Association between Adverse Childhood Experiences and Muscle Strength in Older Age

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Keywords
Childhood misfortune · Grip strength · Health · Aging · Gender

Abstract
Background: Muscle weakness – a biomarker of health – may have its origins in early life and be related to factors such as adverse childhood experiences (ACE), which refer to a set of early-life traumatic and stressful psychosocial events out of the child’s control. To date, evidence of an association between ACE and muscle strength in older age is lacking.

Objective: Here, we assessed the associations between ACE (child in care, parental death, parental mental illness, parental drinking, period of hunger, or property taken away) and the risk of low muscle strength (LMS) in older age.

Methods: We used data from the Survey of Health, Ageing, and Retirement in Europe, a 12-year cohort study with 6 repeated measurements between 2004 and 2015. Muscle strength was measured using a handheld dynamometer. Confounder-adjusted logistic mixed-effect models were used to examine the associations between ACE (child in care, parental death, parental mental illness, parental drinking, period of hunger, or property taken away) and the risk of LMS in older age.

Results: 24,179 participants (96,372 observations; 13,477 women; aged 50–96 years) living in 14 countries were included. LMS increased with age for both genders. For women, there was a gradual increase in the risk of LMS with the number of experienced ACE (ORs = 1.22 for 1 ACE, 1.74 for $\geq 2$ ACE compared to no ACE). However, there was no significant association among men. This association was only slightly attenuated when adjusting for socioeconomic cir-
cumstances and unhealthy behaviors in adulthood. **Conclusions:** ACE are associated with later-life muscle weakness among women. These associations were not compensated by the adoption of healthy behaviors or an improvement in socioeconomic circumstances in adulthood. These results suggest that tackling these early-life risk factors in women could promote long-term grip strength, a biomarker of aging.

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**Introduction**

Muscle weakness, as measured by grip strength, is a biomarker of aging [1] and has been shown to predict a range of adverse health outcomes including future disability [2, 3], morbidity [2, 4], and mortality [5–7]. Because of its high prognostic value, a better understanding of the factors influencing the decline in muscle strength as adults grow older is important.

Muscle weakness may have its origins in early life and, therefore, be associated with risk factors such as childhood socioeconomic circumstances [8–10] and adverse childhood experiences (ACE), which are defined as a set of early-life traumatic and stressful psychosocial events out of the child’s control (e.g., parental separation, mental illness, or death) [10–12]. These early-life events may influence muscle strength through socioeconomic, behavioral, and biological mechanisms [13]. For example, individuals from disadvantaged childhood socioeconomic backgrounds who experienced ACE are more likely to engage in unhealthy and risky behaviors (e.g., smoking or alcohol consumption) [14, 15] and develop physiological wear and tear in response to chronic stress [16, 17]. This stress hypothesis is based on previous studies showing that the physiological stress response and the immune system are developed and calibrated by early-life experiences [18]. For example, children suffering from chronic stress build up higher levels of the stress hormone cortisol, making the body’s response to threats more pronounced [19]. As such, early-life chronic stress has been found to be linked to a host of diseases [20] through mechanisms including the dysregulation of the immune system [21].

Although some studies have found that ACE are associated with poorer objective physical capability [11], very few studies have assessed whether these associations are independent of childhood socioeconomic circumstances [10, 11]. Yet, socioeconomic disadvantage and adverse experiences in childhood co-occur [22–24]. As such, adjusting for socioeconomic circumstances during childhood is required to examine the independent association of ACE with muscle strength. Furthermore, to the best of our knowledge, no study has assessed the associations between ACE and muscle strength or any other objective measure of physical function in adults aged 65 years or more. Accordingly, evidence for an independent association of ACE with objectively measured muscle strength in older age has not been investigated to date.

The objective of this study was to assess the association between ACE and the risk of low muscle strength (LMS) at older age independently of childhood socioeconomic circumstances. We also examined whether this association is explained by socioeconomic circumstances and health-related behaviors in adulthood. We hypothesized, in line with the stress hypothesis, that individuals who experienced ACE would show a higher risk of LMS at older age (H1) and show a higher rate of increased risk of LMS with aging (H2). We further expected that the associations between ACE and LMS would partly be explained by adult-life socioeconomic circumstances and unhealthy lifestyle behaviors (H3).

**Methods**

**Study Population and Design**

Data were retrieved from the Survey of Health, Ageing, and Retirement in Europe (SHARE), a longitudinal (12-year [2004–2015]) European and retrospective database (6 waves of data collected every 2 years) of individuals aged 50 years or older [25]. Muscle strength was assessed at each of the 6 waves. Retrospective life course data including ACE were collected in the third wave. We included data for participants aged 50–96 years who participated in the third wave and had at least 1 observation of muscle strength. Participants with no information on their childhood life history, adult-life socioeconomic circumstances, or healthy behaviors were excluded. Online supplementary Table S1 (for all online suppl. material, see www.karger.com/doi/10.1159/000494972) shows the flow diagram of participant inclusion. Supplemental materials 1 and 2 show more details on the design and all measures described below.

**Measures**

**ACE**

ACE were identified as a set of traumatic events (emotional, physical, or linked to household dysfunction) occurring during childhood and being out of a child’s control [14]. We carefully screened SHARE for variables that matched this definition. The following binary indicators reflecting specific ACE (from 0 to 15 years) were included: child in care (living in a children’s home or with a foster family), parental death (father, mother, or both), parental mental illness, parental drinking abuse, period of hunger, and property taken away. Consistent with previous studies [26], by combining these 6 indicators, we computed a 3-level categorical variable of participants with no ACE (i.e., participants who only answered "no"), 1 ACE (i.e., participants who answered "yes" at 1 indicator

Cheval et al.
Adverse Childhood Experiences and Elderly Muscle Strength

Grip strength was measured twice in both hands, alternating between the hands, using a handheld dynamometer (Smedley, S Dynamometer; TTM, Tokyo, Japan; 100 kg). Participants were instructed to stand (preferably) or sit, with the elbow at a 90° angle, the wrist in neutral position, and to keep the upper arm tight against the trunk. Interviewers applied standardized instructions to ensure that the grip strength was performed with maximum effort. The mean of the maximum values obtained for each hand was used as an indicator of muscle strength [5, 8]. When values for one hand were missing or were equal to 0, the measurement at this time point was excluded from the analysis. Then, consistent with previous literature, the cutoff for LMS was computed based on gender and body mass index (BMI) quartiles [27, 28] following the Fried criterion (i.e., grip strength in the lowest 20% adjusted for gender and BMI). For men with BMI \( \leq 24 \), \( 24–26 \), \( 26–28 \), and \( \geq 28 \), the cutoffs for LMS were \( 26, 29, 30, \) and \( 32 \) kg, respectively. For women with BMI \( \leq 23 \), \( 23–26 \), \( 26–28 \), and \( \geq 28 \) kg/m\(^2\), the cutoffs for LMS were 17, 17.3, 18, and 21 kg, respectively.

Explanatory Variables

Adult-Life Socioeconomic Circumstances

The following variables were included: highest educational attainment, main occupational position during adult life, and satisfaction with household financial situation.

Unhealthy Behaviors

The following self-reported behaviors were included: physical inactivity, unhealthy eating, smoking, and alcohol consumption.

Covariates

We considered childhood socioeconomic circumstances [29], birth cohorts, countries, attrition, childhood health problems (long or multiple hospitalizations, childhood illness, and childhood serious health condition), and height as covariates.

Statistical Analysis

To account for the nested structure of the data (multiple observations within a single individual), logistic mixed-effect models were used [30]. Analyses were stratified by gender because previous studies have shown potential differences between female and male disease development over the life course [26]. Model 0 tested the association between ACE and the risk of LMS, adjusting for height only. Height was controlled for because it has been shown to be strongly correlated with grip strength [3, 5] and to be socioeconomically patterned (low socioeconomic position is associated with short height) [31, 32]. As such, adjusting for height is deemed necessary to ensure that the association observed between ACE and muscle strength did not simply result from differences in height. Model 1 tested the association between ACE and the risk of LMS, adjusting for childhood socioeconomic circumstances, birth cohort, attrition, childhood health problems, and height. Age was centered at the midpoint of the age range (73 years) and divided by 10 so that the coefficient yielded effects of an increased risk of LMS over a 10-year period (model 1). Additionally, an interaction term between ACE and age was added to test whether ACE moderated the effect of age on the risk of LMS (model 1b). Highest educational attainment (model 2), main occupational position during adult life (model 3), satisfaction with household financial situation (model 4), and unhealthy behaviors (model 5) were added as potential explanatory variables. Interaction terms between these variables and age were also tested (models 2b, 3b, 4b, and 5b).

We performed the following 6 sensitivity analyses: (1) assessing ACE exposure until 18 years, (2) assessing ACE exposure until 20 years, (3) controlling for the following chronic conditions: self-reported heart attack, high blood pressure, stroke, diabetes, chronic lung disease, Parkinson disease, or cancer, (4) excluding participants who dropped out for reasons other than death, (5) excluding participants who died during the survey, and (6) modeling grip strength as a continuous variable. The rationale and main results of these sensitivity analyses are presented in the supplemental materials. Statistical analyses were performed using the R language and the lme4 and lmerTest packages [33–35].

Data Sharing

This SHARE dataset is available at http://www.share-project.org/data-access.html.

Transparency

The lead author affirms that the manuscript is a honest, accurate, and transparent account of the study being reported; that no important aspects of the study have been omitted; and that any discrepancies from the study as planned have been explained.

Results

Descriptive Results

Table 1 shows the characteristics of the analytical sample stratified by gender. The sample included 24,179 participants (96,372 observations; 13,477 women [55.7%]) living in 14 European countries. LMS was less frequent in men than women (~3.14%, \( p < 0.001 \)). Men reported slightly more ACE than women, but this difference was not significant. Women had a lower level of educational attainment, lower main occupational position during adult life, a lower satisfaction with household financial situation, and reported less unhealthy-related behaviors than men (\( p < 0.001 \)).

Minimally Adjusted Model (Model 0)

For women, results of the model adjusted for height only showed a gradual increase in the odds of LMS with the number of ACE experienced (ORs = 1.41, \( p = 0.003 \) for having reported 1 ACE, and 1.98, \( p = 0.003 \), for \( \geq 2 \) ACE).

For men, results showed that the odds of LMS were greater for men who had reported 1 ACE (OR = 1.33, \( p = 0.036 \)). The association with having reported \( \geq 2 \) ACE was, however, only marginal (OR = 1.56, \( p = 0.099 \)) (Table 2).
<table>
<thead>
<tr>
<th></th>
<th>Women (n = 13,477)</th>
<th>Men (n = 10,702)</th>
<th>p value</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Outcome</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Muscle weakness</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Yes</td>
<td>1,643 (12.19)</td>
<td>968 (9.04)</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>No</td>
<td>11,834 (87.81)</td>
<td>9,734 (90.96)</td>
<td></td>
</tr>
<tr>
<td><strong>Covariates</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Age (SD), years</td>
<td>62.3 (9.3)</td>
<td>62.8 (8.9)</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td><strong>Countries</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Belgium</td>
<td>1,374 (10.20)</td>
<td>1,138 (10.63)</td>
<td></td>
</tr>
<tr>
<td>Austria</td>
<td>485 (3.60)</td>
<td>334 (3.12)</td>
<td></td>
</tr>
<tr>
<td>Denmark</td>
<td>1,060 (7.87)</td>
<td>891 (8.33)</td>
<td></td>
</tr>
<tr>
<td>France</td>
<td>1,181 (8.76)</td>
<td>894 (8.35)</td>
<td></td>
</tr>
<tr>
<td>Germany</td>
<td>890 (6.60)</td>
<td>778 (7.27)</td>
<td></td>
</tr>
<tr>
<td>Greece</td>
<td>1,490 (11.06)</td>
<td>1,172 (10.95)</td>
<td></td>
</tr>
<tr>
<td>Italy</td>
<td>1,247 (9.25)</td>
<td>1,040 (9.72)</td>
<td></td>
</tr>
<tr>
<td>The Netherlands</td>
<td>1,033 (7.67)</td>
<td>867 (8.10)</td>
<td></td>
</tr>
<tr>
<td>Spain</td>
<td>977 (7.25)</td>
<td>738 (6.90)</td>
<td></td>
</tr>
<tr>
<td>Sweden</td>
<td>921 (6.83)</td>
<td>733 (6.85)</td>
<td></td>
</tr>
<tr>
<td>Switzerland</td>
<td>654 (4.85)</td>
<td>498 (4.65)</td>
<td></td>
</tr>
<tr>
<td>Czech Republic</td>
<td>919 (6.82)</td>
<td>664 (6.20)</td>
<td></td>
</tr>
<tr>
<td>Ireland</td>
<td>342 (2.54)</td>
<td>265 (2.48)</td>
<td></td>
</tr>
<tr>
<td>Poland</td>
<td>904 (6.71)</td>
<td>690 (6.45)</td>
<td></td>
</tr>
<tr>
<td>Birth cohort</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>After 1945</td>
<td>6,155 (45.67)</td>
<td>4,489 (41.95)</td>
<td></td>
</tr>
<tr>
<td>Between 1939 and 1945</td>
<td>3,024 (22.44)</td>
<td>2,617 (24.45)</td>
<td></td>
</tr>
<tr>
<td>Between 1929 and 1938</td>
<td>2,992 (22.20)</td>
<td>2,631 (24.58)</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Between 1919 and 1928</td>
<td>1,306 (9.69)</td>
<td>965 (9.02)</td>
<td></td>
</tr>
<tr>
<td><strong>Attrition</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>No dropout</td>
<td>9,685 (71.86)</td>
<td>7,276 (67.99)</td>
<td></td>
</tr>
<tr>
<td>Dropout</td>
<td>2,851 (21.16)</td>
<td>2,312 (21.60)</td>
<td></td>
</tr>
<tr>
<td>Death</td>
<td>941 (6.98)</td>
<td>1,114 (10.41)</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td><strong>Childhood socioeconomic circumstances</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Most disadvantaged</td>
<td>2,430 (18.02)</td>
<td>2,030 (18.97)</td>
<td></td>
</tr>
<tr>
<td>Disadvantaged</td>
<td>3,409 (25.30)</td>
<td>2,660 (24.86)</td>
<td></td>
</tr>
<tr>
<td>Middle</td>
<td>4,415 (32.76)</td>
<td>3,380 (31.58)</td>
<td></td>
</tr>
<tr>
<td>Advantaged</td>
<td>2,466 (18.30)</td>
<td>2,009 (18.77)</td>
<td></td>
</tr>
<tr>
<td>Most advantaged</td>
<td>757 (5.62)</td>
<td>623 (5.82)</td>
<td>0.1345</td>
</tr>
<tr>
<td><strong>Childhood health problems</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>No</td>
<td>10,129 (75.16)</td>
<td>7,921 (74.01)</td>
<td></td>
</tr>
<tr>
<td>Yes</td>
<td>3,348 (24.84)</td>
<td>2,781 (25.99)</td>
<td>0.0439</td>
</tr>
<tr>
<td><strong>Adverse childhood experiences</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>None</td>
<td>10,674 (79.20)</td>
<td>8,389 (78.39)</td>
<td></td>
</tr>
<tr>
<td>One</td>
<td>2,305 (17.10)</td>
<td>1,904 (17.79)</td>
<td></td>
</tr>
<tr>
<td>Two or more</td>
<td>498 (3.70)</td>
<td>409 (3.82)</td>
<td>0.3046</td>
</tr>
<tr>
<td><strong>Adult socioeconomic circumstances</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Level of education</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>High education</td>
<td>2,332 (17.30)</td>
<td>2,530 (23.64)</td>
<td></td>
</tr>
<tr>
<td>Low education</td>
<td>11,145 (82.70)</td>
<td>8,172 (76.36)</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Main occupation class</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>High skill</td>
<td>2,226 (16.52)</td>
<td>3,351 (31.31)</td>
<td></td>
</tr>
<tr>
<td>Low skill</td>
<td>11,251 (83.48)</td>
<td>7,351 (68.69)</td>
<td>&lt;0.001</td>
</tr>
</tbody>
</table>
Adverse Childhood Experiences and Elderly Muscle Strength

Association of ACE with LMS (Model 1)

For women, the odds of LMS increased 4.77-fold every 10 years ($p < 0.001$). The odds of LMS were greater for women who grew up in disadvantaged childhood socioeconomic circumstances (e.g., advantaged childhood socioeconomic circumstances were associated with a decreased risk of LMS compared with most disadvantaged childhood socioeconomic circumstances; OR = 0.64, $p = 0.006$). Compared to women with no ACE, the odds of LMS were greater for women who had reported 1 ACE (OR = 1.22, $p = 0.033$). For women who had reported 2 ACE, the odds of LMS were even higher (OR = 1.74, $p = 0.002$) (Table 2; Fig. 1). Interactions between ACE and age were not significant (see online suppl. Table S2).

For men, the odds of LMS increased 10.59-fold every 10 years ($p < 0.001$). The odds of LMS were greater for men who grew up in disadvantaged childhood socioeconomic circumstances compared to men who grew up in most disadvantaged childhood socioeconomic circumstances (OR = 0.72, $p = 0.010$). The odds of LMS were not significantly associated with ACE (OR = 1.17, $p = 0.165$), for having reported 1 ACE, and 1.30, $p = 0.230$, for $\geq$ 2 ACE; Table 3; Fig. 1). Interactions between ACE and age were not significant (see online suppl. Table S2).

Associations of Adult-Life Socioeconomic Circumstances and Unhealthy-Related Behaviors with LMS (Models 2–5)

For women, the association between ACE and the odds of LMS was only slightly attenuated with the addition of adult-life socioeconomic circumstances and unhealthy behaviors. The associations remained significant for women who had reported $\geq 2$ ACE (OR = 1.55, $p = 0.011$) but not for women who had reported 1 ACE (OR = 1.16, $p = 0.107$). The odds of LMS were greater for women with lower educational attainment (OR = 1.97, $p < 0.001$; model 2), a disadvantaged main occupational position in adulthood (OR = 1.98, $p < 0.001$; model 3), a lower satisfaction with household financial situation (OR = 1.46, 2.51, 5.45, $p < 0.001$; model 4), and a higher number of unhealthy-related behaviors (OR = 4.65, $p < 0.001$; model 5). The interactions between these explanatory variables and age were not significant.

For men, the associations between ACE and the odds of LMS remained nonsignificant. The odds of LMS were greater for men with a disadvantaged main occupational position in adulthood (OR = 1.83, $p < 0.001$; model 3), a lower satisfaction with household financial situation (OR = 1.85 and 3.22, $p < 0.001$; model 4), and a higher number of unhealthy-related behaviors (OR = 3.08, $p < 0.001$; model 5), but not with educational attainment (OR = 1.16, $p = 0.206$; model 2). The interactions between these explanatory variables and age were not significant.

Sensitivity Analyses

Results of the 6 sensitivity analyses were consistent with those of the main analyses, except for the association between having reported 1 ACE and LMS in women which became marginal in some models (online suppl. Table S3).

Discussion

Main Findings

This study aimed to assess the associations between ACE during the first 15 years of life and the risk of LMS later in life, and whether adult-life socioeconomic circumstances and unhealthy behaviors partly explained these associations. Based on the previous literature and theoretical models, we hypothesized that individuals who experienced ACE...
### Table 2. Predictors of low muscle strength in older age in women

<table>
<thead>
<tr>
<th>Predictor</th>
<th>Model 0</th>
<th>Model 1</th>
<th>Model 2</th>
<th>Model 3</th>
<th>Model 4</th>
<th>Model 5</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>OR (95% CI)</td>
<td>OR (95% CI)</td>
<td>OR (95% CI)</td>
<td>OR (95% CI)</td>
<td>OR (95% CI)</td>
<td>OR (95% CI)</td>
</tr>
<tr>
<td>Age (10 years)</td>
<td>4.77 (4.27–5.33)</td>
<td>4.72 (4.22–5.27)</td>
<td>4.79 (4.28–5.35)</td>
<td>4.81 (4.30–5.37)</td>
<td>4.74 (4.25–5.29)</td>
<td>&lt;0.001 &lt;0.001 &lt;0.001 &lt;0.001</td>
</tr>
<tr>
<td>ACE</td>
<td>None (ref)</td>
<td>One (1.41 (1.12–1.76)</td>
<td>1.22 (1.02–1.47)</td>
<td>1.21 (1.01–1.46)</td>
<td>1.25 (1.04–1.50)</td>
<td>1.17 (0.98–1.40)</td>
</tr>
<tr>
<td>Childhood SEC</td>
<td>None (ref)</td>
<td>One (0.66 (0.50–0.86)</td>
<td>0.67 (0.51–0.87)</td>
<td>0.69 (0.53–0.90)</td>
<td>0.72 (0.55–0.93)</td>
<td>0.72 (0.55–0.93)</td>
</tr>
<tr>
<td>Level of education</td>
<td>High education (ref)</td>
<td>Low education (1.97 (1.57–2.47)</td>
<td>1.51 (1.17–1.93)</td>
<td>1.30 (1.02–1.67)</td>
<td>1.26 (0.99–1.61)</td>
<td>0.001 0.001 0.035 0.063</td>
</tr>
<tr>
<td>Main occupation class</td>
<td>High (ref)</td>
<td>Low (1.98 (1.55–2.53)</td>
<td>1.75 (1.37–2.23)</td>
<td>1.70 (1.34–2.16)</td>
<td>1.70 (1.34–2.16)</td>
<td>&lt;0.001 &lt;0.001 &lt;0.001</td>
</tr>
<tr>
<td>Satisfaction with income</td>
<td>Easy (ref)</td>
<td>Fairly easy (1.46 (1.22–1.76)</td>
<td>&lt;0.001</td>
<td>1.41 (1.18–1.69)</td>
<td>1.41 (1.18–1.69)</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td></td>
<td>With some difficulty (2.51 (2.04–3.09)</td>
<td>&lt;0.001</td>
<td>2.23 (1.82–2.74)</td>
<td>2.23 (1.82–2.74)</td>
<td>&lt;0.001</td>
<td></td>
</tr>
<tr>
<td></td>
<td>With great difficulty (5.45 (4.23–7.02)</td>
<td>&lt;0.001</td>
<td>4.56 (3.55–5.85)</td>
<td>4.56 (3.55–5.85)</td>
<td>&lt;0.001</td>
<td></td>
</tr>
<tr>
<td></td>
<td>Health behaviors (4.65 (3.58–6.05)</td>
<td>&lt;0.001</td>
<td></td>
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ACE, adverse childhood experiences; SEC, socioeconomic circumstances. All models are adjusted for country of residence, birth cohort, attrition, height, and childhood health problems. Interaction terms between ACE and age (model 1b), level of education and age (model 2b), main occupation class and age (model 3b), satisfaction with household income and age (model 4b), and health behaviors and age (models 5b) were also tested. Only the interaction between age and health behaviors was significant. Model 0 was only adjusted for height. Health behaviors included physical inactivity, unhealthy eating, smoking, and alcohol consumption (higher score indicating higher number of unhealthy behaviors).
would show a higher risk of LMS at older age (H1) and would show a higher increased risk of LMS with aging (H2). Additionally, we expected that the associations between ACE and risk of LMS would be only partially explained by adult-life socioeconomic circumstances and unhealthy behaviors (H3). In this large European study, consistent with H1, we found that ACE were associated with an increased risk of LMS in women, independently of socioeconomic circumstances during childhood. However, there was no strong evidence of associations among men. Our H2 was not confirmed, however, since we did not observe an association between ACE and the increased risk of LMS with aging in both men and women. Finally, consistent with H3, adjustment for adult-life socioeconomic circumstances (i.e., education, main occupational position, and satisfaction with household financial situation) and unhealthy behaviors (i.e., physical inactivity, unhealthy eating, smoking, and alcohol consumption) did not explain the association between ACE and risk of LMS among women.

**Comparison with Previous Studies**

Our study showed that ACE was associated with a higher risk of LMS in older women independently of socioeconomic circumstances during childhood. To our knowledge, this study was the first to find evidence for this unique association between ACE and muscle weakness in older age. Indeed, although a few previous studies have found ACE and objective measures of physical capability to be associated, these associations did not remain significant after adjusting for childhood socioeconomic circumstances [10, 11]. Three factors could explain this discrepancy. Our study used a single indicator to assess physical capability, whereas the 2 previous studies used a composite score involving grip strength, chair rise time, one-leg standing balance with eyes closed, and walking speed. Scores derived from a single test (grip strength) are likely more stable than scores derived from multiple tests. Additionally, grip strength may be more strongly associated with ACE than other physical indicators. Finally, the samples of the previous studies were smaller (n = 2,221 and 5,362), younger (midlife and 60–64 years), and involved British adults only.

In our study, ACE were associated with an increased risk of LMS in women only. These results may be explained by gender differences in the puberty period, which occurs later in men and involves different physiological processes [36]. For muscle tissue, puberty induces changes in cell metabolism and typology, associated to increased muscle growth. These muscle maturation occurs at the end of adolescence for women whereas boys achieve it in young adulthood [37, 38].

Muscle tissue maturation occurs earlier for women with a fiber diameter peaking during adolescence, whereas boys achieve this peak in young adulthood [38, 39]. This result suggests that events occurring before this developmental period may have a small influence on muscle growth, whereas events occurring during this period may be particularly influential and have a permanent effect on muscle function for the rest of the life. In addition, gender differences may be linked to the fact that women are more susceptible to cumulative disadvantages over the life course than men [40]. For instance, women generally reach a lower level of education, earn less, and are more likely to sacrifice their professional careers to take care of their child [41].

In line with previous studies, adjustment for adult-life socioeconomic circumstances and health-related behaviors did not explain the association between ACE and physical function [42]. A potential explanation of this result is linked to the chronic stress hypothesis [43, 44]. Stress responses and the immune system are shaped and calibrated by early-life experiences [18]. Chronic stress in childhood can lead to a cumulative wear-and-tear effect on the physiological system that governs individuals’ response to their environment, permanently altering the equilibrium and reactivity of these systems leading to
Table 3. Predictors of low muscle strength in older age in men

<table>
<thead>
<tr>
<th></th>
<th>Model 0</th>
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<th>Model 1</th>
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<th>Model 2</th>
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<td></td>
<td>OR (95% CI)</td>
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<td>OR (95% CI)</td>
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<td>Age (10 years)</td>
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<tr>
<td></td>
<td>10.59 (9.10–12.32)</td>
<td>&lt;0.001</td>
<td>10.59 (9.10–12.33)</td>
<td>&lt;0.001</td>
<td>10.57 (9.08–12.30)</td>
<td>&lt;0.001</td>
<td>10.61 (9.12–12.34)</td>
<td>&lt;0.001</td>
<td>10.64 (9.15–12.37)</td>
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<tr>
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<td>(ref)</td>
<td>0.165</td>
<td>(ref)</td>
<td>0.164</td>
<td>(ref)</td>
<td>0.187</td>
<td>(ref)</td>
<td>0.316</td>
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<tr>
<td>One</td>
<td>1.33 (1.02–1.75)</td>
<td>0.036</td>
<td>1.17 (0.94–1.47)</td>
<td>0.165</td>
<td>1.16 (0.94–1.47)</td>
<td>0.164</td>
<td>1.12 (0.90–1.40)</td>
<td>0.316</td>
<td>1.05 (0.84–1.31)</td>
<td>0.652</td>
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<td>Two or more</td>
<td>1.56 (0.92–2.65)</td>
<td>0.099</td>
<td>1.30 (0.85–1.99)</td>
<td>0.23</td>
<td>1.31 (0.85–2.00)</td>
<td>0.219</td>
<td>1.24 (0.82–1.89)</td>
<td>0.315</td>
<td>1.18 (0.78–1.79)</td>
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<td>Most disadvantaged</td>
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<tr>
<td>Disadvantaged</td>
<td>0.72 (0.55–0.92)</td>
<td>0.01</td>
<td>0.71 (0.55–0.92)</td>
<td>0.009</td>
<td>0.74 (0.58–0.96)</td>
<td>0.023</td>
<td>0.77 (0.60–0.99)</td>
<td>0.045</td>
<td>0.79 (0.62–1.01)</td>
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<td>Middle</td>
<td>0.78 (0.60–1.02)</td>
<td>0.065</td>
<td>0.79 (0.60–1.03)</td>
<td>0.079</td>
<td>0.87 (0.66–1.14)</td>
<td>0.309</td>
<td>0.95 (0.73–1.24)</td>
<td>0.69</td>
<td>0.97 (0.75–1.27)</td>
<td>0.836</td>
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<tr>
<td>Advantaged</td>
<td>0.81 (0.59–1.10)</td>
<td>0.178</td>
<td>0.83 (0.61–1.15)</td>
<td>0.265</td>
<td>1.00 (0.72–1.38)</td>
<td>0.997</td>
<td>1.06 (0.77–1.46)</td>
<td>0.734</td>
<td>1.05 (0.76–1.44)</td>
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<td>Most advantaged</td>
<td>1.05 (0.68–1.63)</td>
<td>0.821</td>
<td>1.11 (0.71–1.75)</td>
<td>0.647</td>
<td>1.41 (0.90–2.23)</td>
<td>0.137</td>
<td>1.50 (0.96–2.36)</td>
<td>0.076</td>
<td>1.51 (0.97–2.37)</td>
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<td>Level of education</td>
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<td>High education</td>
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<tr>
<td>Low education</td>
<td>1.16 (0.92–1.47)</td>
<td>0.206</td>
<td>0.93 (0.72–1.19)</td>
<td>0.57</td>
<td>0.84 (0.66–1.08)</td>
<td>0.181</td>
<td>0.83 (0.65–1.07)</td>
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<td>Main occupation class</td>
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<tr>
<td>Low</td>
<td>1.83 (1.46–2.29)</td>
<td>&lt;0.001</td>
<td>1.72 (1.37–2.15)</td>
<td>&lt;0.001</td>
<td>1.67 (1.34–2.10)</td>
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<td>Satisfaction with income</td>
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<td>Easy</td>
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<tr>
<td>Fairly easy</td>
<td>1.21 (0.97–1.51)</td>
<td>0.087</td>
<td>1.17 (0.94–1.46)</td>
<td>0.152</td>
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<tr>
<td>With some difficulty</td>
<td>1.85 (1.42–2.40)</td>
<td>&lt;0.001</td>
<td>1.72 (1.33–2.23)</td>
<td>0.001</td>
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<tr>
<td>With great difficulty</td>
<td>3.22 (2.30–4.52)</td>
<td>&lt;0.001</td>
<td>2.81 (2.01–3.93)</td>
<td>&lt;0.001</td>
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<td>Health behaviors</td>
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<td>3.08 (2.22–4.29)</td>
<td>&lt;0.001</td>
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</table>

ACE, adverse childhood experiences; SEC, socioeconomic circumstances. All models are adjusted for country of residence, birth cohort, attrition, height, and childhood health problems. Interaction terms between ACE and age (model 1b), level of education and age (model 2b), main occupation class and age (model 3b), satisfaction with household income and age (model 4b), and health behaviors and age (models 5b) were also tested. All the interactions were nonsignificant. Model 0 was only adjusted for height. Health behaviors included physical inactivity, unhealthy eating, smoking, and alcohol consumption (higher score indicating higher number of unhealthy behaviors).
long-lasting consequences on health. Difficult life events, such as parental drinking abuse or separation from caregivers, are associated with higher levels of stress that can permanently disrupt the ability to be healthy in older age.

Disadvantaged socioeconomic circumstances and unhealthy-related behaviors during adulthood were associated with an increased risk of LMS for both genders. These associations between adulthood socioeconomic circumstances and physical performance are consistent with previous studies [45] and with the stress hypothesis, as disadvantaged adulthood socioeconomic circumstances strongly correlate with chronic stress [43, 44]. In addition, the results for unhealthy behaviors were expected as physical inactivity, unhealthy eating, smoking, and alcohol consumption have an adverse impact on the musculoskeletal system. These findings confirm that adopting healthy lifestyle behaviors, such as physical activity, may improve physical function even at older age [46]. They also reinforce the importance of adopting a life course approach to better understand (un)healthy aging [47].

Strengths and Weaknesses
To the best of our knowledge, this study was the first to assess the association between ACE and an objective measure of muscle strength in older age. Our longitudinal study used a large sample of men and women aged 50–96 years from 14 European countries. We examined the risk of LMS in relation to ACE after taking into account childhood socioeconomic circumstances, thereby reducing the effect of confounders related to early-life conditions. In 2050, one in 4 Europeans should be aged 65 years and older [48]. These findings of a unique association between adversities in early life and a biomarker of health in older age are important in this context of an aging population.

However, there are some limitations that should be noted. First, ACE information was retrospectively self-reported during adult life, leading to potential recall bias. Previous studies showed that retrospective recall measures of exposure to adverse life events in childhood showed satisfactory validity [49] and that retrospective measures of adversity during childhood underestimated the association with objectively assessed health outcomes [50]. Therefore, the associations observed between ACE and our objective measure emerged despite this bias associated with the retrospective nature of the design, not because of it. In contrast, this bias requires cautious interpretation of the nonsignificant associations, especially among men. Second, because of the longitudinal design (i.e., 6 waves of measurement over 12 years) and the old age of the participants, a selection bias due to attrition cannot be excluded.

To minimize this bias, we performed statistical analyses adjusted for attrition (i.e., for participants who died or dropped out during the follow-up) and conducted sensitivity analyses without participants who died or dropped out during the follow-up, which yielded similar results. Third, the adverse events available in SHARE to construct the ACE score were not exactly the same as those used in the previous literature [14]. Thus, comparison of our results with previous studies should be done cautiously. Fourth, although grip strength has been shown to predict a range of adverse health outcomes [2–7] and is considered as a biomarker of aging [1], using other indicators of physical performance (e.g., chair stand, walking speed, or balance) should draw a broader picture of the individual’s physical capability. Finally, although main occupational position during adult life is associated with the physical job demands (i.e., higher skill levels require less physical or manual tasks than lower skill levels), the objective level of exercise during daily work was not controlled for in this study. Assessing the association between objective daily-work physical demands and grip strength will be important in future research.

Conclusion and Policy Implications
Our results reveal that ACE are associated with an increased risk of LMS in older women independently of childhood socioeconomic circumstances. This risk is not offset by the adoption of healthy behaviors or an improvement in socioeconomic circumstances in adulthood. These findings reveal that in women, childhood events out of their control are linked to long-term grip strength, a biomarker of healthy aging, thereby suggesting that a tougher start in life has a direct and long-lasting effect on women’s health. They reinforce the importance of considering the public health implications of ACE and the necessity to develop both primary prevention programs aimed at reducing them [12] and secondary prevention interventions to mitigate their negative impact over the life course. However, as this is the first study to assess the association between ACE and an objective measure of muscle strength in older age, further research is needed to confirm our findings.

Acknowledgments
This paper uses data from SHARE waves 1, 2, 3 (SHARELIFE), 4, 5, and 6 (DOI: 10.6103/SHARE.w1.600, 10.6103/SHARE.w2.600, 10.6103/SHARE.w3.600, 10.6103/SHARE.w4.600, 10.6103/SHARE.w5.600, and 10.6103/SHARE.w6.600).
Disclosure Statement

The authors declare no conflict of interest.

Funding Sources

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Statement of Ethics

This study was part of the SHARE study, approved by the relevant research ethics committees in the participating countries, and all participants provided written informed consent.

Author Contributions

B.C. and S.C. designed the analyses. B.C. analyzed the data. B.C., S.C., and M.P.B. drafted the manuscript. All authors critically appraised and approved the final version of the manuscript.

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7 Cooper R, Kuh D, Hardy R; Mortality Review Group; FALCon and HALCyon Study Teams. Objectively measured physical capability levels and mortality: systematic review and meta-analysis. BMJ. 2010 Sep 9;341:c4467.
Adverse Childhood Experiences and Elderly Muscle Strength


