## SHORT COMMUNICATION

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# Cognitive impairment and metabolic syndrome in a population of Brazilian oldest-old

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## INTRODUCTION

The global oldest-old population is projected to increase 351% between 2010 and 2050. In Brazil, approximately 4.3 million people are aged over 80 years, making it the fastest-growing stratum of the country's population. The total number of people with dementia worldwide is set to reach 82 million in 2030 and 152 million in 2050. Approximately 60% of people with dementia live in low-and middle-income countries. The prevalence of dementia doubles for every 5-year increase in age; therefore, dementia is a major health issue among those who aged above 80 years<sup>1,2</sup>.

Metabolic syndrome (MS) is a common clinical condition in the older population. It consists of a set of cardiovascular risk factors and is associated with an increased risk of developing diseases, including cerebrovascular disease. Vascular risk factors are associated with cognitive impairment no dementia (CIND) and dementia, particularly vascular dementia<sup>3,4</sup>.

Recently, the World Health Organization published a document containing key topics related to dementia prevention including management of weight, hypertension, diabetes, and dyslipidemia, all of which are MS criteria<sup>5</sup>.

Cognitive impairment no dementia denotes individuals whose cognitive functioning falls below normal, but who do not meet the criteria for dementia. The follow-up of older people with CIND is important in order to identify those who will progress to dementia<sup>6</sup>.

The literature is conflicting regarding the association between MS and CIND or dementia, with some studies confirming this association and others not, particularly in the oldest-old<sup>7</sup>.

The aim of this study was to determine the association between MS and CIND in a population of functionally independent oldest-old.

## **METHODS**

A population of functionally independent community-dwelling subjects who aged above 80 years participated in this cross-sectional study. The population was part of a cohort ("Projeto Longevos") routinely followed at an outpatient clinic of a university hospital in São Paulo, Brazil. Data were obtained from the first assessment of the individuals at the clinic and included sociodemographic information, cognitive screening results, and presence of clinical factors related to MS.

The clinical and laboratory data used for MS diagnosis were as follows: waist circumference  $\geq 102$  cm (male) and  $\geq 88$  cm (female); triglycerides  $\geq 150$  mg/dL or use of medication for dyslipidemia, high-density lipoprotein (HDL) cholesterol <40 mg/dL (male) and <50 mg/dL (female) or use of lipid-lowering drugs; systemic arterial hypertension reported by the participant or use of antihypertensive drugs; and fasting glycemia  $\geq 100$  mg/dL or use of medication for diabetes. In accordance with the National Cholesterol Education Program-Adult Treatment Panel Guide III (NCEP/ATP III) criteria<sup>8</sup>, three out of these five items needed to be present in order to establish an MS diagnosis.

The instruments used for cognitive screening were the Mini-Mental State Exam<sup>9</sup>, the Clock Drawing Test<sup>10</sup>, and Verbal Fluency<sup>11</sup>. Scores on two of the three instruments had to be below the expected level for this population in order to consider the participant as having CIND.

The study was approved by the Ethics Committee of UNIFESP (1532/09).

#### **Statistical analysis**

Categorical variables were expressed as absolute and relative values, whereas continuous variables were expressed as central tendency measures. The association of categorical variables was

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assessed using the chi-square test and the comparison of continuous variables between two groups was assessed using the Student's *t* test. Logistic regression was used in the multivariate analysis. The level of statistical significance adopted was 0.05.

### RESULTS

The total number of oldest-old in the cohort was 323, of which 68 were excluded due to missing items of information collected for this study. Data from 255 participants were analyzed. There were 171 (67%) women and 84 (33%) men. Overall, 55% had MS in the female group versus 39% in the male group.

The mean age of the total sample was  $85.4 (\pm 4.4)$  years and the mean schooling was  $4.8 (\pm 4.5)$  years. Considering the groups "with and without metabolic syndrome," there was no statistically significant difference for age (p=0.84) and for schooling (p=0.09).

The mean age of the group with CIND was statistically and significantly higher than the group without CIND (p=0.02). No statistically significant difference was observed between the groups with and without CIND considering schooling (p=0.17).

As shown in Table 1, no association was found between CIND and MS (p=0.35).

As shown in Table 2, no association was found between any of the components of MS and CIND. Simple logistic regression was then performed using independent variables with p<0.25 in the bivariate analysis which in this case were age and schooling. "With CIND" was the dependent variable. Only the difference in age was associated with "having CIND" (odds ratio [OR] 1.07; p=0.02; and 95% confidence interval [CI] 1.008–1.13), which means that for each increased year there is 7.07% more chance to have CIND.

## DISCUSSION

The sample population was predominantly women, consistent with the global demographic profile showing feminization of old age. In Brazil, 55.7% of its 20 million older people are women. As expected, there was a statistically significant association between older age and CIND<sup>2</sup>.

There was no association between CIND and MS in the cohort of oldest-old studied. This is in line with the literature, where previous studies have failed to find this association among the oldest-old<sup>7</sup>. An analysis of relevant scientific publications revealed that the association between MS and cognitive impairment depends on the age group of the population studied. In younger older adults, most studies suggest an association between MS and cognitive impairment<sup>12-17</sup>, whereas in the oldest-old, the results fail to confirm this association<sup>18-22</sup>.

 Table 1. Association between cognitive impairment no dementia and metabolic syndrome.

	Metabolic syndrome							
CIND	No		Yes		Total		р	
	n	%	n	%	n	%		
No	78	67	101	73	179	70		
Yes	38	33	38	27	76	30	0.35	
Total	116	45	139	55	255	100		

CIND: cognitive impairment no dementia.

Гаb	le 2.	Association	between eacl	n component	t of me	tabolic sync	frome and CIND.
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	No		Yes		Total			
CIND	n	%	n	%	n	%	р	
High waist circumforance	No	85	47	38	50	123	48	0.71
High waist circumerence	Yes	94	53	38	50	132	52	
Low HDL-cholesterol or use of lipid-	No	64	36	29	38	93	36	0.72
lowering medication	Yes	115	64	47	62	162	64	
Hyperglycemia or use of diabetes	No	97	54	46	61	143	56	0.25
medication	Yes	82	46	30	39	112	44	0.55
Reported hypertension or use of	No	38	21	15	20	53	21	0.79
medication	Yes	141	79	61	80	202	79	
Llupertrialuceridemia	No	142	79	57	75	199	78	0.45
пурегидусениенна	Yes	37	21	19	25	56	22	0.45

CIND: cognitive impairment no dementia.

The existence of this difference raises questions. One explanation may lie in the MS cutoffs — do they apply to the oldest-old as they do to other populations? Another interesting point is the "survival effect," indicating that the oldest-old reach old age despite cardiovascular risks<sup>18</sup>. The antagonistic pleiotropy hypothesis could partially explain the greater deleterious effect of MS in younger ages compared to oldest-old<sup>23</sup>.

The main limitation of this study was its cross-sectional design. However, this is the first study conducted in Brazil investigating the association of CIND with MS in the oldest-old.

## **AUTHORS' CONTRIBUTIONS**

VH: Conceptualization, Data Curation, Writing – Review and Editing. AFJ: Formal Analysis, Writing – Original Draft. LMQA: Writing – Review and Editing. MSC: Writing – Review and Editing. CMAF: Conceptualization, Formal Analysis, Writing – Review and Editing.

## REFERENCES

- 1. World Health Organization. Global health and ageing. 2011. Geneva: World Health Organization; 2011. [cited on Apr. 04, 2020]. Available from: https://www.who.int/ ageing/publications/global\_health/en/
- BRASIL. Instituto Brasileiro de Geografia e Estatística (IBGE). Número de idosos cresce 18% em 5 anos e ultrapassa 30 milhões em 2017. Rio de Janeiro: Agência IBGE Notícias; 2018. [cited on Apr. 04, 2020]. Available from: https:// agenciadenoticias.ibge.gov.br/agencia-noticias/2012-agenciade-noticias/noticias/20980-numero-de-idosos-cresce-18-em-5-anos-e-ultrapassa-30-milhoes-em-2017
- Frisardi V, Solfrizzi V, Seripa D, Capurso C, Santamato A, Sancarlo D, et al. Metabolic-cognitive syndrome: a cross-talk between metabolic syndrome and Alzheimer's disease. Ageing Res Rev. 2010;9(4):399-417. https://doi. org/10.1016/j.arr.2010.04.007
- Yaffe K. Metabolic syndrome and cognitive disorders: is the sum greater than its parts? Alzheimer Dis Assoc Disord. 2007;21(2):167-71. https://doi.org/10.1097/ WAD.0b013e318065bfd6
- World Health Organization. Risk reduction of cognitive decline and dementia. Geneva: World Health Organization; 2019. [cited on Apr. 04, 2020]. Available from: https://www. who.int/mental\_health/neurology/dementia/guidelines\_ risk\_reduction/en/
- Jacova C, Peters KR, Beattie BL, Wong E, Riddehough A, Foti D, et al. Cognitive impairment no dementia neuropsychological and neuroimaging characterization of an amnestic subgroup. Dement Geriatr Cogn Disord. 2008;25(3):238-47. https://doi.org/10.1159/000115848
- Assuncao N, Sudo FK, Drummond C, Felice FG, Mattos P. Metabolic Syndrome and cognitive decline in the elderly: a systematic review. PLoS One. 2018;13(3):e0194990. https://doi.org/10.1371/journal.pone.0194990.
- Penalva DQF. Síndrome metabólica: diagnóstico e tratamento. Rev Med. 2008;87(4):245-50. https://doi. org/10.11606/issn.1679-9836.v87i4p245-250
- Brucki SM, Nitrini R, Caramelli P, Bertolucci PH, Okamoto IH. Sugestões para o uso do mini-exame do estado mental no Brasil. Arq Neuropsiquiatr. 2003;61(3B):777-81. https:// doi.org/10.1590/s0004-282x2003000500014
- Aprahamian I, Martinelli JE, Yassuda MS. Doença de Alzheimer em idosos com baixa escolaridade: o teste do desenho do relógio pode ser útil no rastreio cognitivo? Rev Soc Bras Clín Méd. 2008

[cited on Apr. 04, 2020];6(4):130-4. Available from: http://files. bvs.br/upload/S/1679-1010/2008/v6n4/a130-134.pdf

- Brucki SMD, Rocha MSG. Category fluency test: Effects of age, gender and education on total scores, clustering, in Brazilian Portuguese-speaking subjects. Braz J Med Biol Res. 2004;37(12):1771-7. https://doi.org/10.1590/ s0100-879x2004001200002
- Yaffe K, Kanaya A, Lindquist K, Simonsick EM, Harris T, Shorr RI, et al. The metabolic syndrome, inflammation, and risk of cognitive decline. JAMA. 2004;292(18):2237-42. https://doi.org/10.1001/jama.292.18.2237
- Crane PK, Walker R, Hubbard RA, Li G, Nathan DM, Zheng H, et al. Glucose levels and risk of dementia. N Engl J Med. 2013;369(6):540-8. https://doi.org/10.1056/ NEJMoa1215740
- Solfrizzi V, Scafato E, Capurso C, D'Introno A, Colacicco AM, Frisardi V, et al. Metabolic syndrome, mild cognitive impairment, and progression to dementia. The italian longitudinal study on aging. Neurobiol Aging. 2011;32(11):1932-41. https://doi.org/10.1016/j. neurobiolaging.2009.12.012
- Vanhanen M, Koivisto K, Moilanen L, Helkala EL, Hänninen T, Soininen H, et al. Association of metabolic syndrome with Alzheimer disease: a population-based study. Neurology. 2006;67(5):843-7. https://doi.org/10.1212/01. wnl.0000234037.91185.99
- Vieira JR, Elkind MS, Moon YP, Rundek T, Boden-Albala B, Paik MC, Sacco RL, Wright CB. The metabolic syndrome and cognitive performance: the northern manhattan study. Neuroepidemiology. 2011;37(3-4):153-9. https:// doi.org/10.1159/000332208
- Shigaeff N, Jacinto AF, Franco FGM, Chiochetta G, Cendoroglo MS, Cítero VA. Cognitive assessment in an elderly population with metabolic syndrome in Brazil. Dement Neuropsychol. 2013;7(2):206-9. https://doi. org/10.1590/S1980-57642013DN70200011
- van den Berg E, Biessels GJ, de Craen AJ, Gussekloo J, Westendorp RG. The metabolic syndrome is associated with decelerated cognitive decline in the oldest old. Neurology. 2007;69(10):979-85. https://doi.org/10.1212/01. wnl.0000271381.30143.75
- Katsumata Y, Todoriki H, Higashiuesato Y, Yasura S, Willcox DC, Ohya Y, et al. Metabolic syndrome and cognitive decline among the oldest old in Okinawa: in search of a

mechanism. J Gerontol A Biol Sci Med Sci. 2012;67(2):126-34. https://doi.org/10.1093/gerona/glr189

- 20. Tournoy J, Lee DM, Pendleton N, O'Neill TW, O'Connor DB, Bartfai G, et al. Association of cognitive performance with the metabolic syndrome and with glycaemia in middle-aged and older European men: the European male ageing study. Diabetes Metab Res Rev. 2010;26(8):668-76. https://doi.org/10.1002/dmrr.1144
- 21. Luo L, Yang M, Hao Q, Yue J, Dong B. Cross-sectional study examining the association between metabolic syndrome

and cognitive function among the oldest old. J Am Med Dir Assoc. 2013;14(2):105-8. https://doi.org/10.1016/j. jamda.2012.10.001

- 22. Forti P, Pisacane N, Rietti E, Lucicesare A, Olivelli V, Mariani E, et al. Metabolic syndrome and risk of dementia in older adults. J Am Geriatr Soc. 2010;58(3):487-92. https://doi.org/10.1111/j.1532-5415.2010.02731.x
- Le Couteur DG, Simpson SJ. Adaptive senectitude: the prolongevity effects of aging. J Gerontol A Biol Sci Med Sci. 2011;66(2):179-82. https://doi.org/10.1093/gerona/glq171

