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EXPLORING THE "OBESITY PARADOX" AS A CORRELATE OF COGNITIVE AND PHYSICAL FUNCTION IN COMMUNITY-DWELLING BLACK AND WHITE OLDER ADULTS

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In Memory

With heavy hearts, the co-authors would like to acknowledge the untimely death of our lead author, Dr. Jeannine Skinner. We honor the life and service of Dr. Skinner who touched

the lives of many with her kind spirit as a mentor, colleague, scientist, and advocate for social justice; her energy, grace, and enthusiasm will be truly missed.

INTRODUCTION

Excess adiposity as characterized by overweight status and obesity is a major public health problem for all segments of the US population, including older adults. Recent epidemiological reports show one third of US older adults are obese, with the highest rates of obesity among older adults aged 65-74 years.¹ Among ethno-racial groups, obesity is highest among older African Americans and Hispan-

amine the underlying physiological processes linking adiposity to executive function in older adults. *Ethn Dis.* 2017;27(4):387-394; doi:10.18865/ed.27.4.387.

Keywords: Race; Body Mass Index; Cognition; Physical Function

¹Department of Psychological Science, University of North Carolina at Charlotte, Charlotte, NC ²School of Nursing, University of North Carolina at Charlotte, Charlotte, NC ic/Latinos. Differences in the prevalence of overweight status and obesity in these groups are a key contributor to racial and ethnic health disparities. Given the exponential growth of the minority older adult population and steady increase in overweight and obesity rates across the lifespan, it is important to improve our understanding of the association between excess adiposity and physical and cognitive health outcomes among different segments of the older adult population.

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Objective: The obesity paradox has been documented in aged populations, yet it remains unclear if this paradox persists for physical and cognitive outcomes in community-dwelling older adult populations. Our study examines associations between body mass index (BMI) classification, cognitive function, and physical function. We also investigate whether these associations are modified by race or age.

Design: Cross-sectional study.

Settings: Senior residential sites and community centers in Saint Louis, Missouri.

Participants: Study participants included 331 adults, aged >55 years. Age was stratified into young-old (aged 55-74 years) and older (aged ≥75 years).

Outcome Measures: Physical function was measured using the mini-Physical Performance Test (mini-PPT) and grip strength. Cognitive function was assessed with the Short Blessed Test (SBT) and the Trail Making Tests (TMT-A and TMT-B) performance.

Results: Older adults who were obese had significantly better cognitive flexibility (TMT-B) performance than normal weight older adults (P=.02), and this association was not influenced by age or race. Adiposity was not associated with psychomotor speed (TMT-A), general cognition (SBT), or measures of physical function (Ps>.05).

Conclusion: In a diverse sample of community-dwelling older adults, we found partial support for the controversial obesity paradox. Our results suggest excess adiposity may be protective for executive function processes. Future research is needed to ex-

Physical function encompasses a range of functional abilities including activities of daily living and mobility tasks. Physical function is a robust predictor of clinical outcomes such as hospitalization and mortality.^{2,3} Although excess adiposity in early life and adulthood has been linked to poorer physical function,⁴ associations in late-life have yielded less consistent results.^{5,6} Among aged populations, low body mass index

The aim of this study was to examine the relationship between weight classification, cognitive function and physical function, and to determine whether these associations were modified by race or stratified age in a sample of community-dwelling older adults.

(BMI), weight loss, and obesity are reported to be negatively associated with mobility, while overweight status may be protective against loss of mobility⁷ and disability.⁸ Researchers surmise that factors such as increased bone mineral density and decreased risk for fractures,^{8,9} and muscle strength,⁷ may be important underlying mechanisms contributing to this phenomenon. Moreover, evidence suggests the association between late-life adiposity and physical function may vary by race. For example, evidence suggests older African Americans experience reduced mobility at a heavier weight than older White adults.¹⁰ African Americans and Whites differ in body fat and lean muscle composition^{11,12} and this may explain, in part, studies showing a weaker association between adiposity and mobility in African American elders relative to White elders.

Counterintuitive associations between adiposity and cognitive function are well-documented in aged populations. Several studies report late-life adiposity may confer cognitive benefits including better baseline cognitive function¹³ and decreased risk for cognitive decline and dementia among heavier older adults.^{14,15} Some studies describe a nonlinear relationship between adiposity and cognitive function, such that both low weight and obesity are associated with poorer cognitive function relative to overweight status,13 while others do not.¹⁴ Additional support comes from neurobiological studies reporting lower Alzheimer's disease neuropathology burden in overweight, non-demented and mild cognitive impairment (MCI) older adult populations.^{16,17} It is important to note that this association may be modified by age and anthropometric measurement method¹⁸ and not all studies provide evidence in support of the obesity paradox.¹⁹⁻²¹ However, given the discrepancies in existing literature and the importance of cognitive and physical health in the maintenance of independent living abilities in late-life, it is important to clarify these complex associations.

The complex associations between late-life adiposity and physical and cognitive health have often been explained in the context of the obesity paradox. The obesity paradox describes an association between excess adiposity and favorable health outcomes.²² This paradox has been reported in several aged, clinical populations including patients with coronary heart disease²³ and diabetes.²⁴ There is a paucity of research examining this phenomenon in non-patient, community-dwelling older adult populations. Moreover, the association between excess adiposity and health outcomes may vary by race,²⁵ obesity severity,26 and cardiorespiratory fitness.²⁷ Researchers attempting to explain the obesity paradox have proposed several hypotheses. One explanation is that low weight may indicate weight loss and underlying pathology linked to declining general health and the development of dementia.^{28,29} Another explanation proposes that excess adiposity confers more energy reserves and a stronger inflammatory response that may improve the body's ability to combat acute illness.²⁷ An additional argument asserts that in aged populations, BMI is a robust predictor of skeletal mass,³⁰ which is positively related to cognitive function.³¹ Lastly, persons with excess adiposity may be treated more aggressively because they present with chronic health conditions earlier than persons of lower weight.³² Collectively these explanations suggest that underlying mechanisms of the obesity paradox remain unclear and warrant further investigation.

Based on existing evidence, it remains unclear whether: a) the obesity paradox persists for physical and cognitive outcomes in non-patient, community-dwelling older adult populations; and b) whether the paradox varies by race or stratified age. As previously stated, different patterns in the association between BMI and mortality have been observed between African American and White adults, such that the association is weaker for African Americans than it is for Whites.³² These findings may suggest that parity in BMI may not translate to similar body composition or comparable risk to health outcomes. Addressing these questions may advance our understanding of key contributors to late life health disparities and inform the development of targeted prevention strategies aimed at obesity and obesity-related health conditions. Therefore, the aim of this study was to examine the relationship between weight classification, cognitive function and physical function, and to determine whether these associations were modified by race or stratified age in a sample of community-dwelling older adults.

METHODS

Data Source and Participants

The Collaborative Assessment to Revitalize the Elderly (CARE) program was a health improvement program designed to screen for risk factors for frailty and implement an evidence-based intervention to increase physical function in community-dwelling elders at senior residential sites and community centers in Saint Louis, Missouri. Overall inclusion criteria for participation in the CARE program were: aged >55 years, community-dwelling, and the ability to give informed consent. For the purposes of this study, age was stratified (young-old, aged 55-74 years; older, aged \geq 75 years) and baseline data were analyzed. We stratified age because previous reports have shown distinct associations between BMI quartiles and cognitive outcomes when comparing young-old and older adults.¹⁸

Instruments

Anthropometric Measure

Height and weight were obtained using standardized examination procedures and calibrated equipment, and BMI was computed as the ratio of weight-to-height squared (kg/m²). World Health Organization (WHO) BMI classification was used to categorize participants as underweight, normal weight, overweight, or obese. Underweight participants were excluded due to their small sample size (n=2).

Physical Function

The mini-Physical Performance Test (mini-PPT)³³ was used to measure balance, strength and mobility. This 4-item, validated measure includes 4 components: picking up a penny from the floor, a timed 50-ft walk, chair rises (5 times), and a progressive Romberg test. Higher scores indicated better performance. Handgrip strength was used to measure forearm strength. This performancebased measure is correlated with general muscle strength³⁴ and activities of daily living abilities.³⁵ Participants squeezed the dynamometer with their dominant hand; their performance was measured in pounds. Higher scores indicated greater strength.

Cognitive Function

The Short Blessed Test (SBT) was used to assess general cognition. This cognitive screening measure tests orientation and memory and is designed to discriminate between mild, moderate, and severe cognitive impairment.³⁶ For the SBT, higher scores indicate poorer performance. Trailmaking tests (TMT) measured psychomotor speed and complex attention. Trail-making Test-B (TMT-B) also assesses cognitive flexibility, a key component of executive function.37 For this test, participants drew lines to connect alphanumeric stimuli in ascending order that were randomly placed on a page (TMT-A). In the more difficult condition (TMT-B), participants alternately tracked two sets of stimuli (letters, numbers) while performing the task. Scores reflect time taken to complete the tasks; higher scores reflect poorer performance. In addition to assessing TMT-A and TMT-B performance, a difference score (TMT-B minus TMT-A) was also calculated to reflect the unique task requirements of TMT-B.³⁸

Covariates

As potential predictors of the outcome, covariates were selected based on relevant literature then confirmed based on significant relations with outcome variables. Potential covariates included self-reported general health perception, smoking status, number of medications and depression. General health perception was rated (1=poor to 5=excel-

Variables	Total		White		A	Р		
		Normal	Overweight	Obese	Normal	Overweight	Obese	
Ν	331	44	51	39	32	56	109	
Age	76.9 ± 9.4	84.4 ± 7.8	82.6 ± 6.9	78.3 ± 8.7	77.3 ± 9.0	75.3 ± 8.5	72.1 ± 8.7	<.001
Hispanic, n (%)	6 (1.7)	0 (0)	1 (2.0)	0 (0)	3 (9.4)	1 (0.4)	1 (0.1)	$X^2 = .40$
Female, n (%)	313 (94.6)	44 (100)	46 (90.2)	35 (89.7)	27 (84.4)	48 (85.7)	99 (90.8)	$X^2 = .19$
Smoking, n (%)	29 (8.8)	0 (0)	1 (2.0)	1 (2.6)	6 (18.8)	9 (14.3)	12 (11)	$X^2 < .001$
BMI, kg/m ²	30.5 ± 7.2	22.3 ± 1.5	27.7 ± 1.3	35.8 ± 4.5	22.7 ± 1.6	28.0 ± 1.2	37.3 ± 6.1	$X^2 < .001$
Medications	7.2 ± 4.1	7.4 ± 4.4	7.6 ± 3.2	10.1 ± 5.9	7.0 ± 4.2	5.6 ± 2.9	6.9 ± 3.6	<.001
GHP	$2.8 \pm .8$	$3.0 \pm .7$	$2.9 \pm .8$	$2.8 \pm .8$	$2.7 \pm .8$	$2.8 \pm .8$	$2.6 \pm .7$.03
BPP	3.6 ± 1.4	3.7 ± 1.2	3.9 ± 1.4	3.9 ± 1.3	4.1 ± 1.4	3.9 ± 1.3	3.4 ± 1.3	.76
PHQ-9	3.8 ± 4.0	3.3 ± 3.3	3.2 ± 3.4	3.8 ± 4.4	2.8 ± 3.4	3.8 ± 4.4	3.9 ± 4.1	.70
Mini-PPT	8.9 ± 3.3	8.9 ± 2.8	8.5 ± 2.9	9.2 ± 3.2	9.6 ± 3.5	9.2 ± 3.2	9.0 ± 3.7	.06
TMT-A	69.1 ± 40.6	66.2 ± 41.7	57.6 ± 31.0	82.2 ± 48.3	87.7 ± 45.0	82.2 ± 48.3	70.1 ± 38.1	<.001
TMT-B	146.2 ± 40.1	149.9 ± 32.4	136.7 ± 43.0	152.7 ± 41.0	165.5 ± 24.1	152.7 ± 41.0	150.2 ± 38.8	<.01
SBT	1.6 ± 2.1	2.2 ± 2.3	1.2 ± 1.5	1.7 ± 2.7	1.9 ± 2.1	1.7 ± 2.7	1.5 ± 1.9	.55

Data are means \pm SD unless noted otherwise.

P, statistically significant difference by race.

BMI, body mass index; smoking categorized as current smoker vs previous smoker/never smoked; medications, number of medications; PHQ-9, patient health questionnaire 9-item; GHP, general health perception; BPP, bodily pain perception; Mini-PPT, mini Physical Performance Test; TMT-A, Trail-making Test A; TMT-B, Trail-making Test Part B; SBT, Short Blessed Test

lent), and smoking status was dichotomized as current smoker vs previous smoker/never smoked. The 9-item Patient Health Questionnaire (PHQ-9) was used to determine severity of depression symptoms.

Statistical Analysis

Descriptive statistics were computed for all predictor variables (race, age, BMI category), outcome variables (physical function and cognitive function scores), and prospective covariates. TMT-A, SBT, and PHQ-9 performance were square root transformed to correct positive skewness. Separate hierarchical multiple regression models were employed to determine associations between race (African American vs White), BMI, and stratified age (young-old: aged 55-74 years; old, aged >75 years) on each outcome, while adjusting for covariates. Body mass index was coded with binary values and the obese group served as the referent group. Only covariates that correlated with outcome variables and did not demonstrate multicolinearity were included. Interaction terms were created to assess the combined effect of primary predictors on functional outcomes. Relevant covariates where entered in to the model first, followed by primary predictors, and interaction terms.

RESULTS

Demographic characteristics are summarized in Table 1. Of the 331 participants in the sample, most were women (94.6%), and the average age was 77. White elders were significantly older (P<.001) and reported better general health (P=.03) than African American elders. On average, the sample was obese and African Americans had a higher prevalence of overweight and obesity (P<.001).

Table 2 illustrates results from the hierarchical regression analysis for physical function outcomes and general cognition measure (SBT). Depression scores, number of medications, and general health perception were included as covariates for mini-PPT and grip strength outcomes (Model 1, data not shown). For both models, stratified age was a significant predictor (Model 3: mini-PPT [B=-2.03, P<.001]; grip strength [B=-2.03, P<.001]), indicating that young-old adults performed better than older adults on these measures. Race and BMI classifications were not significant predictors; nor were there significant interactions for this model (Ps>.05). For general cognition (SBT), number of medications and depression scores were included as covariates (Model 1, data not shown). There was a trend for stratified age (B=.31, P=.05) and race (B=-.40, P=.05), but not for BMI classifi-

	Mini-PPT				Grip Strength				SBT			
	Model 2		Model 3		Model 2		Model 3		Model 2		Model 3	
	В	SE	В	SE	В	SE	В	SE	В	SE	В	SE
Age	-2.43ª	.40	-2.03 ^b	.48	-2.43ª	.40	-2.03 ^b	.48	.29ª	.13	.31	.15
Race	.17	.40	.03	.63	.17	.40	.03	.63	25	.13	40	.21
Normal weight	.93	.45	2.66	2.01	.93ª	.46	2.66	2.01	.38ª	.15	18	.66
Overweight	.30	.40	1.25	1.53	.30	.40	1.25	1.53	.03	.14	25	.53
Normal weight x race			.15	1.01			.15	1.01			.52	.33
Normal weight x age			-1.14	1.15			-1.14	1.15			10	.36
Overweight x race			.56	.88			.56	.88			.15	.30
Overweight x age			73	.57			73	.57			.02	.20

Table 2. Hierarchical multiple regression analysis to predict physical function and general cognition. Short Blessed Test

a. P<0.05.

b. P<.001.

Referent was obese group.

Model 1 mini-PPT and grip strength covariates were depression scores, number of medications, general health perception; model 1 SBT covariates were depression scores and number of medications.

Mini-PPT, mini physical performance test; SBT, Short Blessed Test.

 $\begin{array}{l} \text{Mini-PPT-Model 1: } F_{_{(3,249)}} = 11.94, P < .001, R^2 = .10, R^2 \Delta = .10; \text{ Model 2: } F_{_{(4,222)}} = 10.50, P < .001; R^2 = .22, R^2 \Delta = .11; \text{ Model 3: } F_{_{(4,280)}} = .61 P = .65, R^2 = .22, R^2 \Delta = .07. \\ \text{Grip Strength-Model 1: } F_{_{(3,257)}} = 3.53; P = .01; R^2 = .04; R^2 \Delta = .04; \text{ Model 2: } F_{_{(4,245)}} = 20.01, P < .001; R^2 = .27; R^2 \Delta = .25; \text{ Model 3: } F_{_{(4,249)}} = 1.79, P = .13, R^2 = .29, R^2 \Delta = .02. \\ \text{SBT-Model 1: } F_{_{(2,249)}} = .70, P = .48, R^2 = .006; R^2 \Delta = .006; \text{ Model 2: } F_{_{(4,245)}} = 3.65, P = .007; R^2 = .06, R^2 \Delta = .05; \text{ Model 3: } F_{_{(4,241)}} = .63, P = .63, R^2 = .07, R^2 \Delta = .01. \\ \end{array}$

cation or interaction terms (Ps>.05).

Table 3 illustrates the results from the hierarchical regression analysis for TMT performance. Number of medications and depression scores were included in the first block as covariates (data not shown). For psychomotor speed (TMT-A) performance, stratified age (B=1.39, P<.001) and race (B=-1.86, P<.001) were significant predictors of psychomotor speed; indicating young-old and White participants performed better on TMT-A than older adults and African American participants. No significant interactions were observed (Ps>.05). For cognitive flexibility (TMT-B) performance, similar results were

	TMT-A			ТМТ-В				TMT difference				
	Model 2		Model 3		Model 2		Model 3		Model 2		Model 3	
	В	SE	В	SE	В	SE	В	SE	В	SE	В	SE
Age	1.29 ^c	.29	1.39 ^c	.35	21.72 ^c	6.09	24.11 ^b	7.40	6.49	5.81	8.80	7.12
Race	-1.69°	.30	-1.86 ^c	.49	-26.94 ^c	6.20	-33.34 ^c	9.41	-6.44	5.98	-10.17	9.33
Normal weight	0.80*	.36	1.54	1.51	16.03ª	7.23	29.90	28.92	.78	7.04	-6.96	28.02
Overweight	0.51	.31	.09	1.16	4.00	6.34	-15.34	22.84	-7.12	6.12	-2.16	22.06
Normal weight x race			.64	.79			17.97	16.26			13.92	15.84
Normal weight x age			96	.83			-23.36	17.02			-7.98	16.33
Overweight x race			.03	.69			7.10	13.66			2.53	13.30
Overweight x age			.16	.43			3.90	8.62			-3.34	8.26

Table 3. Hierarchical multi	ple regression ana	lysis to predict cog	nitive performance	. Trail-making Test

a. P<.05. b. P<.01. c. P<.001. Referent was obese group. Model 1 covariates were depression scores and number of medications. TMT, Trail-making Test. TMT-A- Model 1: F_(2,243)=.95, P=.38, R²=.008, R²\Delta=.008; Model 2: F_(4,239)=10.08, P<.001; R²=.008, R²\Delta=.008; Model 3: F_(4,235)=.46, P=.76, R²=.15, R²\Delta=.007. TMT-B-Model 1: F_(2,223)=.23; P=.78; R²=.002; R²\Delta=.002; Model 2: F(4,219)=6.59, P<.001; R²=.10; R²\Delta=.10; Model 3: F_(4,215)=.75, P=.55, R²=.10, R²\Delta=.10. TMT difference-Model 1: F_(2,218)=.80, P=.45, R²=.001; R²\Delta=.001; Model 2: F_(4,214)=.93, P=.44; R²=.18 R²\Delta=.01; Model 3: F_(4,210)=.24, P=.91, R²=.02, R²\Delta=.005. observed for stratified age (B=21.72, P<.001) and race (B=-26.94, P<.01). Obese weight status compared with normal weight (B=16.03, P=.02) was associated with better cognitive flexibility. There was no significant difference in cognitive flexibility between normal weight and obese, or obese and overweight participants, nor were there any significant interactions (Ps>.05). Trail-making test difference scores revealed no significant main effects or interactions (Ps>.05).

DISCUSSION

In this sample of communitydwelling older adults, BMI was significantly associated with cognitive flexibility. Older adults who were obese had significantly better cognitive flexibility performance than normal weight older adults, and this association was not influenced by age or race. In addition, excess adiposity was not associated with better general cognition or physical function in our sample.

The obesity paradox remains controversial despite mounting evidence of its existence. Questions remain regarding under what conditions the obesity paradox exists and what underlying processes contribute to this phenomenon. Our findings add to this discussion by demonstrating an association between obesity status and cognitive function. Our results support those found by Fitzpatrick and colleagues.³⁹ In their study, latelife obesity was associated with a reduced risk for dementia, relative to late-life normal weight.³⁹ Although dementia was not our outcome of interest, we did find obesity to be associated with better executive function. Prior research has primarily focused on the association between adiposity and dementia risk,^{18,40,41} while fewer studies have examined how adiposity relates to specific cognitive domains.^{13,19} We did not find a significant association between excess adiposity and processing speed. Our null results were inconsistent with previous studies showing a positive association between adiposity and processing speed,¹³ this may be due to our cursory cognitive battery as stud-

Older adults who were obese had significantly better cognitive flexibility performance than normal weight older adults...

ies yielding positive results administered more robust cognitive measures. With regard to general cognition, our null findings are aligned with previous research.⁴² Our study adds to this literature by documenting an association between excess adiposity and cognitive flexibility. A focus on specific cognitive domains is important because age-related and diseaserelated changes in cognitive processes are typically not uniform;⁴³ therefore, a better understanding of which cognitive domains are most affected by adiposity and adiposity-related health conditions could inform strategies for diagnosis and treatment of cognitive

conditions in older adults. Future research is needed to better delineate the association between late-life excess adiposity and cognitive function.

In our study, BMI was not associated with physical function measures. These findings are inconsistent with previous reports showing paradoxical associations between adiposity and physical function,⁵ and with studies reporting higher BMI to be associated with worse physical function.44 One reason why our results may differ from that of previous reports is our use of a single measure to assess primarily lower body physical performance. The mini-PPT,³³ although validated, tested previously in community settings, and derived from a more extensive physical performance test, has not been compared with other performance measures and therefore may not be comparable to other objective measures of physical function. In addition, our findings may provide support for the argument that BMI is a less than optimal indicator of adiposity and health risk in aged populations, and indices of central adiposity are more robust indicators of health status and predictive of health-related outcomes.45

Our study was limited by a crosssectional design and precludes us from determining causality. We did not collect data on vascular comorbidities, which are known to contribute to cognitive⁴⁶ and physical function⁴⁷ in aged populations; therefore we could not account for the contribution of these factors to our findings. Males were also underrepresented in our study sample. Several researchers have argued that BMI classification may not be the most robust proxy of excess body fat in older adults and other anthropometric measures such as waist-to-hip ratio and other measures of central obesity may be more appropriate.²⁹ However, consensus regarding the best measure of obesity in older adults is lacking.⁴⁸ Also, our use of a limited physical function and cognitive battery hinders the generalizability of our results. Finally, we did not query the educational background of our sample and therefore could not account for any significant educational differences between African Americans and Whites in our sample.

CONCLUSION

Despite these limitations, our study has several strengths. Our study advances existing knowledge on the obesity paradox in diverse, community-dwelling older adult populations. Few studies have explored whether the obesity paradox persists in non-patient populations,²⁰ and even fewer studies have investigated this phenomenon in racially diverse populations.²⁵ For this reason, our study makes a noteworthy contribution to research focused on key contributors to functional health outcomes in aged populations. Future studies with larger sample sizes are needed to replicate our findings. Future work should also include neuroimaging and biomarkers of adipose tissue to examine the underlying biological processes linking latelife adiposity to executive function.

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Author Contributions

Research concept and design: Skinner, Wilkins; Acquisition of data: Wilkins; Data analysis and interpretation: Skinner, Abel, McCoy, Wilkins; Manuscript draft: Skinner, Abel, McCoy, Wilkins; Statistical expertise: Skinner; Acquisition of funding: Wilkins; Administrative: Skinner, Abel, McCoy, Wilkins; Supervision: Wilkins

References

- Fakhouri TOC, Carroll MD, Kit BK, Flegal KM. Prevalence of Obesity Among Older Adults in the United States, 2007–2010. NCHS Data Brief, no 106 Hyattsville. MD: National Center for Health Statistics; 2012.
- Afilalo J, Eisenberg MJ, Morin JF, et al. Gait speed as an incremental predictor of mortality and major morbidity in elderly patients undergoing cardiac surgery. *J Am Coll Cardiol.* 2010;56(20):1668-1676. https://doi.org/10.1016/j.jacc.2010.06.039. PMID:21050978.
- Hardy SE, Perera S, Roumani YF, Chandler JM, Studenski SA. Improvement in usual gait speed predicts better survival in older adults. J Am Geriatr Soc. 2007;55(11):1727-1734. https://doi.org/10.1111/j.1532-5415.2007.01413.x. PMID:17916121.
- Frilander H, Viikari-Juntura E, Heliövaara M, Mutanen P, Mattila VM, Solovieva S. Obesity in early adulthood predicts knee pain and walking difficulties among men: A life course study. *Eur J Pain*. 2016;20(8):1278-1287. https://doi.org/10.1002/ejp.852. PMID:26996726.
- Hardy R, Cooper R, Aihie Sayer A, et al; HALCyon study team. Body mass index, muscle strength and physical performance in older adults from eight cohort studies: the HALCyon programme. *PLoS One.* 2013;8(2):e56483. https://doi.org/10.1371/ journal.pone.0056483. PMID:23437142.
- Vaquero-Cristóbal R, Martínez González-Moro I, Alacid Cárceles F, Ros Simón E. [Strength, flexibility, balance, resistance and flexibility assessment according to body mass index in active older women]. *Rev Esp Geriatr Gerontol.* 2013;48(4):171-176. PMID:23332412.
- Marsh AP, Rejeski WJ, Espeland MA, et al; LIFE Study Investigators. Muscle strength and BMI as predictors of major mobility disability in the Lifestyle Interventions and Independence for Elders pilot (LIFE-P). *J Gerontol A Biol Sci Med Sci.* 2011;66(12):1376-1383. https://doi.org/10.1093/gerona/glr158. PMID:21975090.
- 8. Al Snih S, Ottenbacher KJ, Markides

KS, Kuo YF, Eschbach K, Goodwin JS. The effect of obesity on disability vs mortality in older Americans. *Arch Intern Med.* 2007;167(8):774-780. https:// doi.org/10.1001/archinte.167.8.774. PMID:17452539.

- Lloyd JT, Alley DE, Hawkes WG, Hochberg MC, Waldstein SR, Orwig DL. Body mass index is positively associated with bone mineral density in US older adults. *Arch Osteoporos*. 2014;9(1):175. https://doi.org/10.1007/ s11657-014-0175-2. PMID:24664472.
- Xu B, Houston DK, Gropper SS, Zizza CA. Race/Ethnicity differences in the relationship between obesity and gait speed among older Americans. *J Nutr Elder*. 2009;28(4):372-385. https://doi. org/10.1080/01639360903393515. PMID:21184379.
- Jones A Jr, Shen W, St-Onge MP, et al. Bodycomposition differences between African American and white women: relation to resting energy requirements. *Am J Clin Nutr.* 2004;79(5):780-786. PMID:15113715.
- Araujo AB, Chiu GR, Kupelian V, et al. Lean mass, muscle strength, and physical function in a diverse population of men: a population-based cross-sectional study. *BMC Public Health.* 2010;10(1):508. https:// doi.org/10.1186/1471-2458-10-508. PMID:20727198.
- Kuo HK, Jones RN, Milberg WP, et al. Cognitive function in normal-weight, overweight, and obese older adults: an analysis of the Advanced Cognitive Training for Independent and Vital Elderly cohort. *J Am Geriatr Soc.* 2006;54(1):97-103. https://doi. org/10.1111/j.1532-5415.2005.00522.x. PMID:16420204.
- Hsu CL, Voss MW, Best JR, et al. Elevated body mass index and maintenance of cognitive function in late life: exploring underlying neural mechanisms. *Front Aging Neurosci*. 2015;7:155. https://doi.org/10.3389/ fnagi.2015.00155. PMID:26347646.
- Aslan AK, Starr JM, Pattie A, Deary I. Cognitive consequences of overweight and obesity in the ninth decade of life? *Age Ageing*. 2015;44(1):59-65. https://doi.org/10.1093/ ageing/afu108. PMID:25249169.
- Vidoni ED, Townley RA, Honea RA, Burns JM; Alzheimer's Disease Neuroimaging Initiative. Alzheimer disease biomarkers are associated with body mass index. *Neurol*ogy. 2011;77(21):1913-1920. https://doi. org/10.1212/WNL.0b013e318238eec1. PMID:22105948.
- Buchman AS, Schneider JA, Wilson RS, Bienias JL, Bennett DA. Body mass index in older persons is associated with Alzheimer disease pathology. *Neurol*ogy. 2006;67(11):1949-1954. https://doi. org/10.1212/01.wnl.0000247046.90574.0f. PMID:17159099.

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- Luchsinger JA, Patel B, Tang MX, Schupf N, Mayeux R. Measures of adiposity and dementia risk in elderly persons. *Arch Neurol.* 2007;64(3):392-398. https://doi.org/10.1001/ archneur.64.3.392. PMID:17353383.
- Nilsson LG, Nilsson E. Overweight and cognition. *Scand J Psychol.* 2009;50(6):660-667. https://doi.org/10.1111/j.1467-9450.2009.00777.x. PMID:19930267.
- Strandberg TE, Stenholm S, Strandberg AY, Salomaa VV, Pitkälä KH, Tilvis RS. The "obesity paradox," frailty, disability, and mortality in older men: a prospective, longitudinal cohort study. *Am J Epidemiol.* 2013;178(9):1452-1460. https://doi. org/10.1093/aje/kwt157. PMID:24008903.
- Tobias DK, Pan A, Jackson CL, et al. Body-mass index and mortality among adults with incident type 2 diabetes. *N Engl J Med.* 2014;370(3):233-244. https:// doi.org/10.1056/NEJMoa1304501. PMID:24428469.
- Braun N, Gomes F, Schütz P. "The obesity paradox" in disease--is the protective effect of obesity true? *Swiss Med Wkly*. 2015;145:w14265. PMID:26709887.
- Niedziela J, Hudzik B, Niedziela N, et al. The obesity paradox in acute coronary syndrome: a meta-analysis. *Eur J Epidemiol*. 2014;29(11):801-812. https:// doi.org/10.1007/s10654-014-9961-9. PMID:25354991.
- 24. Doehner W, Erdmann E, Cairns R, et al. Inverse relation of body weight and weight change with mortality and morbidity in patients with type 2 diabetes and cardiovascular co-morbidity: an analysis of the PROactive study population. *Int J Cardiol.* 2012;162(1):20-26. https://doi.org/10.1016/j. ijcard.2011.09.039. PMID:22037349.
- Cohen SS, Signorello LB, Cope EL, et al. Obesity and all-cause mortality among black adults and white adults. *Am J Epidemiol.* 2012;176(5):431-442. https://doi. org/10.1093/aje/kws032. PMID:22822174.
- Flegal KM, Kruszon-Moran D, Carroll MD, Fryar CD, Ogden CL. Trends in Obesity Among Adults in the United States, 2005 to 2014. *JAMA*. 2016;315(21):2284-2291. https://doi.org/10.1001/jama.2016.6458. PMID:27272580.
- Goel K, Lopez-Jimenez F, De Schutter A, Coutinho T, Lavie CJ. Obesity paradox in different populations: evidence and controversies. *Future Cardiol.* 2014;10(1):81-91. https://doi. org/10.2217/fca.13.84. PMID:24344665.
- Buchman AS, Wilson RS, Bienias JL, Shah RC, Evans DA, Bennett DA. Change in body mass index and risk of incident Alzheimer disease. *Neurol*ogy. 2005;65(6):892-897. https://doi. org/10.1212/01.wnl.0000176061.33817.90. PMID:16186530.
- 29. Kalantar-Zadeh K, Horwich TB, Oreopou-

los A, et al. Risk factor paradox in wasting diseases. *Curr Opin Clin Nutr Metab Care*. 2007;10(4):433-442. https://doi.org/10.1097/ MCO.0b013e3281a30594. PMID:17563461.

- Iannuzzi-Sucich M, Prestwood KM, Kenny AM. Prevalence of sarcopenia and predictors of skeletal muscle mass in healthy, older men and women. J Gerontol A Biol Sci Med Sci. 2002;57(12):M772-M777. https:// doi.org/10.1093/gerona/57.12.M772. PMID:12456735.
- Won H, Abdul Manaf Z, Mat Ludin AF, et al. Wide range of body composition measures are associated with cognitive function in community-dwelling older adults. *Geriatr Gerontol Int.* 2017; 17(4):554-560. PMID:27231255.
- 32. Schenkeveld L, Magro M, Oemrawsingh RM, et al. The influence of optimal medical treatment on the 'obesity paradox', body mass index and long-term mortality in patients treated with percutaneous coronary intervention: a prospective cohort study. *BMJ Open*. 2012;2(1):e000535. https://doi.org/10.1136/ bmjopen-2011-000535. PMID:22327630.
- Wilkins CH, Roe CM, Morris JC. A brief clinical tool to assess physical function: the mini-physical performance test. *Arch Gerontol Geriatr.* 2010;50(1):96-100. https:// doi.org/10.1016/j.archger.2009.02.006. PMID:19282039.
- 34. Massy-Westropp NM, Gill TK, Taylor AW, Bohannon RW, Hill CL. Hand Grip Strength: age and gender stratified normative data in a population-based study. *BMC Res Notes*. 2011;4(1):127. https:// doi.org/10.1186/1756-0500-4-127. PMID:21492469.
- Fauth EB, Zarit SH, Malmberg B. Mediating relationships within the Disablement Process model: a cross-sectional study of the oldestold. *Eur J Ageing*. 2008;5(3):161-179. https:// doi.org/10.1007/s10433-008-0092-6.
- Katzman R, Brown T, Fuld P, Peck A, Schechter R, Schimmel H. Validation of a short Orientation-Memory-Concentration Test of cognitive impairment. *Am J Psychiatry*. 1983;140(6):734-739. https://doi. org/10.1176/ajp.140.6.734. PMID:6846631.
- Reitan RM. Validity of the trailmaking test an indicator of organi brain damage. *Percept Mot Skills*. 1958;8(3):271-276. https://doi. org/10.2466/pms.1958.8.3.271.
- Misdraji EL, Gass CS. The Trail Making Test and its neurobehavioral components. J Clin Exp Neuropsychol. 2010;32(2):159-163. https://doi. org/10.1080/13803390902881942. PMID:19459077.
- Fitzpatrick AL, Kuller LH, Lopez OL, et al. Midlife and late-life obesity and the risk of dementia: cardiovascular health study. *Arch Neurol.* 2009;66(3):336-342. https:// doi.org/10.1001/archneurol.2008.582. PMID:19273752.

- Albert SM, Bear-Lehman J, Anderson SJ. Declines in mobility and changes in performance in the instrumental activities of daily living among mildly disabled community-dwelling older adults. *J Gerontol A Biol Sci Med Sci*. 2015;70(1):71-77. https://doi.org/10.1093/ gerona/glu088. PMID:24952575.
- West NA, Haan MN. Body adiposity in late life and risk of dementia or cognitive impairment in a longitudinal communitybased study. *J Gerontol A Biol Sci Med Sci*. 2009;64(1):103-109. https://doi.org/10.1093/ gerona/gln006. PMID:19168781.
- Kanaya AM, Lindquist K, Harris TB, et al; Health ABC Study. Total and regional adiposity and cognitive change in older adults: The Health, Aging and Body Composition (ABC) study. Arch Neurol. 2009;66(3):329-335. https://doi.org/10.1001/archneurol.2008.570. PMID:19273751.
- Wilson RS, Leurgans SE, Boyle PA, Schneider JA, Bennett DA. Neurodegenerative basis of age-related cognitive decline. *Neurol*ogy. 2010;75(12):1070-1078. https://doi. org/10.1212/WNL.0b013e3181f39adc. PMID:20844243.
- Wei L, Wu B. Racial and ethnic differences in obesity and overweight as predictors of the onset of functional impairment. *J Am Geriatr Soc.* 2014;62(1):61-70. https://doi. org/10.1111/jgs.12605. PMID:24384026.
- Jankowski CM, Gozansky WS, Van Pelt RE, et al. Relative contributions of adiposity and muscularity to physical function in community-dwelling older adults. *Obesity (Silver Spring)*. 2008;16(5):1039-1044. https://doi. org/10.1038/oby.2007.84. PMID:18292753.
- Chui HC, Ramirez Gomez L. Vascular Contributions to Cognitive Impairment in Late Life. *Neurol Clin.* 2017;35(2):295-323. https://doi.org/10.1016/j.ncl.2017.01.007. PMID:28410661.
- Lee CG, Schwartz AV, Yaffe K, Hillier TA, LeBlanc ES, Cawthon PM; Study of Osteoporotic Fractures Research Group. Changes in physical performance in older women according to presence and treatment of diabetes mellitus. *J Am Geriatr Soc.* 2013;61(11):1872-1878. https://doi.org/10.1111/jgs.12502. PMID:24219188.
- Kalish VB. Obesity in Older Adults. *Prim Care*. 2016;43(1):137-144, ix. https:// doi.org/10.1016/j.pop.2015.10.002. PMID:26896206.