

National and regional seasonal dynamics of all-cause and cause-specific mortality in the USA from 1980 to 2016

Robbie M Parks^{1,2}, James E Bennett^{1,2,3}, Kyle J Foreman^{1,2,4}, Ralf Toumi⁵, Majid Ezzati^{1,2,3*}

¹ MRC-PHE Centre for Environment and Health, Imperial College London, London, United Kingdom, W2 1PG

² Department of Epidemiology and Biostatistics, School of Public Health, Imperial College London, London, United Kingdom, W2 1PG

³ WHO Collaborating Centre on NCD Surveillance and Epidemiology, Imperial College London, London, United Kingdom, W2 1PG

⁴ Institute for Health Metrics and Evaluation, University of Washington, Seattle, USA, WA 98121

⁵ Space and Atmospheric Physics, Imperial College London, London, United Kingdom, W7 2AZ

Address correspondence to Majid Ezzati (majid.ezzati@imperial.ac.uk)

Abstract

In temperate climates, winter deaths exceed summer ones. However, there is limited information on how the timing and the relative magnitudes of minimum and maximum mortality vary by local climate and how these features have changed over time, in relation to age group, sex and medical cause of death. We used geo-coded mortality data and wavelet analytical techniques to analyse the seasonality of all-cause and cause-specific mortality by age group and sex from 1980 to 2016 in the USA, nationally and in nine subnational climatic regions. Death rates in men and women ≥ 45 years exhibited seasonality with peak in December to February and minimum in June to August, driven by seasonality of cardiorespiratory diseases and injuries. In these ages, percent difference in death rates between peak and minimum months did not vary across climate regions, and was largely unchanged from 1980 to 2016. Under five years of age, seasonality of all-cause mortality largely disappeared after the 1990s. In adolescents and young adults, especially in males, death rates peaked in June/July and were lowest in December/January, driven by seasonality of injury deaths.

Introduction

It is well-established that death rates vary throughout the year, and in temperate climates there tend to be more deaths in winter than in summer (Campbell, 2017; Fowler et al., 2015; Healy, 2003; McKee, 1989). It has therefore been hypothesized that a warmer world may lower winter mortality in temperate climates (Langford & Bentham, 1995; Martens, 1998). In a large country like the USA, which possesses distinct climate regions, the seasonality of mortality may vary geographically, due to geographical variations in mortality, localized weather patterns, and regional differences in adaptation measures such as heating, air conditioning and healthcare (Davis, Knappenberger, Michaels, & Novicoff, 2004; Ferreira

Braga, Zanobetti, & Schwartz, 2001; Kalkstein, 2013; Medina-Ramón & Schwartz, 2007).
The presence and extent of seasonal variation in mortality may also itself change over time
(Bobb, Peng, Bell, & Dominici, 2014; Carson, Hajat, Armstrong, & Wilkinson, 2006;
Seretakis, 1997; Sheridan, Kalkstein, & Kalkstein, 2009).

A thorough understanding of the long-term dynamics of seasonality of mortality, and its
geographical and demographic patterns, is needed to identify at-risk groups, plan responses at
the present time as well as under changing climate conditions. Although mortality seasonality
is well-established, there is limited information on how seasonality, including the timing of
minimum and maximum mortality, varies by local climate and how these features have
changed over time, especially in relation to age group, sex and medical cause of death (Rau,
2004; Rau, Bohk-Ewald, Muszyńska, & Vaupel, 2018).

In this paper, we comprehensively characterize the spatial and temporal patterns of all-cause
and cause-specific mortality seasonality in the USA by sex and age group, through the
application of wavelet analytical techniques, to over three decades of national mortality data.
Wavelets have been used to study the dynamics of weather phenomena (Moy, Seltzer,
Rodbell, & Anderson, 2002) and infectious diseases (Grenfell, Bjørnstad, & Kappey, 2001).
We also used centre of gravity analysis and circular statistics methods to understand the
timing of mortality minimum and maximum. In addition, we identify how the percentage
difference between death rates in maximum and minimum mortality months has changed
over time.

Results

Table 1 presents number of deaths by cause of death and sex. Deaths from cardiorespiratory diseases make up nearly half of all deaths (48.1%), with most deaths from cardiovascular diseases. Next highest during the study period were deaths from cancers (23.2%), followed by injuries (6.8%), with two thirds of those being from unintentional injuries.

All-cause mortality in males had a 12-month seasonality in all age groups, except ages 35-44 years, for whom there was periodicity at 6 months (Figure 2). In females, there was 12-month seasonality in all groups except 5-14 and 25-35 years (p -values=0.20 and 0.24, respectively). While seasonality persisted throughout the entire analysis period in older ages, it largely disappeared after late 1990s in children aged 0-4 years in both sexes and in women aged 15-24 years.

Deaths from all causes of death were seasonal in older adults (above 65 or 75 years depending on cause, p -values<0.06) (Figure 2 and Supplementary Figure 1), except for intentional injuries and substance use disorders. Deaths from cardiorespiratory diseases, and within it respiratory infections, exhibited seasonality throughout the life-course (p -values<0.03) except for males aged 5-24 years and females aged 15-24 years (p -values>0.11). In addition to older ages, injury deaths were seasonal from childhood through 44 years in women and through 64 years in men (p -values<0.08). Unintentional injuries drove the seasonality of injury deaths for females, whereas both unintentional and intentional injuries were seasonal in males in most ages, with the exception of below 15 years and above 85 years when intentional injuries were not seasonal (Supplementary Figure 1). Consistent seasonality in cancer deaths only appeared after 55 years of age (p -values<0.04). No consistent seasonality was evident in maternal conditions or substance use disorders (Supplementary Figure 1).

Centre of gravity analysis showed that death rates in men aged ≥ 45 years and women aged ≥ 35 years peaked in December, January or February and were lowest in June to August, for all-cause mortality as well as for all non-injury and non-maternal causes of death (Figure 3 and Supplementary Figure 2). Deaths from cardiorespiratory diseases, including cardiovascular diseases, chronic respiratory diseases and respiratory infections, were also consistently highest in January and February and lowest in July and August across all ages, except for chronic respiratory diseases in ages 5-24 years where there are few deaths from this cause leading to unstable estimates (p-values for seasonality from wavelet analysis ranged from 0.35 to 0.49 for these ages). A similar temporal pattern was seen for all-cause and non-injury mortality in children younger than five years of age, whose all-cause death rate was highest in February and lowest in August. In contrast, among males aged 5-34 years, all-cause mortality peaked in June or July, as did deaths from injuries, which generally had a summer peak in males and females below 45 years of age.

From 1980 to 2016, the proportional (percent) difference in all-cause death rates between peak and minimum months declined little for people older than 45 years of age (by less than eight percentage points with p-values for declining trend > 0.1) (Figure 4). In contrast, the difference between peak (summer) and minimum (winter) death rates declined in younger ages, by over 25 percentage points in males aged 5-14 years and 15-24 years (p-values < 0.01), largely driven in the declining difference between summer and winter injury deaths. Under five years of age, percent seasonal difference in all-cause death rates declined by 13 percentage points (p-value < 0.01) for boys but only 5 percentage points (p-value = 0.12) for girls. These declines in seasonality of child deaths were a net effect of declining winter-summer difference in cardiorespiratory diseases deaths and increasing summer-winter

115 difference in injury deaths, itself driven by increasing difference in non-intentional injuries
116 (Supplementary Figure 3). Within the cardiorespiratory diseases cluster in under-five
117 children, percent difference declined for cardiorespiratory diseases, cardiovascular diseases,
118 and chronic respiratory diseases while increasing for respiratory infections.

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120 The subnational centre of gravity analysis showed that all-cause mortality peaks and minima
121 in different climate regions are consistent with the national ones (Figure 5), indicating the
122 seasonality is largely independent of geography. The relative homogeneity of the timing of
123 maximum and minimum mortality contrasts with the large variation in seasonal temperatures
124 among climate regions. For example, in men and women aged 65-74 years, all-cause
125 mortality peaked in February in the Northeast and Southeast, even though the average
126 temperatures for those regions were different by over 13 degrees Celsius (9.3 in the Southeast
127 compared with -3.8 in the Northeast). Furthermore, above 45 years of age, there was little
128 inter-region variation in the percent seasonal difference in all-cause mortality, despite the
129 large variation in temperature difference between the peak and minimum months (Figure 6).

131 **Strengths and limitations**

132 The strengths of our study are its innovative methods of characterizing seasonality of
133 mortality dynamically over space and time, by age group and cause of death; using wavelet
134 and centre of gravity analyses; using ERA-Interim data output to compare the association
135 between seasonality of death rates and regional temperature. A limitation of our study is that
136 we did not investigate seasonality of mortality by socioeconomic characteristics which may
137 help with understanding its determinants and planning responses.

139 **Discussion**

We used wavelet and centre of gravity analyses, which allowed systematically identifying and characterizing seasonality of total and cause-specific mortality in the USA, and examining how seasonality has changed over time. We identified distinct seasonal patterns in relation to age and sex, including higher all-cause summer mortality in young men (Feinstein, 2002; Rau et al., 2018). Importantly, we also showed that all-cause and cause-specific mortality seasonality is largely similar in terms of both timing and magnitude across diverse climatic regions with substantially different summer and winter temperatures. Insights of this kind would not have been possible analysing data averaged over time or nationally, or fixed to pre-specified frequencies.

Prior studies have noted seasonality of mortality for all-cause mortality and for specific causes of death in the USA (Feinstein, 2002; Kalkstein, 2013; Rau, 2004; Rau et al., 2018; Rosenwaike, 1966; Seretakis, 1997). Few of these studies have done consistent national and subnational analyses, and none has done so over time, for a comprehensive set of age groups and causes of death, and in relation to regional temperature differences. Our results on strong seasonality of cardiorespiratory diseases deaths and weak seasonality of cancer deaths, restricted to older ages, are broadly consistent with these studies (Feinstein, 2002; Rau et al., 2018; Rosenwaike, 1966; Seretakis, 1997), which had limited analysis on how seasonality changes over time and geography (Feinstein, 2002; Rau et al., 2018; Rosenwaike, 1966). Similarly, our results on seasonality of injury deaths are supported by a few prior studies (Feinstein, 2002; Rau et al., 2018; Rosenwaike, 1966), but our subnational analysis over three decades revealed variations in when injury deaths peaked and in how seasonal differences in these deaths have changed over time in relation to age group which had not been reported before.

A study of 36 cities in the USA, aggregated across age groups and over time, also found that excess mortality was not associated with seasonal temperature range (Kinney et al., 2015). In contrast, a European study found that the difference between winter and summer mortality was lower in the colder Nordic countries than in warmer southern European nations (Healy, 2003; McKee, 1989)(the study's measure of temperature was mean annual temperature which differed from the temperature difference between maximum and minimum mortality used in our analysis although the two measures are correlated). The absence of variation in the magnitude of mortality seasonality indicates that different regions in the USA are similarly adapted to temperature seasonality, whereas Nordic countries may have better environmental (e.g., housing insulation and heating) and health system measures to counter the effects of cold winters than those in southern Europe. If the observed absence of association between the magnitude of mortality seasonality and seasonal temperature difference across the climate regions also persists over time, the changes in temperature as a result of global climate change are unlikely to affect the winter-summer mortality difference.

The cause-specific analysis showed that the substantial decline in seasonal mortality differences in adolescents and young adults was related to the diminishing seasonality of (unintentional) injuries, especially from road traffic crashes, which are more likely to occur in the summer months (National Highway Traffic Safety Administration, 2005) and are more common in men. The weakening of seasonality in boys under five years of age was related to two phenomena: first, the seasonality of death from cardiorespiratory diseases declined, and second, the proportion of deaths from perinatal conditions, which exhibit limited seasonality (Supplementary Figure 1), increased (MacDorman & Gregory, 2015).

In contrast to young and middle ages, mortality in older ages, where death rates are highest, maintained persistent seasonality over a period of three decades (we note that although the percent seasonal difference in mortality has remained largely unchanged in these ages, the absolute difference in death rates between the peak and minimum months has declined because total mortality has a declining long-term trend). This finding demonstrates the need for environmental and health service interventions targeted towards this group irrespective of geography and local climate. Examples of such interventions include enhancing the availability of both environmental and medical protective factors, such as better insulation of homes, winter heating provision and flu vaccinations, for the vulnerable older population (Public Health England, 2017). Social interventions, including regular visits to the isolated elderly during peak mortality periods to ensure that they are optimally prepared for adverse conditions, and responsive and high-quality emergency care, are also important to protect this vulnerable group (Healy, 2003; Lerchl, 1998; Public Health England, 2017). Emergent new technologies, such as always-connected hands-free communications devices with the outside world, in-house cameras, and personal sensors also provide an opportunity to enhance care for the older, more vulnerable groups in the population, especially in winter when the elderly have fewer social interactions (Morris, 2013). Such interventions are important today, and will remain so as the population ages and climate change increases the within- and between-season weather variability.

Materials and methods

Data

We used data on all 85,854,176 deaths in the USA from 1980 to 2016 from the National Center for Health Statistics (NCHS). Age, sex, state of residence, month of death, and underlying cause of death were available for each record. The underlying cause of death was

214 coded according to the international classification of diseases (ICD) system (9th revision of
215 ICD from 1980 to 1998 and 10th revision of ICD thereafter). Yearly population counts were
216 available from NCHS for 1990 to 2016 and from the US Census Bureau prior to 1990
217 (Ingram et al., 2003). We calculated monthly population counts through linear interpolation,
218 assigning each yearly count to July.

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220 We also subdivided the national data geographically into nine climate regions used by the
221 National Oceanic and Atmospheric Administration (Figure 1 and Table 2) (Karl & Koss,
222 1984). On average, the Southeast and South are the hottest climate regions with average
223 annual temperatures of 18.4°C and 18°C respectively; the South also possesses the highest
224 average maximum monthly temperature (27.9°C in July). The lowest variation in temperature
225 throughout the year is that of the Southeast (an average range of 17.5°C). The three coldest
226 climate regions are West North Central, East North Central and the Northwest (7.8°C, 8.0°C,
227 8.1°C respectively). Mirroring the characteristics of the hottest climate regions, the largest
228 variation in temperature throughout the year is that of the coldest region, West North Central
229 (an average range of 30.5°C), which also has the lowest average minimum monthly
230 temperature (-6.5°C in January). The other climate regions, Northeast, Southwest, and
231 Central, possess similar average temperatures (11 to 13°C) and variation within the year of
232 (23 to 26°C), with the Northeast being the most populous region in the United States (with
233 19.8% total population in 2016).

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235 Data were divided by sex and age in the following 10 age groups: 0-4, 5-14, 25- 34, 35-44,
236 45-54, 55-64, 65-74, 75-84, 85+ years. We calculated monthly death rates for each age and
237 sex group, both nationally and for sub-national climate regions. Death rate calculations

accounted for varying length of months, by multiplying each month's death count by a factor that would make it equivalent to a 31-day month.

For analysis of seasonality by cause of death, we mapped each ICD-9 and ICD-10 codes to four main disease categories (Table 1) and to a number of subcategories which are presented in the Supplementary Note. Cardiorespiratory diseases and cancers accounted for 56.4% and 21.2% of all deaths in the USA, respectively, in 1980, and 40.3% and 22.4%, respectively, in 2016. Deaths from cardiorespiratory diseases have been associated with cold and warm temperatures (Basu, 2009; Basu & Samet, 2002; Bennett, Blangiardo, Fecht, Elliott, & Ezzati, 2014; Braga, Zanobetti, & Schwartz, 2002; Gasparrini et al., 2015). Injuries, which accounted for 8% of all deaths in the USA in 1980 and 7.3% in 2016, may have seasonality that is distinct from so-called natural causes. We did not further divide other causes because the number of deaths could become too small to allow stable estimates when divided by age group, sex and climate region.

We obtained data on temperature from ERA-Interim, which combines predictions from a physical model with ground-based and satellite measurements (Dee et al., 2011). We used gridded four-times-daily estimates at a resolution of 80km to generate monthly population-weighted temperature by climate region throughout the analysis period.

Statistical methods

We used wavelet analysis to investigate seasonality for each age-sex group. Wavelet analysis uncovers the presence, and frequency, of repeated maxima and minima in each age-sex-specific death rate time series (Hubbard, 1998; Torrence & Compo, 1998). In brief, a Morlet wavelet, described in detail elsewhere (Cazelles et al., 2008), is equivalent to using a moving

window on the death rate time series and analysing periodicity in each window using a short-
 form Fourier transform, hence generating a dynamic spectral analysis, which allows
 measuring dynamic seasonal patterns, in which the periodicity of death rates may disappear,
 emerge, or change over time. In addition to coefficients that measure the frequency of
 periodicity, wavelet analysis estimates the probability of whether the data are different from
 the null situation of random fluctuations that can be represented with white (an independent
 random process) or red (autoregressive of order 1 process) noise. For each age-sex group, we
 calculated the p-values of the presence of 12-month seasonality for the comparison of
 wavelet power spectra of the entire study period (1980-2016) with 100 simulations against a
 white noise spectrum, which represents random fluctuations. We used the R package
 WaveletComp (version 1.0) for the wavelet analysis. Before analysis, we de-trended death
 rates using a polynomial regression, and rescaled each death rate time series so as to range
 between 1 and -1.

To identify the months of maximum and minimum death rates, we calculated the centre of
 gravity and the negative centre of gravity of monthly death rates. Centre of gravity was
 calculated as a weighted average of months of deaths, with each month weighted by its death
 rate; negative centre of gravity was also calculated as a weighted average of months of
 deaths, but with each month was weighted by the difference between its death rate and the
 year's maximum death rate. In taking the weighted average, we allowed December (month
 12) to neighbour January (month 1), representing each month by an angle subtended from 12
 equally-spaced points around a unit circle. Using a technique called circular statistics, a mean
 ($\bar{\theta}$) of the angles ($\theta_1, \theta_2, \theta_3 \dots, \theta_n$) representing the deaths (with n the total number of deaths
 in an age-sex group for a particular cause of death) is found using the relation below:

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$$\bar{\theta} = \arg \left\{ \sum_{j=1}^n \exp(i\theta_j) \right\},$$

where *arg* denotes the complex number argument and θ_j denotes the month of death in angular form for a particular death j . The outcome of this calculation is then converted back into a month value (Fisher, 1995). Along with each circular mean, a 95% confidence interval (CI) was calculated by using 1000 bootstrap samples. The R package CircStats (version 0.2.4) was used for this analysis.

For each age-sex group and cause of death, and for each year, we calculated the percent difference in death rates between the maximum and minimum mortality months. We fitted a linear regression to the time series of seasonal differences from 1980 to 2016, and used the fitted trend line to estimate how much the percentage difference in death rates between the maximum and minimum mortality months had changed from 1980 to 2016. We weighted seasonal difference by the inverse of the square of its standard error, which was calculated using a Poisson model to take population size of each age-sex group through time into account. This method gives us a p-value for the change in seasonal difference per year, which we used to calculate the seasonal difference at the start (1980) and end (2016) of the period of study. Our method of analysing seasonal differences avoids assuming that any specific month or group of months represent highest and lowest number of deaths for a particular cause of death, which is the approach taken by the traditional measure of Excess Winter Deaths. It also allows the maximum and minimum mortality months to vary by age group, sex and cause of death.

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313 **Author contributions**

314 All authors contributed to study concept, analytical approach, and interpretation of results.

315 RP, KF and ME collated and organised mortality files. RP performed the analysis, with input

316 from JB. RP and ME wrote the first draft of the paper; other authors contributed to revising

317 and finalising the paper.

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319 **Competing financial interests**

320 The authors declare no competing financial interests.

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