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Adult Bone Strength of Children from Single-Parent Families: The Midlife in the U.S. Study

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SSM takes responsibility for the integrity of data analysis.

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Abstract

Purpose—Because peak bone mass is acquired during childhood, bone health may be negatively impacted by childhood socio-environmental disadvantage. The goal of this study was to determine whether being raised in a single-parent household is associated with lower bone strength in adulthood.

Methods—Using dual-energy x-ray absorptiometry data from 708 participants (mean age 57 years) in the Midlife in the United States Biomarker Project, we examined the independent associations of composite indices of femoral neck bone strength relative to load (in three failure modes: compression, bending, and impact) in adulthood with the experience of single-parent childhood and parental death or divorce in childhood.

Results—After adjustment for gender, race, menopause transition stage, age, and body mass index, each additional year of single-parent childhood was associated with 0.02 to 0.03 SD lower indices of adult femoral neck strength. In those with 9-16 years of single-parent childhood, the compression strength index was 0.41 SD lower, bending strength index was 0.31 SD lower, and impact strength index was 0.25 SD lower (all *p*-values < 0.05). In contrast, parental death or divorce during childhood was not by itself independently associated with adult bone strength indices. The magnitudes of these associations were unaltered by additional adjustment for lifestyle factors and socioeconomic status in childhood and adulthood.

Conclusions—Independent of parental death or divorce, growing up in a single-parent household is associated with lower femoral neck bone strength in adulthood, and this association is not entirely explained by childhood or adult socioeconomic conditions or lifestyle choices.

Keywords

single-parent childhood; bone strength; parental divorce; parental death; composite strength indices; strength relative to load

Introduction

Bone strength in older ages is critically dependent on bone acquisition in the growing yearsthe higher the peak bone mass achieved, the lower the likelihood of developing osteoporosis later in life [1-3]. Many environmental factors influence bone accrual during childhood and adolescence [4]. Infancy and childhood are also critical periods that are characterized by increased vulnerability to stressors [5-7]. Exposure to early life stressors has been linked both with poor health and with problematic health behaviors, such as smoking [8, 9], greater alcohol consumption [8, 10], and lower levels of physical activity [8], that are deleterious to bone mass.

However, little is known about how exposure to childhood stress affects adult bone health. We recently reported that childhood socioeconomic disadvantage is associated with lower adult hip strength relative to load in men [11] and with lower adult lumbar spine bone mineral density (BMD) in both men and women [12]. To explore further the potential link between childhood stressors and bone health, we postulated that being raised by a single parent and experiencing parental death or divorce would also be associated with lower adult

bone strength. Indeed, the experience of a parental death or divorce during childhood has been linked with poor mental and physical health (e.g. stroke, smoking- and alcohol-related cancers, obesity, disabling conditions) and decreased survival in adulthood [9, 13-32]. Parental separation during childhood is also associated with several deleterious health behaviors, including smoking [9, 33, 34] and problem drinking [21, 33, 35-37].

However, it is not clear whether links between parental death or divorce and adult health are related to not having two parents for most of one's childhood, or to the experience of parental separation itself. Indeed, the absence of a father figure is associated with earlier menarche in girls [38, 39], and children in single-parent families have poorer physical and psychological health than do children in two-parent families [40-43], a difference that persists into midlife and early old age [44], and that may even decrease life expectancy [45]. Especially germane to the research question posed here is that children in single-parent families have lower bone age [46].

Childhood stressors can also affect body size [47-51], and therefore bone size. Larger bones are stronger, but bone strength has to be commensurate with the load to which it is exposed in a fall (which also increases with body size)[52]. Composite indices of femoral neck bone strength relative to load integrate femoral neck size and BMD with body size to capture their combined influence on fracture risk, and are inversely associated with incident hip fracture risk [53, 54]. These indices are robust indicators of resistance to fracture, and unlike BMD, correctly stratify risk across ethnic groups [55, 56] and between adults with and without diabetes [57].

We hypothesized that being raised by a single parent for most of one's childhood and experiencing parental divorce or death during childhood would each be independently associated with lower adult femoral neck strength relative to load. To test this hypothesis, we analyzed data from The Midlife in the U.S. (MIDUS) National Study of Health and Well-Being [58-60].

Methods

Participants

The Midlife in the U.S. (MIDUS) National Study of Health and Well-Being

We used data from MIDUS [58-60]. The MIDUS study recruited a national sample of adults between ages 25 and 75 residing in the coterminous United States in 1995-1996 and reinterviewed them 9-10 years later (MIDUS II). Details regarding study design, recruitment, and retention are available at http://www.icpsr.umich.edu/icpsrweb/NACDA/. Of 3191 MIDUS II participants who were medically able to travel to the research sites, 1255 consented to participate in the MIDUS II biomarker project. The MIDUS II biomarker project (data collection occurred between July 2004 and May 2009) entailed travel to one of the three clinical research centers: University of California at Los Angeles, Georgetown University, and University of Wisconsin. Reasons given for nonparticipation were travel burden, family obligations, and being too busy. Via self-administered questionnaires and interviews, we obtained medical history information. Using standardized protocols, body

weight and height were measured for calculation of body mass index (BMI, body weight in kilograms divided by the square of body height in meters). Informed consent was provided by each participant. Each MIDUS center obtained institutional review board approval.

The characteristics of the MIDUS II participants were similar to those of the MIDUS I participants [60], and the MIDUS biomarker project participants (e.g. subjective health status, chronic health conditions, exercise, alcohol use) were similar to those of the MIDUS II participants as a whole [59].

Of the 1255 participants in the MIDUS II Biomarkers Project, we excluded data from 348 participants without BMD measurement (which was added to the Biomarkers Project partway into data collection), 94 participants who reported the use of medications known to influence BMD (oral corticosteroids, alendronate, anastrozole, calcitonin, ibandronate, leuprolide, letrozole, raloxifene, risedronate, tamoxifen, zoledronic acid, testosterone, finasteride, dutasteride), 88 female participants whose menopause transition stage was unclassifiable, 11 participants for whom we lacked complete information regarding education and/or childhood socioeconomic status, 2 participants with BMI values >60 kg/m², and 3 participants for whom we lacked information regarding parental death and divorce. One additional participant reported experiencing parental divorce and being raised by a single parent, but did not report the age at which the parental divorce occurred, resulting in an analytic sample of 708 participants.

Outcomes

Bone strength

According to standardized protocols, trained technologists performed DXA scans using GE Healthcare Lunar Prodigy (U. Wisconsin - Madison) or Hologic 4500 (UCLA and Georgetown U.) densitometers. Phantoms were scanned on all densitometers three times per week, and on all days on which scans were obtained. No densitometer shift or drift occurred during the course of this study. BMD and bone size measurements from all DXA scans were adjudicated centrally at the University of Wisconsin DXA center using manufacturerprovided software (GE Lunar, Inc. and Hologic, Inc.).

On the DXA scans, we measured the 2-dimensional (2D) projected areal BMD in the femoral neck, the femoral neck axis length (FNAL) - the distance on the 2D projected plane along the femoral neck axis from the lateral margin of the base of the greater trochanter to the apex of the femoral head, and the femoral neck width (FNW) - the smallest thickness of the femoral neck on the 2D projected plane along a line perpendicular to the femoral neck axis. Using those DXA-based measurements, and measured body height and body weight, we calculated composite indices of femoral neck strength that index bone strength relative to the load during a fall using the following formulas [11, 53], which have been validated against 3-dimensional (3D) methods based on quantitative computed tomography [61]:

 $\begin{array}{l} \text{Compression Strength Index} = \frac{BMD^*FNW}{\text{Weight}} \text{ Bending Strength Index} = \frac{BMD^*FNW^2}{FNAL^*\text{Weight}} \\ \text{Impact Strength Index} = \frac{BMD^*FNW^*FNAL}{\text{Height}^*\text{Weight}} \end{array}$

The compression strength index reflects the ability of the femoral neck to withstand axial compressive loading, the bending strength index reflects its ability to withstand bending, and the impact strength index reflects its ability to absorb the energy of impact in a fall from standing height.

Predictors

Childhood family environment

Information regarding childhood family structure and stability was self-reported in MIDUS I (1995-1996). Participants were asked whether they had lived with both biological parents for most of their childhood until the age of 16. If the response was negative, they were asked the reason (mother died, father died, parents separated/divorced, parents never lived together, adopted). If the reason provided was death or divorce/separation of biological parents or adoption, they were further asked their age when this occurred.

Participants were then asked who was the female head of household for most of their childhood until the age of 16 (biological mother, adoptive mother, stepmother, other female, no female in household) and who was the male head of household for most of their childhood until the age of 16 (biological father, adoptive father, stepfather, other male, no male in household). Participants who reported having either no male head of household or no female head of household for most of their childhood were identified as having a single-parent childhood. For each participant, we calculated the number of years in a single-parent household before reaching age 16 years by subtracting the age (in years) at which parental death or divorce occurred from 16. We assigned a value of 0 to participants who did not have a single-parent childhood, and a value of 16 to those who reported living in a single-parent household without having experienced a parental death or divorce.

Race/ethnicity

Race/ethnicity was self-identified as white, black/African American, other, or multiracial. For this analysis, we classified race as black vs. not black; the latter group was mostly white, but included a small number (n = 31, 4.4%) of participants who reported being neither white nor black/African American. This latter group included participants who reported being Asian, Native American/Pacific Islander, multiracial, or "other".

Menopausal transition stage and age

Menopausal staging in women was based on self-reported menstrual patterns, self-reported use of sex steroid hormones in the past year, and examination of medication bottles brought to the clinical research center. We defined the following menopause transition stage categories: premenopausal (no change in regularity of menses), early perimenopausal (menses in last 3 months with change in regularity of menses), late perimenopausal (last menses 3-12 months previously with change in regularity of menses), postmenopausal (no menses in prior 12 months) not taking menopausal hormone therapy, and postmenopausal taking menopausal women, we combined them with the postmenopausal women not taking menopausal hormone therapy.

Men were categorized by age. The choice of age categories in men was based on previous observations that significant age-related bone loss in men does not start until age 50 years [62], and to age-match the oldest group to the post-menopausal women, because only 0.3% of occurrences of spontaneous menopause take place at or after 59 years of age [63].

In addition to these categories, in order to control for declines in bone strength with aging in later years, age was also included as a continuous variable (in whole years) only in men 60 years and older and in late peri/post-menopausal women not taking menopausal hormone therapy.

Socioeconomic status

Socioeconomic status measures included childhood socioeconomic advantage, education and adult financial advantage. These classifications were based on the categories used in our recent study examining BMD and socioeconomic advantage [12]. Childhood socioeconomic advantage score was calculated by summing three components (possible range 0-6): being on welfare during childhood (0: yes, 2: no), childhood financial level relative to others (0: worse off, 1: same, 2: better), and highest parental education level (0: < high school, 1: high school/general educational development [GED] certificate, 2: some college or more). Scores were calculated only for participants who supplied data regarding at least 2 of the 3 components; the missing component was imputed as the rounded mean of the other two components for 47 participants. The participant's reported educational level was collapsed to a 3-category variable: 1) no college, 2) some college or Associate's degree, or 3) Bachelor's degree or more.

To calculate the adult financial advantage score, we first determined the family-adjusted poverty-to-income ratio (FPIR) for each participant as the ratio of the participant's total household income (sum of self-reported earnings, pension, social security, and government assistance for all household members) to the U.S. Census Bureau poverty threshold specific to the participant's age, presence of a spouse or partner in the household, the number of children under age 18 living in the household, and year of data collection. An FPIR of 3 would correspond to a total household income 3 times the Census Bureau-defined poverty level for his/her family. Adult financial advantage score was calculated by summing 4 components (possible range 0-8): FPIR (0 for FPIR<3, 1 for FPIR 3 but <6, 2 for FPIR 6, reflecting approximate tertiles of its distribution), self-rated current financial situation (0: worst, 1: average, 2: best), sufficient money to meet needs (0: not enough, 1: just enough, 2: more than enough), and degree of difficulty paying bills (0: very, 1: not very, 2: not at all). Scores were calculated only for participants who supplied data regarding at least 3 of the 4 components; the missing component was imputed as the rounded mean of the other three components for 20 participants.

Lifestyle measures

On self-administered questionnaires, participants were asked to report their alcohol intake levels during the period of life at which their alcohol consumption was highest as well as their alcohol intake within the last month.

Smoking was quantified as total pack-years of cigarette smoking (years smoked regularly multiplied by number of cigarettes per day divided by 20) and also as a binary variable of current smoker versus not current smoker. Participants were also asked at what age they started smoking, and we created a dichotomous variable indicating smoking regularly before age 18 years.

Recalled self-reported physical activity during three life stages was recorded: the number of years of participation in competitive sports and recreational sports (separately) during high school (ages 14-18 years); the number of years of light, moderate, and vigorous exercise (self-categorized) during young adulthood (ages 20-35 years), and the average number of minutes per week currently spent doing light, moderate, and vigorous exercise. For each participants' young adulthood and current levels of physical activity, we created summary scores as weighted sums of reported times for light (weight of 1), moderate (weight of 2), and vigorous (weight of 3) activity. Construct validity of summary scores based on recalled self-reports of physical activity has been previously reported [64, 65].

Statistical Analysis

LOESS plots of each of the three indices of femoral neck strength relative to load as a function of the number of childhood years in a single parent household showed that bone strength stayed relatively stable up to about 8 years of exposure to a single parent household, and then trended down steeply (Figure 1). Therefore, we modeled years of single-parent childhood both as a continuous predictor and as a categorical predictor (0, 1-8 years, 9-16 years). We also included a binary indicator for parental death or divorce before age 16 as a second exposure to test whether the experience of parental death or divorce had effects that were independent of subsequently being raised by a single parent. The reference group for this exposure was stable (i.e. continuous), two-parent childhoods, which included those who had either lived with both biological parents until age 16 or were adopted at birth.

The three composite indices of adult femoral neck strength relative to load (bending, compression, and impact) were the dependent variables in three separate linear regression models that included adjustment for gender, race (black vs. non-black), study site, menopausal transition stage (in women), BMI (continuous linear plus quadratic terms), gender by BMI interaction (to model gender-specific effects of BMI on bone strength), and age. Age was operationalized with two continuous variables to capture age-related declines in older adults (one that tracked age in men 60 years and older, and one that tracked age in late perimenopausal and postmenopausal women not taking menopausal hormones), and a 3-category age variable (<50 years, 50-59 years, and 60 years) in men only, to capture age/ cohort differences in younger men. In younger women, age was highly collinear with menopausal transition stage, and was therefore not included separately.

To test whether socioeconomic disadvantage and unhealthy lifestyles accounted for the associations of single parenting and parental death or divorce in childhood with adult bone strength, we added the following potential explanatory variables to the regression models in a second step: childhood socioeconomic advantage score, number of years in recreational sports during ages 14-18 years, number of years in competitive sports during ages 14-18 years, physical activity summary score for ages 20-35 years, smoking initiation at age 18

years, heavy alcohol intake at time of peak alcohol consumption, participant education level, adult financial advantage score, current physical activity score, current smoking, pack-years of smoking, and heavy alcohol intake in the past month. Women reporting any of the following alcohol intake patterns were classified as heavy drinkers: 8 drinks per week, >3 drinks per day on average, or >4 drinks on a single occasion. Men reporting any of the following alcohol intake patterns were classified as heavy drinkers: 15 drinks per week, >4 drinks per day on average, or >4 drinks on a single occasion. These thresholds were taken from the National Institute on Alcohol Abuse and Alcoholism [66].

To test whether associations varied by gender or menopausal transition stage, we added an interaction term between the primary predictors (years of single-parent childhood-continuous, and parental death or divorce experienced before age 16-yes/no) and gender/menopause status (pre-/early peri-menopausal women, late peri-/post-menopausal women, men [reference]) to the regression models.

Results

The characteristics of the analytic sample were similar to those of the overall MIDUS II biomarker project (Table 1). The Madison, WI study site, which had a larger proportion of black participants by design, began DXA scanning prior to the other study sites; thus, the current study sample had a larger proportion of blacks compared with the biomarker project sample. On average, participants were aged 56.7 years; 49% of the analytic sample participants were male, and 24% were black. The majority (72.5%) of participants had a stable two-parent family until age 16 (i.e. they either lived with biological parents until age 16 or were adopted at birth), but 22.7% experienced a parental death or divorce before they were 16 years-old. Only 8.1% reported having a single parent for most of their childhood, 2.0% for 1-8 years, and 6.1% for 9-16 years.

Adjusted associations between childhood family environment and adult bone strength indices

In linear regression models adjusted for race, site, menopausal stage, age, gender, and BMI, and duration of single-parent childhood as a continuous predictor, a longer duration of single-parent childhood was associated with lower indices of femoral neck strength relative to load (Table 2, **Model 1**). For each additional year of single-parent childhood, femoral neck compression strength index was 0.029 SD lower (p < 0.05), bending strength index was 0.021 SD lower (p < 0.05), and impact strength index was 0.017 SD lower (p-value = 0.07). The magnitudes of these associations were similar after further adjustment for childhood and adult socioeconomic status and childhood and adult lifestyle factors (Table 2, **Model 2**).

However, in linear regression with duration of single-parent childhood as a categorical predictor, we found that the associations (adjusted as before for race, site, menopausal stage, age, gender, and BMI) were primarily driven by those who reported 9 or more years of single-parent childhood (Table 3, **Model 1**). Compared with not living in a single parent household before age 16, living in a single parent household for 9-16 years was associated with 0.401 SD lower compression strength index, 0.307 SD lower bending strength index,

and -0.254 SD lower impact strength index (all *p*-values <0.05). The magnitudes of these associations were similar after further adjustment for childhood and adult socioeconomic status and childhood and adult lifestyle factors (Table 3, **Model 2**).

In contrast, parental death or divorce itself was not independently associated with adult bone strength, suggesting that the chronic experience of residing in a single-parent family rather than the acute event of parental marital dissolution contributes to decreased adult bone strength.

Interaction testing revealed that associations of bone strength indices with parental death or divorce prior to age 16 and of the number of childhood years in a single parent did not vary by gender or menopausal transition stage (p-values 0.11 to 0.93, model 1).

Discussion

Independent of parental divorce or parental death, growing up in a single-parent household was associated with lower bone strength in adulthood. There was a strong inverse association between the number of childhood years in a single-parent household and all three indices of adult femoral neck strength relative to load. These associations were not explained by childhood or adult socioeconomic status or by health behaviors over the life course, and add to the growing list of adverse sub-clinical and clinical health outcomes that have been linked to childhood psychosocial disadvantage [17, 67-73]. However, the experience of parental death or divorce during childhood was not independently associated with adult bone strength once we accounted for the number of years spent living in a single-parent household, suggesting that the event of parental death or divorce during childhood does not by itself have direct effects on bone health independent of the subsequent chronic exposure to single parenting.

Although previous studies have documented links between childhood socioeconomic exposures and adult bone health [12, 74-76], as far as we are aware, this is the first investigation of the effect of childhood family stability and structure on adult bone strength. Previous studies have shown that each SD increment in the femoral neck composite strength indices was associated with 34%-41% relative decrement in the rate (hazard) of fracture at any site in women going through the menopausal transition [56], and 57%–66% relative decrement in the risk of hip fracture over 10 years in postmenopausal women [53]. If the differences in the composite strength indices seen in this study between single-parent and two-parent childhoods lead to similar fracture risk differences, women who experienced nine or more years of single parenting in childhood would be at 14%-19% relative increase (relative to women who did not have a single parent for most of childhood) in fracture hazard when going through the menopausal transition, and 31%-41% relative increase in 10-year hip fracture risk in the post-menopause. These findings carry important implications for the bone health of future cohorts of adults. In 2012, only 64% of American children resided with both parents [77]. Thus, large and growing numbers of children who spend much of their first 16 years in a single-parent household may be at elevated risk for poor bone health as adults.

We considered several possible reasons for our findings. First, the physical absence of a father or mother figure could influence adult bone health via the adoption of maladaptive health behaviors such as smoking and underage or heavy alcohol intake. Single-parent childhood is associated with increased smoking [34, 78-81] and alcohol intake [79, 81-84] by adolescents; in turn, smoking and alcohol intake have known adverse effects on bone health [85-88]. Second, decreased physical activity is more common in children in single-parent families [89-92], and skeletal growth is negatively influenced by childhood, especially adolescent, physical inactivity [4, 93-97]. Finally, many children living with single parents are economically disadvantaged [40, 98-106], which also has been linked to poor bone health [11, 12, 107, 108]. However, in our study, the associations between the single-parent childhood and lower adult bone strength were not explained by childhood or adult socioeconomic status or health behaviors over the life course.

Independent of socioeconomic and behavioral pathways, the environmental and psychological stresses of growing up with a single parent could directly affect the hormonal milieu in the child, thereby affecting bone accrual. This is supported by the following observations: 1) children living with single parents face more emotional and environmental stressors than children from two-parent families [40, 98]; 2) basal cortisol levels in children vary by the magnitude of environmental stress, and these gradients are stronger in younger (below age 10) than in older children [109]; 3) early life adversity has been linked to dysregulation of the sympathetic nervous system, as well to chronic inflammation [110-113]; 4) the activation of sympathetic nervous signaling by stress and increased levels of inflammatory markers are both associated with low BMD [114-123].

Our study has several strengths, including the focus on the independent effects of family disruption and single parenting, careful attention to the duration of for which a child resided in a single-parent household, the national sample recruited from across the U.S. coterminous states, availability of socioeconomic and lifestyle data over the life course, and the focus on hip strength relative to load, which is better at fracture risk stratification in a diverse population than is BMD on its own.

Limitations of our study include its observational design, which does not allow attribution of causality, and the lack of information on childhood diet and nutrition. Further, our ability to examine possible racial and gender differences in associations between childhood family environment and adult bone strength was limited by the small number of non-white MIDUS participants and the small number of participants with the primary exposures. The number of participants who were raised by a single parent from birth was also very small; this meant that we could not disentangle the effect of long exposure to single parenting from that of experiencing parental separation at a young age, and the findings from this study may not be applicable to the children of mothers who choose to be single parents from the start. Finally, the phrasing of the MIDUS questions about male and female heads of household "for most of their childhood" could have failed to capture short, time-limited exposure to single parenting; such underreporting and the resulting misclassification would have weakened our power to find an effect for 1-8 years of single-parent childhood.

Conclusions

In conclusion, independent of childhood and adult socioeconomic status, being raised in a single-parent household for most of one's childhood was associated with lower levels of femoral neck strength in adulthood, whereas the event of parental death or divorce experienced during childhood was not independently associated with bone strength. Further research is needed to elucidate the mechanisms underlying these associations and, ultimately, to develop targeted strategies aimed at decreasing the fracture burden and other adverse health effects of childhood stresses.

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References

- Hansen MA, Overgaard K, Riis BJ, Christiansen C. Role of peak bone mass and bone loss in postmenopausal osteoporosis: 12 year study. BMJ. 1991; 303:961–964. [PubMed: 1954420]
- Riis BJ, Hansen MA, Jensen AM, Overgaard K, Christiansen C. Low bone mass and fast rate of bone loss at menopause: equal risk factors for future fracture: a 15-year follow-up study. Bone. 1996; 19:9–12. [PubMed: 8830981]
- 3. Javaid MK, Cooper C. Prenatal and childhood influences on osteoporosis. Best practice & research Clinical endocrinology & metabolism. 2002; 16:349–367. [PubMed: 12064897]
- Holroyd C, Harvey N, Dennison E, Cooper C. Epigenetic influences in the developmental origins of osteoporosis. Osteoporos Int. 2012; 23:401–410. [PubMed: 21656266]
- Charmandari E, Achermann JC, Carel JC, Soder O, Chrousos GP. Stress response and child health. Sci Signal. 2012; 5:mr1. [PubMed: 23112343]
- Hallqvist J, Lynch J, Bartley M, Lang T, Blane D. Can we disentangle life course processes of accumulation, critical period and social mobility? An analysis of disadvantaged socio-economic positions and myocardial infarction in the Stockholm Heart Epidemiology Program. Soc Sci Med. 2004; 58:1555–1562. [PubMed: 14759698]
- Ben-Shlomo Y, Kuh D. A life course approach to chronic disease epidemiology: conceptual models, empirical challenges and interdisciplinary perspectives. Int J Epidemiol. 2002; 31:285–293. [PubMed: 11980781]
- van de Mheen H, Stronks K, Looman CW, Mackenbach JP. Does childhood socioeconomic status influence adult health through behavioural factors? Int J Epidemiol. 1998; 27:431–437. [PubMed: 9698131]
- Melchior M, Moffitt TE, Milne BJ, Poulton R, Caspi A. Why do children from socioeconomically disadvantaged families suffer from poor health when they reach adulthood? A life-course study. Am J Epidemiol. 2007; 166:966–974. [PubMed: 17641151]
- Poulton R, Caspi A, Milne BJ, Thomson WM, Taylor A, Sears MR, Moffitt TE. Association between children's experience of socioeconomic disadvantage and adult health: a life-course study. Lancet. 2002; 360:1640–1645. [PubMed: 12457787]
- Karlamangla AS, Mori T, Merkin SS, Seeman TE, Greendale GA, Binkley N, Crandall CJ. Childhood Socioeconomic Status and Adult Femoral Neck Bone Strength: Findings from The Midlife in the United States Study. Bone. (in press).

- Crandall CJ, Merkin SS, Seeman TE, Greendale GA, Binkley N, Karlamangla AS. Socioeconomic status over the life-course and adult bone mineral density: the Midlife in the U.S. Study. Bone. 2012; 51:107–113. [PubMed: 22543227]
- Luecken LJ, Roubinov DS. Pathways to lifespan health following childhood parental death. Soc Personal Psychol Compass. 2012; 6:243–257. [PubMed: 23555319]
- 14. Neeleman J, Sytema S, Wadsworth M. Propensity to psychiatric and somatic ill-health: evidence from a birth cohort. Psychol Med. 2002; 32:793–803. [PubMed: 12171374]
- 15. Agid O, Shapira B, Zislin J, Ritsner M, Hanin B, Murad H, Troudart T, Bloch M, Heresco-Levy U, Lerer B. Environment and vulnerability to major psychiatric illness: a case control study of early parental loss in major depression, bipolar disorder and schizophrenia. Mol Psychiatry. 1999; 4:163–172. [PubMed: 10208448]
- Felitti VJ, Anda RF, Nordenberg D, Williamson DF, Spitz AM, Edwards V, Koss MP, Marks JS. Relationship of childhood abuse and household dysfunction to many of the leading causes of death in adults. The Adverse Childhood Experiences (ACE) Study. Am J Prev Med. 1998; 14:245–258. [PubMed: 9635069]
- 17. Krause N. Early parental loss, recent life events, and changes in health among older adults. J Aging Health. 1998; 10:395–421. [PubMed: 10346692]
- Lowman BC, Drossman DA, Cramer EM, McKee DC. Recollection of childhood events in adults with irritable bowel syndrome. J Clin Gastroenterol. 1987; 9:324–330. [PubMed: 3611687]
- Mustonen U, Huurre T, Kiviruusu O, Haukkala A, Aro H. Long-term impact of parental divorce on intimate relationship quality in adulthood and the mediating role of psychosocial resources. J Fam Psychol. 2011; 25:615–619. [PubMed: 21639631]
- Larson K, Halfon N. Parental divorce and adult longevity. Int J Public Health. 2013; 58:89–97. [PubMed: 22674375]
- Kuh D, Maclean M. Women's childhood experience of parental separation and their subsequent health and socioeconomic status in adulthood. J Biosoc Sci. 1990; 22:121–135. [PubMed: 2298757]
- Angarne-Lindberg T, Wadsby M. Psychiatric and somatic health in relation to experience of parental divorce in childhood. Int J Soc Psychiatry. 2012; 58:16–25. [PubMed: 20851827]
- Huurre T, Junkkari H, Aro H. Long-term psychosocial effects of parental divorce: a follow-up study from adolescence to adulthood. Eur Arch Psychiatry Clin Neurosci. 2006; 256:256–263. [PubMed: 16502211]
- Gilman SE, Kawachi I, Fitzmaurice GM, Buka SL. Family disruption in childhood and risk of adult depression. Am J Psychiatry. 2003; 160:939–946. [PubMed: 12727699]
- Kessler RC, Davis CG, Kendler KS. Childhood adversity and adult psychiatric disorder in the US National Comorbidity Survey. Psychol Med. 1997; 27:1101–1119. [PubMed: 9300515]
- 26. Novak M, Ahlgren C, Hammarstrom A. A life-course approach in explaining social inequity in obesity among young adult men and women. Int J Obes (Lond). 2006; 30:191–200. [PubMed: 16172616]
- Fuller-Thomson E, Dalton AD. Gender differences in the association between parental divorce during childhood and stroke in adulthood: findings from a population-based survey. Int J Stroke. 2012
- Hemminki K, Chen B. Lifestyle and cancer: effect of parental divorce. Eur J Cancer Prev. 2006; 15:524–530. [PubMed: 17106333]
- Blackburn CM, Spencer NJ, Read JM. Is the onset of disabling chronic conditions in later childhood associated with exposure to social disadvantage in earlier childhood? A prospective cohort study using the ONS Longitudinal Study for England and Wales. BMC pediatrics. 2013; 13:101. [PubMed: 23802581]
- Schwartz JE, Friedman HS, Tucker JS, Tomlinson-Keasey C, Wingard DL, Criqui MH. Sociodemographic and psychosocial factors in childhood as predictors of adult mortality. Am J Public Health. 1995; 85:1237–1245. [PubMed: 7661231]
- Martin LR, Friedman HS, Clark KM, Tucker JS. Longevity following the experience of parental divorce. Soc Sci Med. 2005; 61:2177–2189. [PubMed: 15936133]

- Remes HM, Martikainen PT. Living arrangements and external causes of deaths in early adulthood. J Adolesc Health. 2012; 50:164–171. [PubMed: 22265112]
- Wolfinger NH. The effects of parental divorce on adult tobacco and alcohol consumption. J Health Soc Behav. 1998; 39:254–269. [PubMed: 9785697]
- 34. O'Loughlin J, Karp I, Koulis T, Paradis G, Difranza J. Determinants of first puff and daily cigarette smoking in adolescents. Am J Epidemiol. 2009; 170:585–597. [PubMed: 19635735]
- 35. Hope S, Power C, Rodgers B. The relationship between parental separation in childhood and problem drinking in adulthood. Addiction. 1998; 93:505–514. [PubMed: 9684389]
- Umberson D, Williams K, Anderson K. Violent behavior: a measure of emotional upset? J Health Soc Behav. 2002; 43:189–206. [PubMed: 12096699]
- Marks NF, Jun H, Song J. Death of Parents and Adult Psychological and Physical Well-Being: A Prospective U.S. National Study. Journal of family issues. 2007; 28:1611–1638. [PubMed: 19212446]
- 38. Hoier S. Father absence and age at menarche. Hum Nat. 2003; 14:209–233.
- Neberich, Wwnsh-bd; Penke, L.; Lehnart, J.; Asendorpf, JB. Family of origin, age at menarche, and reproductive strategies: A test of four evolutionary-developmental models. European Journal of Developmental Psychology. 2010; 7:153–177.
- 40. Waldfogel J, Craigie TA, Brooks-Gunn J. Fragile families and child wellbeing. The Future of children / Center for the Future of Children, the David and Lucile Packard Foundation. 2010; 20:87–112.
- East L, Jackson D, O'Brien L. Father absence and adolescent development: a review of the literature. Journal of child health care : for professionals working with children in the hospital and community. 2006; 10:283–295. [PubMed: 17101621]
- 42. Avison WR. Single motherhood and mental health: implications for primary prevention. CMAJ: Canadian Medical Association journal = journal de l'Association medicale canadienne. 1997; 156:661–663.
- Kitsantas P, Kornides ML, Cantiello J, Wu H. Chronic physical health conditions among children of different racial/ethnic backgrounds. Public health. 2013; 127:546–553. [PubMed: 23583033]
- 44. Maier E,H, Lachman ME. Consequences of early parental loss and separation for health and wellbeing in midlife. International Journal of Behavioral Development. 2000; 24:183–189.
- Hayward MD, Gorman BK. The long arm of childhood: the influence of early-life social conditions on men's mortality. Demography. 2004; 41:87–107. [PubMed: 15074126]
- Cole TJ, Cole AJ. Bone age, social deprivation, and single parent families. Arch Dis Child. 1992; 67:1281–1285. [PubMed: 1444529]
- Dixon B, Pena MM, Taveras EM. Lifecourse approach to racial/ethnic disparities in childhood obesity. Adv Nutr. 2012; 3:73–82. [PubMed: 22332105]
- Godfrey KM, Inskip HM, Hanson MA. The long-term effects of prenatal development on growth and metabolism. Semin Reprod Med. 2011; 29:257–265. [PubMed: 21769765]
- Gundersen C, Mahatmya D, Garasky S, Lohman B. Linking psychosocial stressors and childhood obesity. Obes Rev. 2011; 12:e54–63. [PubMed: 21054757]
- Batty GD, Shipley MJ, Gunnell D, Huxley R, Kivimaki M, Woodward M, Lee CM, Smith GD. Height, wealth, and health: an overview with new data from three longitudinal studies. Econ Hum Biol. 2009; 7:137–152. [PubMed: 19628438]
- Montgomery SM, Bartley MJ, Wilkinson RG. Family conflict and slow growth. Arch Dis Child. 1997; 77:326–330. [PubMed: 9389237]
- 52. Robinovitch SN, Hayes WC, McMahon TA. Prediction of femoral impact forces in falls on the hip. J Biomech Eng. 1991; 113:366–374. [PubMed: 1762432]
- Karlamangla AS, Barrett-Connor E, Young J, Greendale GA. Hip fracture risk assessment using composite indices of femoral neck strength: the Rancho Bernardo study. Osteoporos Int. 2004; 15:62–70. [PubMed: 14605798]
- Yu N, Liu YJ, Pei Y, et al. Evaluation of compressive strength index of the femoral neck in Caucasians and chinese. Calcif Tissue Int. 2010; 87:324–332. [PubMed: 20814670]

- 55. Ishii S, Cauley JA, Greendale GA, Danielson ME, Safaei Nili N, Karlamangla A. Ethnic differences in composite indices of femoral neck strength. Osteoporos Int. 2012; 23:1381–1390. [PubMed: 21927926]
- 56. Ishii S, Greendale GA, Cauley JA, Crandall CJ, Huang MH, Danielson ME, Karlamangla AS. Fracture risk assessment without race/ethnicity information. J Clin Endocrinol Metab. 2012; 97:3593–3602. [PubMed: 22865903]
- 57. Ishii S, Cauley JA, Crandall CJ, Srikanthan P, Greendale GA, Huang MH, Danielson ME, Karlamangla AS. Diabetes and femoral neck strength: findings from the Hip Strength Across the Menopausal Transition Study. J Clin Endocrinol Metab. 2012; 97:190–197. [PubMed: 22072739]
- 58. Brim, OG.; Ryff, CD.; Kessler, RC. How healthy are we? : a national study of well-being at midlife. University of Chicago Press; Chicago: 2004.
- Dienberg Love G, Seeman TE, Weinstein M, Ryff CD. Bioindicators in the MIDUS national study: protocol, measures, sample, and comparative context. J Aging Health. 2010; 22:1059–1080. [PubMed: 20876364]
- 60. Radler BT, Ryff CD. Who participates? Accounting for longitudinal retention in the MIDUS national study of health and well-being. J Aging Health. 2010; 22:307–331. [PubMed: 20103686]
- Danielson ME, Beck TJ, Karlamangla AS, et al. A comparison of DXA and CT based methods for estimating the strength of the femoral neck in post- menopausal women. Osteoporos Int. 2013; 24:1379–1388. [PubMed: 22810918]
- 62. Riggs BL, Wahner HW, Dunn WL, Mazess RB, Offord KP, Melton LJ 3rd. Differential changes in bone mineral density of the appendicular and axial skeleton with aging: relationship to spinal osteoporosis. J Clin Invest. 1981; 67:328–335. [PubMed: 7462421]
- 63. Treloar AE. Menstrual cyclicity and the pre-menopause. Maturitas. 1981; 3:249–264. [PubMed: 7334935]
- 64. Bernstein L, Patel AV, Ursin G, et al. Lifetime recreational exercise activity and breast cancer risk among black women and white women. J Natl Cancer Inst. 2005; 97:1671–1679. [PubMed: 16288120]
- 65. Friedenreich CM, Courneya KS, Bryant HE. The lifetime total physical activity questionnaire: development and reliability. Med Sci Sports Exerc. 1998; 30:266–274. [PubMed: 9502356]
- 66. National Institute on Alcohol Abuse and Alcoholism. Updated 2007. National Institutes of Health; 2005. Helping patients who drink too much: A clinician's guide.
- Tamayo T, Christian H, Rathmann W. Impact of early psychosocial factors (childhood socioeconomic factors and adversities) on future risk of type 2 diabetes, metabolic disturbances and obesity: a systematic review. BMC Public Health. 2010; 10:525. [PubMed: 20809937]
- Haas S. Trajectories of functional health: the 'long arm' of childhood health and socioeconomic factors. Soc Sci Med. 2008; 66:849–861. [PubMed: 18158208]
- Osler M, Andersen AM, Lund R, Holstein B. Effect of grandparent's and parent's socioeconomic position on mortality among Danish men born in 1953. Eur J Public Health. 2005; 15:647–651. [PubMed: 16093305]
- Galobardes B, Lynch JW, Smith GD. Is the association between childhood socioeconomic circumstances and cause-specific mortality established? Update of a systematic review. J Epidemiol Community Health. 2008; 62:387–390. [PubMed: 18413449]
- Goodman E, McEwen BS, Huang B, Dolan LM, Adler NE. Social inequalities in biomarkers of cardiovascular risk in adolescence. Psychosom Med. 2005; 67:9–15. [PubMed: 15673618]
- 72. Murali R, Chen E. Exposure to violence and cardiovascular and neuroendocrine measures in adolescents. Ann Behav Med. 2005; 30:155–163. [PubMed: 16173912]
- 73. Johnston-Brooks CH, Lewis MA, Evans GW, Whalen CK. Chronic stress and illness in children: the role of allostatic load. Psychosom Med. 1998; 60:597–603. [PubMed: 9773764]
- 74. Karlamangla AS, Mori T, Merkin SS, Seeman TE, Greendale GA, Binkley N, Crandall CJ. Childhood socioeconomic status and adult femoral neck bone strength: findings from the Midlife in the United States Study. Bone. 2013; 56:320–326. [PubMed: 23810840]
- Pearce MS, Birrell FN, Francis RM, Rawlings DJ, Tuck SP, Parker L. Lifecourse study of bone health at age 49-51 years: the Newcastle thousand families cohort study. J Epidemiol Community Health. 2005; 59:475–480. [PubMed: 15911643]

- 76. Karlamangla, A.; Crandall, C.; ryff, CD. Psychological well-being is positively associated with adult bone mineral density. Findings from the Study of Midlife in the United States. 2012 Annual Meeting of the American Society for Bone and Mineral Research; Minneapolis, MN. American Society for Bone and Mineral Research; 2012. al e
- 77. Statistics FIFoCaF. America's Children: Key National Indicators of Well-Being, 2013. U.S. Government Printing Office; Washington, DC: 2013.
- Razaz-Rahmati N, Nourian SR, Okoli CT. Does household structure affect adolescent smoking? Public health nursing. 2012; 29:191–197. [PubMed: 22512420]
- Frech A. Healthy Behavior Trajectories between Adolescence and Young Adulthood. Advances in life course research. 2012; 17:59–68. [PubMed: 22745923]
- Chang HY, Wu WC, Wu CC, Cheng JY, Hurng BS, Yen LL. The incidence of experimental smoking in school children: an 8-year follow-up of the child and adolescent behaviors in long-term evolution (CABLE) study. BMC Public Health. 2011; 11:844. [PubMed: 22051222]
- Brown SL, Rinelli LN. Family Structure, Family Processes, and Adolescent Smoking and Drinking. Journal of research on adolescence : the official journal of the Society for Research on Adolescence. 2010; 20:259–273. [PubMed: 20543893]
- Bratek A, Beil J, Banach M, Jarzabek K, Krysta K. The impact of family environment on the development of alcohol dependence. Psychiatria Danubina. 2013; 25(Suppl 2):S74–77. [PubMed: 23995149]
- Donovan JE, Molina BS. Childhood risk factors for early-onset drinking. Journal of studies on alcohol and drugs. 2011; 72:741–751. [PubMed: 21906502]
- Wang J, Simons-Morton BG, Farhat T, Luk JW. Socio-demographic variability in adolescent substance use: mediation by parents and peers. Prevention science : the official journal of the Society for Prevention Research. 2009; 10:387–396. [PubMed: 19582581]
- National Osteoporosis Foundation. Clinician's Guide to Prevention and Treatment of Osteoporosis. National Osteoporosis Foundation; Washington, DC: 2013.
- 86. Taes Y, Lapauw B, Vanbillemont G, Bogaert V, De Bacquer D, Goemaere S, Zmierczak H, Kaufman JM. Early smoking is associated with peak bone mass and prevalent fractures in young, healthy men. J Bone Miner Res. 2010; 25:379–387. [PubMed: 19653814]
- Rudang R, Darelid A, Nilsson M, Nilsson S, Mellstrom D, Ohlsson C, Lorentzon M. Smoking is associated with impaired bone mass development in young adult men: a 5-year longitudinal study. J Bone Miner Res. 2012; 27:2189–2197. [PubMed: 22653676]
- Eisman JA, Kelly PJ, Morrison NA, Pocock NA, Yeoman R, Birmingham J, Sambrook PN. Peak bone mass and osteoporosis prevention. Osteoporos Int. 1993; 3(Suppl 1):56–60. [PubMed: 8499027]
- Quarmby T, Dagkas S, Bridge M. Associations between children's physical activities, sedentary behaviours and family structure: a sequential mixed methods approach. Health education research. 2011; 26:63–76. [PubMed: 21059801]
- 90. Gibson LY, Byrne SM, Davis EA, Blair E, Jacoby P, Zubrick SR. The role of family and maternal factors in childhood obesity. Med J Aust. 2007; 186:591–595. [PubMed: 17547550]
- Sweeney NM, Glaser D, Tedeschi C. The eating and physical activity habits of inner-city adolescents. Journal of pediatric health care : official publication of National Association of Pediatric Nurse Associates & Practitioners. 2007; 21:13–21.
- Lindquist CH, Reynolds KD, Goran MI. Sociocultural determinants of physical activity among children. Prev Med. 1999; 29:305–312. [PubMed: 10547056]
- Nilsson M, Ohlsson C, Eriksson AL, Frandin K, Karlsson M, Ljunggren O, Mellstrom D, Lorentzon M. Competitive physical activity early in life is associated with bone mineral density in elderly Swedish men. Osteoporos Int. 2008; 19:1557–1566. [PubMed: 18373050]
- MacKelvie KJ, Khan KM, McKay HA. Is there a critical period for bone response to weightbearing exercise in children and adolescents? a systematic review. Br J Sports Med. 2002; 36:250– 257. discussion 257. [PubMed: 12145113]
- 95. Kannus P, Haapasalo H, Sankelo M, Sievanen H, Pasanen M, Heinonen A, Oja P, Vuori I. Effect of starting age of physical activity on bone mass in the dominant arm of tennis and squash players. Ann Intern Med. 1995; 123:27–31. [PubMed: 7762910]

- 96. Bass S, Pearce G, Bradney M, Hendrich E, Delmas PD, Harding A, Seeman E. Exercise before puberty may confer residual benefits in bone density in adulthood: studies in active prepubertal and retired female gymnasts. J Bone Miner Res. 1998; 13:500–507. [PubMed: 9525351]
- Lorentzon M, Mellstrom D, Ohlsson C. Association of amount of physical activity with cortical bone size and trabecular volumetric BMD in young adult men: the GOOD study. J Bone Miner Res. 2005; 20:1936–1943. [PubMed: 16234966]
- Amato PR. The impact of family formation change on the cognitive, social, and emotional wellbeing of the next generation. The Future of Children. 2005; 15:75–96. [PubMed: 16158731]
- 99. Tarasuk V, Vogt J. Household food insecurity in Ontario. Canadian journal of public health = Revue canadienne de sante publique. 2009; 100:184–188. [PubMed: 19507719]
- 100. Seguin L, Nikiema B, Gauvin L, Lambert M, Thanh Tu M, Kakinami L, Paradis G. Tracking exposure to child poverty during the first 10 years of life in a Quebec birth cohort. Canadian journal of public health = Revue canadienne de sante publique. 2012; 103:e270–276. [PubMed: 23618640]
- 101. Gisle L, Van Oyen H. Household composition and suicidal behaviour in the adult population of Belgium. Soc Psychiatry Psychiatr Epidemiol. 2013; 48:1115–1124. [PubMed: 23151963]
- Sieh DS, Visser-Meily JM, Oort FJ, Meijer AM. The diurnal salivary cortisol pattern of adolescents from families with single, ill and healthy parents. J Psychosom Res. 2012; 72:288– 292. [PubMed: 22405223]
- 103. Osborne C, Berger LM, Magnuson K. Family structure transitions and changes in maternal resources and well-being. Demography. 2012; 49:23–47. [PubMed: 22215507]
- 104. Hummer RA, Hamilton ER. Race and ethnicity in fragile families. The Future of children / Center for the Future of Children, the David and Lucile Packard Foundation. 2010; 20:113–131.
- 105. Lerman RI. Capabilities and contributions of unwed fathers. The Future of children / Center for the Future of Children, the David and Lucile Packard Foundation. 2010; 20:63–85.
- 106. Kalil A, Ryan RM. Mothers' economic conditions and sources of support in fragile families. The Future of children / Center for the Future of Children, the David and Lucile Packard Foundation. 2010; 20:39–61.
- 107. Crandall CJ, Miller-Martinez D, Greendale GA, Binkley N, Seeman TE, Karlamangla AS. Socioeconomic status, race, and bone turnover in the Midlife in the US Study. Osteoporos Int. 2012; 23:1503–1512. [PubMed: 21811862]
- 108. Howe LD, Lawlor DA, Propper C. Trajectories of socioeconomic inequalities in health, behaviours and academic achievement across childhood and adolescence. J Epidemiol Community Health. 2013; 67:358–364. [PubMed: 23322849]
- 109. Hertzman C, Boyce T. How experience gets under the skin to create gradients in developmental health. Annual review of public health. 2010; 31:329–347. 323p following 347.
- 110. Wilkinson PO, Goodyer IM. Childhood adversity and allostatic overload of the hypothalamicpituitary-adrenal axis: a vulnerability model for depressive disorders. Dev Psychopathol. 2011; 23:1017–1037. [PubMed: 22018079]
- 111. Blair C, Raver CC, Granger D, Mills-Koonce R, Hibel L. Allostasis and allostatic load in the context of poverty in early childhood. Dev Psychopathol. 2011; 23:845–857. [PubMed: 21756436]
- 112. Evans GW. A multimethodological analysis of cumulative risk and allostatic load among rural children. Dev Psychol. 2003; 39:924–933. [PubMed: 12952404]
- 113. Evans GW, English K. The environment of poverty: multiple stressor exposure, psychophysiological stress, and socioemotional adjustment. Child Dev. 2002; 73:1238–1248. [PubMed: 12146745]
- 114. Elefteriou F. Regulation of bone remodeling by the central and peripheral nervous system. Arch Biochem Biophys. 2008; 473:231–236. [PubMed: 18410742]
- 115. Schweiger U, Deuschle M, Korner A, Lammers CH, Schmider J, Gotthardt U, Holsboer F, Heuser I. Low lumbar bone mineral density in patients with major depression. Am J Psychiatry. 1994; 151:1691–1693. [PubMed: 7943462]
- 116. Michelson D, Stratakis C, Hill L, Reynolds J, Galliven E, Chrousos G, Gold P. Bone mineral density in women with depression. N Engl J Med. 1996; 335:1176–1181. [PubMed: 8815939]

- 117. Yirmiya R, Goshen I, Bajayo A, Kreisel T, Feldman S, Tam J, Trembovler V, Csernus V, Shohami E, Bab I. Depression induces bone loss through stimulation of the sympathetic nervous system. Proc Natl Acad Sci U S A. 2006; 103:16876–16881. [PubMed: 17075068]
- 118. Cizza G, Ravn P, Chrousos GP, Gold PW. Depression: a major, unrecognized risk factor for osteoporosis? Trends Endocrinol Metab. 2001; 12:198–203. [PubMed: 11397644]
- 119. Dennison E, Hindmarsh P, Fall C, Kellingray S, Barker D, Phillips D, Cooper C. Profiles of endogenous circulating cortisol and bone mineral density in healthy elderly men. J Clin Endocrinol Metab. 1999; 84:3058–3063. [PubMed: 10487665]
- 120. Kann P, Laudes M, Piepkorn B, Heintz A, Beyer J. Suppressed levels of serum cortisol following high-dose oral dexamethasone administration differ between healthy postmenopausal females and patients with established primary vertebral osteoporosis. Clin Rheumatol. 2001; 20:25–29. [PubMed: 11254236]
- 121. Reynolds RM, Dennison EM, Walker BR, Syddall HE, Wood PJ, Andrew R, Phillips DI, Cooper C. Cortisol secretion and rate of bone loss in a population-based cohort of elderly men and women. Calcif Tissue Int. 2005; 77:134–138. [PubMed: 16151676]
- 122. Papanicolaou DA, Wilder RL, Manolagas SC, Chrousos GP. The pathophysiologic roles of interleukin-6 in human disease. Ann Intern Med. 1998; 128:127–137. [PubMed: 9441573]
- 123. Ganesan K, Teklehaimanot S, Tran TH, Asuncion M, Norris K. Relationship of C-reactive protein and bone mineral density in community-dwelling elderly females. J Natl Med Assoc. 2005; 97:329–333. [PubMed: 15779496]

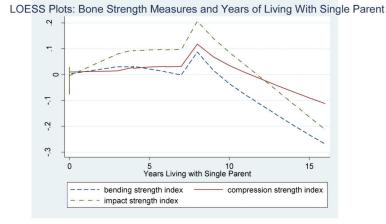


Figure 1.

LOESS Plots of numbers of years of living with a single parent versus bone strength indices, where bone strength indices are in standardized units

Table 1

Descriptive statistics for the study sample and the Midlife in the United States (MIDUS) II Biomarker Sample

	Study Sample N=708 ¹	MIDUS II Biomarke Sample (n=1255) ²
	% (n) or median [interquartile range]	% (n) or median [interquartile range]
Age (years)	56.0 [48.0-64.0]	57.0 [48.0-65.0]
Gender, male	48.9% (346)	43.2%
Gender, female	51.1% (362)	56.8% (713)
Race black	23.9% (169)	17.7% (222)
Race white	72.0% (539)	77.2% (1031)
Body mass index (kg/meters ²)	29.0 [25.3-33.6]	28.6 [25.2-33.0]
Males		
Age <50 years	32.4% (112)	28.8% (156)
Age 50-59 years	30.1% (104)	28.2% (153)
Age 60 years	37.6% (130)	43.0% (233)
Females		
Premenopausal	17.4% (63)	12.7% (72)
Early perimenopausal	14.4% (52)	9.9% (56)
Late peri-/post-menopausal, no menopausal hormone therapy	58.8% (213)	65.9% (374)
Post-menopausal, taking menopausal hormone therapy	9.4% (34)	11.6% (66)
Smoking (pack-years)	0 [0-11.3]	0 [0-11.3]
Smoking currently	16.7% (118)	14.9% (187)
Started Smoking < age	18 26.8% (190)	25.1% (314)
Heavy drinker at peak alcohol consumption	33.1% (232)	29.6% (368)
Heavy drinker in the past month	18.2% (128)	15.4% (192)
Physical Activity Scores		
Number of years doing recreational sports ages 14-18 years	0 [0-4.0]	0 [0-4.0]
Number of years doing competitive sports ages 14-18 years	1.0 [0-4.0]	1 [0-4.0]
Summary score for ages 20-35 years 3	30.0 [15.0-45.0]	30.0 [15.0-45.0]
Summary score for current (at time of MIDUS biomarker project visit) ⁴	312.5 [67.5-780.0]	320.0 [70.0-720.0]
Education		
High School	29.8% (211)	27.7% (344)
Some College	28.8% (204)	29.9% (371)
College	41.4% (293)	42.4% (527)
Childhood Socioeconomic Advantage Score ⁵	4.0 [3.0-5.0]	4.0 [3.0-5.0]

	Study Sample N=708 ¹	MIDUS II Biomarker Sample (n=1255) ²
Financial Adult Advantage Score ⁶	4.0 [2.0-6.0]	4.0 [2.0-6.0]
Bone Strength Measures		
Bending strength index, g/kg-m	1.2 [1.0-1.4]	1.2 [1.0-1.4]
Compression strength index, g/kg-m	3.5 [3.1-4.1]	3.5 [3.1-4.0]
Impact strength index, g/kg-m	0.2 [0.17-0.23]	0.2 [0.17-0.23]
Parental death/divorce before age 16 years		
Lived with both biological parents until age 16 years, or adopted at birth	72.5%	74.8%
Parental death/divorce when participant was under age 16 years	22.7%	21.1%
Other reason for not living with both biological parents until age 16	3.0%	2.6%
Number of years living in a single-parent household before age 16 years	0 [0-0]	0 [0-0]
0 years	92.0%* (651)	92.3% (1154)
1-8 years	2.0% (14)	2.0% (25)
9-16 years	6.1% (43)	5.7% (71)

IMajor reason for exclusion was unavailability of bone mineral density measurement. Sample sizes are reduced for some of the measures, including: financial advantage (n=706), smoking pack-years (n=642), physical activity ages 14-18 (n=644), physical activity ages 20-35 (n=640), current physical activity (n=706), heavy drinker at peak (n=701), and heavy drinker in past month (n = 703).

²Sample sizes are reduced for some of the measures, including race (n=1253), menopause group (n=1110), started smoking <age18 (n=1253), heavy drinker at peak (n=1245), heavy drinker in past month (n = 1246), education (n=1242), parental death/divorce (n=1251), years in a single-parent household (n = 1250) body mass index (n=1254), financial advantage (n=1252), childhood advantage (n=1249), smoking pack years (n=842), physical activity ages 14-18 (n=845, physical activity ages 20-35 (n=841), physical activity at p4 year (n=1250), bone strength measures (n=907).

³Summary score = (number of years doing light exercise * 1) + (number of years doing moderate exercise * 2) + (number of years doing vigorous exercise * 3)

⁴Summary score = (average number of minutes per week doing light exercise * 1) + average number of minutes doing moderate exercise * 2) + (average number of minutes doing vigorous exercise * 3)

 5 Childhood socioeconomic advantage score = being on welfare during childhood + childhood financial level relative to others + highest parental education. Possible range of score was 0-6.

 6 Adult financial advantage score = family-adjusted poverty-to-income ratio + self-rated current financial situation + sufficient money to meet needs + degree of difficulty paying bills. Possible range of score was 0-8.

Table 2

Adjusted associations of childhood family environment with indices of femoral neck strength relative to load; number of single parenting years as a continuous exposure⁷

		Model 1 (N=708) ⁸	Model 2 (N = $619)^9$
		Compression strength index Effect size (95% CI) in SD units	Compression strength index Effect size (95% CI) in SD units
Parental death or divorce (ref: Stable 2-parent family) IO	Parental death or divorce until age 16	0.171 (-0.021, 0.362)~	0.190 (-0.030, 0.410)~
	Other reason for not living with biological parents under age 16^{II}	0.381 (-0.234, 0.996)	0.330 (-0.321, 0.982)
Single parent childhood	Per additional year in single parent household	$-0.029\left(-0.053,-0.006 ight)^{*}$	$-0.028 \left(-0.051, -0.004 ight)^{*}$
		Bending strength index Effect size (95% CI) in SD units	Bending strength index Effect size (95% CI) in SD units
Parental death or divorce (ref: Stable 2-parent family)	Parental death or divorce until age 16	0.039 (-0.112, 0.190)	0.044 –0.128, 0.216)
	Other reason for not living with biological parents under age 16	0.103 (-0.244, 0.450)	0.017 (-0.309, 0.343)
Single parent childhood	Per additional year in single parent household	$-0.021 \left(-0.039, -0.004 ight)^{*}$	$-0.021 \left(-0.041, -0.002 ight)^{*}$
		Impact strength index Effect size (95% CI) in SD units	Impact strength index Effect size (95% CI) in SD units
Parental death or divorce (ref: Stable 2-parent family) 12	Parental death or divorce until age 16	0.105 (-0.037, 0.247)	0.096 (-0.068, 0.260)
	Other reason for not living with biological parents under age 16	-0.006 (-0.330, 0.318)	-0.070 (-0.398, 0.258)
Single parent childhood	Per additional year in single parent household	-0.017 (-0.034, 0.001)~	-0.018 (-0.037, 0.001)~
* n-value () ()5-() ()1·			

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p-value 0.05-0.01;

 $^{**}_{0.01-0.001;}$

*** <0.001; ~0.05-0.1

 $^7\mathrm{SD}$ denotes standard deviation, 95% CI denotes 95% confidence interval

 $\frac{8}{Model 11}$. Adjusted for race, site, menopausal stage, age, gender, body mass index (BMI), and BMI-gender interaction

9 <u>Model 2:</u> Adjusted for all covariates in Model 1 + education, adult financial advantage, childhood socioeconomic advantage, smoking pack years, recreational physical activity at ages 14-18, competitive sports at ages 14-18, physical activity at ages 20-35, current physical activity, smoking initiation before age 18, current smoking, heavy peak alcohol use, and heavy alcohol use in past month

 $^{IO}\!\!\!\!\!\!$ Lived with both biological parents until age 16, or adopted at birth

11 Lived in single-parent household without death or divorce, or gave other reason for not living with biological parents until age 16

 12 Lived with both biological parents until age 16, or adopted at birth

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Table 3

Adjusted associations of childhood family environment with indices of femoral neck strength relative to load; years of single parenting as a categorical exposure¹³

	-		
		Model 1 (N=708) ¹⁴	Model 2 (N = 619) ¹⁵
		Compression strength index Effect size (95% CI) in SD units	Compression strength index Effect size (95% CI) in SD units
Parental death or divorce (ref: Stable 2-parent family) ¹⁶	Parental death or divorce under age 16	0.168 (-0.032, 0.367)~	0.179 (-0.051, 0.410)
	Other reason for not living with biological parents until age 16^{17}	0.356 (-0.231, 0.944)	0.308 (-0.315, 0.932)
Single parent childhood (ref: Did not have single parent for most of childhood)	Single parent household for 1-8 years	-0.067 (-0.568, 0.435)	0.012 (-0.524, 0.548)
	Single parent household for 9-16 years	$-0.401 \ (-0.701, -0.100)^{*}$	$-0.376 \left(-0.675, -0.077\right)^{*}$
		Bending strength index Effect size (95% CI) in SD units	Bending strength index Effect size (95% CI) in SD units
Parental death or divorce (ref: Stable 2-parent family)	Parental death or divorce under age 16	0.027 (-0.124, 0.178)	0.024 (-0.151, 0.198)
	Other reason for not living with biological parents until age 16	0.090 (-0.247, 0.427)	0.008 (-0.310, 0.325)
Single parent childhood (ref: Did not have single parent for most of childhood)	Single parent household for 1-8 years	0.089 (-0.469, 0.647)	0.177 (-0.425, 0.780)
	Single parent household for 9-16 years	$-0.307 (-0.537, -0.076)^{*}$	$-0.307 (-0.558, -0.057)^{*}$
		Impact strength index Effect size (95% CI) in SD units	Impact strength index Effect size (95% CI) in SD units
Parental death or divorce (ref: Stable 2-parent family)	Parental death or divorce under age 16	0.091 (-0.050, 0.232)	0.076 (-0.091, 0.243)
	Other reason for not living with biological parents under age 16	-0.010 (-0.321, 0.301)	-0.073 (-0.385, 0.240)
Single parent childhood (ref: Did not have single parent for most of childhood)	Single parent household for 1-8 years	0.155 (-0.463, 0.772)	0.213 (-0.456, 0.883)

	Model 1 (N=708) ¹⁴	Model 2 (N = $619)^{I5}$
Single parent household for 9-16 years	$-0.254 \left(-0.484, -0.023\right)^{*}$	$-0.271 (-0.521, -0.020)^{*}$

* p-value 0.05-0.01;

 $^{**}_{0.01-0.001;}$

*** <0.001; ~0.05-0.1 13 SD denotes standard deviation, 95% CI denotes 95% confidence interval

 I_4^{I} Model 1: Adjusted for race, site, menopausal stage, age, gender, body mass index (BMI), and BMI-gender interaction

15 Model 2: Adjusted for all covariates in Model 1 + education, adult financial advantage, childhood socioeconomic advantage, smoking pack years, recreational physical activity at ages 14-18, competitive sports at ages 14-18, physical activity at ages 20-35, current physical activity, smoking initiation before age 18, current smoking, heavy peak alcohol use, and heavy alcohol use in past month

 I_6^{-16} Lived with both biological parents until age 16, or adopted at birth

17 Lived in single-parent household without death or divorce, or gave other reason for not living with biological parents until age 16