JAMA | Special Communication

Life Expectancy and Mortality Rates in the United States, 1959-2017

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IMPORTANCE US life expectancy has not kept pace with that of other wealthy countries and is now decreasing.

OBJECTIVE To examine vital statistics and review the history of changes in US life expectancy and increasing mortality rates; and to identify potential contributing factors, drawing insights from current literature and an analysis of state-level trends.

EVIDENCE Life expectancy data for 1959-2016 and cause-specific mortality rates for 1999-2017 were obtained from the US Mortality Database and CDC WONDER, respectively. The analysis focused on midlife deaths (ages 25-64 years), stratified by sex, race/ethnicity, socioeconomic status, and geography (including the 50 states). Published research from January 1990 through August 2019 that examined relevant mortality trends and potential contributory factors was examined.

FINDINGS Between 1959 and 2016, US life expectancy increased from 69.9 years to 78.9 years but declined for 3 consecutive years after 2014. The recent decrease in US life expectancy culminated a period of increasing cause-specific mortality among adults aged 25 to 64 years that began in the 1990s, ultimately producing an increase in all-cause mortality that began in 2010. During 2010-2017, midlife all-cause mortality rates increased from 328.5 deaths/100 000 to 348.2 deaths/100 000. By 2014, midlife mortality was increasing across all racial groups, caused by drug overdoses, alcohol abuse, suicides, and a diverse list of organ system diseases. The largest relative increases in midlife mortality rates occurred in New England (New Hampshire, 23.3%; Maine, 20.7%; Vermont, 19.9%) and the Ohio Valley (West Virginia, 23.0%; Ohio, 21.6%; Indiana, 14.8%; Kentucky, 14.7%). The increase in midlife mortality during 2010-2017 was associated with an estimated 33 307 excess US deaths, 32.8% of which occurred in 4 Ohio Valley states.

CONCLUSIONS AND RELEVANCE US life expectancy increased for most of the past 60 years, but the rate of increase slowed over time and life expectancy decreased after 2014. A major contributor has been an increase in mortality from specific causes (eg, drug overdoses, suicides, organ system diseases) among young and middle-aged adults of all racial groups, with an onset as early as the 1990s and with the largest relative increases occurring in the Ohio Valley and New England. The implications for public health and the economy are substantial, making it vital to understand the underlying causes.

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Supplemental content

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ife expectancy at birth, a common measure of a population's health, has decreased in the United States for 3 consecutive years. This has attracted recent public attention, but the core problem is not new—it has been building since the 1980s. Although life expectancy in developed countries has increased for much of the past century, US life expectancy began to lose pace with other countries in the 1980s. And, by 1998, had declined to a level below the average life expectancy among Organisation for Economic Cooperation and Development countries. While life expectancy in these countries has continued to increase, US life expectancy stopped increasing in 2010 and has been decreasing since 2014. Despite excessive spending on health care, vastly exceeding that of other countries, the United States has a long-standing health disadvantage relative to other high-income countries that extends

beyond life expectancy to include higher rates of disease and cause-specific mortality rates. $^{6.7,10,13}$

This Special Communication has 2 aims: to examine vital statistics and review the history of changes in US life expectancy and increasing mortality rates; and to identify potential contributing factors, drawing insights from current literature and from a new analysis of state-level trends.

Methods

Data Analysis

Measures

This report examines longitudinal trends in life expectancy at birth and mortality rates (deaths per 100 000) in the US population,

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with a focus on *midlife*, defined here as adults aged 25 to 64 years. This age range was chosen because the literature has reported increases in mortality rates among both young adults (as young as 25 years) and middle-aged adults (up to age 64 years); in this article, midlife mortality refers to mortality in both age groups combined (25-64 years). Life expectancy at birth is an estimate of the number of years a newborn is predicted to live, based on period life table calculations that assume a hypothetical cohort is subject throughout its lifetime to the prevailing age-specific death rates for that year.¹⁴ All-cause mortality and cause-specific mortality rates for key conditions were examined, using the International Statistical Classification of Diseases and Related Health Problems (ICD-10)¹⁵ codes detailed in the Supplement. Age-specific rates were examined for age groups of 10 years or fewer, whereas age-adjusted rates were examined for broader age groups. Age-adjustment rates were provided, and calculated, by the National Center for Health Statistics, using methods described elsewhere. 16

Data Sources

Life expectancy data were obtained from the National Center for Health Statistics¹⁷ and US Mortality Database.¹⁸ The latter was used for long-term trend analyses because it provided complete life tables for each year from 1959 to 2016 and at multiple geographic levels. 19 The analysis examined 2 periods. First, life expectancy was examined from a long-term perspective (from 1959 onward) to identify when life expectancy trajectories began to change in the United States and the 50 states. Second, knowing from the literature that mortality rates for specific causes (eg, drug overdoses) began increasing in the 1990s, a detailed analysis of cause-specific mortality trends was conducted for 1999-2017. Mortality rates were obtained from CDC WONDER.²⁰ Pre-1999 mortality data, although available, were not examined because the priority was to understand the conditions responsible for current mortality trends and because changes in coding in the transition from the ICD-9 (International Classification of Diseases, Ninth Revision) to the ICD-10¹⁵ could introduce artifactual changes in mortality rates. Methods available to make these conversions were therefore not pursued.

Analytic Methods

Life expectancy and mortality data were stratified by sex and across the 5 racial/ethnic groups used by the US Census Bureau²¹: non-Hispanic American Indian and Alaskan Native, non-Hispanic Asian and Pacific Islander, non-Hispanic black (or African American), non-Hispanic white, and Hispanic. Mortality rates were stratified by geography, including rates for the 9 US Census divisions (New England, Middle Atlantic, East North Central, West North Central, South Atlantic, East South Central, West South Central, Mountain, and Pacific), the 50 states, and urban and rural counties as defined in the Supplement. Data for the District of Columbia and US territories were not examined.

Changes in mortality rates between 2 years (2-point comparisons) were deemed significant based on 95% confidence intervals. Trends in life expectancy and mortality over time were examined to identify changes in slope and points of retrogression—defined as a period of progress (increasing life expectancy or decreasing mortality) followed by stagnation

(slope statistically equivalent to zero) or a significant reversal. Temporal trends were analyzed using Joinpoint Regression Program version 4.7.0.0,²² which models consecutive linear segments on a log scale, connected by joinpoints where the segments meet (ie, years when slopes changed significantly). A modification of the program's Bayesian Information Criteria method (called BIC3²³) was substituted for the Monte Carlo permutation tests to reduce computation time. Slopes (annual percent rate change [APC]) were calculated for the line segments linking joinpoints, and the weighted average of the APCs (the average annual percent change [AAPC]) was calculated for 3 periods: 1959-2016, 2005-2016, and 2010-2016 for life expectancy and 1999-2017, 2005-2017, and 2010-2017 for mortality rates. Slopes were considered increasing or decreasing if the estimated slope differed significantly from zero. The statistical significance of the APCs and the change in APCs between consecutive segments was determined by 2-sided t testing ($P \le .05$). Specific model parameters are available in the Supplement.

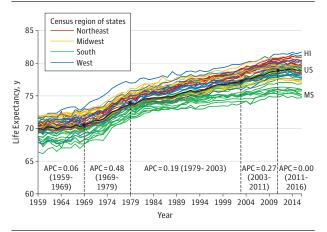
Excess deaths attributed to the increase in midlife mortality during 2010-2017 were estimated by multiplying the population denominator for each year by the mortality rate of the previous year, repeating this for each year from 2011 to 2017, and summing the difference between expected and observed deaths. ²⁴⁻²⁶ Excess deaths were estimated for each state and census division, allowing for estimates of their relative contribution to the national total.

Literature Review

To add context to the vital statistics described above and more fully characterize what is known about observed trends, the epidemiologic literature was examined for other research on US and state life expectancy and mortality trends. Using PubMed and other bibliographic databases, studies published between January 1990 and August 2019 that examined life expectancy or midlife mortality trends or that disaggregated data by age, sex, race/ethnicity, socioeconomic status, or geography were examined, along with the primary sources they cited. Research on the factors associated with the specific causes of death (eg, drug overdoses, suicides) responsible for increasing midlife mortality was also reviewed. Research on the methodological limitations of epidemiologic data on mortality trends was also examined.

To review contextual factors that may explain observed mortality trends and the US health disadvantage relative to other high-income countries, epidemiologic research was augmented by an examination of relevant literature in sociology, economics, political science, history, and journalism. A snowball technique^{27,28} was used to locate studies and reports on (1) the history and timing of the opioid epidemic; (2) the contribution of modifiable risk factors (eg, obesity) to mortality trends; (3) changes in the prevalence of psychological distress and mental illness; (4) the evidence linking economic conditions and health; (5) relevant economic history and trends in income and earnings, wealth inequality, and austerity during the observation period; (6) changes in subjective social status (eg, financial insecurity) and social capital; and (7) relevant federal and state social and economic policies, including the role of geography (eg, rural conditions) and state-level factors. The study was exempt from institutional review under 45 CFR 46.101(b)(4).

Figure 1. Life Expectancy for United States and 50 States, Grouped by Census Region, 1959-2016



Black curve indicates US life expectancy; bolded data points on black curve note joinpoint years, when the linear trend (slope) changed significantly based on joinpoint analysis. APC indicates the average annual percent change for the 5 periods identified on joinpoint analysis. Northeast census region (red) includes Connecticut, Maine, Massachusetts, New Hampshire, Rhode Island, and Vermont [New England division] and New York, New Jersey, and Pennsylvania [Middle Atlantic division]). Midwest census region (yellow) includes Ohio, Indiana, Illinois, Michigan, and Wisconsin [East North Central division] and Iowa. Kansas, Minnesota, Missouri, Nebraska, North Dakota, and South Dakota [West North Central division]). South census region (green) includes Delaware, Florida, Georgia, Maryland, North Carolina, Virginia, South Carolina, and West Virginia [South Atlantic division]; Alabama, Kentucky, Mississippi, and Tennessee [East South Central division]; and Arkansas, Louisiana, Oklahoma, and Texas [West South Central division]). West census region (blue) includes Arizona, Colorado, Idaho, Montana, Nevada, New Mexico, Wyoming, and Utah [Mountain division] and Alaska, California, Hawaii, Oregon, and Washington [Pacific division]). Life expectancy data obtained from the US Mortality Database. $^{\rm 18}$

Results

Life Expectancy

Between 1959 and 2016, US life expectancy increased by almost 10 years, from 69.9 years in 1959 to 78.9 years in 2016, with the fastest increase (highest APC) occurring during 1969-1979 (APC = 0.48, P < .01) (Figure 1). Life expectancy values for 1959-2016 are reported in eTable 1 in the Supplement for the United States, 9 census divisions, and the 50 states.

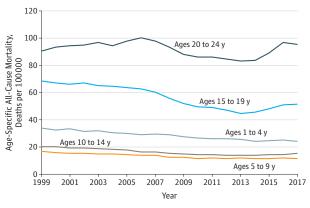
Life expectancy began to advance more slowly in the 1980s and plateaued in 2011 (after which the APC differed nonsignificantly from zero). The National Center for Health Statistics reported that US life expectancy peaked (78.9 years) in 2014 and subsequently decreased significantly for 3 consecutive years, reaching 78.6 years in 2017.^{2,9} The decrease was greater among men (0.4 years) than women (0.2 years) and occurred across racial-ethnic groups; between 2014 and 2016, life expectancy decreased among non-Hispanic white populations (from 78.8 to 78.5 years), non-Hispanic black populations (from 75.3 years to 74.8 years), and Hispanic populations (82.1 to 81.8 years).¹⁷

All-Cause Mortality

1998

The recent decrease in US life expectancy was largely related to increases in all-cause mortality among young and middle-aged

Figure 2. Age-Specific, All-Cause Mortality Rates Among US Youth, Aged 1-24 Years, 1999-2017



Source: CDC WONDER.20

adults. During 1999-2017, infant mortality decreased from 736.0 deaths/100 000 to 567.0 deaths/100 000, mortality rates among children and early adolescents (1-14 years) decreased from 22.9 deaths/100 000 to 16.5 deaths/100 000 (Figure 2), and age-adjusted mortality rates among adults aged 65 to 84 years decreased from 3774.6 deaths/100 000 to 2875.4 deaths/100 000.²⁰

eTable 2 in the Supplement presents age-specific, all-cause mortality rates for infants, children aged 1 to 4 years, and subsequent age deciles. Individuals aged 25 to 64 years experienced retrogression: all-cause mortality rates were in decline in 2000, reached a nadir in 2010, and increased thereafter. Retrogression even occurred among those aged 15 to 24 years (Figure 2). However, the increase was greatest in midlife—among young and middle-aged adults (25-64 years), whose age-adjusted all-cause mortality rates increased by 6.0% during 2010-2017 (from 328.5 deaths/100 000 to 348.2 deaths/100 000) (Figure 3). The relative increase in midlife mortality was greatest among younger adults (25-34 years), whose age-specific rates increased by 29.0% during this period (from 102.9 deaths/100 000 to 132.8 deaths/100 000).²⁰ The increases in death rates among middleaged adults (45-64 years) were less related to mortality among those aged 45 to 54 years, which decreased (from 407.1 deaths/ 100 000 to 401.5 deaths per 100 000), than among those aged 55 to 64 years, whose age-specific rates increased during 2010-2017 (from 851.9 deaths/100 000 to 885.8 deaths/100 000).²⁰

Cause-Specific Mortality

Although all-cause mortality in midlife did not begin increasing in the United States until 2010, midlife mortality rates for a variety of specific causes (eg, drug overdoses, hypertensive diseases) began increasing earlier (Figure 4).^{29,30} eTable 3 in the Supplement presents absolute and relative changes in agespecific mortality rates by cause of death between 1999 and 2017 (and between 2010 and 2017) for every age group (by age decile), from infancy onward, and shows that mortality rates increased primarily in midlife for 35 causes of death. The increase in cause-specific mortality was not always restricted to midlife; younger and older populations were often affected, although

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B Ages 45-64 y A All age groups^a 700 Age-Adjusted Mortality, Deaths per 100 000 Age-Adjusted Mortality, Deaths per 100 000 Ages 45-64 y (581.7 [2012]; 591.8 [2017]) 600 660 500 620 400 Ages 25-64 y (328.5 [2010]; 348.2 [2017]) 300 580 200 Ages 25-44 y (139.8 [2010]; 166.8 [2017]) 540 100 2001 2005 2007 2009 2011 2013 2015 2001 2007 2009 2011 2013 Year Year **C** Ages 25-64 y **D** Ages 25-44 y 200 Age-Adjusted Mortality, Deaths per 100 000 Age-Adjusted Mortality, Deaths per 100 000 360 160 340 140 320 120 2001 2007 2009 2011 2013 2015 2001 2007 2009 2011 2013 2015 1999 2003 2005 1999 2003 2005 Year Year

Figure 3. Age-Adjusted, All-Cause Mortality Rates, US Adults Aged 25-64 Years, 25-44 Years, and 45-64 Years, 1999-2017

Source: CDC WONDER.20

^a The lowest mortality rates per 100 000 (and the years they were achieved) are listed first in parentheses; mortality rates for 2017 listed second.

typically not as greatly (in relative or absolute terms) as those aged 25 to 64 years.

Year-by-year midlife mortality rates by cause for 1999-2017 (eTable 4 in the Supplement) show that retrogression occurred across multiple causes of death, in which progress in lowering midlife mortality was reversed. From 1999 to 2009, these cause-specific increases were not reflected in all-cause mortality trends because they were offset by large, co-occurring reductions in mortality from ischemic heart disease, cancer, HIV infection, motor vehicle injuries, and other leading causes of death. ³¹⁻³³ However, increases in cause-specific mortality rates before 2010 slowed the rate at which all-cause mortality decreased (and life expectancy increased) and eventually culminated in a reversal. The end result was that all-cause mortality increased after 2010 (and life expectancy decreased after 2014). ^{34,35}

Drug Overdoses, Alcoholic Liver Disease, and Suicides

A major cause of increasing midlife mortality was a large increase in fatal drug overdoses, beginning in the 1990s. 30,35,36 Between 1999 and 2017, midlife mortality from drug overdoses increased by 386.5% (from 6.7 deaths/100 000 to 32.5 deaths/100 000). 20 Age-specific rates increased for each age

subgroup: rates increased by 531.4% (from 5.6 deaths/100 000 to 35.1 deaths/100 000) among those aged 25-34 years, by 267.9% (from 9.5 deaths/100 000 to 35.0 deaths/100 000) among those aged 35-44 years, and by 350.9% (from 7.2 deaths/ 100 000 to 32.7 deaths/100 000) among those aged 45-54 years. The largest relative increase in overdose deaths (909.2%, from 2.3 deaths/100 000 to 23.5 deaths/100 000) occurred among those aged 55 to 64 years.²⁰ Midlife mortality rates also increased for chronic liver disease and cirrhosis^{31,34,37,38}; during 1999-2017, age-adjusted death rates for alcoholic liver disease increased by 40.6% (from 6.4 deaths/100 000 to 8.9 deaths/ 100 000); age-specific rates among young adults aged 25 to 34 years increased by 157.6% (from 0.6 deaths/100 000 to 1.7 deaths/100 000).20 The age-adjusted suicide rate at ages 25 to 64 years increased by 38.3% (from 13.4 deaths/100 000 to 18.6 deaths/100 000) and by 55.9% (from 12.2 deaths/100 000 to 19.0 deaths/100 000) among individuals aged 55 to 64 years.²⁰ As others have reported,³⁹ suicide rates also increased among those younger than age 25 years. eTable 3 in the Supplement shows that, across all age groups, the largest relative increase in suicide rates occurred among children aged 5 to 14 years (from 0.6 deaths/100 000 to 1.3 deaths/100 000).

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Drug poisoning Diabetes Hypertensive diseases A Ages 25-34 y **B** Ages 35-44 y Mental/behavioral disorders involving psychoactive substances Alcoholic liver disease Deaths per 100 000 Age-Specific Mortality, Deaths per 100 000 Assault (homicide) Other heart diseasea Other transport accidents^a Suicide Chronic lower respiratory diseases Age-Specific Mortality, Year Year **C** Ages 45-54 y **D** Ages 55-64 y Age-Specific Mortality, Deaths per 100 000 Age-Specific Mortality, Deaths per 100 000

Figure 4. US Age-Specific Mortality Rates for Selected Causes, by Age Decile, 1999-2017

Causes of death (and corresponding International Statistical Classification of Diseases and Related Health Problems, Tenth Revision [ICD-10] codes) include diabetes mellitus (E10-E14), mental and behavioral disorders due to psychoactive substance use (F10-F19), hypertensive diseases (I10-I15), unintentional drug poisoning (X40-X44), intentional

self-harm (suicide) (X60-X84), and assault (homicide) (X85-Y09). Source: CDC WONDER. 20

^a Other heart disease (I30-I51) includes arrhythmias and heart failure; other transport accidents (V80-V99) include land, water, air, space, and other transport accidents.

Organ System Diseases and Injuries

The increase in deaths caused by drugs, alcohol, and suicides was accompanied by significant increases in midlife mortality from organ system diseases and injuries, some beginning in the 1990s. ^{26,31,34} Data for several examples are reported in eTables 3 and 4 in the Supplement. For example, between 1999 and 2017, age-adjusted midlife mortality rates for hypertensive diseases increased by 78.9% (from 6.1 deaths/100 000 to 11.0 deaths/100 000) and for obesity increased by 114.0% (from 1.3 deaths/100 000 to 2.7 deaths/100 000) (eTable 4 in the Supplement). ²⁰ The increase in mortality from hypertension is consistent with other reports. ⁴⁰

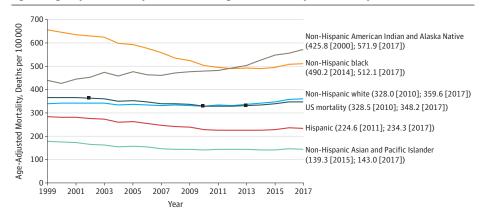
Early studies reported increasing midlife mortality from heart disease and lung (notably chronic pulmonary) disease, hypertension, stroke, diabetes, and Alzheimer disease, ^{31,34,41} but the trend appears to be even broader. According to 1 study, the increase in midlife mortality among non-Hispanic white populations during 1999-2016 was associated with an estimated 41 303 excess deaths

due to drug overdoses (n = 33 003) and suicides (n = 8300) but also more than 30 000 excess deaths due to organ system diseases (eg, hypertensive diseases [n = 5318], alcoholic liver disease [n = 3901], infectious diseases [n = 2149], liver cancer [n = 1931]), mental and behavioral disorders, obesity, pregnancy, and injuries (eg, pedestrian-vehicle collisions). 26 eTable 3 in the Supplement shows that the increase in organ disease mortality extended beyond midlife and, for certain diseases, was more pronounced in older age groups. For example, the largest increases in mortality from degenerative neurologic diseases (eg, Alzheimer disease) occurred among individuals 75 years and older.

Decomposition analyses, which quantify the relative contribution of specific causes of death to mortality patterns, have confirmed the large role played by organ system diseases. ^{10,31,33} For example, a decomposition analysis of the decline in US life expectancy between 2014 and 2015 found that respiratory and cardiovascular diseases contributed almost as much as external causes (including drug overdoses) among US women; among

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Figure 5. Age-Adjusted Mortality Rates, US Adults Aged 25-64 Years, by Race/Ethnicity, 1999-2017



Black curve indicates age-adjusted mortality for all US adults aged 25 to 64 years; bolded data points indicate joinpoint years, when the linear trend (slope) changed significantly based on joinpoint analysis. The lowest mortality rates per 100 000 (and the years they were achieved) are listed first in parentheses; mortality rates for 2017 listed second. Source: CDC WONDER.²⁰

men, drug overdoses explained almost all of the life expectancy decline. ¹⁰ In a more recent decomposition analysis, Elo et al³³ examined changes in life expectancy among US white populations between 1990-1992 and 2014-2016, stratifying the results by sex and geography. Deaths from mental and nervous system disorders were second only to deaths from drug overdoses in influencing changes in life expectancy and were the leading contributors to decreased life expectancy among white females. Among white females, respiratory disease mortality was a larger contributor to changes in life expectancy than either suicides or alcohol-related causes and accounted for more deaths in rural areas than drug overdoses (eTable 5 in the Supplement).

Sex-Related Patterns

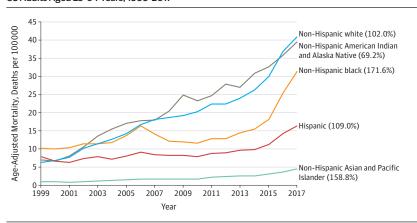
Absolute and relative increases in midlife mortality rates were higher among men than women.²⁰ Between 2010 and 2017, men aged 25 to 44 years experienced a larger relative increase in agespecific mortality rates than did women of that age, whereas women aged 45 to 64 years experienced a slightly larger relative increase in mortality than men of their age (eFigure 1 in the Supplement). Similarly, although men across age groups generally had higher cause-specific mortality rates and larger relative increases in mortality than did women, a pronounced female disadvantage emerged for certain major causes of death. For example, between 1999 and 2017, the relative increase in midlife fatal drug overdoses was 485.8% among women (from 3.5 deaths/100 000 to 20.2 deaths/100 000), 1.4 times higher than among men (350.6%, from 10.0 deaths/100 000 to 44.8 deaths/100 000). The relative increase in midlife mortality among women was 3.4 times higher for alcoholic liver disease (increasing from 3.2 deaths/100 000 to 5.8 deaths/100 000 among women and from 9.8 deaths/100 000 to 12.2 deaths/100 000 among men) and 1.5 times higher for suicide (increasing from 5.8 deaths/100 000 to 8.7 deaths/100 000 among women and 21.3 deaths/100 000 to 28.6 deaths/100 000 among men).20 This is consistent with reports elsewhere of gender-specific influences on mortality and a growing health disadvantage among US women, including smaller gains in life expectancy than among US men, larger relative increases in mortality from certain causes, and inferior health outcomes in comparison with women in other high-income countries. 11,33,42-46

Racial and Ethnic Patterns

Figure 5 stratifies all-cause mortality rates by race/ethnicity for adults aged 25 to 64 years; eFigure 2 in the Supplement similarly stratifies rates by age subgroups (25-44 years and 45-64 years). Midlife mortality rates among non-Hispanic American Indian and Alaskan Native and non-Hispanic black adults exceeded rates among other racial/ethnic groups, 20 consistent with other reports. 47,48 During 1999-2017, retrogression occurred in all racial/ ethnic groups except non-Hispanic American Indian and Alaskan Native adults, who experienced steady increases in midlife mortality rates on a larger relative scale than any other group. 5,20,34,38 Retrogression in the non-Hispanic white population, beginning in 2010, preceded its occurrence in non-Hispanic black (2014) and Hispanic (2011) populations (Figure 5), perhaps explaining why early studies reported that midlife mortality rates had not increased in these groups and focused their research on the white population. 31,34,37,38,41 Mortality patterns varied significantly by race/ethnicity and age, as illustrated in eFigure 3 in the Supplement, where absolute and relative changes in age-specific mortality rates for men and women are plotted separately for 20 combinations of race and age. Among the findings are that rates generally decreased after 1999 among non-Hispanic Asian and Pacific Islander adults older than 35 years and Hispanic adults older than 45 years and—as Masters et al reported³⁰—that rates increased after 2010 among non-Hispanic white women aged 45 to 54 years but not among men of that age.

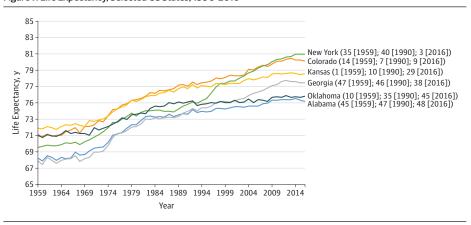
Consistent with the larger US population, populations of color began experiencing increases in cause-specific mortality rates long before experiencing the retrogression in all-cause mortality. ^{20,31,38} Midlife death rates in these populations increased across multiple, diverse conditions. One study reported that midlife mortality rates increased for 12 causes in the non-Hispanic American Indian and Alaskan Native population, 17 causes in the non-Hispanic black population, 12 causes in the Hispanic population, and 6 causes in the non-Hispanic Asian and Pacific Islander population. ²⁶ Each of these groups experienced large increases in fatal drug overdoses; between 2010 and 2017, the largest relative increase (171.6%) occurred among the non-Hispanic black population (**Figure 6**). ^{29,49} As shown online in eTable 6 in the Supplement, each of the 5 racial and ethnic groups also experienced increases in midlife deaths

Figure 6. Age-Adjusted Mortality From Unintentional Drug Overdoses, by Race/Ethnicity, US Adults Aged 25-64 Years, 1999-2017



Values in parentheses indicate relative increases in age-adjusted mortality rates by race/ethnicity between 2010 and 2017.
Source: CDC WONDER.²⁰

Figure 7. Life Expectancy, Selected US States, 1990-2016



Graph illustrates divergences in state-level life expectancy that began in the 1990s, including among neighboring states (Alabama/Georgia and Colorado/Kansas). Values in parentheses refer to state life expectancy rankings (among the 50 states in 1959, 1990, and 2016). As the 1990s began, life expectancy in Oklahoma exceeded that of New York. Source: US Mortality Database. 18

from alcoholic liver diseases, suicides, and hypertensive diseases, among others. ^{26,48} For example, in the non-Hispanic black population, midlife mortality from neurologic diseases increased from 10.2 deaths/100 000 to 14.1 deaths/100 000 between 1999 and 2017. The reversal (retrogression) in mortality rates that occurred among non-Hispanic black and Hispanic populations erased years of progress in lowering mortality rates (and reducing racial/ethnic disparities). The increase intensified recently for certain conditions (notably drug overdoses⁵⁰), with nonwhite populations experiencing larger relative and absolute year-to-year increases in death rates than white populations. ²⁶

Socioeconomic Patterns

Although an extensive literature links health to education, wealth, and employment, 51-58 direct evidence of their association with changes in life expectancy or mortality is limited, hampered by limited data to link deaths and socioeconomic history at the individual level. A growing body of evidence, however, indicates that the decline in US life expectancy and mortality risks have been greater among individuals with limited education (eg, less than high school) and income. 35,37,59-67 The gradient in life expectancy based on income has also widened over time, 68 with outcomes at the lower end of the distribution explaining much of the US disadvantage relative to other countries. 69

Geographic Patterns

Census Divisions and States

The range in life expectancy across the 50 states widened after 1984, reaching 7.0 years in 2016 (Figure 1). ¹⁸ States' life expectancy rankings also shifted over time, as illustrated in Figure 7. In 1959, Kansas had the nation's highest life expectancy (71.9 years), but its ranking declined over time, to 29th by 2016. In 1959, life expectancy in Oklahoma (71.1 years), 10th highest in the nation, exceeded that of New York (69.6 years), which ranked 35th. By 2016, New York's life expectancy (80.9 years) was third in the nation and Oklahoma's life expectancy (75.8 years) ranked 45th. State life expectancy trajectories often changed acutely after the 1990s, a finding that was more apparent when it occurred in adjacent states. For example, life expectancy in Colorado and Kansas differed by only 0.3 years in 1990, but that difference increased to 1.5 years in 2016; the difference between Alabama and Georgia increased from 0.1 years to 2.3 years. ¹⁸

The recent decrease in US life expectancy and increase in midlife mortality rates were concentrated in certain states, with the largest changes observed in New England and East North Central states and smaller changes in the Pacific and West South Central divisions (Figure 8). The chart book in the Supplement contains 120 graphs of life expectancy (and all-cause mortality) trends for the United States, 9 census divisions, and 50 states, as modeled by the Joinpoint Regression Program. It shows that, in the years leading up to 2016,

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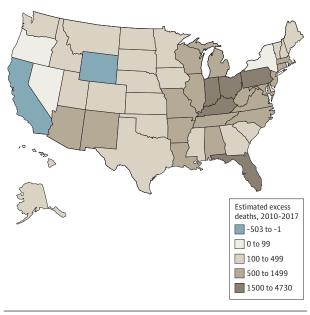
life expectancy trended downward in 4 census divisions and 31 states—beginning in 2009 (n = 3), 2010 (n = 4), 2011 (n = 6), 2012 (n = 9), 2013 (n = 6), and 2014 (n = 3)—and decreased significantly (based on APC) in Kentucky, Ohio, and New Hampshire.

The **Table** presents APC and AAPC data for life expectancy trends in the United States, 9 census divisions, and 50 states. The table displays the APC for the 2 most recent time intervals, how the slope changed between intervals, and the AAPC for 2010-2016. For example, life expectancy in New Hampshire increased significantly (APC = 0.2) from 1978 to 2012 but decreased significantly thereafter (APC = -0.4), with the joinpoint year of 2012 marking a statistically significant ($P \le .05$) unfavorable reduction in slope (-0.63%). For the period of 2010-2016, the slope was significantly negative (AAPC = -0.20). Unfavorable reductions in slope occurred from 2009 onward in 38 states—ie, life expectancy either decreased more rapidly or increased more slowly—and the slope change was significant (P < .05) in every census division and in 29 states. The largest decreases in life expectancy (based on AAPC for 2010-2016) occurred in New Hampshire, Kentucky, Maine, Ohio, West Virginia, South Dakota, New Mexico, Utah, Indiana, Mississippi, and Tennessee. Other states did not experience decreases in life expectancy; for example, life expectancy increased significantly in the Pacific division and in 13 states (Virginia, Delaware, South Carolina, Texas, Hawaii, New York, Oregon, New Jersey, Montana, Wyoming, Alabama, Arkansas, and Oklahoma).

Figure 9, Figure 10, and Figure 11 show the increase in midlife mortality rates during 1999-2017. Blue cells indicate favorable (negative APC) mortality trends; brown cells indicate unfavorable (positive APC) trends, based on joinpoint analysis (gray cells represent joinpoints). Many states experienced retrogressiondeclining mortality followed by a mortality reversal. For example, in Connecticut, a period of decreasing midlife mortality during 1999-2008 (blue shading) was followed by a statistically stable period in 2008-2014 (no shading)—during which the lowest mortality rate (253.7 deaths/100 000) was reached in 2011—and then by a significant increase in midlife mortality during 2014-2017 (brown shading). The remaining columns explain that midlife mortality in Connecticut increased by 9.0% between 2010 and 2017 ($P \le .05$), that what appeared to be a long-term decrease in mortality during 1999-2017 (AAPC = -0.6) obscured progressively less favorable trends in recent periods (AAPC = 0.2 [2005-2017] and 1.3 [2010-2017]), and that the increase in mortality (APC = 3.2) in the most recent period (2014-2017) differed significantly both from zero (footnote a) and from the slope of the prior segment (footnote b). The final column notes that year-to-year changes in mortality during 1999-2017 caused an estimated 441 excess midlife deaths in Connecticut.

The increase in midlife mortality was geographically wide-spread. Figures 9 through 11 show that the AAPC for 2010-2017 was positive in 8 census divisions and all but 4 states (California, New York, Oregon, and Texas). Thirty-seven states experienced statistically significant increases in midlife mortality (positive APC) in the years leading up to 2017. However, the trend was concentrated in certain states. Between 2010 and 2017, the largest relative increases in mortality occurred in New England (New Hampshire, 23.3%; Maine, 20.7%; Vermont, 19.9%, Massachusetts 12.1%) and the Ohio Valley (West Virginia, 23.0%; Ohio, 21.6%; Indiana, 14.8%; Kentucky, 14.7%), as well as in New Mexico (17.5%), South Dakota (15.5%), Pennsylvania (14.4%), North Dakota (12.7%), Alaska (12.0%),

Figure 8. Estimated Excess Deaths From Increasing Midlife Mortality, United States, 2010-2017



Estimated number of deaths caused by year-to-year increases in age-adjusted mortality rates among adults ages 25 to 64 years during 2010-2017. The map displays the number of deaths in each state without consideration of the state's population, to emphasize which states contributed the largest absolute number of deaths and exerted the largest influence on national trends. For example, although New Hampshire experienced a large (23.3%) relative increase in midlife mortality rates between 2010 and 2017, that state had a relatively small population (0.4% of US population) and therefore accounted for only 1.2% of excess deaths in the United States. States in blue had "negative" excess deaths, indicating a favorable net trend in which midlife mortality generally decreased. Estimated excess deaths for the remaining states are grouped for convenient interpretation. Divided by quintiles, the ranges were –503 to 151 (quintile 1), 152 to 369 (quintile 2), 370 to 563 (quintile 3), 564 to 1023 (quintile 4), and 1024 to 4730 (quintile 5). See text for more details about methods used to calculate excess deaths.

and Maryland (11.0%). In contrast, the nation's most populous states (California, Texas, and New York) experienced relatively small increases in midlife mortality.

Five states (Iowa, New Mexico, Oklahoma, West Virginia, and Wyoming) experienced a nearly continuous increase in midlife mortality (only positive APC segments) throughout 1999-2017, the largest (33.8%) occurring in West Virginia. Thirty-eight states experienced progress (declining mortality) as the millennium began, followed by retrogression (time segments beginning in 1999-2003 with negative APCs, followed by periods of increasing mortality with positive APCs). These reversals occurred earlier in some states than others; for example, midlife mortality rates in Iowa and North Dakota reached a nadir in 2004, whereas nadirs in New Jersey and New York did not occur until 2014 and 2015, respectively. Causespecific mortality trends also varied by state, sometimes in opposite directions. For example, whereas rates of firearm-related suicides increased nationwide during 1999-2017, they remained stable or decreased in California, Connecticut, Maryland, New Jersey, and New York.20

Between 2010 and 2017, year-to-year changes in midlife mortality accounted for an estimated 33 307 excess US deaths

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Table. Joinpoint Analysis of Life Expectancy Trends—United States, Census Divisions, and States

	Trends in Last 2	2 Segments, y ^a				
	Penultimate AF	PC	Last APC		Change in Slope,	AAPC, 2010-2016
	Period	APC (95% CI)	Period	APC (95% CI)	% (95% CI)	(95% CI) ^a
United States	2003 to 2011	0.27 (0.21 to 0.34)	2011 to 2016	0.00 (-0.12 to 0.12)	-0.27 (-0.40 to -0.14)	0.05 (-0.05 to 0.14)
Division 1: New England	1977 to 2012	0.22 (0.22 to 0.23)	2012 to 2016	-0.09 (-0.32 to 0.14)	-0.32 (-0.54 to -0.10)	0.01 (-0.14 to 0.16)
Connecticut	2000 to 2010	0.31 (0.26 to 0.36)	2010 to 2016	0.04 (-0.06 to 0.14)	-0.27 (-0.38 to -0.16)	0.04 (-0.06 to 0.14)
Maine	1978 to 2011	0.19 (0.18 to 0.20)	2011 to 2016	-0.16 (-0.34 to 0.02)	-0.35 (-0.52 to -0.17)	-0.10 (-0.25 to 0.04)
Massachusetts	1977 to 2013	0.23 (0.22 to 0.24)	2013 to 2016	-0.17 (-0.59 to 0.24)	-0.40 (-0.81 to 0.00)	0.03 (-0.17 to 0.23)
New Hampshire	1978 to 2012	0.22 (0.21 to 0.23)	2012 to 2016	-0.41 (-0.66 to -0.15)	-0.63 (-0.88 to -0.38)	-0.20 (-0.36 to -0.03)
Rhode Island	1979 to 2012	0.20 (0.19 to 0.21)	2012 to 2016	-0.06 (-0.36 to 0.24)	-0.26 (-0.55 to 0.03)	0.03 (-0.16 to 0.22)
Vermont	1977 to 2009	0.24 (0.23 to 0.25)	2009 to 2016	-0.05 (-0.15 to 0.06)	-0.28 (-0.39 to -0.18)	-0.05 (-0.15 to 0.06)
Division 2: Middle Atlantic	1988 to 2010	0.31 (0.29 to 0.32)	2010 to 2016	0.10 (-0.01 to 0.21)	-0.21 (-0.31 to -0.10)	0.10 (-0.01 to 0.21)
New Jersey	1988 to 2012	0.30 (0.28 to 0.32)	2012 to 2016	0.05 (-0.16 to 0.25)	-0.25 (-0.45 to -0.05)	0.13 (0.00 to 0.26)
New York	1998 to 2012	0.31 (0.28 to 0.34)	2012 to 2016	0.10 (-0.07 to 0.27)	-0.21 (-0.38 to -0.05)	0.17 (0.06 to 0.28)
Pennsylvania	1981 to 2014	0.19 (0.19 to 0.20)	2014 to 2016	-0.41 (-1.03 to 0.21)	-0.60 (-1.21 to 0.00)	-0.01 (-0.21 to 0.19)
Division 3: East North Central	1980 to 2012	0.19 (0.18 to 0.20)	2012 to 2016	-0.10 (-0.28 to 0.08)	-0.29 (-0.46 to -0.11)	0.00 (-0.12 to 0.11)
Illinois	1994 to 2012	0.29 (0.26 to 0.31)	2012 to 2016	-0.03 (-0.23 to 0.18)	-0.31 (-0.51 to -0.11)	0.08 (-0.05 to 0.21)
Indiana	1981 to 2011	0.14 (0.13 to 0.15)	2011 to 2016	-0.10 (-0.24 to 0.05)	-0.24 (-0.38 to -0.10)	-0.06 (-0.18 to 0.06)
Michigan	1978 to 2011	0.20 (0.19 to 0.20)	2011 to 2016	-0.05 (-0.16 to 0.06)	-0.25 (-0.36 to -0.14)	-0.01 (-0.10 to 0.08)
Ohio	1983 to 2012	0.15 (0.14 to 0.16)	2012 to 2016	-0.22 (-0.41 to -0.04)	-0.37 (-0.55 to -0.19)	-0.10 (-0.22 to 0.02)
Wisconsin	2002 to 2009	0.25 (0.17 to 0.34)	2009 to 2016	-0.03 (-0.09 to 0.04)	-0.28 (-0.39 to -0.18)	-0.03 (-0.09 to 0.04)
Division 4: West North Central	2001 to 2010	0.20 (0.15 to 0.25)	2010 to 2016	-0.02 (-0.09 to 0.06)	-0.21 (-0.30 to -0.13)	-0.02 (-0.09 to 0.06)
Iowa	1980 to 2010	0.16 (0.15 to 0.16)	2010 to 2016	-0.04 (-0.13 to 0.05)	-0.20 (-0.28 to -0.11)	-0.04 (-0.13 to 0.05)
Kansas	2001 to 2011	0.16 (0.12 to 0.20)	2011 to 2016	-0.03 (-0.14 to 0.08)	-0.19 (-0.30 to -0.08)	0.00 (-0.08 to 0.09)
Minnesota	2000 to 2009	0.26 (0.20 to 0.32)	2009 to 2016	0.00 (-0.07 to 0.08)	-0.26 (-0.35 to -0.16)	0.00 (-0.07 to 0.08)
Missouri	1999 to 2012	0.18 (0.16 to 0.21)	2012 to 2016	-0.11 (-0.28 to 0.06)	-0.29 (-0.46 to -0.12)	-0.01 (-0.12 to 0.10)
Nebraska	1979 to 2011	0.17 (0.16 to 0.18)	2011 to 2016	-0.04 (-0.18 to 0.11)	-0.20 (-0.34 to -0.06)	0.00 (-0.12 to 0.12)
North Dakota	2002 to 2005	0.39 (-0.69 to 1.48)	2005 to 2016	0.01 (-0.04 to 0.06)	-0.37 (-1.43 to 0.68)	0.01 (-0.04 to 0.06)
South Dakota	1982 to 2013	0.16 (0.15 to 0.18)	2013 to 2016	-0.32 (-0.80 to 0.16)	-0.48 (-0.95 to -0.01)	-0.08 (-0.31 to 0.16)
Division 5: South Atlantic	2003 to 2012	0.30 (0.23 to 0.37)	2012 to 2016	-0.06 (-0.28 to 0.16)	-0.36 (-0.59 to -0.13)	0.06 (-0.09 to 0.20)
Delaware	1969 to 1977	0.60 (0.46 to 0.73)	1977 to 2016	0.21 (0.20 to 0.22)	-0.39 (-0.52 to -0.25)	0.21 (0.20 to 0.22)
Florida	2005 to 2013	0.33 (0.22 to 0.44)	2013 to 2016	-0.14 (-0.55 to 0.26)	-0.47 (-0.88 to -0.06)	0.09 (-0.11 to 0.30)
Georgia	2003 to 2012	0.31 (0.22 to 0.40)	2012 to 2016	-0.12 (-0.39 to 0.16)	-0.42 (-0.71 to -0.14)	0.02 (-0.16 to 0.21)
Maryland	2001 to 2012	0.36 (0.31 to 0.41)	2012 to 2016	-0.11 (-0.33 to 0.12)	-0.47 (-0.69 to -0.24)	0.05 (-0.10 to 0.20)
North Carolina	2000 to 2012	0.26 (0.22 to 0.30)	2012 to 2016	-0.09 (-0.30 to 0.12)	-0.35 (-0.56 to -0.14)	0.02 (-0.11 to 0.16)
South Carolina	1972 to 1979	0.79 (0.61 to 0.96)	1979 to 2016	0.20 (0.19 to 0.21)	-0.59 (-0.76 to -0.41)	0.20 (0.19 to 0.21)
Virginia	1969 to 1979	0.53 (0.47 to 0.60)	1979 to 2016	0.22 (0.21 to 0.23)	-0.31 (-0.37 to -0.25)	0.22 (0.21 to 0.23)
West Virginia	1996 to 2014	0.05 (0.03 to 0.06)	2014 to 2016	-0.37 (-1.07 to 0.35)	-0.41 (-1.11 to 0.28)	-0.09 (-0.32 to 0.14)
Division 6: East South Central	2005 to 2011	0.20 (0.07 to 0.33)	2011 to 2016	-0.10 (-0.23 to 0.03)	-0.31 (-0.49 to -0.13)	-0.05 (-0.16 to 0.06)
Alabama	1969 to 1982	0.49 (0.45 to 0.53)	1982 to 2016	0.09 (0.09 to 0.10)	-0.39 (-0.43 to -0.35)	0.09 (0.09 to 0.10)
Kentucky	1992 to 2013	0.09 (0.08 to 0.11)	2013 to 2016	-0.38 (-0.73 to -0.03)	-0.47 (-0.81 to -0.13)	-0.14 (-0.31 to 0.03)
Mississippi	2005 to 2010	0.31 (0.06 to 0.55)	2010 to 2016	-0.06 (-0.18 to 0.06)	-0.36 (-0.63 to -0.10)	-0.06 (-0.18 to 0.06)
Tennessee	2003 to 2011	0.23 (0.14 to 0.32)	2011 to 2016	-0.10 (-0.26 to 0.06)	-0.33 (-0.51 to -0.16)	-0.05 (-0.17 to 0.08)
Division 7: West South Central	2003 to 2011	0.25 (0.19 to 0.31)	2011 to 2016	0.03 (-0.08 to 0.14)	-0.22 (-0.34 to -0.10)	0.07 (-0.02 to 0.15)
Arkansas	1973 to 1978	0.68 (0.44 to 0.92)	1978 to 2016	0.09 (0.08 to 0.10)	-0.58 (-0.82 to -0.35)	0.09 (0.08 to 0.10)
Louisiana	2005 to 2010	0.42 (0.18 to 0.66)	2010 to 2016	0.02 (-0.10 to 0.14)	-0.40 (-0.66 to -0.14)	0.02 (-0.10 to 0.14)
Oklahoma	1987 to 1995	-0.03 (-0.10 to 0.05)	1995 to 2016	0.06 (0.05 to 0.07)	0.09 (0.01 to 0.16)	0.06 (0.05 to 0.07)
Texas	1972 to 1977	0.65 (0.42 to 0.88)	1977 to 2016	0.19 (0.19 to 0.20)	-0.45 (-0.67 to -0.23)	0.19 (0.19 to 0.20)

(continued)

Table. Joinpoint Analysis of Life Expectancy Trends—United States, Census Divisions, and States (continued)

	Trends in Last 2	Segments, y ^a				
	Penultimate AP	С	Last APC		Change in Slope,	AAPC, 2010-2016
	Period	APC (95% CI)	Period	APC (95% CI)	% (95% CI)	(95% CI) ^a
Division 8: Mountain	2005 to 2010	0.31 (0.15 to 0.48)	2010 to 2016	0.01 (-0.07 to 0.10)	-0.30 (-0.48 to -0.12)	0.01 (-0.07 to 0.10)
Arizona	2006 to 2009	0.60 (-0.22 to 1.42)	2009 to 2016	0.05 (-0.05 to 0.16)	-0.55 (-1.34 to 0.25)	0.05 (-0.05 to 0.16)
Colorado	2002 to 2012	0.24 (0.19 to 0.29)	2012 to 2016	-0.07 (-0.28 to 0.13)	-0.31 (-0.52 to -0.11)	0.03 (-0.10 to 0.16)
Idaho	1991 to 2010	0.16 (0.14 to 0.17)	2010 to 2016	-0.02 (-0.11 to 0.08)	-0.17 (-0.27 to -0.08)	-0.02 (-0.11 to 0.08)
Montana	1977 to 1988	0.34 (0.27 to 0.41)	1988 to 2016	0.12 (0.11 to 0.13)	-0.22 (-0.29 to -0.15)	0.12 (0.11 to 0.13)
Nevada	2005 to 2010	0.40 (0.20 to 0.61)	2010 to 2016	0.06 (-0.05 to 0.16)	-0.34 (-0.56 to -0.12)	0.06 (-0.04 to 0.16)
New Mexico	1983 to 2013	0.13 (0.12 to 0.15)	2013 to 2016	-0.27 (-0.70 to 0.16)	-0.40 (-0.82 to 0.02)	-0.07 (-0.28 to 0.14)
Utah	2004 to 2009	0.28 (0.10 to 0.45)	2009 to 2016	-0.07 (-0.14 to 0.00)	-0.34 (-0.53 to -0.16)	-0.07 (-0.13 to 0.00)
Wyoming	1971 to 1984	0.57 (0.49 to 0.64)	1984 to 2016	0.12 (0.11 to 0.14)	-0.44 (-0.51 to -0.37)	0.12 (0.11 to 0.14)
Division 9: Pacific	1988 to 2013	0.27 (0.26 to 0.28)	2013 to 2016	0.01 (-0.24 to 0.26)	-0.26 (-0.51 to -0.02)	0.14 (0.02 to 0.26)
Alaska	1984 to 2010	0.21 (0.18 to 0.23)	2010 to 2016	-0.04 (-0.27 to 0.18)	-0.25 (-0.47 to -0.03)	-0.04 (-0.27 to 0.18)
California	1988 to 2014	0.30 (0.29 to 0.31)	2014 to 2016	-0.12 (-0.70 to 0.45)	-0.42 (-0.98 to 0.14)	0.16 (-0.03 to 0.34)
Hawaii	1977 to 1997	0.14 (0.12 to 0.16)	1997 to 2016	0.18 (0.16 to 0.20)	0.04 (0.02 to 0.07)	0.18 (0.16 to 0.20)
Oregon	1971 to 1979	0.46 (0.35 to 0.58)	1979 to 2016	0.17 (0.16 to 0.18)	-0.29 (-0.40 to -0.18)	0.17 (0.16 to 0.18)
Washington	1980 to 2013	0.19 (0.19 to 0.20)	2013 to 2016	-0.03 (-0.32 to 0.26)	-0.22 (-0.51 to 0.06)	0.08 (-0.06 to 0.22)

Abbreviations: AAPC, average annual percent change; APC, annual percent change.

change for 2010-2016. The AAPC over any fixed interval is calculated using a weighted average of the slope coefficients of the underlying joinpoint regression line, with weights equal to the length of each segment over the interval. The final step of the calculation transforms the weighted average of slope coefficients to an annual percent change. Life expectancy data obtained from the US Mortality Database. ¹⁸

(Figures 9-11). Population sizes influenced states' individual contribution to national mortality trends. For example, although several New England states (New Hampshire, Maine, and Vermont) experienced large (20%-23%) relative increases in midlife mortality during 2010-2017, these states accounted for only 3.0% of excess deaths, owing to their small populations. The East North Central division accounted for 28.6% of excess deaths, and Ohio, Pennsylvania, Indiana, and Kentucky (which include 10.8% of the US population) accounted for the largest number of excess deaths: these 4 states accounted for approximately one-third (32.8%) of excess deaths. Eight of the 10 states with the largest number of excess deaths were in the industrial Midwest or Appalachia.

Counties and Cities

As a group, rural US counties experienced larger increases in all-cause midlife mortality than did metropolitan counties, ^{31,33} but more complex patterns emerged when county data were disaggregated by population size, sex, race/ethnicity, age, and causes of death. For example, although the relative increase in midlife drug overdose deaths during 1999-2017 among non-Hispanic white populations was higher in rural counties (749.4%, from 4.0 deaths/ 100 000 to 33.8 deaths/100 000) than metropolitan counties (531.2%, from 6.7 deaths/100 000 to 42.5 deaths/100 000), the largest relative increase in overdose deaths (857.8%, from 4.7 deaths/100 000 to 45.2 deaths/100 000) occurred in the suburbs of large cities (populations ≥1 million), where Hispanic populations also experienced their largest increase in midlife overdose deaths.²⁰ Among non-Hispanic black populations, the largest increase in overdose deaths occurred in small cities (populations

<250 000), but the largest increase among black adults aged 55 to 64 years was in large cities.³¹ The largest increase in midlife suicides among non-Hispanic American Indian and Alaskan Native and Hispanic adults was in metropolitan areas, whereas the largest increase among non-Hispanic black and white adults occurred in rural counties.²⁰ Among young white adults (25-34 years), the largest increase in suicides occurred in the suburbs.³¹ Mortality patterns for men and women also varied significantly across urban and rural areas, with residents of large cities experiencing the greatest increases in life expectancy.³³

Geographic disparities in mortality were associated with demographic characteristics and with community contextual factors independent of individual and household characteristics. For example, a multivariate analysis of drug-related mortality in 2006-2015 found that drug deaths were higher in counties with certain demographic characteristics (eg, older adults, active duty military or veterans, Native Americans) and in counties with miningdependent economies, high economic and family distress indices, vacant housing, or high rent. Mortality rates were lower in counties with more religious establishments, recent in-migrants, and dependence on public sector (ie, government) employment. 70 Similarly, studies in 5 states (California, Kansas, Missouri, Minnesota, and Virginia) found that increases in midlife mortality from "stressrelated conditions" (drug overdoses, alcohol poisoning, alcoholic liver disease, and suicides) were highest in counties with prolonged exposure to high poverty, unemployment, and stagnant household income. Examples included the Central Valley and northern rural counties of California, 71 the Ozark and Bootheel regions of Missouri, 72 and the southwestern coalfields of Virginia.⁷³

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^a Table presents slopes for the 2 most recent linear trends in life expectancy during 1959-2016. For the penultimate and most recent periods, the table presents the annual percent change, the years marking the beginning and end of each period, and the degree to which the slope changed between the 2 comparison periods. The table also presents the average annual percent

Figure 9. Age-Adjusted All-Cause Mortality Rates, Adults Aged 25-64 Years, for the United States, Census Divisions 1-3 (New England, Middle Atlantic, East North Central), and States (1999-2017)

No. Section No.																							Decreasin Increasing	Decreasing mortality (P <.05)	P <.05)
OJ. by Feat 201. by Feat Col. by Feat Recent Trend (APC, 2011) Recent Trend (APC, 2011) 201. by Feat 2011 2012 2013 2014 2015 2016 2017 30 339 339 339 339 339 339 339 339 339						•	-		1		1	0	,						Relative	Long-ter	m Trends	(AAPC)			Excess
2011 2012 2013 2014 2015 2017 2017 2017 yof Last Segment) 329.7 329.1 330.1 33.8 34.0 348.2 6.0° -0.3 0.0 0.9 1.5° 2013-2017 255.8 265.8 264.2 296.3 296.3 1.3° 0.0 0.9 1.5° 2013-2017 253.7 250.1 360.0 276.8 266.2 286.3 20.7° 0.0° 0.0° 1.3° 2.9° 2014-2017 300.1 300.1 310.5 310.1 23.9 0.0° 1.3° 3.2° 2014-2017 259.3 266.1 267.2 200.0 284.4 53.8 0.7 1.5 3.3° 0.0 0.0° 1.3° 2012-2017 269.1 300.1 32.9 314.7 23.3° 0.7 1.5 3.3° 2012-2017 269.2 303.1 309.1 31.0 0.0° 1.3° 20.2 2012-2017						4	vge-Aaju	핆	rality K.	ates (Dec	atus per	100000), by rea						(%, 2017	1999-	2002	2010-	Recent Trer		(2010-
329.1 329.1 380.9 33.8 347.0 348.2 6.0a -0.3 0.0 0.9b 1.5b 2013-2017 255.2 264.1 265.2 265.2 265.2 265.2 265.2 1.30a 0.0b 1.8b 2.9bc 2013-2017 255.2 265.1 266.2 266.2 266.2 267.2 1.7b 2.9bc 2013-2017 259.2 265.2 270.6 280.2 1.4b 2.6b 4.9bc 2013-2017 259.3 266.4 280.1 280.4 380.1 380.1 380.1 380.2		1999 20	000 20	02 100	02 200	3 200	4 200.	7	2007		2009	2010 2		012 20			15 2016		vs 2010)	2017	2017	2017	y of Last Se	_	2017)
265.8 264.1 269.1 275.8 286.3 296.3 13.0°a 0.0 0.0 b 1.8 b 2.9 bc 2012-2017 253.7 259.2 260.0 256.8 267.4 275.6 280.9 9.0°a -0.6 b 1.8 b 2.9 bc 2014-2017 380.1 333.2 260.2 262.2 272.6 280.0 284.4 291.8 12.1°a -0.6 b 1.4 b 2.6 b 4.9 bc 2013-2017 259.3 256.2 272.2 280.0 284.4 291.8 12.1°a -0.1 b 1.7 b 2.9 bc 2013-2017 269.1 266.2 260.2 272.6 284.4 291.8 12.1°a -0.7 b 1.7 b 2.9 bc 2012-2017 284.0 265.3 260.3 310.4 312.2 19.9°a 0.4 1.1 b 2.6 bc 2012-2017 284.0 265.3 284.3 310.2 284.3 310.2 2012-2017 2012-2017 285.5 286.9 290.4 </td <td></td> <td>365.4 36</td> <td>65.3 36</td> <td>54.8 36</td> <td>2.0 360</td> <td>.2 350.</td> <td>1 351.</td> <td>4 347.0</td> <td>340.3</td> <td>338.1</td> <td>335.5</td> <td>328.5</td> <td>29.7 3.</td> <td>29.1 33</td> <td></td> <td>3.9 338</td> <td>.2 347.</td> <td>0 348.2</td> <td>e.0a</td> <td>-0.3</td> <td>0.0</td> <td>0.9^b</td> <td></td> <td>13-2017</td> <td>33 307</td>		365.4 36	65.3 36	54.8 36	2.0 360	.2 350.	1 351.	4 347.0	340.3	338.1	335.5	328.5	29.7 3.	29.1 33		3.9 338	.2 347.	0 348.2	e.0a	-0.3	0.0	0.9 ^b		13-2017	33 307
253.2 259.2 260.0 256.8 60.6 b 0.0 b 1.3 b 3.2 bc 2014-2017 300.1 33.4 30.4 358.7 20.7 a 0.8 b 1.4 b 2.6 b 4.9 bc 2014-2017 259.3 256.2 272.2 272.6 280.0 287.4 29.7 a 12.1 a -0.1 b 1.4 b 2.6 b 4.9 bc 2013-2017 259.3 256.2 272.6 280.0 287.4 29.1 a 12.1 a -0.1 b 0.4 b 1.7 b 2.9 bc 2012-2017 259.2 250.2 272.6 280.1 23.3 a 0.7 1.5 a 3.9 bc 2012-2017 259.2 260.2 250.2 313.1 309.1 5.1 0.0 b 1.1 b 2.6 b 2012-2017 259.5 289.8 289.4 309.0 310.7 6.4 a -0.2 b -0.2 b 0.0 c 20.0 20.0 20.0 20.0 20.0 20.0 20.0 20.1 20.1 20.1 <th< td=""><td></td><td>302.1 30</td><td>07.3 30</td><td>14.4 304</td><td>1.3 301</td><td>.1 286.</td><td>1 283.</td><td>2 280.1</td><td>275.4</td><td>269.9</td><td>269.6</td><td>265.0 2</td><td>65.8 2</td><td>54.1 26</td><td>9.1 275</td><td>5.8 286</td><td>.9 296.</td><td></td><td>13.0^{a}</td><td>0.0</td><td>0.6^b</td><td></td><td></td><td>12-2017</td><td>2718</td></th<>		302.1 30	07.3 30	14.4 304	1.3 301	.1 286.	1 283.	2 280.1	275.4	269.9	269.6	265.0 2	65.8 2	54.1 26	9.1 275	5.8 286	.9 296.		13.0^{a}	0.0	0.6 ^b			12-2017	2718
300.1 303.4 304.1 36.9 1.4b 2.6b 4.9bc 2013-2017 259.3 265.2 262.2 272.6 280.0 287.4 291.8 12.1a -0.1 0.4b 1.7b 2.9bc 2013-2017 263.2 265.2 272.2 280.1 280.1 13.1 30.1 23.3a 0.7 1.5 3.3b 2.0 2015-2017 269.1 266.2 266.3 303.2 294.3 304.4 312.2 1.9b 0.7 1.1b 2.6b 2.0bc 2012-2017 284.0 205.4 302.1 31.2 1.9pa 0.7 1.1b 2.6b 2.0bc 2012-2017 284.0 201.8 202.9 30.4 312.2 1.9pa 0.2b 0.2b 2.0bc 2.0b		303.4 31	15.7 30	12.1 29.	7.4 292	.8 283.	8 274.	3 269.9	1 265.0	261.5	260.0	257.7 2	53.7 2:	59.2 26	0.0	5.8 267			9.0a	-0.6 ^b	0.2			14-2017	441
259.3 26.2 26.2.2 <td></td> <td>312.8 31</td> <td>13.4 30</td> <td>7.7 320</td> <td>0.4 314</td> <td>.8 298.</td> <td>7 317.</td> <td>9 314.4</td> <td>1 299.1</td> <td>300.3</td> <td>294.3</td> <td>297.3 3</td> <td>00.1</td> <td>33.4 30</td> <td></td> <td>333</td> <td>.0 344.</td> <td>6 358.7</td> <td>20.7a</td> <td>0.8^b</td> <td>1.4^b</td> <td></td> <td></td> <td>13-2017</td> <td>437</td>		312.8 31	13.4 30	7.7 320	0.4 314	.8 298.	7 317.	9 314.4	1 299.1	300.3	294.3	297.3 3	00.1	33.4 30		333	.0 344.	6 358.7	20.7a	0.8 ^b	1.4 ^b			13-2017	437
263.2 266.7 263.1 263.2 369.1 314.7 23.3° 0.7 1.5 3.3° 2.0 2015-2017 296.1 265.2 367.7 367.4 312.1 309.1 5.1 0.0 0.4° 0.9° 1.3° 2015-2017 296.1 261.8 271.6 294.4 309.4 312.2 19.9° 0.4 1.1° 2.6° 2.6° 2010-2017 295.5 286.9 270.6 310.7 6.4° -0.8° -0.2 0.9 2.8° 2010-2017 279.4 270.5 280.6 297.0 7.6° -0.8° -0.2 0.9 2.8° 2010-2017 279.4 270.5 280.6 297.3 3.6° 3.6° 3.0° 3.6° 3.0° 3.8° 3.0° <td></td> <td>300.3 30</td> <td>07.3 30</td> <td>9.6</td> <td>3.4 306</td> <td>.2 289.</td> <td>9 282.</td> <td>8 279.4</td> <td>1 274.8</td> <td>264.4</td> <td>268.9</td> <td>260.2</td> <td>59.3</td> <td>56.2 26</td> <td>2.2 272</td> <td>2.6 280</td> <td>.0 287.</td> <td>4 291.8</td> <td>12.1^a</td> <td>-0.1</td> <td>0.4^b</td> <td></td> <td></td> <td>12-2017</td> <td>1155</td>		300.3 30	07.3 30	9.6	3.4 306	.2 289.	9 282.	8 279.4	1 274.8	264.4	268.9	260.2	59.3	56.2 26	2.2 272	2.6 280	.0 287.	4 291.8	12.1 ^a	-0.1	0.4 ^b			12-2017	1155
296.1 285.3 30.7 299.3 30.4 31.1 30.1 5.1 0.0 0.4b 0.9b 1.3bc 2012-2017 284.0 261.8 276.8 271.6 294.4 309.4 312.2 19.9a 0.4 1.1b 2.6b 2.6bc 2010-2017 225.5 289.9 200.8 20.2 0.2 0.2 0.9 2.6bc 2010-2017 279.4 270.1 270.2 270.2 271.6 288.5 297.0 7.6a -0.9b -0.2 1.1 5.1 2014-2017 279.4 270.2 270.2 271.6 288.5 297.0 7.6a -0.9b -0.2 1.1 5.1 2014-2017 279.4 270.5 270.2 281.6 275.3 1.1 4.1 4.2bc 2014-2017 280.8 280.2 280.2 287.1 1.14a 0.1 0.7b 1.9bc 2.9bc 2014-2017 380.4 380.8 380.8 380.8		285.4 28	82.6 27	8.0 28.	1.6 276	.5 260.	1 275.	9 264.1	265.3	261.0	261.1	255.1 2	63.2 2	50.7 26		309	.1 329.	0 314.7	23.3a	0.7	1.5	3.3 ^b		15-2017	429
284.0 261.8 276.8 271.6 294.4 309.4 312.2 19.9a 0.4 1.1b 2.6b 2.6bx 2010-2017 295.5 289.9 290.8 289.5 291.4 300.0 310.7 6.4a -0.8b -0.2 0.9 2.8bx 2014-2017 279.4 270.1 271.6 288.5 297.0 7.6a -0.9b -0.2 1.1 5.1 2014-2017 273.4 270.5 280.6 275.5 280.7 7.6a -0.9b -0.2 1.1 5.1 2014-2017 340.8 237.0 286.0 275.3 360.2 377.1 14.4a 0.1 0.7b 1.0p 4.2bx 2014-2017 340.8 339.2 346.6 355.6 366.2 374.5 11.6a 0.1 0.6b 1.3b 4.2bx 2014-2017 330.4 308.8 304.6 351.3 361.3 367.1 41.8a 0.6b 1.1b 1.6b 2.9bx 2014-20		318.7 31	14.7 32	2.4 32	9.7 326	.4 302.	2 299.	1 297.7	, 299.6	303.9	297.1	294.1 2	96.1	35.3 30	3.7 299	9.3 305	.4 313.	1 309.1	5.1	0.0	0.4 ^b	0.9 ^b		12-2017	84
295.5 289.9 290.8 289.5 289.4 300.0 310.7 6.4a -0.8b -0.2 0.9 2.8b.c 2014-2017 273.4 73.1 270.2 271.6 288.5 297.0 7.6a -0.9b -0.2 1.1 5.1 2015-2017 273.4 270.5 286.0 275.5 39.0 7.6a -0.9b -0.1 0.7b 2012-2017 340.8 335.0 286.2 377.1 14.4a 0.1 0.7b 1.9b 4.2bc 2014-2017 380.9 335.2 345.3 346.6 37.4 11.6a 0.1 0.6b 1.9b 4.2bc 2014-2017 380.1 390.4 310.4 31.1 31.1 31.1 41.4a 0.1 0.6b 1.9b 4.2bc 2014-2017 380.2 340.4 308.1 31.1 31.1 31.1 41.4a 0.6b 1.0b 1.3bc 2012-2017 380.2 380.4 31.1 31.1		299.1 28	89.6 28	34.9 27	5.8 274	.7 267.	0 253.	4 278.5	272.3	271.1	251.1	260.3	84.0 2	51.8 27	6.8 271	1.6 294	.4 309.		19.9ª	0.4	1.1b			10-2017	171
279.4 273.1 274.6 288.5 297.0 7.6** -0.9* -0.2 1.1 5.1 2015-2017 273.4 270.5 280.5 275.5 275.3 0.1 -0.3* -0.1 0.7* 2012-2017 340.8 332.2 335.0 385.2 387.1 14.4* 0.1 0.7* 1.9* 4.7* 2012-2017 330.9 332.2 345.3 366.8 374.5 11.6* 0.1 0.6* 1.9* 4.7* 2014-2017 380.4 380.5 340.5 367.1 31.4 4.7* -0.8* -0.4* 0.5* 1.3* 2014-2017 381.2 382.5 304.5 31.7 31.1 4.7* -0.8* -0.4* 0.5* 1.3* 2014-2017 382.4 382.5 382.5 382.1 31.4 1.4.8* 0.6* 1.1 1.8* 3.0* 2014-2017 382.5 382.6 382.6 382.7 382.1 41.8* 1.0*		353.9 35	50.4 35	6.8 33	9.8 331	.4 323.	1 320.	2 317.0	305.8	302.4	298.3	292.1 2	95.5 2	39.9 29	0.8 289	<u>3.5</u> 293	.4 309.	0 310.7	6.4a	-0.8 ^b	-0.2			14-2017	4221
273.4 270.5 270.2 268.0 267.5 280.5 275.3 0.1 -1.3b -0.8b -0.1 0.7bc 2012-2017 340.8 332.2 335.0 336.8 349.5 369.2 377.1 14.4a 0.1 0.7b 1.9b 4.2bc 2014-2017 330.9 3392.2 345.3 346.6 377.1 11.6a 0.1 0.6b 1.6b 2.9bc 2014-2017 330.4 340.8 340.5 317.1 321.6 4.7a -0.8b -0.4b 0.5b 1.3bc 2014-2017 370.4 318.1 31.1 31.1 41.8 14.8a 0.6b 1.1b 1.8b 3.0bc 2014-2017 370.2 340.6 36.1 361.1 418.4 14.8a 0.0 0.0 1.1b 1.8b 3.0bc 2014-2017 370.2 340.6 356.8 361.1 418.4 14.8a 0.0 0.5b 1.0b 1.9bc 2014-2017 370.2		344.2 34	42.1 35	50.8 328	3.1 318	.6 309.	9 307.	1 309.5	1 287.0	286.0	276.4	276.1 2	79.4	73.1 27	4.0 270	271	.6 288.	5 297.0	7.6a	-0.9 ^b	-0.2	1.1		15-2017	1004
340.8 332.2 335.0 336.8 349.5 369.2 377.1 14.4° 0.1 0.7 1.9° 4.2° 2014-2017 339.9 339.2 347.3 346.6 358.3 374.5 11.6° 0.1 0.6° 1.6° 2.9° 2014-2017 305.4 308.4 31.2 11.6° 0.1 0.6° 1.6° 1.3° 2014-2017 370.4 318.2 304.2 307.3 317.1 321.6 4.7° -0.8° -0.4° 0.5° 1.3° 2014-2017 370.2 370.2 370.2 380.1 41.8 14.8° 0.6° 1.1° 1.8° 3.0° 2014-2017 370.2 370.2 367.1 378.0 378.2 6.4° 0.0 0.5° 1.0° 1.0° 2014-2017 380.2 317.2 383.6 421.5 40.4 21.6° 0.0 0.5° 1.0° 1.0° 2014-2017 380.2 381.2 382.0 421.5		349.8 34	41.9 35	3.6 330	0.8 321	.5 311.	7 304.	5 298.4	1 292.8	286.8	282.5	275.0 2	73.4 2	70.5 27	0.2 268	3.0 267	.5 280.		0.1	-1.3 ^b	-0.8 ^b			12-2017	38
339.9 339.2 342.3 346.6 353.6 368.3 374.5 11.6 ^a 0.1 0.6 ^b 1.6 ^b 2.9 ^b 2014-2017 305.4 308.8 304.5 307.9 311.7 321.6 321.9 4.7 ^a -0.8 ^b -0.4 ^b 0.5 ^b 1.3 ^b 2012-2017 373.7 375.7 378.5 387.1 391.8 407.1 418.4 14.8 ^a 0.6 ^b 1.1 ^b 1.8 ^b 3.0 ^b 2014-2017 358.1 340.2 361.3 367.1 378.0 378.2 64. ^a 0.0 0.5 ^b 1.0 ^b 1.5 ^b 2012-2017 370.2 371.9 377.2 383.6 398.0 421.5 440.4 21.6 ^a 1.0 ^b 1.5 ^b 2.8 ^b 4.9 ^b 2014-2017 383.6 389.3 30.6 30.6 30.6 36.9 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6	1 7	366.8 36	69.3 36	55.5 36.	2.2 355.	.9 350.	5 353.	9 350.7	339.1	337.8	338.2		40.8 3.	32.2 33	5.0 336		.5 369.		14.49	0.1	0.7 ^b			14-2017	3179
321.9 4.7a -0.8b -0.4b 0.5b 1.3bx 2012-2017 418.4 14.8a 0.6b 1.1b 1.8b 3.0bx 2014-2017 378.2 6.4a 0.0 0.5b 1.0b 1.5bx 2012-2017 440.4 21.6a 1.0b 1.5b 2014-2017 305.6 9.6a -0.1 0.4b 1.3b 2010-2017	1 7	366.2 36	64.0 36	51.8 36.	2.6 355	.3 346.	0 350.	7 345.8	341.1	343.7	340.5	335.7	39.9 3.	39.2 34	2.3 346	5.6 353	.6 368.		11.6^{a}	0.1	0.6 ^b			14-2017	9531
418.4 14.8° 0.6° 1.1° 1.8° 3.0°s 2014-2017 378.2 6.4° 0.0 0.5° 1.0° 1.5°s 2012-2017 440.4 21.6° 1.0° 1.5°s 4.9°s 2014-2017 305.6 9.6° -0.1 0.4° 1.3°s 2010-2017		372.9 36	64.7 36	3.3 36.	1.8 347	.5 341.	5 337.	5 330.6	323.4	326.6	318.0	307.4 3	05.4 3	38.8	4.5 307	7.9 311	.7 321.	6 321.9	4.7a	-0.8 ^b	-0.4 ^b			12-2017	686
378.2 6.4a 0.0 0.5b 1.0b 1.5bx 2012-2017 440.4 21.6a 1.0b 1.5b 2.8b 4.9bx 2014-2017 305.6 9.6a -0.1 0.4b 1.3b 1.3bx 2010-2017	. 7	373.9 37	77.7 37	75.9 37.	3.0 379	.0 362.	9 365.	2 374.0	358.4	366.7	368.6	364.4 3	73.7 3	75.7 37	8.5 387	7.1 391	.8 407.		14.8ª	q9'0	1.1b			14-2017	1839
440.4 21.6 ^a 1.0 ^b 1.5 ^b 2.8 ^b 4.9 ^b ^c 2014-2017 305.6 9.6 ^a -0.1 0.4 ^b 1.3 ^b 1.3 ^b ^c 2010-2017		382.3 37	76.4 37	3.1 37.	3.5 364	.9 355.	1 364.	7 359.5	356.3	356.6	354.0	355.5 3	58.1 3	<u> 19.6</u>	6.8 361	1.3 367	.1 378.	0 378.2	6.4a	0.0	0.5 ^b			12-2017	1167
305.6 9.6a -0.1 0.4b 1.3b 1.3bc 2010-2017		370.8 37	71.6 36	59.8 37.	3.7 369.	.9 362.	2 371.	2 364.4	1 362.6	368.7	368.6	362.2	70.2	71.9 37	7.2 383	3.6 398	.0 421.		21.6^{a}	1.0 ^b	1.5 ^b	_		14-2017	4730
	1	301.5 30	06.6 3C	14.3 308	3.9 297	.2 286.	6 296.	7 284.9	1 289.9	282.6	278.6	278.9 2	85.9 2	34.1 29	2.1 289	9.3 293	.7 306.	3 305.6	e9.6	-0.1	0.4 ^b	_	_	10-2017	908

Age-adjusted, all-cause mortality rates (per 100 000) among US adults ages 25-64 years for 1999-2017, along with the relative increase in mortality rates between 1999 and 2017 and the slopes modeled by the Joinpoint Regression Program (see Methods and Supplement). Slopes include the average annual percent change (AAPC) for 3 periods–1999-2017, 2005-2017, and 2010-2017—and the annual percent change (APC) for the most recent linear trend in the joinpoint model. Also shown are the estimated number of excess deaths in the United States caused by year-to-year changes in midlife mortality rates between 1999 and 2017. Blue shading indicates years during which mortality rates decreased (statistically significant [P < .05] negative APC); brown shading, years of increasing mortality (statistically significant [P < .05] positive APC); no shading, periods when the APC did not

differ significantly from zero; black cells, joinpoint years, when changes occurred in the modeled linear trends. Underlined mortality rates denote the lowest mortality rates (nadir) for 1999-2017. NA indicates not applicable (the Joinpoint program plotted a single trend line for 1999-2017; thus, no last segment).

³ Statistically significant (P < .05).

 $^{^{\}rm b}$ Slope (APC or AAPC) that differed significantly from zero (P < .05)

Statistically significant slope change estimate, a measure of the change in slope from that of the previous period;
 see e Table 7 in the Supplement for 95% confidence intervals. Mortality rates obtained from CDC WONDER.²⁰

Figure 10. Age-Adjusted All-Cause Mortality Rates, Adults Aged 25-64 Years, for US Census Divisions 4-6 (West North Central, South Atlantic, East South Central) and States (1999-2017)

																								Decr	Decreasing mortality (P <.05)	y (P <.05)
							-			į			2						æ 5	Relative	Long-ter	Long-term Trends (AAPC)	(AAPC)			Excess
US Census Divisions	1999 2000 2001 2002 2003	000	001	, , , 00		Age-Adjus	Age-Adjusted		auty Kat	Mortality Kates (Deaths per 100000), by Year	2009 3	2010	0), by Yes	. 5	2013 2	2014 2	2015 2	2016 20	(%)	(%, 2017	1999-	2005-	2010-	Recent	Recent Trend (APC,	(2010-
Division 4: West North Central 329.8 321.7 320.9 322.9 324.0 312.3 317.7 3	329.8 32	21.7 3	20.9 3	22.9 3	24.0 3	12.3	317.7	15.6			12.3 3		11.4 3	_	14.8 3.	18.8	1	_		7.7a	0.1	0.4b	1.0b	1.0b,c	2010-2017	2625
lowa	296.9 297.1 288.8 287.2 304.7 285.8 291.3 2	37.1 2	88.8 28	87.2 30	04.7	85.8	91.3	9.88	300.5	300.9	93.9 2	95.4 2	93.9 2	88.6 290.5 300.9 293.9 295.4 293.9 298.5 302.5	02.5	99.0	99.0	299.0 299.0 314.8 314.3	14.3	6.4 _a	0.5b	0.6 ^b	1.0	2.1	2014-2017	296
Kansas	342.7 329.5 337.2 342.0 337.3 334.5 339.4 3	29.5	37.2 34	42.0 3	37.3 3	34.5	339.4	336.6 326.0		323.8 328.4 328.4	128.4 3	28.4	333.1 3	333.3 329.7		335.9 3	348.0 342.3		353.6	7.7a	0.2	0.2	0.8b	0.8b,c	2008-2017	369
Minnesota	274.3 269.5 264.8 262.4 257.5 253.3 254.7 2	59.5	64.8 26	52.4 2	57.5 2	53.3	254.7	251.9	242.9	247.4 2	49.6	41.6	44.8 2	51.9 242.9 247.4 249.6 241.6 244.8 245.1 247.1 242.1 249.5 255.3	47.1 2	42.1 2	49.5 2		258.8	7.1a	-0.3	0.2	0.8b	2.0b,c	2014-2017	496
Missouri	395.4 387.9 387.5 391.8 394.7 380.9 387.4 3	37.9 3	87.5 39	91.8	94.7 3	80.9	387.4	386.6 369.2	369.2	387.4 380.9 378.0	80.9		380.7	381.6 386.4	86.4 3	95.0 3	395.0 399.9 404.9		407.5	7.8a	0.2b	0.5 ^b	1.1b	1.4b,c	2011-2017	931
Nebraska	312.5 291.7 298.3 314.4 295.2 284.6 286.2 289.1 285.7 290.5 285.4 279.3	91.7 2	98.3 3.	14.4	95.2 2	84.6	286.2	289.1	285.7	290.5	85.4		87.7 2	287.7 284.6 281.6	81.6 30	03.1 3	303.1 303.0 292.3		300.9	7.8a	0.0	0.3	1.0 ^b	1.0b,c	2010-2017	206
North Dakota	303.4 289.1 299.2 293.0 300.6 267.0 286.6 2	39.1 2	99.2	93.0 3	00.6 2	67.0 2	286.6	286.0 269.7		295.7 2	299.6 280.4		8.06	290.8 313.0 322.1	22.1 30	02.5 3	302.5 328.7 318.1		316.1	12.7a	0.5	1.2 ^b	1.4b	1.4b,c	2006-2017	128
South Dakota	339.8 321.5 323.1 316.0 342.1 309.5 333.4 3	21.5 3	23.1 3.	16.0 3	42.1 3	09.5	333.4	317.3 2	17.3 292.1	292.8 303.0 301.1	303.0	01.1	302.4	302.4 310.9 306.0	0.90	37.5 3	337.5 330.8 333.5	33.5 34	347.9	15.5a	0.3	0.5	1.9 ^b	1.9b	2008-2017	200
Division 5: South Atlantic	399.8 404.0 398.8 394.5 395.7 385.1 383.7 3	0.40	98.8	94.5	95.7	85.1	383.7	379.6 372.2	372.2	366.3 363.5 356.2	363.5		352.0 3	352.0 351.5 352.4	52.4 3.	55.9	355.9 361.1 375.2		375.8	5.5a	-0.3	-0.1	6.0	2.3 ^b	2014-2017	9029
Delaware	370.3 384.1 398.0 365.9 376.1 366.4 368.3 3	34.1 3	98.0 36	65.9	76.1 3	66.4	368.3	373.5	358.6	374.5 3	3.95	8.09	51.5 3	63.2 3	68.5 3	47.5 3	58.8 3	73.5 358.6 374.5 356.8 360.8 351.5 363.2 368.5 <u>347.5</u> 358.8 362.4 385.8	35.8	6.9a	0.0	0.3	8.0	4.1	2015-2017	125
Florida	389.1 393.6 388.9 386.5 389.3 382.1 377.6	93.6	88.9 38	86.5	89.3	82.1	377.6	372.6 364.5	364.5	363.6 3	358.6 350.7	50.7	344.7	342.9	339.6	343.2 3	348.8 367.8	67.8 36	364.6	3.9a	-0.3 ^b	-0.2	0.7 ^b	2.3 ^{b,c}	2013-2017	1513
Georgia	426.9 428.0 425.6 425.1 424.0 414.7 408.5 4	28.0 4	25.6 42	25.1 4.	24.0 4	14.7	108.5	102.2	388.6	02.2 388.6 382.2 383.6 377.8 371.3 368.3	183.6	77.8	171.3	68.3	376.5 37	375.4 3	377.8 387.8		383.2	1.4	-0.6 ^b	-0.4 ^b	0.4b	oʻq6.0	2012-2017	298
Maryland	381.5 383.9 379.5 375.8 370.0 358.9 356.2	33.9 3	79.5 3.	75.8 3	70.0	58.9	356.2	351.2	344.5	51.2 344.5 331.0 334.2 315.2	34.2 3		307.4	311.2 312.4		315.7	320.6 347.5	47.5 34	349.8	11.0a	-0.5	-0.1	1.6	4.2 ^b	2014-2017	1123
North Carolina	401.2 412.2 398.4 393.4 394.2 386.8 385.6 3	12.2 3	98.4 39	93.4 3	94.2 3	8.98	385.6	385.6	376.5	85.6 376.5 369.1 364.4 358.5	64.4	58.5	357.2 3	360.0 360.7 364.8 368.6 378.6	60.7	64.8	68.6	78.6 38	383.7	7.0a	-0.3	0.0	1.1 ^b	2.0 ^b	2014-2017	1330
South Carolina	470.0 465.9 463.2 454.9 462.1 447.2 445.9	55.9 4	63.2 4	54.9 4.	62.1 4	47.2 4	145.9	145.5 439.3		423.8 4	420.6 428.4 417.9	28.4 4	117.9	417.2 4	420.9 427.5	27.5 4	437.7 441.9	41.9 44	446.0	4.1a	-0.3 ^b	0.1	0.8 ^b	1.6 ^{b,c}	2012-2017	453
Virginia	347.9 352.2 344.5 337.3 338.5 319.9 329.4 3	52.2 3	44.5 33	37.3 3.	38.5 3	19.9	329.4	319.4	19.4 317.4	311.5 3	308.0	9.00	8.00	311.5 308.0 300.6 300.8 299.1 300.7	00.7	07.0	307.0 310.6 317.6	17.6 32	320.4	e.6a	-0.5 ^b	-0.1	0.8 ^b	1.7 ^{b,c}	2012-2017	890
West Virginia	424.4 431.0 440.8 438.4 446.1 438.2 444.5	31.0 4	40.8 4.	38.4 4.	46.1 4	38.2 4	444.5 4	155.0 473.1	173.1	460.6 472.1 461.8	172.1 4	61.8 4	193.1 4	493.1 492.0 489.0	89.0 4	499.3 5	518.3 548.7		567.9	23.0a	1.7 ^b	2.0 ^b	2.6 ^b	4.7b,c	2014-2017	1023
Division 6: East South Central 457.0 458.9 461.2 467.2 472.4 463.3 469.6	1 457.0 45	58.9 4	61.2 40	57.2 4	72.4 4	63.3 4	169.6 4	467.5 455.2	155.2	156.3 4	158.6	57.0	162.5 4	456.3 458.6 457.0 462.5 466.4 467.4 478.0 486.5 496.3	67.4 4.	78.0 4	86.5 4	96.3 49	497.3	8.8 _a	0.5 ^b	9.0	1.3b	1.3b,c	2010-2017	3991
Alabama	469.7 467.8 470.3 471.0 482.9 478.6 485.3	57.8 4	70.3 4.	71.0 4.	82.9 4	78.6	485.3 4	187.3 472.1		478.5 4	475.0 468.5		173.5 4	473.5 479.1 483.3		482.2 4	493.9 501.9		497.4	6.2a	0.4 ^b	0.3 ^b	1.0 ^b	oʻq6.0	2010-2017	729
Kentucky	425.5 <u>424.9</u> 439.2 443.7 442.4 429.2 434.5	24.9 4	39.2 4	43.7 4.	42.4 4	29.2	134.5	130.0	128.2	129.9 4	148.5 4	45.9	149.7 4	30.0 428.2 429.9 448.5 445.9 449.7 462.9 448.8 470.8 490.4 510.3	48.8 4	70.8 4	90.4 5	10.3 51	511.7	14.7a	1.2 ^b	1.6 ^b	2.2 ^b	3.1 _{b,c}	2013-2017	1524
Mississippi	516.9 519.0 512.6 521.5 518.4 515.8 531.0	19.0	12.6 5	21.5 5	18.4 5	15.8 5	531.0	527.2 506.0		508.1 492.1	92.1	497.6 5	05.2 4	505.2 499.7 517.7	17.7 5.	25.2 5	525.2 525.3 532.7		529.2	6.3 _a	0.2	0.2	1.1 ^b	1.1^{b}	2009-2017	482
Tennessee	441.8 448.1 445.8 456.0 464.4 451.5 454.3	48.1 4	45.8 4.	56.0 4	64.4	51.5	154.3	151.8	138.5	51.8 438.5 434.7 438.4 437.8	138.4 4	137.8 4	144.3 4	444.3 445.0 446.4 459.0 461.6 467.3	46.4 4.	59.0 4	61.6 4		474.2	8.3 _a	0.4 ^b	0.4 ^b	1.1 ^b	1.1 _{b,c}	2009-2017	1257

during which mortality rates decreased (statistically significant negative APC); brown shading, years of increasing mortality (statistically significant positive APC); no shading, periods when the APC did not differ significantly from Regression Program (see Methods and Supplement). Slopes include the average annual percent change (AAPC) for 3 periods—1999-2017, 2005-2017, and 2010-2017—and the annual percent change (APC) for the most recent linear trend in the joinpoint model. Also shown are the estimated number of excess deaths in the United States caused by year-to-year changes in midlife mortality rates between 1999 and 2017. Blue shading indicates years Age-adjusted, all-cause mortality rates (per 100 000) among US adults ages 25-64 years for 1999-2017, along with the relative increase in mortality rates between 1999 and 2017 and the slopes modeled by the Joinpoint

zero; black cells, joinpoint years, when changes occurred in the modeled linear trends. Underlined mortality rates denote the lowest mortality rates (nadir) for 1999-2017. NA indicates not applicable (the Joinpoint program plotted a single trend line for 1999-2017; thus, no last segment).

 $^{^{\}rm a}$ Statistically significant (P < .05).

 $^{^{\}rm b}$ Slope (APC or AAPC) that differed significantly from zero (P < .05)

^c Statistically significant slope change estimate, a measure of the change in slope from that of the previous period; see e Table 7 in the Supplement for 95% confidence intervals. Mortality rates obtained from CDC WONDER. ²⁰

Decreasing mortality (P <.05)

Figure 11. Age-Adjusted All-Cause Mortality Rates, Adults Aged 25-64 Years, for US Census Divisions 7-9 (West South Central, Mountain, Pacific) and States (1999-2017)

																			Relative	_	,	()	_		Excess
						Age-	Age-Adjusted	i Mortal	ity Rate	Mortality Rates (Deaths per 100000), by Year	is per 10	(0000)	, by Yea						Increase	٠,	Ē.	ds (AAPC)		-	Deaths
US Census Divisions and States	1999	2000	1999 2000 2001 2002 2003 2004 2005	2002	2003	2004 2		2006 2	2007 2	2008 20	2009 20	2010 20	2011 20	2012 20	2013 20	2014 20	2015 20	2016 2017	— (%, 201 <i>)</i> 7 vs 2010)	7 1999 7 2017	. 2005- 2017	2010- 2017		Recent Trend (APC, y of Last Segment)	(2010- 2017)
Division 7: West South Central 398.5 405.9 404.5 406.6 403.4 396.1 398.6 3	398.5	405.9	404.5	06.6	03.4 3	96.1	398.6	89.8 387.0	87.0 3	388.5 37	379.9 373.0 370.3	73.0 37		372.4 377.0	7.0 380	380.9 378.3		378.3 380.1	1 1.9 ^a	-0.3b	-0.3 ^b	0.2	0.4b,c	2011-2017	1910
Arkansas	439.5	451.6	439.5 451.6 447.0 461.8 457.5 461.5 457.0	61.8 4	57.5	61.5 4	4	60.8 4	56.4 4	60.8 456.4 473.7 463.7 449.6 457.5	53.7 44	19.6	57.5 46	1.4 46	461.4 462.1 475.2	5.2 47	479.5 485.1	5.1 486.8	8 8.3 _a	0.5	0.5	1.0 ^b	1.3 ^b	2011-2017	563
Louisiana	481.0	491.4	481.0 491.4 494.2 487.4 497.1 489.7 509.5	87.4 4	97.1	89.7	4	89.4 477.9	77.9 4	472.0 460.1 453.8 <u>442.4</u> 453.9	50.1 45	3.8	12.4	3.9 46	468.4 474.0 472.1	4.0 47.		475.1 481.8	8 6.2 ^a	-0.1	-0.4	8.0	0.3	2014-2017	681
Oklahoma	431.1	443.9	431.1 443.9 430.4 452.8 450.9 446.8 463.0	52.8 4	50.9 4	46.8 4	4	50.1 4	52.9 4	50.1 452.9 472.9 461.3 458.4 461.4 462.5 477.8	51.3 45	18.4 46	51.4 46	12.5 47	7.8 47.	7.7 48.	2.7 48.	477.7 482.7 481.2 480.7	7 4.9a	9.0	0.6 ^b	0.6 ^b	0.6 ^b	NA	438
Texas	368.7	373.9	368.7 373.9 375.0 374.1 368.5 359.7 357.5	74.1 3	68.5	59.7	~	52.1 350.9	50.9	349.2 34	342.3 33	15.8 3	33.0	335.8 <u>333.0</u> 334.0 335.8	5.8 33	339.7 33	335.7 335	335.6 337.4	4 0.5	-0.5 ^b	-0.5 ^b	0.0	0.2€	2011-2017	228
Division 8: Mountain	343.6	340.2	343.6 340.2 342.6 344.4 347.7 334.4 337.3	44.4	47.7	34.4 3	~	32.8 3	28.2 3	32.8 328.2 324.0 324.4 <u>316.2</u>	24.4 31		21.7 32	3.0 32	321.7 323.0 325.4 328.5 334.0 339.9	3.5 33	4.0 33	9.9 335.3	3 6.0a	0.0	0.1	0.9 ^b	0.9b,c	2010-2017	2372
Arizona	374.8	369.4	374.8 369.4 365.5 372.5 367.7 356.1 363.9	72.5	67.7	56.1 3	3	58.4 343.3	43.3 3	333.5 333.5	33.5 32	329.8 32	329.6 336.3	6.3 34	343.0 338	338.1 346	346.8 355	355.4 350.	.4 6.2 ^a	-0.3	-0.2	1.0 ^b	1.0 ^{b,c}	2009-2017	691
Colorado	297.0	295.5	297.0 295.5 296.9 300.2 301.8 289.6 285.4	00.2	01.8	89.6	7	79.4	81.6 2	83.7 28	35.1 27	71.8 28	32.9 28	0.2 27	5.0 28.	2.9 28.	1.2 29.	79.4 281.6 283.7 285.1 271.8 282.9 280.2 275.0 282.9 281.2 291.7 288.0	0 6.0 ^a	-0.1	0.1	9.0	1.1	2013-2017	469
Idaho	300.0	299.4	300.0 299.4 304.8 302.6 316.6 299.0 292.3	02.6	16.6 2	99.0	10	93.1 285.6	85.6 2	287.9 30	304.5 29	291.1 29	294.7 29	290.4 315.5	5.5 30	304.7 310	310.6 307	307.1 307.2	2 5.5	0.3	0.5	0.7b	0.7b	2006-2017	129
Montana	330.1	330.2	330.1 330.2 337.5 351.3 349.2 326.4 344.7	51.3 3	49.2	126.4 3	3	32.8 3	32.9 3	32.8 332.9 333.4 345.9	15.9 32	3.9 3	12.3 34	323.9 342.3 343.4 348.2	8.2 33.	337.7 380.7	0.7 36.	361.5 355.6	9.8 _a	0.4 ^b	0.7 ^b	1.3 ^b	1.3b,c	2010-2017	166
Nevada	415.3	418.4	415.3 418.4 404.8 403.8 412.7 390.2 407.4	03.8 4	12.7	90.2	ന	86.0 381.2	81.2 3	374.3 370.4 361.9	70.4 36	51.9 3t	365.3 36	0.1 36	360.1 365.0 361.8 365.8	1.8 36		371.9 367.2	2 1.5	-0.7 ^b	-0.6 ^b	0.3	0.3€	2010-2017	79
New Mexico	371.5	362.3	371.5 362.3 384.5 386.3 388.7 379.0 379.5	86.3	88.7	179.0	\sim	89.6	85.3 3	96.8 3	35.6 37	74.4 3	90.6	96.8 39	89.6 385.3 396.8 385.6 374.4 390.6 396.8 395.6 424.5 427.5 443.2	4.5 42	7.5 44.	3.2 439.9	9 17.5a	1.0 ^b	1.3b	2.1 ^b	2.4 _{b,c}	2011-2017	694
Utah	296.7	291.2	296.7 291.2 302.0 287.5 303.8 300.4 284.5	87.5 3	3.83	00.4 2		91.4 2	97.3 2	291.4 297.3 280.8 284.8	34.8 28	280.3	283.7 28	287.1 28	282.6 293.9	3.9 29.	295.9 298	298.4 291.5	5 4.0	0.0	0.2	0.7b	0.9b,c	2011-2017	151
Wyoming	346.3	339.0	346.3 339.0 354.4 348.5 367.1 331.1 349.9	48.5	67.1	31.1	c	52.0 3	57.7 3	48.4 3.	28.9 35	3.4 3	10.2 3	5.5 34	0.7 35	5.8 37	1.2 35.	52.0 357.7 348.4 <u>328.9</u> 353.4 340.2 345.5 340.7 356.8 371.2 353.7 350.7	7 -0.8	0.1	0.1	0.1	0.1	NA	-7
Division 9: Pacific	319.7	316.4	319.7 316.4 316.8 <mark> 313.7 314.6 </mark> 303.7 304.0 2	13.7	14.6	03.7	304.0 2	99.4 2	94.4 2	99.4 294.4 286.3 285.2	35.2 27	272.3 27	74.4 27	3.2 27	274.4 273.2 272.4 272.7 274.5 275.6	2.7 27	4.5 27:	5.6 273.5	5 0.5	-0.8 ^b	-0.8 ^b	-0.1	0.1℃	2011-2017	318
Alaska	340.2	343.3	340.2 343.3 357.3 378.4 340.0 327.4 332.7	78.4 3	40.0	27.4	m	26.5 3	26.5 343.4 335.3		331.9 331.1	11.1 34	16.2 34	5.5 35	346.2 345.5 356.7 361.6 369.8 379.3	1.6 36	9.8 37	9.3 370.8	8 12.0 ^a	9.0	1.3 ^b	1.3 ^b	1.3 ^b	2005-2017	159
California	323.0	319.1	323.0 319.1 319.5 313.1 315.9 304.6 304.4	13.1 3	15.9	04.6	ന	0.00 293.4	93.4 2	283.4 281.0 267.3	31.0 26		58.9 26	7.7 26	268.9 267.7 266.0 263.9 266.2 269.4	3.9 26	5.2 26	9.4 264.9	6.0-	-1.1 ^b	-1.1 ^b	-0.2	-0.2c	2010-2017	-503
Hawaii	302.4	300.8	302.4 300.8 301.0 296.1 285.6 288.1 295.9	96.1 2	85.6	88.1 2	7	93.3 2	81.5 2	93.3 281.5 277.8 287.5		75.6 28	33.2 28	34.3 28	0.1 29t	5.5 28	5.8 281	275.6 283.2 284.3 280.1 296.5 286.8 280.3 296.7	7 7.7a	-0.2	0.0	9.0	0.6℃	2010-2017	156
Oregon	324.6	317.5	324.6 317.5 326.5 329.4 331.9 318.6 320.3	29.4	31.9 3	18.6		16.1 3	16.4 3	09.2 3.	14.3 30	30	0.90	14.6 30	316.1 316.4 309.2 314.3 301.8 306.0 304.6 304.7 313.0 312.2 308.2	3.0 31	2.2 308	3.2 305.5	5 1.2	-0.3	-0.4	-0.1	-1.2	2015-2017	72
Washington	301.5	301.9	301 5 301 9 296 0 304 7 301 3 292 1 291 7	10473	01 3 2	92 1 2	2	86.0 7	85 5 7	86.0 285.5 286.3 285.2 275.9	15 2 27	15 0 7	76 7 370	77	7 0 79.	196 65	5 0 270	1 786 1 976 0 385 0 385 0 376 1 387 1	1 4 1a	-0 4a	-0.2	0 3	0 JC	7105-0106	121

Age-adjusted, all-cause mortality rates (per 100 000) among US adults ages 25-64 years for 1999-2017, along with the relative increase in mortality rates between 1999 and 2017 and the slopes modeled by the Joinpoint Regression Program (see Methods and Supplement). Slopes include the average annual percent change (AAPC) for 3 periods—1999-2017, 2005-2017, and 2010-2017—and the annual percent change (AAPC) for the most recent linear trend in the joinpoint model. Also shown are the estimated number of excess deaths in the United States caused by year-to-year changes in middlife mortality rates between 1999 and 2017. Blue shading indicates years during which mortality rates decreased (statistically significant positive APC); no shading, periods when the APC did not differ significantly from

zero; black cells, joinpoint years, when changes occurred in the modeled linear trends. Underlined mortality rates denote the lowest mortality rates (nadir) for 1999-2017. NA indicates not applicable (the Joinpoint program plotted a single trend line for 1999-2017; thus, no last segment).

 $^{^{1}}$ Statistically significant (P < .05).

 $^{^{\}rm b}$ Slope (APC or AAPC) that differed significantly from zero (P < .05)

Statistically significant slope change estimate, a measure of the change in slope from that of the previous period;
 see e Table 7 in the Supplement for 95% confidence intervals. Mortality rates obtained from CDC WONDER. 20

Discussion

US life expectancy increased from 1959 to 2014, but the rate of increase was greatest in 1969-1979 and slowed thereafter, losing pace with other high-income countries, plateauing in 2011, and decreasing after 2014. A major contributor was an increase in all-cause mortality among young and middle-aged adults, which began in 2010, and an increase in cause-specific mortality rates in this midlife age group, which began as early as the 1990s. The increase in causespecific mortality involved deaths from drug overdoses, alcohol abuse, and suicides and from diverse organ system diseases, such as hypertensive diseases and diabetes. Although non-Hispanic white populations experienced the largest absolute number of deaths, all racial groups and the Hispanic population were affected. For certain causes of death (eg, fatal drug overdoses, alcoholic liver disease, and suicide), women experienced larger relative increases in mortality than men, although the absolute mortality rates for these causes were higher in men than women.

By 2010, increases in cause-specific mortality rates at ages 25 to 64 years had reversed years of progress in lowering mortality from other causes (eg, ischemic heart disease, cancer, HIV infection)—and all-cause mortality began increasing. The trend began earlier (eg, the 1990s) in some states and only recently in others (eg, New York, New Jersey). Gaps in life expectancy across states began widening in the 1980s, with substantial divergences occurring in the 1990s. Changes in life expectancy and midlife mortality were greatest in the eastern United States—notably the Ohio Valley, Appalachia, and upper New England—whereas many Pacific states were less affected. The largest relative increases in midlife mortality occurred among adults with less education and in rural areas or other settings with evidence of economic distress or diminished social capital.

Potential Explanations

The increase in midlife mortality after 1999 was greatly influenced by the increase in fatal drug overdoses. Heroin use increased substantially in the 1960s and 1970s, as did crack cocaine abuse in the 1980s, disproportionately affecting (and criminalizing) the black population. 74,75 Mortality from drug overdoses increased exponentially from the 1970s onward.⁷⁶ The sharp increase in overdose deaths that began in the 1990s primarily affected white populations and came in 3 waves: (1) the introduction of OxyContin in 1996 and overuse of prescription opioids, followed by (2) increased heroin use, often by patients who had become addicted to prescription opioids,⁷⁷ and (3) the subsequent emergence of potent synthetic opioids (eg, fentanyl analogues)—the latter triggering a large post-2013 increase in overdose deaths. ^{29,78,79} That white populations first experienced a larger increase in overdose deaths than nonwhite populations may reflect their greater access to health care (and thus prescription drugs). 5,80 That non-Hispanic black and Hispanic populations experienced the largest relative increases in fentanyl deaths after 201181 may explain the retrogression in overdose deaths observed in these groups. 49 Geographic differences in the promotion and distribution of opioids may also explain the concentration of midlife deaths in certain states.82

However, the increase in opioid-related deaths is only part of a more complicated phenomenon and does not fully explain the in-

crease in midlife mortality rates from other causes, such as alcoholic liver disease or suicides (85.2% of which involve firearms or other nonpoisoning methods⁸³). Opioid-related deaths also cannot fully explain the US health disadvantage, which began earlier (in the 1980s) and involved multiple diseases and nondrug injuries.⁵⁻⁷ Two recent studies estimated that drug overdoses accounted for 15% or less of the gap in life expectancy between the United States and other high-income countries in 2013 and 2014, respectively.^{84,85}

The National Research Council examined the US health disadvantage in detail and identified 9 domains in which the United States had poorer health outcomes than other high-income countries: these included not only drug-related deaths but also adverse birth outcomes, injuries and homicides, adolescent pregnancy and sexually transmitted infections, HIV and AIDS, obesity and diabetes, heart disease, chronic lung disease, and disability. Compared with the average mortality rates of 16 other high-income countries, the United States has lower mortality from cancer and cerebrovascular diseases but higher mortality rates from most other major causes of death, including circulatory disorders (eg, ischemic heart and hypertensive diseases), external causes (eg, drug overdoses, suicide, homicide), diabetes, infectious diseases, pregnancy and childbirth, congenital malformations, mental and behavioral disorders, and diseases of the respiratory, nervous, genitourinary, and musculoskeletal systems. 86 According to one estimate, if the slow rate of increase in US life expectancy persists, it will take the United States more than a century to reach the average life expectancy that other high-income countries had achieved by 2016.¹⁰

Tobacco Use and Obesity

Exposure to behavioral risk factors could explain some of these trends. Although tobacco use in the United States has decreased, higher smoking rates in prior decades could have produced delayed effects on current tobacco-related mortality and life expectancy patterns, especially among older adults. ^{6,87,88} For example, a statistical model that accounted for the lag between risk factor exposure and subsequent death estimated that much of the gap in life expectancy at age 50 years that existed in 2003 between the United States and other high-income countries—41% of the gap in men and 78% of the gap in women—was attributable to smoking. ⁸⁹ Smoking explained 50% or more of the geographic differences in mortality within the United States in 2004. ^{88,90} However, it is unclear whether smoking, which has declined in prevalence, continues to have as large a role in current life expectancy patterns or in explaining increases in mortality among younger adults.

The obesity epidemic, a known contributor to the US health disadvantage, 6 could potentially explain an increase in midlife mortality rates for diseases linked to obesity, such as hypertensive heart disease 91 and renal failure. 92 As long ago as 2005, the increasing prevalence of obesity prompted Olshansky et al 93 to predict a forthcoming decrease in US life expectancy. By 2011, Preston et al 94 estimated that increases in obesity had reduced life expectancy at age 40 years by 0.9 years. Elo et al 33 noted that changes in obesity prevalence had the largest correlation with geographic variations in life expectancy of any variable they examined.

However, neither smoking nor obesity can fully explain current mortality patterns, such as those among younger adults and increasing mortality from conditions without known causal links to these risk factors. Suggesting that other factors may be at play,

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Muennig and Glied noted that Australia and other countries with patterns of smoking and obesity similar to those in the United States achieved greater gains in survival between 1975 and 2005.¹³

Deficiencies in Health Care

Deficiencies in the health care system could potentially explain increased mortality from some conditions. Although the US health care system excels on certain measures, countries with higher life expectancy outperform the United States in providing universal access to health care, removing costs as a barrier to care, care coordination, and amenable mortality. 95-97 In a difficult economy that imposes greater costs on patients, 98 adults in midlife may have greater financial barriers to care than children and older adults, who benefit from the Children's Health Insurance Program and Medicare coverage, respectively. 99 Although poor access or deficiencies in quality could introduce mortality risks among patients with existing behavioral health needs or chronic diseases, these factors would not account for the underlying precipitants (eg, suicidality, obesity), which originate outside the clinic. Physicians contributed to the overprescription of opioids, 100 and iatrogenic factors could potentially explain increases in midlife mortality from other causes, but empirical evidence is limited. Nor would systemic deficiencies in the health care system explain why midlife death rates increased for some chronic diseases while decreasing greatly for others (eg. ischemic heart disease, cancer, and HIV infection).

Psychological Distress

Despair has been invoked as a potential cause for the increase in deaths related to drugs, alcohol, and suicide (referred to by some as "deaths of despair"). 31,37,66,101 Some studies suggest that psychological distress, anxiety, and depression have increased in the United States, especially among adolescents and young adults. ^{66,102-108} Psychological distress and mental illness are risk factors for substance abuse and suicides^{83,109,110} and may complicate organ system diseases, as when hopelessness erodes motivation to pursue health care or manage chronic illnesses. 111 Chronic stress has neurobiological and systemic effects on allostatic load and end organs and may increase pain sensitivity (and thus analgesic needs). 112-115 However, the evidence that the prevalence of psychological distress or mental illness increased during the relevant period is inconclusive. Epidemio $logic\,data\,about\,mental\,illness\,have\,methodological\,limitations, {}^{116,117}$ and some surveillance studies report no increase in prevalence rates. 118,119 Moreover, even if the prevalence of certain mental illnesses did increase, a causal link to the full spectrum of midlife mortality deaths has not been established.

Socioeconomic Conditions

2010

Three lines of evidence suggest a potential association between mortality trends and economic conditions, the first being timing. The US health disadvantage and increase in midlife mortality began in the 1980s and 1990s, a period marked by a major transformation in the nation's economy, substantial job losses in manufacturing and other sectors, contraction of the middle class, wage stagnation, and reduced intergenerational mobility. 120-125 Income inequality widened, surpassing levels in other countries, concurrent with the deepening US health disadvantage. 126-132 The second line of evidence concerns affected populations: those most

vulnerable to the new economy (eg, adults with limited education, women) experienced the largest increases in death rates. The third line of evidence is geographic: increases in death rates were concentrated in areas with a history of economic challenges, such as rural US areas^{133,134} and the industrial Midwest, ^{135,136} and were lowest in the Pacific division and populous states with more robust economies (eg, Texas, New York). One theory for the larger life expectancy gains in metropolitan areas is an increase in the population with college degrees. ³³

Socioeconomic pressures and unstable employment could explain some of the observed increases in mortality spanning multiple causes of death. Financial hardship and insecurity limit access to health care and the social determinants of health (eg, education, food, housing, transportation) and increase the risk of chronic stress, disease, disability, pessimism, and pain. 101,137-144 One study estimated that a 1% increase in county unemployment rates was associated with a 3.6% increase in opioid deaths. 145 However, the evidence to date has not proven that economic conditions are responsible for the recent increase in midlife mortality 146; correlations with state and local indicators (eg, employment, poverty rates) are not always consistent, and the causal link between income inequality and health is debated. 147,148

The causes of economic despair may be more nuanced; perceptions and frustrated expectations may matter as much as absolute income or net worth. 149,150 For example, ethnographers describe the dismay among the working-class white population over their perceived loss of social position and uncertain future, 101,151-155 a popular (but unsubstantiated) thesis for why this historically privileged population experienced larger increases in midlife deaths than did minority groups (eg, non-Hispanic black populations) with greater social and economic disadvantages. 156-159 Also unclear is the extent to which household socioeconomic status acts as a proxy for important contextual conditions in communities (eg, social environment, services infrastructure, economy, labor market) that also shape health. 5,82,150,160-162

The above explanations are not independent and collectively shape mortality patterns; major contributors like smoking, drug abuse, and obesogenic diets are shaped by environmental conditions, psychological distress, and socioeconomic status. The same economic pressures that force patients to forego medical care can also induce stress and unhealthy coping behaviors and can fracture communities. Fenelon, whose research quantified the contribution of smoking to mortality, also noted that smoking may "represent one critical piece of a broader cultural, socioeconomic, and behavioral puzzle that has implications for numerous health-related behaviors and outcomes." ⁹⁰

Methodologic Considerations

Any theory for decreasing life expectancy—whether related to opioids, despair, poverty, or social division—must account for the timing of exposure and lagged effects on outcomes. Whereas observed increases in mortality could occur shortly after increased exposure to certain causes, such as fentanyl or lethal firearms, increases in premature mortality from chronic conditions may require decades of prolonged exposure (eTable 8 in the Supplement). Some mortality patterns exhibit period effects, such as the increase in opioid deaths that began in the 1990s, and affected multiple age cohorts, whereas other causes show

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cohort-based variation. For example, Masters et al identified a specific cohort—non-Hispanic white individuals born in the 1950s—at heightened risk of midlife mortality from obesity, heart disease, diabetes, and hypertension.³⁰ Zang et al documented a heightened mortality risk among cohorts born during 1973-1991.¹⁶³

Any theory for decreasing US life expectancy must explain why this trend is less pronounced in other industrialized countries. ¹⁰ A National Research Council panel, charged with this question, focused its research on how the United States differs in terms of health care, unhealthy behaviors, socioeconomic factors, the physical and social environment, and public policies and priorities. ⁷ Social protection policies deserve special attention: countries with higher life expectancy spend more of their budgets on social services ^{144,164} and outperform the United States in terms of education, child poverty, and other measures of well-being. ^{5,7}

Causal theories must also explain why US mortality trends have affected some states (and counties) more than others and why their trajectories often diverged in the 1990s. The causes of geographical disparities may be compositional, as when states became more populated by people with risk factors for midlife mortality (eg, rural white individuals with limited education) or by large, growing cities that skew state averages. State statistics are also influenced by demographic shifts (eg, immigration, depopulation, and in-migration) and economic trends. For example, the divergence in life expectancy between Oklahoma and New York (Figure 7) may reflect the fate of different economies, one reliant on agriculture and mining and the other on service industries (eg, finance, information technology). The clustering of midlife deaths in certain states, such as recent increases in upper New England states or rural areas, may reflect differences in drug abuse rates and in the distribution and marketing of illicit drugs. 29,82,146,165-167

To some extent, however, divergent state health trajectories may reflect different policy choices. 168 Policy differences seem more likely to explain disparities between adjacent states (eg, Colorado/Kansas, Alabama/Georgia) (Figure 7), where marked regional differences in demography or economies are uncommon. Many states diverged in the 1990s, soon after neoliberal policies aimed at free markets and devolution shifted resources (eg, block grants) and authorities to the states. 121,169-171 States enacted different policies on the social determinants of health, such as education spending, minimum wage laws, earned income tax credits, economic development, mass transit, safety net services, and public health provisions (eg, tobacco taxes, Medicaid expansion, preemption laws, gun control). 172-177 These decisions may have had health implications.¹⁷⁸ For example, Dow et al found that changes in state policies on minimum wage and earned income tax credits predicted nondrug suicide trends.¹⁷⁹ In this study, the 5 states that experienced stable or reduced rates of firearm-related suicides during 1999-2017, countering the national trend, were those with stricter gun control laws. 180

Research and Policy Considerations

Moving from speculation to evidence about root causes will require innovative research methods, including cohort studies, multivariate modeling, investigation of migration effects, and the application of machine learning to historical data sets. Fully understanding the timing of US mortality trends will also require

interdisciplinary research involving epidemiology, demography, sociology, political science, history, economics, and the law. Clarifying the role of state policies may be especially important, given the divergent state trajectories reported here.

The implications of increasing midlife mortality are broad, affecting working-age adults and thus employers, the economy, health care, and national security. The trends also affect children, whose parents are more likely to die in midlife and whose own health could be at risk when they reach that age, or sooner. Recent data suggest that all-cause mortality rates are increasing among those aged 15 to 19 years and 20 to 24 years (increasing from 44.8 deaths/100 000 to 51.5 deaths per 100 000 and from 83.4 deaths/100 000 to 95.6 deaths/100 000, respectively, during 2013-2017) (Figure 2). Evidence-based strategies to improve population health seem warranted, such as policies to promote education, increase household income, invest in communities, and expand access to health care, affordable housing, and transportation. 181-185 The increase in mortality from substance abuse, suicides, and organ system diseases argues for strengthening of behavioral health services and the capacity of health systems to manage chronic diseases. 186

Limitations

This review and analysis have several limitations. First, mortality data are subject to errors, among them inaccurate ascertainment of cause of death, race misclassification and undercounting, and numerator-denominator mismatching. 187,188 These are especially problematic in interpreting mortality rates in the American Indian and Alaska Native population, although disparities persist in this population even in studies that circumvent these challenges. 189 Other limitations include the weak statistical power of annual state mortality rates and their inability to account for substate variation, the limits of age adjustment, age-aggregation bias, and the omission of cause-specific mortality data from before 1999.¹⁹⁰ Purported rate increases may also reflect lagged selection bias. 191 Second, errors in coding, such as the misclassification of suicides as overdoses, 192 or changes (or geographic differences) in coding practices could also introduce errors. For example, some increases in maternal mortality rates may reflect heightened surveillance and the addition of a pregnancy checkbox on death certificates. 193-195 Changes in coding or awareness may partly explain the increase in age-adjusted mortality rates from mental and nervous system disorders, an international trend. 196 Third, state mortality rates may also reflect demographic changes, such as immigration patterns (and the immigrant paradox¹⁹⁷⁻¹⁹⁹) or the out-migration of highly educated, healthy individuals.⁵

Conclusions

US life expectancy increased for most of the past 60 years, but the rate of increase slowed over time and life expectancy decreased after 2014. A major contributor has been an increase in mortality from specific causes (eg, drug overdoses, suicides, organ system diseases) among young and middle-aged adults of all racial groups, with an onset as early as the 1990s and with the largest relative increases occurring in the Ohio Valley and New England. The implications for public health and the economy are substantial, making it vital to understand the underlying causes.

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