Cardiovascular health decline in adolescent girls in the NGHS cohort, 1987–1997

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ARTICLE INFO

Keywords:
Cardiovascular disease
Risk factors
Pediatrics
Women

ABSTRACT

Adolescence is a critical time for the preservation or loss of cardiovascular health. We aimed to describe trajectories of cardiovascular health in adolescent girls and identify early adolescent factors associated with cardiovascular health in young adulthood. We used data from the National Growth and Health Study, a longitudinal cohort of 2,379 girls followed annually from ages 9–19 years. We classified participants as having ideal, intermediate, or poor levels of the seven cardiovascular health metrics at four developmental stages: early (ages 9–11), middle (ages 12–14), and late (ages 15–17) adolescence, and early young adulthood (ages ≥ 18). We calculated total cardiovascular health scores (range 0–14) at each stage and empirically identified patterns of cardiovascular health trajectories. We examined associations between trajectory group membership and various demographic, behavioral, and physiological factors. Mean cardiovascular health scores declined with age from 10.8 to 9.4 in white girls and 10.3 to 8.9 in black girls; 17% of white girls and 23% of black girls had low cardiovascular health (score < 8) by early young adulthood. We identified five cardiovascular health trajectories: high-stable (14% of participants), high-to-moderate (48%), high-to-low (20%), moderate-stable (10%), and moderate-to-low (8%). Exceeding 14 h per week of television in early adolescence and teen pregnancy were associated with higher odds of being in several less healthy trajectory groups. In conclusion, cardiovascular health declines during adolescence and black-white disparities begin before early adolescence. Key targets for improving cardiovascular health in adolescent girls may include reductions in sedentary behavior and prevention of teen pregnancy.

1. Introduction

Adults who reach middle age with ideal cardiovascular health (CVH) live longer, healthier lives (Wilkins et al., 2012) free from cardiovascular disease (CVD) (Fang et al., 2016) and other chronic illnesses (Fang et al., 2016; Kulshreshtha et al., 2013; Muntner et al., 2013; Colangelo et al., 2011; Ogunmoroti et al., 2017). Ideal CVH includes seven metrics: optimal dietary patterns, regular physical activity, abstinence from tobacco, a healthy body mass index (BMI), and optimal levels of blood pressure, blood cholesterol, and blood glucose (Lloyd-Jones et al., 2010). While 45% of children in the US under age 20 meet ideal levels of at least five CVH metrics, the prevalence drops to 31% for adults ages 20–39 and 10% for adults over age 40 (Benjamin et al., 2019). A pooled analysis of five prospective cohort studies spanning from childhood to adulthood recently identified distinct trajectories of CVH loss, with earlier loss of CVH associated with subclinical atherosclerosis in middle age (Allen et al., 2016).

In this same pooled cohort analysis, late adolescence appeared to be a key period for CVH loss (Allen et al., 2018). Many changes occur during adolescence that affect CVH including changes in fat distribution...
adolescence. The cohort was established to identify the emergence of CVH trajectories in adolescent girls (Obesity and cardiovascular disease, 1992). Key findings from NGHS include greater increases in body fat (Kimm et al., 2004) and greater declines in physical activity (Kimm et al., 2002) in black compared to white girls. In this study, we aimed to leverage the rich repeated measures data in NGHS in order to 1) describe trajectories of CVH in adolescent girls ages 9–19 years in the NGHS cohort, and 2) identify early and late adolescent factors associated with CVH trajectories.

2. Methods

The NGHS enrolled 1213 black girls and 1166 white girls from Cincinnati, OH, Richmond, CA, and Washington, DC in 1987–1988. The original NGHS protocol was approved by the institutional review board at each participating center; parents provided informed consent and participants provided informed assent. Girls were ages 9–10 at recruitment and were followed annually with in-person examinations for ten years. All NGHS visits were scheduled at least 4 months post-partum for participants experiencing pregnancy during the study. The current study uses publicly available data from the National Heart, Lung, and Blood Institute’s Biolincc repository; the Boston Children’s Hospital Committee on Clinical Investigation and Biolincc approved the use of the data for this study. Further study information is available at https://biolincc.nlhbi.nih.gov/studies/nghs/.

2.1. Definitions of cardiovascular health metrics

We used the American Heart Association (AHA) definitions for CVH metrics for children (Lloyd-Jones et al., 2010), with minor adaptations based on variables collected in NGHS (supplementary Table 1). Girls were asked whether they had ever tried a cigarette starting at visits 1–5, with more detailed smoking histories taken at visits 6–10. Physical activity was measured at visits 1, 3, 5, 7–10 using a Habitual Activity Questionnaire; the metabolic equivalent (MET) and frequency of each reported activity were used to calculate an average score of MET-times per week. Girls were classified as having ideal (≥20 MET-times per week), intermediate (10 ≤ MET-times per week < 20), or poor (<10 MET-times per week) physical activity based on previous NGHS publications defining these groups as active, moderately active, and inactive, respectively (Kimm et al., 2005). Dietary intake was measured with a three-day food diary after participant training by a certified nutritionist at visits 1–5, 7, 8, 10 and analyzed for nutrient composition using the Nutrition Data System from the University of Minnesota’s Nutrition Coordinating Center. Food codes were matched to the United States Department of Agriculture database to produce daily intakes of major food groups used to calculate a 2015 Health Eating Index (HEI-2015) (Krebs-Smith et al., 2018). We used the top decile of HEI-2015 score for the study population from the first study visit to set the score for ideal diet throughout the remaining visits, given that <1% of adolescents report an ideal diet based on the AHA definition (Benjamin et al., 2019); the 2nd-5th deciles defined the intermediate diet score and the bottom five deciles defined the poor diet score.

Height and weight were measured at each visit with a standard stadiometer and electronic scale; BMI was calculated as kg/m² and classified according to the Centers for Disease Control 2000 BMI percentiles for children. Blood pressure was measured annually by a trained technician using a standard mercury sphygmomanometer; the mean of the second and third measurement was entered as the blood pressure for that visit. Fasting total cholesterol was measured at visits 1, 3, 5, 7, and 10; fasting glucose was measured at visits 7 and 10.

We classified each participant as having ideal (score = 2), intermediate (score = 1), or poor (score = 0) levels of each of the seven metrics (Supplementary Table 1) at each available study visit. Given the high proportion of girls (≥97%) with ideal glucose levels at study visit 7, we assigned ideal glucose status to all girls at earlier visits when glucose was not measured. For each of the other six metrics, if the metric was unavailable at a given visit we imputed the metric by substituting the lowest of the two surrounding visits. If no data were available for the given metric at the visit before or after, we considered the metric missing. The number of girls with measured, imputed, and missing data for each of the seven metrics at each visit year is reported in Supplementary Table 2. We then summed the scores on the seven individual metrics to create a total CVH score ranging from 0 to 14 for each participant at each visit. We included participants who had at least one summed total CVH score during any study visit, giving an analytic sample size of 2259 girls.

We created four developmental stage groupings representing early (ages 9–11), middle (ages 12–14), and late (ages 15–17) adolescence and early young adulthood (ages ≥ 18). If a participant had a total CVH score recorded at more than one study visit during a given developmental stage, we used the lower (less ideal) CVH score as the score for that stage. We classified the total CVH score for each developmental stage as low (0–7), moderate (8–11), or high (12–14) based on previous studies in adults demonstrating a graded relationship between these classifications and subclinical atherosclerosis (Polonsky et al., 2017).

2.2. Definitions of covariates

We defined early menarche as occurrence of the first menstrual period before age 12 (Kaplowitz and Oberfield, 1999). We defined participants as regular breakfast eaters in early adolescence if they reported “yes” to the question “On school days do you usually eat breakfast?” at the first study visit. We defined participants as exceeding the recommended amount of television viewing in early adolescence if they reported watching more than 14 h of television per week at the first study visit, based on American Academy of Pediatrics recommendations to limit screen time to <2 h per day (Education, 1999). We defined depressive symptoms as a score of >20 on the Center for Epidemiological Studies Depression scale (Vilagut et al., 2016) when assessed at visits 8 or 10. We defined teen pregnancy as any pregnancy occurring during the study period. Parents reported their highest level of education and total family income at the baseline study visit.

2.3. Statistical approach

We used latent class trajectory modeling to identify and categorize NGHS participants based on patterns of longitudinal change in the total CVH score during the four pre-specified developmental periods. A total of 1971 participants with a CVH score from at least three study visits were eligible for the trajectory analysis. Using a customized SAS macro, PROC TRAJ, we explored the distinct patterns in longitudinal change in the total CVH score over time. First, we determined CVH score trajectories as a function of age using PROC TRAJ. Next, we used Bayesian Information Criterion (BIC) posterior probabilities to evaluate the fit of the most parsimonious model, choosing the best model as the one with the smallest negative BIC. We assigned participants to the trajectory
class for which they had the highest Posterior Predicted Probability (PPP). Our trajectory model had high PPPs ranging from 0.74 to 0.90, indicating that each study participant had a high probability of belonging to one particular trajectory, and a low probability of belonging to the others (Supplementary Table 3).

We compared demographic characteristics and covariates for each of the CVH trajectory groups using chi-square tests and t-tests. We created multinomial logistic regression models to estimate the odds of CVH trajectory group membership based on each of the covariates of interest, with the most favorable CVH trajectory group serving as the referent. We further adjusted the models for demographic characteristics and formally tested for interaction between each covariate and race. A p-value < 0.05 was considered statistically significant. All analyses used SAS® 9.4 (SAS Institute, Cary, N.C., USA).

3. Results

Demographic characteristics and key health indicators for study participants are available in Table 1; all comparisons by race were statistically significant. The mean CVH score declined from 10.6 (SD 1.6) in early adolescence to 9.9 (SD 1.7) in middle adolescence, to 9.5 (SD 1.9) in late adolescence and finally to 9.1 (SD 2.1) in young adulthood. At each developmental stage, black girls had a mean CVH score approximately 0.5 points lower than white girls (p < 0.001 for comparison by race at each time point). Only 30% of girls entered adolescence with high CVH scores of 12–14 and the prevalence of high CVH declined at each subsequent developmental stage (Fig. 1). This loss of total CVH was related to worsening of all CVH metrics except cholesterol, with the greatest loss of ideal CVH seen in the blood pressure, physical activity, and smoking metrics (supplemental Fig. 1a–g). Compared to white participants, black participants were more likely to lose ideal status on each metric except smoking.

Based on observed patterns of changes in the CVH score over time, we identified and labeled five CVH trajectories (Fig. 2). The most common trajectory was high-to-moderate (45.1%) followed by high-to-low (20.1%) and high-stable (14.9%). The decline in CVH in the high-to-low group was most prominent in middle to late adolescence (ages 14–17) whereas the moderate-to-low group had a more prominent decline in CVH in early to middle adolescence (ages 11–14). The high-to-moderate and moderate-stable groups, and the high-to-low and moderate-to-low groups, both ended up with similar CVH scores in young adulthood (10 and 7, respectively). Only the high-stable group retained a mean ideal CVH score ≥ 11 into young adulthood.

We found statistically significant differences in trajectory group membership for all demographic and health indicators tested with the exception of high depression scores (p = 0.051) (Table 2). Girls who retained the highest CVH scores throughout adolescence (those in the high-stable group) were more likely to be white (72%), have parents with the highest education (65%) and income (63%) levels, eat breakfast regularly in early adolescence (90%), and were least likely to experience a teen pregnancy (15%). Girls with the lowest CVH scores at the end of adolescence (those in the high-to-low and moderate-to-low groups) had the highest mean hours of television watching in early adolescence (34.6 and 35.9 h per week), the lowest mean age at menarche (12.8 years), and the highest depression scores (17.2 and 16.9).

In unadjusted models, girls who identified as black, whose parents did not graduate from college, and whose family income was < $40,000 had higher odds of being in any of the less favorable CVH trajectory groups relative to the high-stable group (Table 3), as did girls who exceeded 14 h per week of television in early adolescence, who experienced menarche before age 12, and who experienced a teen pregnancy. In models adjusted for race, parental education and income, exceeding 14 h per week of television in early adolescence and experiencing a teen pregnancy remained associated with higher odds of being in the high-to-moderate, high-to-low, and moderate-stable CVH groups relative to the high-stable CVH group. Early menarche remained associated with higher odds of being in the high-to-low CVH group. The associations with television viewing, teen pregnancy, and early menarche were similar for both black and white participants and were not modified by race.

4. Discussion

Our analysis demonstrates that adolescence is an important developmental period for the loss or preservation of CVH in girls. Notably, almost all adolescents in NGHS demonstrated some decline in CVH. We empirically identified five distinct patterns of CVH loss in this historical cohort of over 2000 black and white girls. Mean CVH scores for each trajectory group varied widely at age 9, highlighting the importance of even earlier life influences on CVH including the perinatal period and early childhood. The largest declines in CVH were seen from early to middle adolescence, coinciding with the physical transition of puberty and the social transition from primary to secondary school. Our results are consistent with prior studies in NGHS documenting dramatic declines in physical activity (Kimm et al., 2002) and increases in the prevalence of obesity (Kimm et al., 2004), sugar sweetened beverage intake (Striegel-Moore et al., 2006), and poor dietary quality throughout adolescence (Moore et al., 2012).

Unsurprisingly, girls who maintained the most favorable CVH scores were more likely to have high parental income and education, as well as better breakfast eating and lower television viewing in early adolescence. Girls who entered adolescence with moderate CVH and declined

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Table 1: Characteristics of 2259 girls enrolled in the National Growth and Health Study, 1987–1997.

| Characteristic                      | White Participants N = 1105 | Black Participants N = 1154 | p-value  |
|------------------------------------|-----------------------------|-----------------------------|----------|
| Demographics                        |                             |                             |          |
| Age at enrollment in years (mean, 95% CI) | 10.0 (9.9–10.0)            | 10.1 (10.0–10.1)            | <0.0001  |
| Years of follow-up (mean, 95% CI)   | 8.6 (8.5–8.7)               | 8.8 (8.7–8.8)               | 0.0127   |
| Parent’s Highest Level of Education, n (%) |                           |                             | <0.0001  |
| High school or less                 | 209 (19.0%)                 | 360 (31.2%)                 |          |
| Some college                        | 337 (30.3%)                 | 547 (47.5%)                 |          |
| College graduate                    | 558 (50.5%)                 | 246 (21.3%)                 |          |
| Family Income, n (%)                |                             |                             | <0.0001  |
| Less than $10,000                   | 74 (7.0%)                   | 295 (27.3%)                 |          |
| $10,000–19,999                      | 97 (9.2%)                   | 193 (16.7%)                 | <0.0001  |
| $20,000–39,999                      | 343 (32.6%)                 | 320 (29.6%)                 |          |
| More than $40,000                   | 539 (51.2%)                 | 257 (23.8%)                 |          |

Early Adolescent Health Variables

| Reports eating breakfast regularly, n (%) | 988 (89.4%) | 881 (76.3%) | <0.0001  |
| Hours of TV viewing per week (mean, 95% CI) | 24.9 (22.9–45.9) | 36.4 (35.4–37.3) | <0.0001  |
| Exceeds 14 h per week of TV viewing, n (%) | 831 (76.7%) | 992 (89.9%) | <0.0001  |
| Age at menarche in years (mean, 95% CI) | 13.6 (13.5–13.7) | 12.7 (12.6–12.8) | <0.0001  |
| Early menarche before age 12, n (%) | 115 (10.7%) | 302 (26.5%) | <0.0001  |

Late Adolescent Health Variables

| Pregnant during study period, n (%) | 217 (19.6%) | 509 (44.1%) | <0.0001  |
| CESD Depression score at age 15 years (mean, 95% CI) | 16.5 (15.7–17.4) | 14.7 (13.8–15.6) | 0.0044  |
| CESD Depression score ≥ 20 at age 15 years, n (%) | 166 (33.4%) | 117 (24.7%) | 0.0028  |

TV = television. CESD = Center for Epidemiological Studies Depression scale.

* Missing observations were not included in calculating the percentage for these characteristics.
further into the low CVH range were more likely to be from lower socioeconomic strata and report less healthy behaviors. Girls in the moderate-stable group shared some features with both the high-declining and moderate-declining groups and represent an intriguing group for future study, as they were the only group to maintain their CVH score throughout adolescence.

Race was associated with both CVH score at the beginning of adolescence and CVH trajectories. Black girls were more likely to enter adolescence with only moderate CVH, and to end up with moderate or low CVH, even if they started adolescence with high CVH. These differences may be due to disparities in income and education between black and white families, as well as unmeasured factors such as access to healthy environments, rooted in differential treatment by race. These remain important areas for future investigation.

Excessive television viewing was associated with CVH decline. Extensive research shows associations between television viewing and lower physical activity (Marshall et al., 2004), poorer dietary habits (Wiecha et al., 2006), and obesity (Marshall et al., 2004) in youth. Furthermore, media use introduces exposure to advertising for unhealthy food (Wiecha et al., 2006) and tobacco products (B.F. New media and tobacco control, 2012). Thus it is not surprising that this behavior was associated with worsening CVH trajectories. The impact of the current media landscape on adolescent CVH - including ubiquitous smartphone use, streaming video, and social media – is an area ripe for investigation.

Earlier menarche, an important marker of later life CVD risk in women, was also associated with lower CVH in early adolescence and with being in the trajectory group with the most CVH loss. Similar associations between early menarche and lower CVH have been shown in post-menopausal women (Cao et al., 2015). Early menarche is

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**Fig. 1.** Total Cardiovascular Health (CVH) Score in Girls enrolled in the National Growth and Health Study, 1987–1997. Total CVH score (0–14) reflects the summation of scores on the seven individual cardiovascular health metrics for each participant during each developmental period.

**Fig. 2.** Empirically-derived Trajectories of the Cardiovascular Health Score over 10 years for Adolescent Girls in the National Growth and Health Study, 1987–1997. Five trajectory classes identified and labeled.
Pregnancy led to accelerated loss of CVH in this population. Teen pregnancy was also associated with less optimal CVH trajectories. Teen pregnancy was also associated with less optimal CVH trajectories.

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Table 2
Association of adolescent characteristics with five cardiovascular health trajectories identified in girls enrolled in the National Growth and Health Study, 1987–1997

| Characteristic | Trajectory Group | p-value |
|---------------|-----------------|---------|
|                | High-Stable n (%) | High-to-Moderate n (%) | High-to-Low n (%) | Moderate-Stable n (%) | Moderate-to-Low n (%) |
| Overall N, % of total sample | 275 (14%) | 954 (48%) | 392 (20%) | 198 (10%) | 152 (8%) |
| **Demographic factors** | | | | | |
| Race | | | | | <0.001 |
| White | 197 (72%) | 490 (51%) | 154 (39%) | 71 (36%) | 43 (28%) |
| Black | 78 (28%) | 464 (49%) | 238 (61%) | 127 (64%) | 109 (72%) |
| Parent’s Highest Level of Education | | | | | <0.001 |
| High school or less | 25 (9%) | 223 (23%) | 119 (30%) | 44 (22%) | 58 (38%) |
| Some college | 72 (26%) | 376 (40%) | 185 (47%) | 82 (41%) | 63 (42%) |
| College graduate | 178 (65%) | 354 (37%) | 88 (23%) | 72 (37%) | 31 (20%) |
| Family Income | | | | | <0.001 |
| Less than $10000 | 16 (6%) | 134 (15%) | 86 (23%) | 27 (14%) | 29 (20%) |
| $10000–19999 | 19 (7%) | 118 (13%) | 67 (18%) | 18 (10%) | 39 (27%) |
| $20000–39990 | 64 (24%) | 314 (35%) | 110 (30%) | 60 (32%) | 42 (30%) |
| More than $40000 | 165 (63%) | 340 (37%) | 104 (29%) | 84 (44%) | 33 (23%) |
| Early Adolescent Health Variables | | | | | |
| Reports eating breakfast regularly | 247 (90%) | 805 (84%) | 313 (80%) | 157 (79%) | 121 (80%) | 0.0026 |
| Mean hours of TV viewing per week (95% CI) | 24.1 (22.2–26.1) | 30.2 (29.1–31.3) | 34.6 (32.9–36.3) | 30.5 (28.2–32.8) | 35.9 (33.2–38.7) | <0.001 |
| Exceeds 14 h/week of TV viewing | 188 (69%) | 778 (84%) | 328 (88%) | 172 (88%) | 135 (92%) | <0.001 |
| Mean age at menarche (95% CI) | 13.4 (12.3–13.6) | 13.0 (12.9–13.1) | 12.8 (12.7–12.9) | 13.0 (12.8–13.2) | 12.8 (12.6–13.1) | <0.001 |
| Menarche < 12y | 31 (11%) | 173 (18%) | 105 (27%) | 32 (16%) | 36 (24%) | <0.001 |
| Late Adolescent Health Variables | | | | | |
| Teen pregnancy | 42 (15%) | 322 (34%) | 152 (39%) | 67 (34%) | 45 (30%) | <0.001 |
| Mean CESD Depression score (95% CI) | 14.9 (13.1–16.7) | 15.0 (14.0–15.9) | 17.2 (15.7–18.7) | 15.6 (13.6–17.6) | 16.9 (14.5–19.2) | 0.0067 |
| CESD Depression score ≥ 20 (n, %) | 34 (28%) | 107 (25%) | 65 (37%) | 29 (30%) | 23 (32%) | 0.0507 |

CESD = Center for Epidemiological Studies Depression; CI = Confidence Interval.

associated with childhood obesity (Rosenfield et al., 2009) and may be a marker of poor diet, physical activity, and BMI in pre-adolescence. The termination of linear growth soon after menarche in those already predisposed to weight gain may result in further weight gain and accelerated loss of ideal status for blood pressure, cholesterol, and glucose.

Teen pregnancy was also associated with less optimal CVH trajectories. Pregnancy is associated with numerous physiological changes that may affect later CVH. Post-partum weight retention is common and may affect later CVH. Post-partum weight retention is common and may affect later CVH.

The prevalence of ideal CVH metrics in this study differ somewhat from contemporary cross-sectional data from adolescents ages 12–19 years old participating in the 2015–2016 National Health and Nutrition Examination Surveys (NHANES) (Benjamin et al., 2019). The prevalence of ideal physical activity in adolescents ages 12–19 years old in NHANES (25.5%) is similar to the prevalence of ideal physical activity in the early and mid adolescent years of NGHS but markedly different than in late adolescence in NGHS. Similar declines in physical activity in girls are seen in contemporary NHANES cohorts, with black females reporting the least physical activity by young adulthood (Armstrong et al., 2018). Examining longitudinal patterns by developmental stage in cohorts such as NGHS is necessary to identify critical windows for intervention to promote maintenance of CVH metrics.

Other historical longitudinal cohorts have investigated the key role of CVH in adolescence on development of cardiovascular risk factors and surrogate markers of CVD in adulthood. The International Childhood Cardiovascular Cohort Consortium (i3C), which includes data from 5785 participants in five international cohort studies, found that the number of ideal CVH metrics in adolescence was inversely associated with carotid intima media thickness in young adulthood (Oikonen et al., 2013). The Special Turku Coronary Risk Factor Intervention Project for Children (STRIP) study found that a randomized dietary intervention from infancy through adolescence led to higher CVH and lower aortic intima-media thickness in young adulthood, demonstrating that CVH is not static and can be improved at this key stage in the life course (Pahkala et al., 2013).

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trajectories as adolescents age into young adulthood are needed to understand the impact of the many transitions in education, housing, relationships, and employment that occur in emerging adults.

Limitations to this study include the reliance on self-reported diet, physical activity, and smoking data and the collection of data approximately 30 years ago, when, for example, teen pregnancy rates were significantly higher than currently (Pazol et al., 2011). We used different diet and physical activity metrics from those in the standard AHA definitions of CVH, although we chose metrics previously shown to have high construct validity for moderate to vigorous physical activity (Kimm et al., 2012) and exemplary dietary patterns (Reedy et al., 2018) as defined by the AHA. We chose to impute missing data based on the lower of the next available metrics given prior studies showing overall loss of CVH in adolescence; this may have biased our results toward greater OR for girls identifying as white or black from three urban communities in the United States; in contrast to nationally representative studies, our sample is limited to young women with both ESRD and mortality. J. Am. Soc. Nephrol. 24 (7), 1159–1801.

5. Conclusion

Despite these limitations, NGHS contains one of the most detailed, frequent measurements of the CVH metrics in adolescence and thus we believe these findings are still relevant today. Future research is needed on modern cohorts that span from early childhood through adolescence to young adulthood in order to better define critical windows for CVH loss. These findings could shape both population and individual level interventions to improve the cardiovascular health of youth.

Sources of Funding

This work was supported by a NIH National Heart, Lung, and Blood Institute Career Development Award to Dr. Gooding [grant number K23-HL12236]. Dr Marma is supported by NIH/NHLBI K23HL145101.

Disclosures

None.

CRediT authorship contribution statement

Holly C. Gooding: Conceptualization, Writing - original draft, Visualization, Funding acquisition. Hongyan Ning: Methodology, Formal analysis, Visualization, Writing - review & editing. Amanda M. Pera: Writing - review & editing. Norrina Allen: Methodology, Writing - review & editing. Donald Lloyd-Jones: Resources, Writing - review & editing, Supervision. Lynn L. Moore: Resources, Writing - review & editing. Martha R. Singer: Software, Data curation. Sarah D. Ferranti: Writing - review & editing, Supervision.

Declaration of Competing Interest

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

Appendix A. Supplementary data

Supplementary data to this article can be found online at https://doi.org/10.1016/j.pmedr.2020.101276.

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