknown time-invariant confounders are inherently adjusted for by design. We compared the temperature on the day of the ED visit (case) with up to four control periods on the same day of the week within the same month and year to control for time trends and day of the week.

In previous studies of apparent temperature and mortality (Kovats RS, Kristie LE, 2016) and hospital visits (Moghadamnia MT, 2017), acute effects of same-day temperature were found to have the best model fit. Thus, our a priori focus for this analysis was the effect of same-day temperature (lag0). We did, however, examine other single-day lags up to three days before the ER visit on lag days 0,1, 2 and 3

Using R studio, conditional logistic regression models were used to obtain estimates of odds ratios (ORs) and 95% confidence intervals (CIs) associated with a 1 degree increase in temperature ($^{\circ}\text{C}$). In our main analyses we assumed a linear effect, but also explored non-linearity by estimating effects for each temperature quartile.

We had an approved Emory IRB for studying environmental exposures and acute morbidity and we listed the CDW data as the data source for this study.

Results

During the 3 year study period from 2016 – 2018, there were 220,210 adult ED visits. Of these, 33,565 (15.2%) were patients with a previous history of hypertension (Table 1). The cause for the visit was hypertension in twelve percent of the visits. Those who presented with hypertension and had a hypertensive history were 15,159 (6.8%) of total visits. About a quarter of all visits were in the elderly.

Gender-based presentation to the ED were similar, Females had 50.3% with males 49.7% of the total visits for hypertension with 128,817 and 93227 respectively. These levels increase a bit when compared to those who presented with hypertension comorbidity with females having 55% and males 45% (33,011 and 27,658 respectively).

Table 1. Descriptive statistics of emergency department visits (HTN=hypertension).

ED visits	Total number	Total %	% Male
All visits	220210	100%	42%
HTN visits	26,454	12%	49.7%
Visits with HTN history	33,565	15.20%	45.87%
HTN visits with HTN history	15,159	6.80%	49.6%
All Visits > 65 years	53,124	24.12%	42.2%
HTN visits >65 years	13,004	5.90%	46.8%
All visits >65 years	16,816	7.60%	43%

with history			
Visits >65 years with H TN and a history of HTN	7,983	3.60%	46.5%

Over the three-summer study period, the median daily maximum temperature was 31.7°C with an interquartile range of 13°C, a minimum of 13°C and a maximum of 38°C. June was the hottest month and May was the coolest month. A temperature time-series can be found in Figure 1.

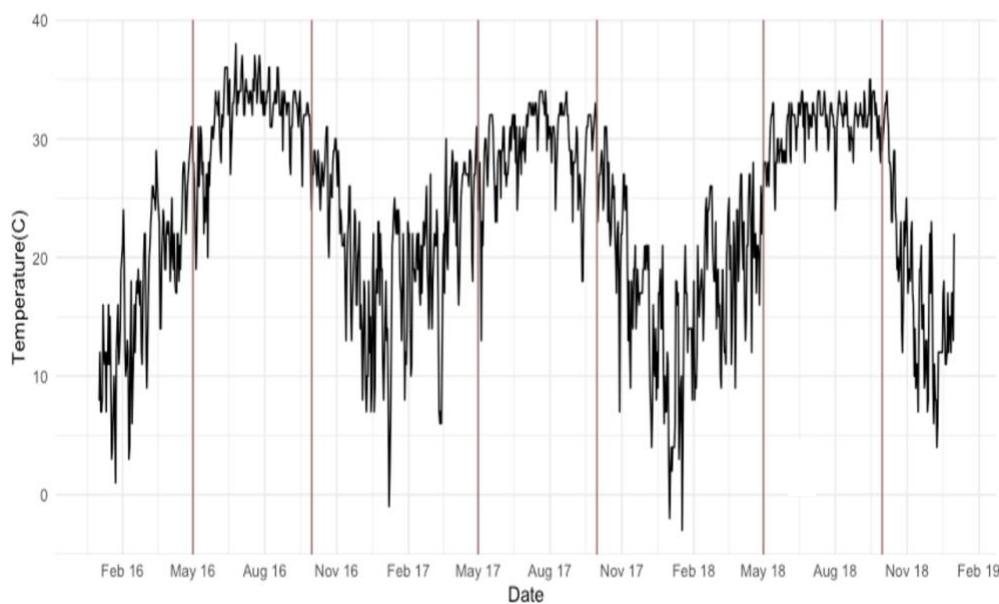


Figure 1. Temperature time series; Study is period between May and September 2016-2018 shown within the highlighted lines

Figure 2. shows the odds ratios for the association between temperature and ED visits for each outcome for each lag. We observed no significant associations except for all visits with lag 0, lag 1, and lag 2.”

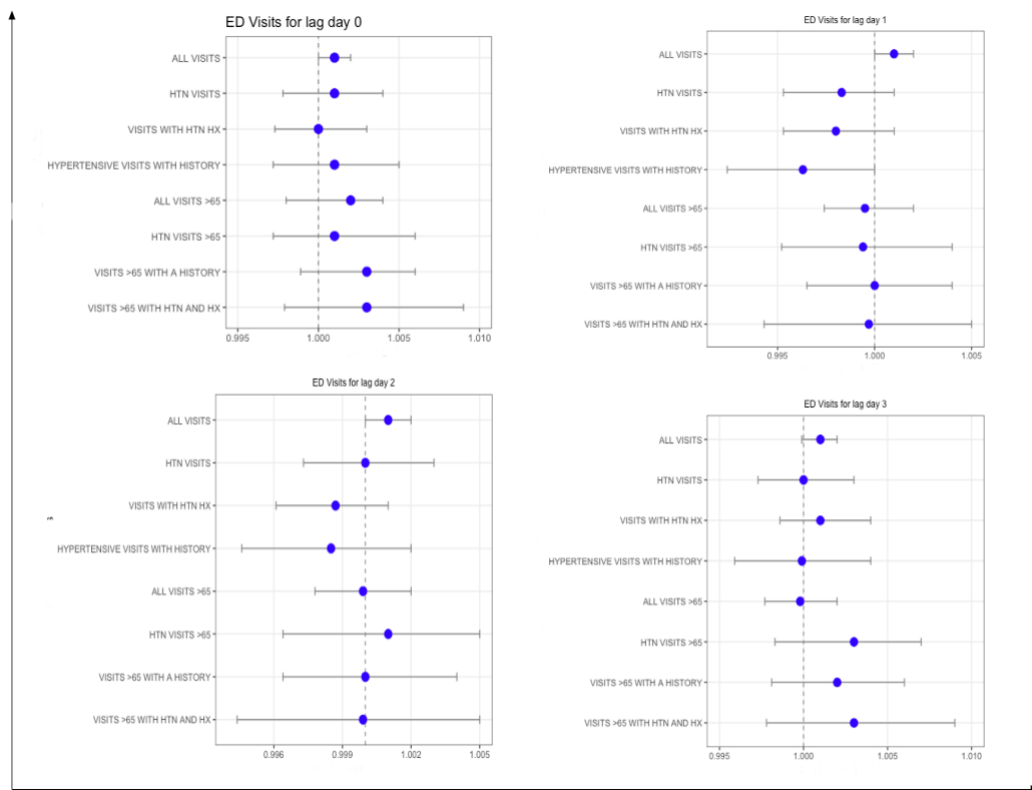


Figure 2. Odds ratios for the association between temperature and ED visits for various outcomes on each lag days

To analyze the potential non-linearity in the temperature-morbidity relationship, we evaluated the association for each temperature quartile (Figure 3). There were no statistically significant associations observed.

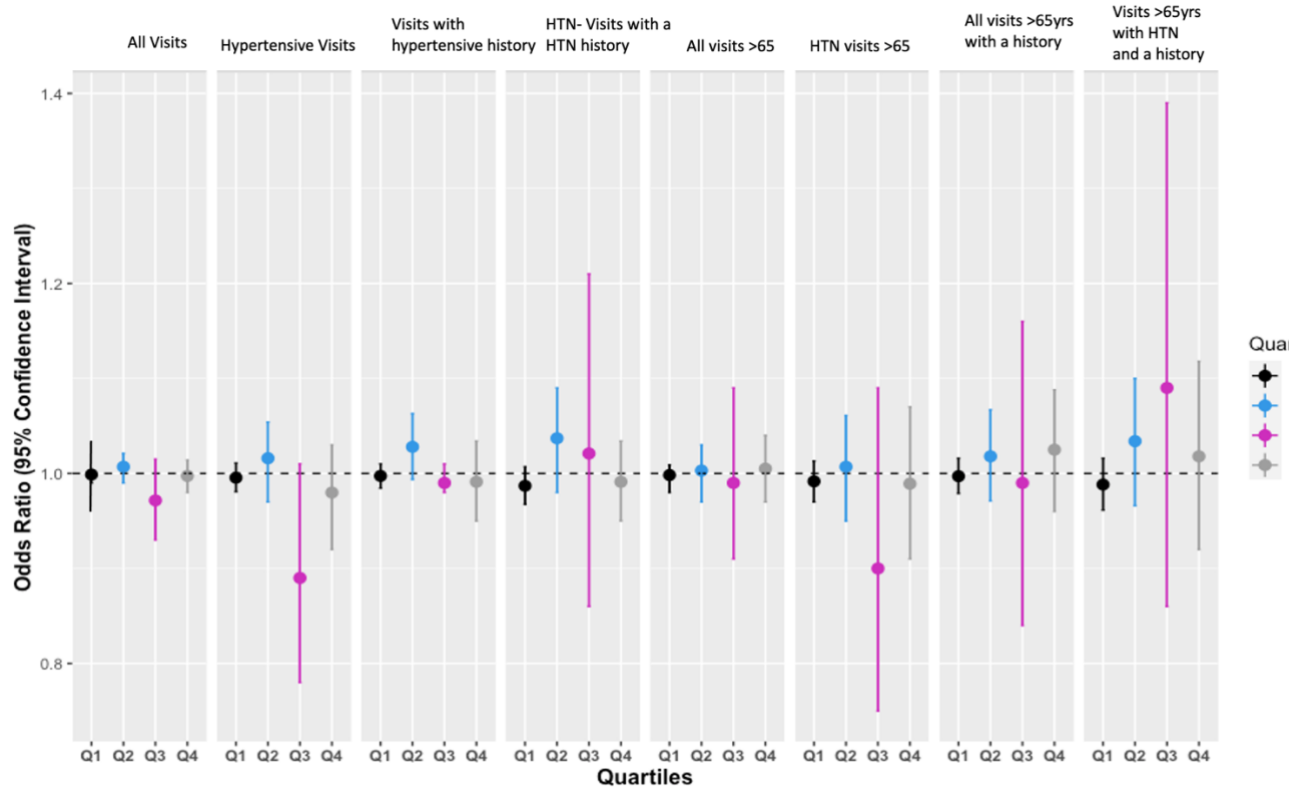


Figure 3. Associations by temperature quartile at lag day 0.

Discussion

We found a statistically significant result for all ED visits and daily ambient temperature in Atlanta during the 2016-2018 warm seasons at lag days 0,1, and 2. We did not find any significant associations with hypertension, history of hypertension, hypertensive visits with a history of hypertension, all visits greater than 65, hypertensive visits greater than 65, hypertensive history among patients greater than 65, and visits with hypertension and history in patients greater than 65 years. This was also true when we explored temperature quartiles.

The finding of a significant association with all-age all-cause ED visits has been reported in a number of other studies (Dong et al, 2016; Mengxuan et al,2019; Basu et al,2012; Kenney et al,2014; Zheng S, et al,2016; Mai Li et al,2016; Huang J, et al, 2014). However, it was surprising that we did not find an association with all visits over age 65, which has also been commonly reported (Fuhrmann CM, et al., 2011; Phung, D et al., 2016; Dadbakhsh M et al., 2012). Most other studies have also shown a positive statistically significant relationship between elderly hypertensive visits on hot days especially during the summer (Zheng S et al.,2016; Mai Li et al.,2016; Huang J et al.,2014; Kim YM et al.,2016 Kenny L.P et al.2010)In terms of hypertension specifically, the literature is mixed. (Basu et al., 2012; Kim S.E. et al., 2019; Lin S et al., 2009) found no association or a negative association between temperature and ED visits for hypertension, as did (Winqvist A., et al. 2016) in a study in Atlanta using different data from ours. Other studies, however, have observed positive associations (Phung D et al., 2016 McNaughton et al., 2013; Candace D et al. 2012)

When we analyzed the ED visits for patients with a prior history of hypertension we found a non-statistically significant result for this association (OR: 1.00, p-value: 0.97, C.I: 0.99 – 1.003), similar to a study by Basu et al. 2012;(Basu et al on the effects of ambient temperatures on emergency room visits in 2012.) That study did find associations when with other cardiovascular diseases, which, along with our findings, may suggest that observed cardiovascular effects are being driven by diseases other than hypertension.

The lack of an association – or of a possible protective association – may have mechanistic support. As suggested by Winkler, A et al., 2016; They studied warm season temperatures and emergency department visits in Atlanta. For example, studies have reported an inverse linear relationship between increasing temperature and increased blood pressure (Chen et al., 2013; Chen et al., 2015; Lanzinger et al., 2014; Q. Wang et al., 2017; S. Wang et al., 2017). This may only be true in warmer seasons (Hozawa et al., 2011). However, Madaniyazi et al. found a V-shaped relationship between daily temperature and blood pressure (Madaniyazi et al., 2016).

Our lack of association could also be because we only looked at outdoor temperatures. Many studies have reported statistically significant associations between indoor temperatures and ED visits for cardiovascular events. (Jung CC et al., 2020) compared indoor temperatures on cardiovascular disease related emergency department visits, where they found significant results for increased indoor temperature and hypertensive visits to the ED. This is similar to studies by (Zhao et al., 2019; Saeki K et al., 2014; Kim S et al., 2019; Glen P Kenney et al., 2014; BW Waugh et al., 2021).

Our study area is an urban setting with a generally warm climate and mild winters, with an air conditioning prevalence of over 94% (U.S. Department of Commerce, 2011). This could be responsible for the largely non-significant results we observed, if local populations are adapted to heat. Studies carried out by Arbuthnot et al on the changes in population susceptibility to heat and

cold over time showed that the presence of air conditioning and other adaptive measures reduced the burdens of heat among these populations. In 2010 Ostro et al discovered that increased air conditioning could lead to a decreased association between temperature and hospital admissions. Other studies by (Anderson GB et al., 2015; Hatvani-Kovacs G, et al. 2016; Waugh et al., 2021) also reported similar findings.

Studies that compared heat waves and morbidity showed a statistically significant association between heat waves and cardiovascular events (Winquist et al.,2016; Hajat et al., 2015; Sun et al.,2014)

Several studies have shown some reduced association between increased temperatures and hypertensive visits (Bai L et al.,2016 Ponjoan A et al.,2017; Hu J et al.,2014; Janke et al., 2013) in the southern part of the united states compared to the northern states, (Wang Y et al.,2017; Anderson and Bell., 2009; Saha et al., 2015), for reasons related to the population's ability to adapt to high temperatures from different adaptive measures, either increased compliance with medication, proper control of the disease, planting of trees, Air conditioning use or more cooling centers.

Strength and limitations

This is one of the few studies that evaluated acute exposures of heat and hypertension among adults in Atlanta using Emergency department data. This data provides information on early acute cardiovascular events on presentation. Our case-crossover study design controlled for confounders that may have been present during the study period by using each individual as both cases and controls. Hence age, sex, race, or ethnicity were controlled for. However we did not control for Air pollution and humidity like other studies, this could also have affected the strength of the association between heat and increased temperature. We also didn't control for holidays, time trends and participation periods which was all done in the (Winqvist et al., 2016; Chen T et al., 2017) studies that showed significant associations.

Disease severity could be a possible confounder, studies have shown that most patients with existing cardiovascular diseases don't present with hypertension but other diagnosis like a stroke, myocardiac infarction and other internal causes (Winqvist A et al.,2016; Basu et al.,2012 Zheng S et al.,2016 Mai Li et al., 2016;Chen T et al., 2017).

Because heat as a cause of Hypertension has not been fully explored as the primary cause of elevated blood pressure there could have been some under-reporting of hypertensive visits, or possible it could have been mis diagnosed, as a handful of medical practitioners don't immediately correlate the association between heat and hypertensive visits. A report, Medical alert; Climate change is harming our health in Wisconsin, by Jonathan pats in October 2020 reported that medical practitioners in the state underreported the

effects of heat on chronic diseases including cardiovascular events like hypertension.

Another reason could be because we only had a subset of Atlanta hospitals (Emory University Hospital) which is not a complete representation of all ED hospital visits for cardiovascular events in Atlanta.

Georgia is one of the slowest-warming states in the US over the years Atlanta have developed a green sustainable environment, most green spaces reduce the detrimental effects of increased temperatures on human health.

(climaterealityproject.org) This could be another reason why we didn't have a statistically significant relationship in our study.

Most people living in Atlanta have adapted to the weather as their physiology have fully adjusted to the humid hot summers and not show a significant effect on their cardiovascular health. Studies have shown that colder states have shown more heat related ED visits compared to warmer states (Medical alert; Climate change is harming our health in Wisconsin, by Jonathan Pats in October 2020; Medina-Ramon et al., 2007; Gasparrini et al., 2015; Curriero et al., 2002; Ponjoan A et al., 2017). Also most people are more likely to be indoors fully air conditioned rather than go out during an extremely hot day. We also didn't incorporate outdoor workers, athletes, military personnel's and people without air conditioning who are the most vulnerable to outdoor temperatures (EPA). Despite not finding a significant relationship among the older adults for hypertensive visits we didn't look at people on medications that could make them vulnerable to increased temperatures. (Layton JB et al., 2020) studied the effect of heat adaptation and medication use on heat related hospitalization for chronic diseases.

The Period of the study was relatively short compared to most studies that found statistically significant results between heat and hypertension were over significantly longer periods of time greater than 10-15 years of study. That showed a greater relationship between heat and cardiovascular diseases. (Mengxuan et al.,2019; McNaughton et al.,2013; Candace D et al., 2013; Gronlund CJ et al., 2014).

We also compared hot days to hypertension which is a broad cardiovascular event that many people over the years have learned to control through physical exercise, environmental sustainability and medications.

Conclusion

We observed no significant association between increased temperature and hypertensive visits to the ED, including in patients with a history of hypertension. However, we did find a statistically significant association for all visits to the ED for all causes, which shows that increased temperatures increased the rate of visits to the ED but not for hypertensive visits, Future studies could help establish public health preparedness and interventions to reduce the adverse health effects of high temperatures especially among vulnerable populations all across Atlanta being affected by increasing temperatures. Identifying the mechanism by which this happens can be pivotal in reducing the global burden of cardiovascular diseases in the US which is the leading cause of death in the US.

References

1. Harikrishna, H et al 2020. Heat exposure and cardiovascular health : Climate and health technical report series USA, July 2020.
2. Ponjoan, A., Blanch, J., Alves-Cabratos, L. et al. Effects of extreme temperatures on cardiovascular emergency hospitalizations in a Mediterranean region: a self-controlled case series study. *Environ Health* 16, 32 (2017).
<https://doi.org/10.1186/s12940-017-0238-0>
3. Zhang, B., et al., Projection of temperature-related mortality due to cardiovascular disease in beijing under different climate change, population, and adaptation scenarios. *Environmental Research*, 2018. 162: p. 152-159.

4. Balbus, J., A. Crimmins, J.L. Gamble, D.R. Easterling, K.E. Kunkel, S. Saha, and M.C. Sarofim, 2016: Ch. 1: Introduction: Climate Change and Human Health. The Impacts of Climate Change on Human Health in the United States: A Scientific Assessment. U.S. Global Change Research Program, Washington, DC, 25–42. <http://dx.doi.org/10.7930/JOVX0DFW>
5. Fall, S., D. Niyogi, A. Gluhovsky, R. A. Pielke, Sr., E. Kalnay, and G. Rochon, 2010: Impacts of land use land cover on temperature trends over the continental United States: Assessment using the North American Regional Reanalysis. *International Journal of Climatology*, 30, 1980-1993, doi:10.1002/joc.1996.
6. Holli, R. et al, 2010, Global. Warming Earths observatory , NASA
7. Greg fischer et al, USCM-Low-Carbon-Future-report-April-2021-WEB.pdf <https://www.c2es.org/site/assets/uploads/2021/04/>
8. John Walsh, et al., Ch. 2: Our Changing Climate. Climate Change Impacts in the United States: The Third National Climate Assessment (U.S. Global Change Research Program, 2014)
9. Anderson GB, Bell ML. Heat waves in the United States: mortality risk during heat waves and effect modification by heat wave characteristics in 43 U.S. communities. *Environ Health Perspect*. 2011;119(2):210-218. doi:10.1289/ehp.1002313
10. Medina-Ramón, M., & Schwartz, J. (2007). Temperature, temperature extremes, and mortality: a study of acclimatisation and effect modification in

50 US cities. *Occupational and environmental medicine*, 64(12), 827–833.

<https://doi.org/10.1136/oem.2007.033175>

11. Moghadamnia MT, Ardalan A, Mesdaghinia A, Keshtkar A, Naddafi K, Yekaninejad MS. Ambient temperature and cardiovascular mortality: a systematic review and meta-analysis. *PeerJ*. 2017;5:e3574. Published 2017 Aug 4. doi:10.7717/peerj.3574
12. Fares A. (2013). Winter Hypertension: Potential mechanisms. *International journal of health sciences*, 7(2), 210–219. <https://doi.org/10.12816/0006044>
13. Baaghideh M, Mayvaneh F. Climate Change and Simulation of Cardiovascular Disease Mortality: A Case Study of Mashhad, Iran. *Iran J Public Health*. 2017;46(3):396-407.
14. Bouchama, A. and J.P. Knochel, Heat Stroke. *New England Journal of Medicine*, 2002. 346(25): p. 1978-1988.
15. Wilker EH, Yeh G, Wellenius GA, Davis RB, Phillips RS, Mittleman MA. Ambient temperature and biomarkers of heart failure: a repeated measures analysis. *Environ Health Perspect*. 2012;120(8):1083-1087. doi:10.1289/ehp.1104380
16. Chen, B., et al., Heat risk of residents in different types of communities from urban heat-exposed areas. *Science of the Total Environment*, 2021. 768.

17. Kovats RS, Kristie LE. Heatwaves and public health in Europe. *Eur J Public Health*. 2006 Dec;16(6):592-9. doi: 10.1093/eurpub/ckl049. Epub 2006 Apr 27. PMID: 16644927.
18. Schifano P, Cappai G, De Sario M, et al. Susceptibility to heat wave-related mortality: a follow-up study of a cohort of elderly in Rome. *Environ Health*. 2009;8:50. Published 2009 Nov 12. doi:10.1186/1476-069X-8-50
19. Kovats RS, Kristie LE. Heatwaves and public health in Europe. *Eur J Public Health*. 2006 Dec;16(6):592-9. doi: 10.1093/eurpub/ckl049. Epub 2006 Apr 27. PMID: 16644927.
20. Hajat S, Armstrong B, Baccini M, Biggeri A, Bisanti L, Russo A, Paldy A, Menne B, Kosatsky T. Impact of high temperatures on mortality: is there an added heat wave effect? *Epidemiology*. 2006 Nov;17(6):632-8. doi: 10.1097/01.ede.0000239688.70829.63. PMID: 17003686.
21. Knowlton K, Rotkin-Ellman M, King G, et al. The 2006 California heat wave: impacts on hospitalizations and emergency department visits. *Environ Health Perspect*. 2009;117(1):61-67. doi:10.1289/ehp.11594
22. Xu Z, Sheffield PE, Hu W, et al. Climate change and children's health--a call for research on what works to protect children. *Int J Environ Res Public Health*. 2012;9(9):3298-3316. Published 2012 Sep 10. doi:10.3390/ijerph9093298

23. Vanos JK. Children's health and vulnerability in outdoor microclimates: A comprehensive review. *Environ Int.* 2015 Mar;76:1-15. doi: 10.1016/j.envint.2014.11.016. Epub 2014 Dec 11. PMID: 25497108.
24. Anderson GB, Bell ML, Peng RD. Methods to calculate the heat index as an exposure metric in environmental health research. *Environ Health Perspect.* 2013;121(10):1111-1119. doi:10.1289/ehp.1206273
25. Bonan, G.B. Effects of Land Use on the Climate of the United States. *Climatic Change* 37, 449–486 (1997). <https://doi.org/10.1023/A:1005305708775>
26. Habeeb, D., Vargo, J. & Stone, B. Rising heat wave trends in large US cities. *Nat Hazards*76, 1651–1665 (2015).
27. Ebi KL, Schmier JK. A stitch in time: improving public health early warning systems for extreme weather events. *Epidemiol Rev.* 2005;27:115-21. doi: 10.1093/epirev/mxi006. PMID: 15958432.
28. Frumkin H, Hess J, Lubber G, Malilay J, McGeehin M. Climate change: the public health response. *Am J Public Health.* 2008 Mar;98(3):435-45. doi: 10.2105/AJPH.2007.119362. Epub 2008 Jan 30. PMID: 18235058; PMCID: PMC2253589.
29. Balbus, J., A. Crimmins, J.L. Gamble, D.R. Easterling, K.E. Kunkel, S. Saha, and M.C. Sarofim, 2016: Ch. 1: Introduction: Climate Change and Human Health. *The Impacts of Climate Change on Human Health in the United States: A Scientific Assessment.* U.S. Global Change Research Program, Washington, DC, 25–42.

30. Fisher, J.A., et al., Case-crossover analysis of short-term particulate matter exposures and stroke in the health professionals follow-up study. *Environment International*, 2019. 124: p. 153-160.
31. Bhaskaran, K., et al., Heat and risk of myocardial infarction: hourly level case-crossover analysis of MINAP database. *BMJ : British Medical Journal*, 2012. 345: p. e8050.
32. Cassandra R et al 2020, Case-Crossover Analysis of Indoor Heat Exposure on Mortality and Hospitalizations among the Elderly in Houston, Texas
33. <https://en.wikipedia.org/wiki/Atlanta#Climate>
34. <https://datausa.io/profile/geo/atlanta-ga/>
35. McNaughton CD, Self WH, Zhu Y, Janke AT, Storrow AB, Levy P. Incidence of Hypertension-Related Emergency Department Visits in the United States, 2006 to 2012. *Am J Cardiol*. 2015;116(11):1717-1723.
doi:10.1016/j.amjcard.2015.09.007
36. Li, Mengxuan et al. "Impact of Extremely Hot Days on Emergency Department Visits for Cardiovascular Disease among Older Adults in New York State." *International journal of environmental research and public health* vol. 16,12 2119. 14 Jun. 2019, doi:10.3390/ijerph16122119
37. Bai L, et al Hospitalizations from Hypertensive Diseases, Diabetes, and Arrhythmia in Relation to Low and High Temperatures

38. Basu R, Pearson D, Malig B, Broadwin R, Green R. The effect of high ambient temperature on emergency room visits. *Epidemiology*. 2012 Nov;23(6):813-20. doi: 10.1097/EDE.0b013e31826b7f97. PMID: 23007039.
39. McNaughton, Candace D et al. "Incidence of Hypertension-Related Emergency Department Visits in the United States, 2006 to 2012
40. Phung D, Thai PK, Guo Y, Morawska L, Rutherford S, Chu C. Ambient temperature and risk of cardiovascular hospitalization: An updated systematic review and meta-analysis. *Sci Total Environ*. 2016 Apr 15;550:1084-1102. doi: 10.1016/j.scitotenv.2016.01.154. Epub 2016 Feb 9. PMID: 26871555.
41. Kim, S.E., Lee, H., Kim, J. et al. Temperature as a risk factor of emergency department visits for acute kidney injury: a case-crossover study in Seoul, South Korea. *Environ Health* **18**, 55 (2019). <https://doi.org/10.1186/s12940-019-0491-5>
42. Lin S, Luo M, Walker RJ, Liu X, Hwang SA, Chinery R. Extreme high temperatures and hospital admissions for respiratory and cardiovascular diseases. *Epidemiology*. 2009 Sep;20(5):738-46. doi: 10.1097/EDE.0b013e3181ad5522. PMID: 19593155.
43. Winquist, A., et al., Warm season temperatures and emergency department visits in Atlanta, Georgia
44. Gronlund CJ, Zanobetti A, Schwartz JD, Wellenius GA, O'Neill MS. Heat, heat waves, and hospital admissions among the elderly in the United States, 1992-2006. *Environ Health Perspect*. 2014 Nov;122(11):1187-92. doi: 10.1289/ehp.1206132. Epub 2014 Jun 6. PMID: 24905551; PMCID: PMC4216145.
45. Kim S, Kim SY, Oh J, Chae Y, Park J, Kim D, Kim YM. Effects of the 2018 heat wave on health in the elderly: implications for adaptation strategies to climate change. *Environ Anal Health Toxicol*. 2020 Dec;35(4):e2020024-0.

- doi: 10.5620/eaht.2020024. Epub 2020 Dec 21. PMID: 33434424; PMCID: PMC7829408.
46. Chen T, Sarnat SE, Grundstein AJ, Winqvist A, Chang HH. Time-series Analysis of Heat Waves and Emergency Department Visits in Atlanta, 1993 to 2012. *Environ Health Perspect*. 2017 May 31;125(5):057009. doi: 10.1289/EHP44. PMID: 28599264; PMCID: PMC5730512.
47. Zheng S, Wang M, Shang K, He S, Yin L, Li T, Wang S. [A case-crossover analysis of heat wave and hospital emergency department visits for cardiovascular diseases in 3 hospitals in Beijing]. *Wei Sheng Yan Jiu*. 2016 Mar;45(2):246-51. Chinese. PMID: 27301223.
48. Huang J, Wang J, Yu W. The lag effects and vulnerabilities of temperature effects on cardiovascular disease mortality in a subtropical climate zone in China. *Int J Environ Res Public Health*. 2014 Apr 11;11(4):3982-94. doi: 10.3390/ijerph110403982. PMID: 24733034; PMCID: PMC4025023.
49. NOAA National Centers for Environmental Information, State of the Climate: Global Climate Report for Annual 2020, online January 2021, retrieved on March 15, 2021 from <https://www.ncdc.noaa.gov/sotc/global/202013>.
50. Lan L, Cui G, Yang C, Wang J, Sui C, Xu G, Zhou D, Cheng Y, Guo Y, Li T. Increased mortality during the 2010 heat wave in Harbin, China. *Ecohealth*. 2012 Sep;9(3):310-4. doi: 10.1007/s10393-012-0790-6. Epub 2012 Aug 15. PMID: 22893101.
51. Can G, Şahin Ü, Sayılı U, Dubé M, Kara B, Acar HC, İnan B, Aksu Sayman Ö, Lebel G, Bustinza R, Küçükali H, Güven U, Gosselin P. Excess Mortality

- in Istanbul during Extreme Heat Waves between 2013 and 2017. *Int J Environ Res Public Health*. 2019 Nov 7;16(22):4348. doi: 10.3390/ijerph16224348. PMID:
52. Aboubakri O, Khanjani N, Jahani Y, Bakhtiari B. Attributable risk of mortality associated with heat and heat waves: A time-series study in Kerman, Iran during 2005-2017. *J Therm Biol*. 2019 May;82:76-82. doi: 10.1016/j.jtherbio.2019.03.013. Epub 2019 Mar 22. PMID: 31128662.
53. Åström DO, Forsberg B, Rocklöv J. Heat wave impact on morbidity and mortality in the elderly population: a review of recent studies. *Maturitas*. 2011 Jun;69(2):99-105. doi: 10.1016/j.maturitas.2011.03.008. Epub 2011 Apr 8. PMID: 21477954.
54. Sohail H, Kollanus V, Tiittanen P, Schneider A, Lanki T. Heat, Heatwaves and Cardiorespiratory Hospital Admissions in Helsinki, Finland. *Int J Environ Res Public Health*. 2020 Oct 28;17(21):7892. doi: 10.3390/ijerph17217892. PMID: 33126485; PMCID: PMC7663418.
55. Cheng J, Xu Z, Bambrick H, Prescott V, Wang N, Zhang Y, Su H, Tong S, Hu W. Cardiorespiratory effects of heatwaves: A systematic review and meta-analysis of global epidemiological evidence. *Environ Res*. 2019 Oct;177:108610. doi: 10.1016/j.envres.2019.108610. Epub 2019 Jul 26. PMID: 31376629.

56. Dong W, Zeng Q, Ma Y, Li G, Pan X. Impact of Heat Wave Definitions on the Added Effect of Heat Waves on Cardiovascular Mortality in Beijing, China. *Int J Environ Res Public Health*. 2016 Sep 21;13(9):933. doi: 10.3390/ijerph13090933. PMID: 27657103; PMCID: PMC5036765.
57. Kenney WL, Craighead DH, Alexander LM. Heat waves, aging, and human cardiovascular health. *Med Sci Sports Exerc*. 2014 Oct;46(10):1891-9. doi: 10.1249/MSS.0000000000000325. PMID: 24598696; PMCID: PMC4155032.
58. Fuhrmann CM, Sugg MM, Konrad CE 2nd, Waller A. Impact of Extreme Heat Events on Emergency Department Visits in North Carolina (2007-2011). *J Community Health*. 2016 Feb;41(1):146-56. doi: 10.1007/s10900-015-0080-7. PMID: 26289379.
59. Dadbakhsh M, Khanjani N, Bahrapour A. The relation between mortality from cardiovascular diseases and temperature in Shiraz, Iran, 2006-2012. *ARYA Atheroscler*. 2018 Jul;14(4):149-156. doi: 10.22122/arya.v14i4.1341. PMID: 30627190; PMCID: PMC6312568.
60. Hatvani-Kovacs G, Belusko M, Skinner N, Pockett J, Boland J. Drivers and barriers to heat stress resilience. *Sci Total Environ*. 2016 Nov 15;571:603-14. doi: 10.1016/j.scitotenv.2016.07.028. Epub 2016 Jul 16. PMID: 27432732.

61. Anderson GB, Oleson KW, Jones B, Peng RD. Projected trends in high-mortality heatwaves under different scenarios of climate, population, and adaptation in 82 US communities. *Clim Change*. 2018 Feb;146(3-4):455-470. doi: 10.1007/s10584-016-1779-x. Epub 2016 Aug 30. PMID: 29628541; PMCID: PMC5881935.
62. Arbuthnott K, Hajat S, Heaviside C, Vardoulakis S. Changes in population susceptibility to heat and cold over time: assessing adaptation to climate change. *Environ Health*. 2016 Mar 8;15 Suppl 1(Suppl 1):33. doi: 10.1186/s12940-016-0102-7. PMID: 26961541; PMCID: PMC4895245.
63. Zhao H, Jivraj S, Moody A. 'My blood pressure is low today, do you have the heating on?' The association between indoor temperature and blood pressure. *J Hypertens*. 2019 Mar;37(3):504-512. doi: 10.1097/HJH.0000000000001924. PMID: 30134311.
64. Saeki K, Obayashi K, Iwamoto J, Tone N, Okamoto N, Tomioka K, Kurumatani N. The relationship between indoor, outdoor and ambient temperatures and morning BP surges from inter-seasonally repeated measurements. *J Hum Hypertens*. 2014 Aug;28(8):482-8. doi: 10.1038/jhh.2014.4.
65. Epub 2014 Feb 20. PMID: 2455 Waugh DW, He Z, Zaitchik B, Peng RD, Diette GB, Hansel NN, Matsui EC, Breyse PN, Breyse DH, Koehler K,

- Williams D, McCormack MC. Indoor heat exposure in Baltimore: does outdoor temperature matter? *Int J Biometeorol*. 2021 Apr;65(4):479-488. doi: 10.1007/s00484-020-02036-2. Epub 2020 Oct 21. PMID: 33089367. 3634.
66. Kim YM, Kim S, Cheong HK, Ahn B, Choi K. Effects of heat wave on body temperature and blood pressure in the poor and elderly. *Environ Health Toxicol*. 2012;27:e2012013. doi:10.5620/eh.2012.27.e2012013
67. Kenny GP, Yardley J, Brown C, Sigal RJ, Jay O. Heat stress in older individuals and patients with common chronic diseases. *CMAJ*. 2010;182(10):1053-1060. doi:10.1503/cmaj.081050
68. Chen T, Sarnat SE, Grundstein AJ, Winkquist A, Chang HH. Time-series Analysis of Heat Waves and Emergency Department Visits in Atlanta, 1993 to 2012. *Environ Health Perspect*. 2017 May 31;125(5):057009. doi: 10.1289/EHP44. PMID: 28599264; PMCID: PMC5730512.
69. Wang YC, Lin YK. Association between temperature and emergency room visits for cardiorespiratory diseases, metabolic syndrome-related diseases, and accidents in metropolitan Taipei. *PLoS One*. 2014 Jun 16;9(6):e99599. doi: 10.1371/journal.pone.0099599. PMID: 24932702; PMCID: PMC4059706.
70. Hu J, He G, Luo J, Xu Y, Xu X, Song X, Chen S, Ji G, Chen Z, Jiang Q, Liu T, Hu J, Xiao J, Zeng W, Guo L, Lin L, Lin P, Ma W. Temperature-adjusted hypertension prevalence and control rate: a series of cross-sectional studies in

- Guangdong Province, China. *J Hypertens*. 2021 May 1;39(5):911-918. doi: 10.1097/HJH.0000000000002738. PMID: 33273194.
71. Janke AT, McNaughton CD, Brody AM, Welch RD, Levy PD. Trends in the Incidence of Hypertensive Emergencies in US Emergency Departments From 2006 to 2013. *J Am Heart Assoc*. 2016 Dec 5;5(12):e004511. doi: 10.1161/JAHA.116.004511. PMID: 27919932; PMCID: PMC5210448.
72. Gasparini, A., Y. Guo, M. Hashizume, E. Lavigne, A. Zanobetti, J. Schwartz, A. Tobias, S. Tong, J. Rocklöv, B. Forsberg, M. Leone, M. DeSario, M.L. Bell, Y.-L. Leon Guo, C.-F. Wu, H. Kan, S.-M. Yi, M. de Sousa Zanotti Stagliorio Coelho, P. Hilario Nascimento Saldiva, Y. Honda, H. Kim, and B. Armstrong. 2015. Mortality risk attributable to high and low ambient temperature: A multicountry observational study. *The Lancet* 386(9991):369–375.
73. Medina-Ramón, M., and J. Schwartz. 2007. Temperature, temperature extremes, and mortality: A study of acclimatization and effect modification in 50 U.S. cities. *Occup. Environ. Med.* 64(12):827–833.
74. Curriero FC, Heiner KS, Samet JM, Zeger SL, Strug L, Patz JA. Temperature and mortality in 11 cities of the eastern United States. *Am J Epidemiol*. 2002 Jan 1;155(1):80-7. doi: 10.1093/aje/155.1.80. PMID: 11772788.
75. Chen T, Sarnat SE, Grundstein AJ, Winquist A, Chang HH. Time-series Analysis of Heat Waves and Emergency Department Visits in Atlanta, 1993 to 2012. *Environ Health Perspect*. 2017 May 31;125(5):057009. doi: 10.1289/EHP44. PMID: 28599264; PMCID: PMC5730512.
76. Layton JB, Li W, Yuan J, Gilman JP, Horton DB, Setoguchi S. Heatwaves, medications, and heat-related hospitalization in older Medicare beneficiaries with chronic conditions. *PLoS One*. 2020 Dec 10;15(12):e0243665. doi: 10.1371/journal.pone.0243665. PMID: 33301532; PMCID: PMC7728169.
77. Ponjoan A, Blanch J, Alves-Cabratos L, Martí-Lluch R, Comas-Cufí M, Parramon D, Del Mar Garcia-Gil M, Ramos R, Petersen I. Effects of extreme

temperatures on cardiovascular emergency hospitalizations in a Mediterranean region: a self-controlled case series study. *Environ Health*. 2017 Apr 4;16(1):32. doi: 10.1186/s12940-017-0238-0. PMID: 28376798; PMCID: PMC5379535.

Abbreviation

1. CDW- Clinical Data Warehouse
2. C-Celsius
3. CI– Confidence intervals
4. ED- Emergency Department
5. F- Fahrenheit
6. HX- History
7. HTN- hypertension
8. ICD- International Classification of Disease
9. OR- Odds Ratio
10. Q- Quartiles