



Relationships Between Depressive Symptoms, Dietary Inflammatory Potential, and Sarcopenia: Mediation Analyses

Guo-Qiang Chen^{1,2}, Gang-Pu Wang³ and Ying Lian^{1,2*}

¹ Department of Health Management and Shandong Engineering Laboratory for Health Management, The First Affiliated Hospital of Shandong First Medical University & Shandong Provincial Qianfoshan Hospital, Jinan, China, ² Department of Medical Record Management and Statistics, Shandong Provincial Qianfoshan Hospital, The First Affiliated Hospital of Shandong First Medical University, Jinan, China, ³ Department of General Surgery, The Fourth People's Hospital of Jinan City, Jinan, China

Background: Sarcopenia is a major public health problem. Depressive symptoms and dietary inflammatory potential play important roles in the development of sarcopenia. We aimed to disentangle the relationships between depressive symptoms, dietary inflammatory potential, and sarcopenia.

OPEN ACCESS

Edited by:

Roberta Zupo, National Institute of Gastroenterology S. de Bellis Research Hospital (IRCCS), Italy

Reviewed by:

Madia Lozupone, University of Bari Aldo Moro, Italy Sara De Nucci, National Institute of Gastroenterology "Saverio de Bellis" Research Hospital, Italy

> *Correspondence: Ying Lian lianying525@sina.com

Specialty section:

This article was submitted to Nutritional Epidemiology, a section of the journal Frontiers in Nutrition

Received: 29 December 2021 Accepted: 14 January 2022 Published: 17 February 2022

Citation:

Chen G-Q, Wang G-P and Lian Y (2022) Relationships Between Depressive Symptoms, Dietary Inflammatory Potential, and Sarcopenia: Mediation Analyses. Front. Nutr. 9:844917. doi: 10.3389/fnut.2022.844917 **Methods:** A total of 6,082 participants from the National Health and Nutrition Examination Survey (NHANES) were included in the analyses. Sarcopenia was defined according to the Foundation for the National Institutes for Health (FNIH) criteria. The Depressive symptoms were assessed using the nine-item Patient Health Questionnaire (PHQ-9). Dietary Inflammatory Index (DII) was calculated based on 24-h dietary recall interview. Two sets of mediation models were constructed separately.

Results: Depressive symptoms and DII were associated with sarcopenia, with odds ratios [ORs] (95% CIs) 2.54 (1.27, 5.13) and 1.17 (1.00, 1.37), respectively. DII score mediated the association of depressive symptoms with low muscle mass, explaining a total of 10.53% of the association (indirect effect = 0.004). Depressive symptoms had a significant mediating effects on the association between DII with low muscle mass, explaining a total of 12.50% of the association (indirect effect = 0.001).

Conclusions: Our findings suggested that both depressive symptoms and dietary inflammatory potential had direct effects, and indirect effects on low muscle mass, handgrip strength, muscle mass, through each other. It provides important insights into integrated nutritional and psychological intervention strategies in preventing sarcopenia.

Keywords: mediation analyses, sarcopenia, diet, inflammation, depressive symptoms

INTRODUCTION

Sarcopenia, a syndrome characterized by a progressive and general loss of muscle mass and strength, is associated with increased adverse outcomes, such as falls, functional decline, frailty, and mortality (1–3). It occurs commonly as an age-related process, with the prevalence of sarcopenia rising in the era of aging (4). Muscle mass gradually decreases from 30 years old and accelerates

1

gradually as individual ages, accompanied by low muscle strength or poor physical performance (5). A systematic review indicated that sarcopenia often co-occurs with depression (6). Both conditions, separately, are associated with a range of deleterious consequences, whereas the comorbidity can result in particularly worse outcomes (7). Therefore, it is imperative to identify the exact association between depressive symptoms and sarcopenia and explore the potential mechanisms underlying the association.

Existing epidemiological studies have explored the association between depressive symptoms and sarcopenia, with some reporting an increased risk associated with depressive symptoms (8). Regarding the potential mechanisms, depressive symptoms and sarcopenia share common pathophysiological pathways related to neurotrophins, inflammation, and oxidative stress, in which inflammation is one of the most frequently cited mechanisms (9, 10). For example, an increased level of inflammatory cytokines could lead to the neuroinflammation, which contributes to depression (11). Meanwhile, increased inflammatory cytokine levels could trigger the neuroinflammation pathways involved in pathophysiological processes of sarcopenia (12). Evidence is accumulating that both depressive symptoms and inflammation are potentially modifiable status and can be improved through effective strategies (13, 14). Thus, contextualizing chronic inflammation within broader biopsychosocial models of sarcopenia would provide further insights into the development of preventive and therapeutic targets (15).

Levels of inflammation can be modulated by lifestyle behaviors, such as diet. Recent studies have indicated that nutritional assessment and management may have a potential implication in preventing sarcopenia (16). The Dietary Inflammatory Index (DII) was designed to determine the inflammatory potential of the overall diet (17, 18). Moreover, existing studies conducted in older adults have demonstrated that a higher DII score positively correlates with an increased risk of sarcopenia (19). Simultaneously, the inflammatory potential of diet may associate bi-directionally with depressive symptoms. One recent longitudinal study showed that an anti-inflammatory diet is associated with a lower risk of depression (18). In addition, mental health, such as depressive symptoms, has been found to affect the dietary choice or preference, causing more consumption of nutrients related to low diet quality and higher DII (20, 21). Given these findings on the close relationships between depressive symptoms and DII and between DII and sarcopenia, it is reasonable to hypothesize that there may be mediating effects of depressive symptoms and DII on the development of sarcopenia and differential relationships between depressive symptoms and sarcopenia among individuals with different DII level.

Therefore, data from the National Health and Nutrition Examination Survey (NHANES) were used to (1) report the relationships between depressive symptoms, DII, and sarcopenia, (2) explore the mediating effect of DII and depressive symptoms on the development of sarcopenia, and (3) examine the relationships between depressive symptoms and sarcopenia among individuals with different DII level.

METHODS

Study Sample

The NHANES is a nationally representative survey on the US population aimed to assess health and nutrition status. The periodic cross-sectional surveys were conducted every 2 years using a stratified multistage clustered probability sampling approach by the US National Center for Health Statistics (NCHS). Participants completed questionnaires, underwent a medical examination, and provided fasted blood samples. The details of NHANES are available elsewhere (22). The survey was approved by the NCHS Institutional Review Board (Protocol #2011-17). All informed consents had been obtained from participants (23).

Data from two NHANES cycles 2011–2014 were enrolled in the present study. Participants being under 18 years old and over 60 years old (n = 11,586) were excluded. We further excluded participants with missing data on sarcopenia (n =1,757), depressive symptoms (n = 341), and DII (n = 165). The final analytical sample thus included a total of 6,082 participants. The flow chart of the study sample was shown in **Supplementary Figure 1**.

Assessment of Sarcopenia

Sarcopenia was defined as the presence of weakness and low muscle mass according to the Foundation for the National Institutes for Health (FNIH) criteria (24). The appendicular lean mass (ALM) was assessed by dual-energy x-ray absorptiometry (QDR-4500 Hologic Scanner Bedford, MA, USA). ALM was the sum of lean mass from both arms and legs. Low muscle mass was identified as the ALM adjusted for body mass index (BMI) <0.512 for women and 0.789 for men. Handgrip strength was assessed by using a handheld dynamometer. Participants were asked to exert maximum effort three times for each hand, and the highest value measured was used. Relative grip strength was calculated as the relative grip strength <1.00 for men and 0.56 for women.

Assessment of Depressive Symptoms

Depressive symptoms were assessed using the nine-item Patient Health Questionnaire (PHQ-9). The PHQ-9 is composed of items related to symptoms of depression, which has been well-validated in previous studies (25). Each item was scored from "0" (not at all) to "3" (nearly every day), and the total score ranged from 0 to 27, with higher scores indicating higher levels of depressive symptoms. A cutoff score of ≥ 10 was used to define the presence of depressive symptoms.

Assessment of DII

The DII was computed based on the dietary intake data gathered by 24-h dietary recall. We calculated the DII score for 27 food parameters available, such as carbohydrate, energy, protein, fat, fiber, cholesterol, saturated fatty acids, monounsaturated fatty acids, polyunsaturated fatty acids, β -carotene, vitamins A, B1, B2, B6, B12, C, D, and E, folic acid, iron, magnesium, zinc, selenium,



omega-3, and omega-6 polyunsaturated fatty acids, alcohol, and caffeine. The detail of the DII calculation method developed by Shivappa et al. is available elsewhere (17).

Covariates

The following sociodemographic health-related and characteristics were included in analyses: age, sex, race (non-Hispanic White, Non-Hispanic Black, Mexican American, Other Hispanic, and other race), educational level (below high school, high school and above), marriage status (married/living with partner, widowed/divorced/separated/never married), family poverty income ratio, smoking status (never, current, and former), drinking status (no, yes), physical activity level (low, moderate, and high), BMI status (underweight/normal: \leq 24.9 kg/m², overweight/obese: >25 kg/m²), diabetes, and hypertension. Diabetes was defined as (1) a self-reported previous diagnosis by healthcare professionals, (2) fasting plasma glucose level of 7.0 mmol/L or higher, (3) HbA1c concentration of 6.5% or higher, or (4) use of glucose-lowering medications (insulin or oral hypoglycemic medications). Hypertension was defined as the average systolic blood pressure ${\geq}140$ mmHg or diastolic blood pressure ${\geq}90$ mmHg, or use of anti-hypertensive medication.

Statistical Analyses

The characteristics of the study population were analyzed using Student's t-tests and Chi-square tests for continuous and categorical variables, respectively. The correlations between DII score and depressive symptoms score with the sarcopenia clinical measures were assessed by the Spearman correlation coefficients. Logistic regression models were used to explore the relationships between depressive symptoms and DII with sarcopenia. Linear regression models were used to explore the relationships between depressive symptoms and DII with the measurements of sarcopenia. For dichotomous outcome variables, the models worked out odds ratios (ORs) and 95% CIs, and for continuous variables, the models worked out ßs and 95% CIs. Several confounders, such as age, sex, race, educational level, marriage status, family poverty income ratio, smoking status, drinking status, physical activity level, BMI status, diabetes, and hypertension, were controlled in multivariable regression models. Two sets of mediation models

TABLE 1 | Sample characteristics stratified by depressive symptoms.

Age (years) 37.22 ± 12.52 39.89 ± 12.72 36.96 ± 12.47 < 0.01 Sex (%) < 0.01 Male $3,162 (51.99)$ $197 (37.24)$ $2,965 (53.39)$ Female $2,920 (48.01)$ $332 (62.76)$ $2,588 (46.61)$ Non-hispanic white $2,392 (39.33)$ $247 (46.69)$ $2,145 (38.63)$ Non-hispanic black 1400 (23.02) 120 (22.68) $1,280 (23.05)$ Mexican-American 779 (12.80) $49 (9.26)$ $730 (13.14)$ Other rispanic 541 (8.90) 59 (11.15) 482 (8.68) Other risce $970 (15.95)$ 54 (10.22) 916 (16.50) Educational level (%) < 0.01 Mariage status (%) < 0.01 Mariage status (%) < < 0.01 Mariage visitus (%) < < 0.01 Mariage status (%) < < 0.01 Sinsking status (%) <t< th=""><th>Characteristics</th><th>Total <i>N</i> = 6,082</th><th>Depressive symptoms n = 529</th><th>Non-depressive symptoms n = 5,553</th><th>Р</th></t<>	Characteristics	Total <i>N</i> = 6,082	Depressive symptoms n = 529	Non-depressive symptoms n = 5,553	Р
Sex (%) -0.01 Male 3,162 (51.99) 197 (37.24) 2,965 (63.39) Female 2,902 (80.10) 2,958 (46.61) -0.01 Race (%) -0.01 -0.01 -0.01 Non-hispanic white 2,392 (39.33) 247 (46.69) 2,145 (38.63) -0.01 Non-hispanic black 1400 (23.02) 120 (22.68) 1,280 (23.05) -0.01 Mexican-American 779 (12.80) 49 (9.26) 730 (13.14) -0.01 Other rispanic 30 (15.93) 64 (10.22) 916 (16.50) -0.01 Educational level (%) 50 (11.15) 482 (8.68) -0.01 Bidy school and above 1,010 (17.0) 145 (27.47) 833 (16.08) -0.01 Mariage status (%)	Age (years)	37.22 ± 12.52	39.89 ± 12.72	36.96 ± 12.47	<0.01
Male 3,162 (51.99) 197 (37.24) 2,965 (53.39) Female 2,920 (48.01) 332 (62.76) 2,588 (46.61) Race (%)	Sex (%)				<0.01
Fendel 2,920 (48.01) 323 (62.76) 2,588 (46.10) Race (%) <.001	Male	3,162 (51.99)	197 (37.24)	2,965 (53.39)	
Race (%) -<	Female	2,920 (48.01)	332 (62.76)	2,588 (46.61)	
Non-hispanic white 2,392 (39.33) 247 (46.69) 2,145 (38.63) Non-hispanic black 1400 (23.02) 120 (22.68) 1,280 (23.05) Mexican-American 779 (12.80) 49 (9.26) 730 (13.14) Other hispanic 541 (8.09) 561 (10.2) 482 (8.68) Other race 970 (15.95) 54 (10.2) 482 (8.68) Educational level (%) 504 (82.90) 384 (72.53) 4.660 (83.92) Below high school 1,010 (17.10) 145 (27.47) 833 (16.08) Marriage status (%) 5042 (82.90) 384 (72.53) 4.660 (83.92) Marriage status (%) 5042 (82.90) 384 (72.53) 4.660 (83.92) Marriage status (%) 3.487 (57.34) 230 (43.43) 3.260 (58.70) Mexican-ducly (%) 2.595 (42.66) 299 (56.57) 2.233 (41.30) Marriage status (%) 3.487 (57.34) 230 (43.43) 3.260 (58.70) Seminy poverty income ratio (%) 2.595 (42.66) 2.98 (39.24) 1,361 (24.51) 1-1.84 1,340 (22.03) 156 (29.58) 1,183 (21.31) 21.85	Race (%)				<0.01
Non-hispanic black 1400 (23.02) 120 (22.68) 1,280 (23.05) Mexican-American 779 (12.80) 49 (9.26) 730 (13.14) Other hispanic 541 (8.90) 59 (11.15) 482 (8.68) Other race 970 (15.95) 51 (10.22) 916 (16.50) Educational level (%) <	Non-hispanic white	2,392 (39.33)	247 (46.69)	2,145 (38.63)	
Mexican-American 779 (12.80) 49 (9.26) 730 (13.14)Other hispanic 541 (8.90) 59 (11.15) 482 (8.68)Other race 970 (15.95) 54 (10.22) 916 (16.50)Educational level (%) 1010 (17.10) 145 (27.47) 893 (16.08)High school and above 502 (28.20) 384 (72.53) 4.660 (83.92)Marriage status (%) $2,595$ (28.06) 299 (65.7) $2,293$ (4.30)Marriage status (%) $2,595$ (26.68) 299 (65.7) $2,293$ (4.30)Marriage line partner $3,487$ (57.34) 230 (43.43) $3,260$ (58.70)Widowed/divorced/separated/never $2,595$ (26.68) 299 (65.7) $2,293$ (4.30)married $3,407$ (26.20) 299 (65.7) $2,293$ (4.30)1-1.84 $1,569$ (25.80) 208 (39.24) $1,361$ (24.51)1-1.84 $1,340$ (22.03) 166 (29.58) $1,183$ (21.31)21.85 $3,713$ (52.17) 156 (31.18) $3,909$ (54.18)21.86 $3,672$ (60.38) 210 (39.77) $3,463$ (62.38)Smoking status (%) $1,262$ (63.84) 210 (39.77) $3,463$ (62.38)Current 844 (16.18) 95 (17.83) 889 (16.00)Former $1,426$ (23.44) 224 (23.01) $1,201$ (21.62)Dirking status (%) $1,262$ (63.84) $1,201$ (21.62)No $4,565$ (49.29) 413 (78.14) $4,143$ (74.61)Yang $3,260$ (56.70) $3,200$ (56.70)Yang $3,652$ (56.08) 162 (13.80) $3,200$ (56.70)	Non-hispanic black	1400 (23.02)	120 (22.68)	1,280 (23.05)	
Other hispanic 541 (8.90) 59 (11.15) 482 (8.68)Other race970 (15.95) 54 (10.22)916 (16.50)Educational level (%)1.010 (17.10) 145 (27.47) 893 (16.08)High school and above 5.042 (82.90) 384 (72.53) $4,660$ (83.92)Marriage status (%) 5.042 (82.90) 384 (72.53) $4,660$ (83.92)Marriage status (%) 5.042 (82.90) 230 (43.43) 3.260 (58.70)Marriad/living with partner 3.487 (57.34) 230 (43.43) 3.260 (58.70)Widowed/separated/never 2.595 (42.66) 299 (56.7) 2.933 (41.30)married1 596 (22.03) 1.66 (29.58) 1.361 (24.51)1-1.84 1.340 (22.03) 156 (29.58) 1.183 (21.31) ≥ 1.85 3.73 (52.17) 165 (31.18) 3.096 (54.18)Smoking status (%) 517.93 889 (16.00)Current 984 (16.18) 95 (17.93) 889 (16.00)Formar 1.426 (23.44) 224 (42.30) 1.201 (21.62)Diriking status (%) 517.93 889 (16.00)Formar 1.426 (23.44) 224 (42.30) 1.201 (21.62)Diriking status (%) 517.93 889 (16.00)Formar 1.426 (23.44) 224 (42.30) 1.201 (21.62)Diriking status (%) 506 (19.2) 1.016 (25.08) 1.410 (25.39)	Mexican-American	779 (12.80)	49 (9.26)	730 (13.14)	
Other race 970 (15.96) 54 (10.22) 916 (16.50) Educational level (%) 1,010 (17.10) 145 (27.47) 893 (16.08) High school and above 5,042 (82.90) 384 (72.53) 4,660 (83.92) Marriage status (%) 3,487 (57.34) 230 (43.43) 3,260 (58.70) Vidrowed/divorced/separated/never 2,595 (42.66) 299 (56.57) 2,233 (41.30) Marriage status (%)	Other hispanic	541 (8.90)	59 (11.15)	482 (8.68)	
Educational level (%)<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<<< <td>Other race</td> <td>970 (15.95)</td> <td>54 (10.22)</td> <td>916 (16.50)</td> <td></td>	Other race	970 (15.95)	54 (10.22)	916 (16.50)	
Below high school1,010 (17.10)145 (27.47)893 (16.08)High school and above5,042 (82.90)384 (72.53)4,660 (83.92)Mariage status (%)230 (43.43)3,260 (58.70)Marined/living with partner3,487 (57.34)230 (43.43)3,260 (58.70)Vidowed/divoreed/separated/never2,595 (42.66)299 (56.57)2,293 (41.30)married2299 (56.57)2,293 (41.30)Family poverty income ratio (%) </td <td>Educational level (%)</td> <td></td> <td></td> <td></td> <td><0.01</td>	Educational level (%)				<0.01
High school and above 5,042 (82.90) 384 (72.53) 4,660 (83.92) Marriage status (%) <0.01	Below high school	1,010 (17.10)	145 (27.47)	893 (16.08)	
Marriage status (%) <0.01	High school and above	5,042 (82.90)	384 (72.53)	4,660 (83.92)	
Married/living with partner 3,487 (57.34) 230 (43.43) 3,260 (58.70) Widowed/divorced/separated/never 2,595 (42.66) 299 (56.57) 2,293 (41.30) married - - - - Family poverty income ratio (%) - - - - - ≤1 1,569 (25.80) 208 (39.24) 1,361 (24.51) -	Marriage status (%)				<0.01
Widowed/divorced/separated/never 2,595 (42.66) 299 (56.57) 2,293 (41.30) Family poverty income ratio (%)	Married/living with partner	3,487 (57.34)	230 (43.43)	3,260 (58.70)	
Family poverty income ratio (%)<<< </td <td>Widowed/divorced/separated/never married</td> <td>2,595 (42.66)</td> <td>299 (56.57)</td> <td>2,293 (41.30)</td> <td></td>	Widowed/divorced/separated/never married	2,595 (42.66)	299 (56.57)	2,293 (41.30)	
≤11,569 (25.80)208 (39.24)1,361 (24.51)1-1.841,340 (22.03)156 (29.58)1,183 (21.31)≥1.853,173 (52.17)165 (31.18)3,009 (54.18)Smoking status (%)<	Family poverty income ratio (%)				<0.01
1-1.841,340 (22.03)156 (29.58)1,183 (21.31)≥1.853,173 (52.17)165 (31.18)3,009 (54.18)Smoking status (%) </td <td>≤1</td> <td>1,569 (25.80)</td> <td>208 (39.24)</td> <td>1,361 (24.51)</td> <td></td>	≤1	1,569 (25.80)	208 (39.24)	1,361 (24.51)	
≥1.853,173 (52.17)165 (31.18)3,009 (54.18)Smoking status (%)<0.01	1–1.84	1,340 (22.03)	156 (29.58)	1,183 (21.31)	
Smoking status (%) < <<0.01 Never 3,672 (60.38) 210 (39.77) 3,463 (62.38) Current 984 (16.18) 95 (17.93) 889 (16.00) Former 1,426 (23.44) 224 (42.30) 1,201 (21.62) Drinking status (%) 0.07 No 4,556 (74.92) 413 (78.14) 4,143 (74.61) Yes 1,526 (25.08) 116 (21.86) 1,410 (25.39)	≥1.85	3,173 (52.17)	165 (31.18)	3,009 (54.18)	
Never 3,672 (60.38) 210 (39.77) 3,463 (62.38) Current 984 (16.18) 95 (17.93) 889 (16.00) Former 1,426 (23.44) 224 (42.30) 1,201 (21.62) Drinking status (%) 0.07 No 4,556 (74.92) 413 (78.14) 4,143 (74.61) Yes 1,526 (25.08) 116 (21.86) 1,410 (25.39)	Smoking status (%)				<0.01
Current 984 (16.18) 95 (17.93) 889 (16.00) Former 1,426 (23.44) 224 (42.30) 1,201 (21.62) Drinking status (%) 0.07 No 4,556 (74.92) 413 (78.14) 4,143 (74.61) Yes 1,526 (25.08) 116 (21.86) 1,410 (25.39)	Never	3,672 (60.38)	210 (39.77)	3,463 (62.38)	
Former 1,426 (23.44) 224 (42.30) 1,201 (21.62) Drinking status (%)	Current	984 (16.18)	95 (17.93)	889 (16.00)	
Drinking status (%) 0.07 No 4,556 (74.92) 413 (78.14) 4,143 (74.61) Yes 1,526 (25.08) 116 (21.86) 1,410 (25.39)	Former	1,426 (23.44)	224 (42.30)	1,201 (21.62)	
No4,556 (74.92)413 (78.14)4,143 (74.61)Yes1,526 (25.08)116 (21.86)1,410 (25.39)	Drinking status (%)				0.07
Yes 1,526 (25.08) 116 (21.86) 1,410 (25.39)	No	4,556 (74.92)	413 (78.14)	4,143 (74.61)	
	Yes	1,526 (25.08)	116 (21.86)	1,410 (25.39)	
Physical activity level (%) <0.01	Physical activity level (%)				<0.01
Low 2,549 (41.91) 291 (55.13) 2,257 (40.65)	Low	2,549 (41.91)	291 (55.13)	2,257 (40.65)	
Moderate 1,905 (31.33) 139 (26.24) 1,767 (31.81)	Moderate	1,905 (31.33)	139 (26.24)	1,767 (31.81)	
High 1,628 (26.76) 99 (18.63) 1,529 (27.54)	High	1,628 (26.76)	99 (18.63)	1,529 (27.54)	
BMI status <0.01	BMI status				<0.01
Underweight/normal 2,044 (33.61) 147 (27.79) 1,897 (34.16)	Underweight/normal	2,044 (33.61)	147 (27.79)	1,897 (34.16)	
Overweight/obese 4,038 (66.39) 382 (72.21) 3,656 (65.84)	Overweight/obese	4,038 (66.39)	382 (72.21)	3,656 (65.84)	
Hypertension (%) 640 (10.53) 73 (13.80) 567 (10.21) 0.01	Hypertension (%)	640 (10.53)	73 (13.80)	567 (10.21)	0.01
Diabetes (%) 593 (9.75) 87 (16.45) 506 (9.16) <0.01	Diabetes (%)	593 (9.75)	87 (16.45)	506 (9.16)	< 0.01
Muscle mass 0.82 ± 0.21 0.74 ± 0.20 0.83 ± 0.21 <0.01	Muscle mass	0.82 ± 0.21	0.74 ± 0.20	0.83 ± 0.21	< 0.01
Handgrip strength 1.46 ± 0.49 1.27 ± 0.48 1.48 ± 0.49 <0.01	Handgrip strength	1.46 ± 0.49	1.27 ± 0.48	1.48 ± 0.49	< 0.01
Sarcopenia (%) 70 (1.15) 16 (3.02) 54 (0.97) <0.01	Sarcopenia (%)	70 (1.15)	16 (3.02)	54 (0.97)	< 0.01
Low muscle mass (%) 536 (8.81) 74 (13.99) 462 (8.32) <0.01	Low muscle mass (%)	536 (8.81)	74 (13.99)	462 (8.32)	< 0.01
Weakness (%) 133 (2.19) 26 (4.91) 107 (1.93) <0.01	Weakness (%)	133 (2.19)	26 (4.91)	107 (1.93)	< 0.01
DII score 0.70 ± 1.89 1.42 ± 1.68 0.63 ± 1.90 <0.01	DII score	0.70 ± 1.89	1.42 ± 1.68	0.63 ± 1.90	< 0.01

were constructed and analyzed separately. In a mediation model, we included sarcopenia and its measurements as dependent variables, depressive symptoms as the independent variable, and the DII score as the mediating variable (see **Figure 1A**). We also investigated the mediating effect of

depressive symptoms on the association of DII with sarcopenia in another mediation model (see **Figure 1B**). We examined the association between depressive symptoms and sarcopenia stratified by DII categories. All analyses were conducted using Stata 14.0 software (StataCorp LP, College Station, TX, USA).

RESULTS

Descriptive and Correlation Analyses

Of the 6,082 participants, 51.99% were men and 48.01% were women, with a mean age of 37.22 years. The prevalence rate of depressive symptoms was 8.70% and sarcopenia 1.15%. **Table 1** shows the characteristics of participants according to depressive symptoms. Participants having depressive symptoms were likely to be older, women, having low education levels, living alone, and low family poverty income ratio. Regarding the health-related variables, lower physical activity, lower handgrip strength, lower muscle mass, sarcopenia, higher DII, and having diabetes and hypertension were significantly associated with depressive symptoms.

Table 2 presents the correlation coefficients between each pair of variables, such as sarcopenia measurements, depressive symptoms scores, and DII. Depressive symptoms scores were negatively correlated with the muscle mass (r = -0.14, p < 0.05) and handgrip strength (r = -0.17, p < 0.05). DII was negatively correlated with the muscle mass (r = -0.23, p < 0.05) and handgrip strength (r = -0.21, p < 0.05). While, depressive symptom scores were positively correlated with DII (r = 0.11, p < 0.05).

Relationships Between Depressive Symptoms, DII, and Sarcopenia

After adjustments for potential confounders, depressive symptoms were significantly associated with sarcopenia (OR = 2.54, 95% CI = 1.27, 5.13) and measurements of sarcopenia, such as weakness (OR = 1.89, 95% CI = 1.11, 3.21), low muscle mass (OR = 1.43, 95% CI = 1.05, 1.94), handgrip strength ($\beta = -0.05$, 95% CI = -0.08, -0.02), and muscle mass ($\beta = -0.02$, 95% CI = -0.03, -0.01). Furthermore, DII was also associated with sarcopenia (OR = 1.17, 95% CI = 1.00, 1.37) and the measurements of sarcopenia. More details are shown in **Table 3**.

Mediation Analyses

In the mediation analyses, the DII score mediated the association between depressive symptoms and low muscle mass, explaining a total of 10.53% of the association (indirect effect = 0.004). Meanwhile, significant mediating effects were established in the handgrip strength and muscle mass. Furthermore, there was a significant mediating effect of depressive symptoms on the association between DII and low muscle mass, explaining a total of 12.50% of the association (indirect effect = 0.001). Meanwhile, significant mediating effects were established for handgrip strength and muscle mass. The results of the mediation analyses are presented in **Table 4**.

Stratified Analyses

In the stratified analyses, depressive symptoms were significantly associated with sarcopenia among participants with the second tertile of DII, with the OR (95% CI) 7.99 (2.50, 25.55).

TABLE 2 | Correlations between variables.

Variables	Depressive symptoms scores	DII score	Muscle mass	Handgrip strength	
Depressive symptoms scores	1				
DII score	0.11*	1			
Muscle mass	-0.14*	-0.23*	1		
Handgrip strength	-0.17*	-0.21*	0.82*	1	

*p < 0.05.

Depressive symptoms were significantly associated with muscle mass, one of the measurements of sarcopenia, among participants with the highest tertile of DII, with the β (95% CI) -0.02 (-0.03, -0.01). The interaction test was not significant (p > 0.05). **Supplementary Table 1** shows the association between depressive symptoms and sarcopenia stratified by categories of DII.

DISCUSSION

This study disentangled the complex relationships between depressive symptoms, dietary inflammatory potential, and sarcopenia using a nationally representative sample of the US middle-aged adults. Depressive symptoms and dietary inflammatory potential were found to be significantly associated with sarcopenia and its clinical measurements. Of particular importance, there is a novel finding that DII score significantly mediated the association between depressive symptoms and low muscle mass, and there was a significant mediating effect of depressive symptoms on the association between DII and low muscle mass.

Our findings on the relationships between dietary inflammatory potential, depressive symptoms, and sarcopenia supported and extended those from previous studies. Most of the existing studies indicated that depression may be related to sarcopenia, whereas there are also inconsistent results (26-28). Meanwhile, there was a relative scarcity of research on the relationship between depressive symptoms with sarcopenia conducted in middle-aged adults compared to elderly adults previously (26), most of which had limitations related to the assessment of sarcopenia, such as only using one measurement of sarcopenia (29). Notably, the present study not only extended existing evidence but further highlighted the association between depression and sarcopenia in middle-aged adults, thereby further strengthening the possible application of our observation to a wider range of population. With respect to nutritional factors, though the link between diet and sarcopenia has been more widely studied, mainly focusing on nutritional status and specific nutrients (30), little is known about the role of dietary inflammatory potential, presenting a distinct biological mechanism. Our results suggested that the pro-inflammatory

TABLE 3 | Relationships between depressive symptoms, DII, and sarcopenia.

Variables	Crude	model	Adjusted model		
	Depressive symptoms	DII score	Depressive symptoms	DII score	
Sarcopenia OR (95%CI)	2.94 (1.66,5.21)*	1.11(0.97,1.27)	2.54 (1.27,5.13)*	1.17(1.00,1.37)*	
Weakness OR (95%Cl)	2.51 (1.62,3.92)*	1.11(0.96,1.17)	1.89 (1.11,3.21)*	1.05(0.94,1.18)	
Low muscle mass OR (95%Cl)	1.66 (1.27,2.16)*	1.11(1.06,1.17)*	1.43 (1.05,1.94)*	1.11(1.05,1.18)*	
Handgrip strength β (95%Cl) Muscle mass β (95%Cl)	-0.17 (-0.22, -0.13)* -0.07 (-0.09, -0.05)*	-0.05(-0.06, -0.04)* -0.02(-0.03, -0.02)*	-0.05 (-0.08, -0.02)* -0.02 (-0.03, -0.01)*	-0.01(-0.02, -0.004)* -0.01(-0.02, -0.005)*	

Adjusted model adjusted for age, sex, race, educational level, marriage status, family poverty income ratio, smoking status, drinking status, physical activity level, BMI status, diabetes, and hypertension. p < 0.05. DII, Dietary Inflammatory Index.

TABLE 4 | Mediation pathways among depressive symptoms, DII, and sarcopenia.

Independent variables	Mediator	Dependent variable	Exposure to mediator	Mediator to outcome	Direct effect	Indirect effect	Total effect	Proportion Mediated (%)
Depressive symptoms	DII	Sarcopenia	0.492*	0.002	0.015*	0.001	0.016*	6.25
		Weakness	0.492*	0.001	0.019*	0.001	0.020*	5.00
		Low muscle mass	0.492*	0.007*	0.034*	0.004*	0.038*	10.53
		Handgrip strength	0.492*	-0.006*	-0.047*	-0.003*	-0.050*	6.00
		Muscle mass	0.492	-0.006*	-0.016*	-0.003*	-0.019*	15.78
DII	Depressive symptoms	Sarcopenia	0.01*	0.02*	0.002	0.001*	0.003	33.33
		Weakness	0.01*	0.02*	0.002	0.001*	0.003	33.33
		Low muscle mass	0.01*	0.04*	0.007*	0.001*	0.008*	12.50
		Handgrip strength	0.01*	-0.05*	-0.007*	-0.001*	-0.008*	12.50
		Muscle mass	0.01*	-0.02*	-0.006*	-0.001*	-0.007*	14.29

Adjusted for age, sex, race, educational level, marriage status, family poverty income ratio, smoking status, drinking status, physical activity level, BMI status, diabetes, and hypertension p < 0.05. DII, Dietary Inflammatory Index.

diet, indicated by a higher DII score, was related to a greater odd of sarcopenia and its component of muscle mass. Dietrelated inflammation may affect muscle proteolysis and myocyte apoptosis, in turn leading to muscle loss and dysfunction (31). Overall, our findings provide convincing evidence on the hypothesized complex relationships and the necessity of clarifying the mechanisms underlying the relationships.

Mediation analyses suggested that there were significant mediating effects of depressive symptoms on the association between DII and low muscle mass, muscle mass, and hand grip strength. The finding indicated that a higher DII score was related to the higher risk of depressive symptoms, which in turn, led to sarcopenia independent of the direct effects of DII on sarcopenia. Similarly, our previous meta-analysis showed that a pro-inflammatory diet was associated with an increased risk of common mental health outcomes, such as depressive symptoms, anxiety, distress, and the association presented in a dose-response manner (32). Potential mechanisms

underlying the association between DII and depressive symptoms may include oxidant-antioxidant imbalance and modified gut microbiota composition and activity (33, 34). Furthermore, we also observed that DII significantly mediated the association of depressive symptoms with low muscle mass and handgrip strength. Even though inflammation has often been mentioned as a potential mechanism in the relationship between depressive symptoms and musculoskeletal health, the mediating effect is the first time to be quantified. Likewise, our finding also coincides with previous literature suggesting that a chronic pro-inflammatory signaling has been identified as a key pathway from depression to a range of health outcomes (35, 36). Reportedly, psychological health has consistently influenced one's dietary intake (37). For instance, depression is thought to disrupt the hypothalamic-pituitary-adrenal axis system, which elevates cortisol levels, encouraging increased consumption of energy-dense foods, which influences the dietary quality and the overall inflammatory potential (38). In line with the psycho-neuro-inflammatory theory, the present results improve our understanding the role of dietary and psychological factors in the pathogenesis of sarcopenia (39). In this sense, resolving both factors through efficient nutritional and psychological intervention might have potential to mitigate neuroinflammatory processes and prevent the development of sarcopenia.

Additionally, depressive symptoms were significantly associated with sarcopenia in the subgroup with a higher level of DII score in the stratified analyses. Previous studies suggested the moderating effect of inflammation on the association between depression and adverse health outcomes (40), such as all-cause mortality (41), to some certain degree supporting our findings. Due to the relatively small number of participants with sarcopenia affecting the statistical power, stratified analyses did not demonstrate significant association in individuals with the highest tertile of DII. Further research is necessary to replicate these findings in a larger sample. Although this result warrants further research, the findings suggest that the existence of depressive symptoms, together with a diet-induced inflammatory state, may be associated with higher odds of sarcopenia. Thus, it may be legitimate that integrated utilization of modulating dietary inflammatory potential and addressing depressive symptoms have important public health benefits in the prevention of sarcopenia.

Our study has several important strengths. First, this study included a nationally representative of the general population with a large sample size, which helps to provide convincing support of the hypothesized relationships. Second, the proposed models using mediation and stratified analyses incorporated psychological factors and lifestyle behaviors related to sarcopenia, which enables an insightful understanding of the inter-relationship of these factors with sarcopenia from a comprehensive perspective, thereby facilitating the development of targeted interventions aimed to prevent sarcopenia. Nevertheless, this study has some limitations that need to be mentioned. First, given the cross-sectional design, it is not possible to elucidate the causal associations among dietary inflammatory potential, depressive symptoms, and sarcopenia. Future studies with longitudinal design should seek to clarify the directionality and temporal relationships. Second, although several important confounding variables have been included in the models, there might be unmeasured covariates having a potential confounding effect on the associations. Future studies incorporating these measures are recommended to verify the present conclusion. Third, the mediation effects in our study were generally small. Considering that the etiology of sarcopenia is complex and multifactorial, other biological mechanisms, such as neuroendocrine responses (42, 43) and epigenetic changes (44), might also play a role in the pathophysiology of sarcopenia. It would be important to understand how these different biological processes influence each other. Moreover, future research is needed to incorporate other intermediate mechanisms in the biopsychosocial model of sarcopenia.

CONCLUSIONS

From a practical point of view, the findings of the present study during middle-aged adults may have significant clinical and public health implications. The observed associations provide a useful perspective for prevention and management of sarcopenia as it implies that sarcopenia is not only an age-related problem, but also one associated with a range of long-term conditions even early in mid-life. Comprehensive psychological and behavioral interventions, such as promoting an anti-inflammatory diet and improving mental health, have the potential for prevention and intervention of sarcopenia at earlier stages of life.

In summary, as hypothesized, depressive symptoms, and dietary inflammatory potential were found to be significantly associated with sarcopenia and its individual component low muscle mass. Specifically, both depressive symptoms and dietary inflammatory potential had significant direct effects, and indirect (mediation) effects on low muscle mass, handgrip strength, muscle mass, through each other in the adjusted mediation analyses. Therefore, integrated intervention strategies, such as promoting an anti-inflammatory diet and improving mental health, are suggested in the prevention of sarcopenia.

DATA AVAILABILITY STATEMENT

The raw data supporting the conclusions of this article will be made available by the authors, without undue reservation.

ETHICS STATEMENT

The studies involving human participants were reviewed and approved by the National Center for Health Statistics (NCHS) and approved by the NCHS Institutional Review Board (IRB). The patients/participants provided their written informed consent to participate in this study.

AUTHOR CONTRIBUTIONS

YL contributed to study concept, acquisition of data, and participated in critical revision of the manuscript. YL and G-QC participated in the analysis and interpretation of data. G-QC and G-PW contributed to drafting of the manuscript. All authors read and approved the final manuscript.

FUNDING

This work was supported by Shandong Provincial Key Research and Development Program (Grant no. 2019GSF108196), Center of China–US Sports Economics and Health Engineering of Shandong (Grant no. SDCA20191013), Academic Promotion Programme of Shandong First Medical University (Grant no. 2019QL013), and Shandong Provincial Soft Science Research Program (Grant no. 2020RKB14163). The funding sources had no role in study design, data analysis and interpretation of data, the writing of the manuscript, or the decision to submit the paper for publication.

ACKNOWLEDGMENTS

The authors thank the NHANES research team, the field team, and every respondent for the time and efforts that they have devoted to the NHANES project.

REFERENCES

- 1. Cruz-Jentoft AJ, Sayer AA. Sarcopenia. Lancet. (2019) 393:2636– 46. doi: 10.1016/S0140-6736(19)31138-9
- Brown JC, Harhay MO, Harhay MN. Sarcopenia and mortality among a population-based sample of community-dwelling older adults. J Cachexia Sarcopenia Muscle. (2016) 7:290–8. doi: 10.1002/jcsm.12073
- Zhang XM, Jiao J, Zhu C, Guo N, Liu Y, Lv D, et al. Association between low handgrip strength and 90-day mortality among older chinese inpatients: a national multicenter prospective cohort study. *Front Nutr.* (2021) 8:628628. doi: 10.3389/fnut.2021.628628
- Pang BWJ, Wee SL, Lau LK, Jabbar KA, Seah WT, Ng DHM, et al. Prevalence and associated factors of sarcopenia in Singaporean adults-the Yishun Study. J Am Med Dir Assoc. (2021) 22:885.e881– 885.e810. doi: 10.1016/j.jamda.2020.05.029
- Wilkinson DJ, Piasecki M, Atherton PJ. The age-related loss of skeletal muscle mass and function: measurement and physiology of muscle fibre atrophy and muscle fibre loss in humans. *Ageing Res Rev.* (2018) 47:123– 32. doi: 10.1016/j.arr.2018.07.005
- Chang KV, Hsu TH, Wu WT, Huang KC, Han DS. Is sarcopenia associated with depression? A systematic review and meta-analysis of observational studies. *Age Ageing*. (2017) 46:738–46. doi: 10.1093/ageing/afx094
- Park S, Cho J, Kim D, Jin Y, Lee I, Hong H, et al. Handgrip strength, depression, and all-cause mortality in Korean older adults. *BMC Geriatr.* (2019) 19:127. doi: 10.1186/s12877-019-1140-0
- Lian Y, Wang GP, Chen GQ, Jia CX. Bidirectional associations between handgrip strengthand depressive symptoms: a longitudinal cohort study. *J Am Med Dir Assoc.* (2021) 22:1744–50.e1741. doi: 10.1016/j.jamda.2021.04.006
- Picca A, Calvani R, Sirago G, Coelho-Junior HJ, Marzetti E. Molecular routes to sarcopenia and biomarker development: per aspera ad astra. *Curr Opin Pharmacol.* (2021) 57:140–7. doi: 10.1016/j.coph.2021.02.006
- Pasco JA, Williams LJ, Jacka FN, Stupka N, Brennan-Olsen SL, Holloway KL, et al. Sarcopenia and the common mental disorders: a potential regulatory role of skeletal muscle on brain function? *Curr Osteoporos Rep.* (2015) 13:351– 7. doi: 10.1007/s11914-015-0279-7
- Kim YK, Na KS, Myint AM, Leonard BE. The role of pro-inflammatory cytokines in neuroinflammation, neurogenesis and the neuroendocrine system in major depression. *Prog Neuropsychopharmacol Biol Psychiatry*. (2016) 64:277–84. doi: 10.1016/j.pnpbp.2015.06.008
- Budui SL, Rossi AP, Zamboni M. The pathogenetic bases of sarcopenia. *Clin Cases Miner Bone Metab.* (2015) 12:22–6. doi: 10.11138/ccmbm/2015.12.1.022
- Vikberg S, Sörlén N, Brandén L, Johansson J, Nordström A, Hult A, et al. Effects of resistance training on functional strength and muscle mass in 70year-old individuals with pre-sarcopenia: a randomized controlled trial. *J Am Med Dir Assoc.* (2019) 20:28–34. doi: 10.1016/j.jamda.2018.09.011
- 14. Lu Y, Niti M, Yap KB, Tan CTY, Zin Nyunt MS, Feng L, et al. Assessment of sarcopenia among community-dwelling at-risk frail adults aged 65 years and older who received multidomain lifestyle interventions: a secondary analysis of a randomized clinical trial. *JAMA Netw Open.* (2019) 2:e1913346. doi: 10.1001/jamanetworkopen.201 9.13346
- Engel GL. The clinical application of the biopsychosocial model. Am J Psychiatry. (1980) 137:535–44. doi: 10.1176/ajp.137.5.535
- 16. Zupo R, Castellana F, Bortone I, Griseta C, Sardone R, Lampignano L, et al. Nutritionaldomains in frailty tools: working towards an

SUPPLEMENTARY MATERIAL

The Supplementary Material for this article can be found online at: https://www.frontiersin.org/articles/10.3389/fnut.2022. 844917/full#supplementary-material

Supplementary Figure 1 | Flow chart of the study sample.

Supplementary Table 1 | Association between depressive symptoms and sarcopenia stratified by categories of DII.

operational definition of nutritional frailty. *Ageing Res Rev.* (2020) 64:101148. doi: 10.1016/j.arr.2020.101148

- Shivappa N, Steck SE, Hurley TG, Hussey JR, Hébert JR. Designing and developing a literature-derived, populationbased dietary inflammatory index. *Public Health Nutr.* (2014) 17:1689–96. doi: 10.1017/S1368980013002115
- Shivappa N, Schoenaker DA, Hebert JR, Mishra GD. Association between inflammatory potential of diet and risk of depression in middle-aged women: the Australian Longitudinal Study on Women's Health. *Br J Nutr.* (2016) 116:1077–86. doi: 10.1017/S0007114516002853
- Bagheri A, Soltani S, Hashemi R, Heshmat R, Motlagh AD, Esmaillzadeh A. Inflammatory potential of the diet and risk of sarcopenia and its components. *Nutr J.* (2020) 19:129. doi: 10.1186/s12937-020-00649-2
- Boutté AK, Turner-McGrievy GM, Wilcox S, Liu J, Eberth JM, Kaczynski AT. Associations of maternal stress and/or depressive symptoms with diet quality during pregnancy: a narrative review. *Nutr Rev.* (2021) 79:495– 517. doi: 10.1093/nutrit/nuaa019
- Yim HR, Yun HJ, Lee JH. An investigation on Korean adolescents' dietary consumption: focused on sociodemographic characteristics, physical health, and mental health. *Int J Environ Res Public Health*. (2021) 18:9773. doi: 10.3390/ijerph18189773
- Schorgg P, Bärnighausen T, Rohrmann S, Cassidy A, Karavasiloglou N, Kühn T. Vitamin B6 status among vegetarians: findings from a population-based survey. *Nutrients*. (2021) 13:1627. doi: 10.3390/nu13051627
- National Health and Nutrition Examination Survey (NHANES): Ethics Review Board (ERB) Approval. Available online at: https://www.cdc.gov/nchs/nhanes/ irba98.htm
- Dam TT, Peters KW, Fragala M, Cawthon PM, Harris TB, McLean R, et al. An evidence-based comparison of operational criteria for the presence of sarcopenia. A Biol Sci Med Sci. (2014) 69:584–90. doi: 10.1093/gerona/glu013
- Cassiani-Miranda CA, Cuadros-Cruz AK, Torres-Pinzón H, Scoppetta O, Pinzón-Tarrazona JH, López-Fuentes WY, et al. Validity of the Patient Health Questionnaire-9 (PHQ-9) for depression screening in adult primary care users in Bucaramanga, Colombia. *Rev Colomb Psiquiatr.* (2021) 50:11– 21. doi: 10.1016/j.rcpeng.2019.09.002
- Wang LT, Huang WC, Hung YC, Park JH. Association between depressive symptoms and risk of sarcopenia in Taiwanese older adults. J Nutr Health Aging. (2021) 25:790–4. doi: 10.1007/s12603-021-1631-5
- Endo T, Akai K, Kijima T, Kitahara S, Abe T, Takeda M, et al. An association analysis between hypertension, dementia, and depression and the phases of pre-sarcopenia to sarcopenia: a cross-sectional analysis. *PLoS ONE.* (2021) 16:e0252784. doi: 10.1371/journal.pone.0252784
- Byeon CH, Kang KY, Kang SH, Kim HK, Bae EJ. Sarcopenia is not associated with depression in Korean adults: results from the 2010-2011 Korean National Health and Nutrition Examination Survey. *Korean J Fam Med.* (2016) 37:37– 43. doi: 10.4082/kjfm.2016.37.1.37
- 29. Geng J, Deng L, Qiu S, Bian H, Cai B, Jin K, et al. Dietary inflammatory potential and risk of sarcopenia: data from national health and nutrition examination surveys. *Aging*. (2020) 13:1913–28. doi: 10.18632/aging.202141
- Liu X, Xia X, Hu F, Hou L, Jia S, Liu Y, et al. Nutrition status mediates the association between cognitive decline and sarcopenia. *Aging*. (2021) 13:8599– 610. doi: 10.18632/aging.202672
- Wang J, Leung KS, Chow SK, Cheung WH. Inflammation and age-associated skeletal muscle deterioration (sarcopaenia). J Orthop Translat. (2017) 10:94– 101. doi: 10.1016/j.jot.2017.05.006

- Chen GQ, Peng CL, Lian Y, Wang BW, Chen PY, Wang GP. Association between dietary inflammatory index and mental health: a systematic review and dose-response meta-analysis. *Front Nutr.* (2021) 8:662357. doi: 10.3389/fnut.2021.662357
- 33. Zheng J, Hoffman KL, Chen JS, Shivappa N, Sood A, Browman GJ, et al. Dietary inflammatory potential in relation to the gut microbiome: results from a cross-sectional study. *Br J Nutr.* (2020) 124:931–42. doi: 10.1017/S0007114520001853
- 34. Ruiz-Saavedra S, Salazar N, Suárez A, de Los Reyes-Gavilán CG, Gueimonde M, González S. Comparison of different dietary indices as predictors of inflammation, oxidative stress and intestinal microbiota in middle-aged and elderly subjects. *Nutrients*. (2020) 12:3828. doi: 10.3390/nu12123828
- 35. Kang J, Moser DK, Biddle MJ, Lennie TA, Smyth SS, Vsevolozhskaya OA. Inflammatory properties of diet mediate the effect of depressive symptoms on Framingham risk score in men and women: results from the National Health and Nutrition Examination Survey (2007-2014). *Nutr Res.* (2020) 74:78–86. doi: 10.1016/j.nutres.2019.11.008
- 36. Jeon YJ, Cho SMJ, Lee YJ, Kim HC, Jung SJ. Depressive symptoms, its sub-factors, and augmentation index: the modifying effects according to inflammatory markers. J Affect Disord. (2020) 272:380–7. doi: 10.1016/j.jad.2020.03.174
- Keck MM, Vivier H, Cassisi JE, Dvorak RD, Dunn ME, Neer SM, et al. Examining the role of anxiety and depression in dietary choices among college students. *Nutrients*. (2020) 12:2061. doi: 10.3390/nu12072061
- Errisuriz VL, Delfausse L, Villatoro AP, McDaniel MD, Esparza L, Parra-Medina D. Depression and physical activity affect diet quality of foreignborn latina women living on the U.S.-Mexico border. *Nutrients*. (2019) 11:1254. doi: 10.3390/nu11061254
- 39. Schoufour JD, Tieland M, Barazzoni R, Ben Allouch S, van der Bie J, Boirie Y, et al. The relevance of diet, physical activity, exercise, and persuasive technology in the prevention and treatment of sarcopenic obesity in older adults. *Front Nutr.* (2021) 8:661449. doi: 10.3389/fnut.2021.661449
- Wu Y, Zhang H, Wang C, Broekman BFP, Chong YS, Shek LP, et al. Inflammatory modulation of the associations between prenatal maternal depression and neonatal brain. *Neuropsychopharmacology*. (2021) 46:470– 7. doi: 10.1038/s41386-020-0774-0

- Castro-Costa E, Diniz BS, Firmo JOA, Peixoto SV, de Loyola Filho AI, Lima-Costa MF, et al. Diabetes, depressive symptoms, and mortality risk in old age: The role of inflammation. *Depress Anxiety*. (2019) 36:941– 9. doi: 10.1002/da.22908
- 42. Waters DL, Qualls CR, Dorin RI, Veldhuis JD, Baumgartner RN. Altered growth hormone, cortisol, and leptin secretion in healthy elderly persons with sarcopenia and mixed body composition phenotypes. J Gerontol A Biol Sci Med Sci. (2008) 63:536–41. doi: 10.1093/gerona/63. 5.536
- Petermann-Rocha F, Gray SR, Pell JP, Celis-Morales C, Ho FK. Biomarkers profile of people with sarcopenia: a crosssectional analysis from UK biobank. J Am Med Dir Assoc. (2020) 21:2017.e2011–e2019. doi: 10.1016/j.jamda.2020.05.005
- Brown DM, Goljanek-Whysall K. microRNAs: Modulators of the underlying pathophysiology of sarcopenia? Ageing Res Rev. (2015) 24:263-73. doi: 10.1016/j.arr.2015. 08.007

Conflict of Interest: The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

Publisher's Note: All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.

Copyright © 2022 Chen, Wang and Lian. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY). The use, distribution or reproduction in other forums is permitted, provided the original author(s) and the copyright owner(s) are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.