A systematic review of cross-sectional studies on the association of sedentary behavior with cardiometabolic diseases and related biomarkers in South American adults

Abstract

Introduction: Sedentary behavior (SB) has been independently associated with detrimental health outcomes in different regions worldwide. The aim of this systematic review was to examine whether domain-specific SB is associated with cardiometabolic diseases (CMD) and related biomarkers in South American adults.

Methods: Nine electronic databases were searched to identify all studies that analyzed the association between SB and CMD – e.g., obesity, diabetes, hypertension, metabolic syndrome (MetS) and clustering of chronic diseases (CCD) – and related biomarkers in South American adults. Two independent reviewers performed the necessary abstract/full-text screening, data abstraction, and quality assessments. The review protocol was registered in the PROSPERO database (CRD42018099319).

Results: From the 1,262 articles identified in the search, 262 were reviewed in full and 20 were used in the analysis in accordance with the inclusion criteria. High SB (mainly sitting and TV time) was associated with an increased likelihood of obesity (n = 8), diabetes (n = 6), and CCD (n = 3), as well as high values of BMI (n = 8), WC (n = 7), % BF (n = 4), plasma lipids (n = 4), and glycemia (n = 5). Eleven out of 20 studies were of higher quality.

Conclusion: Long time spent in SB, mainly sitting and TV time, was positively associated with the occurrence of CMD and related biomarkers in South American adults.

Keywords:
Obesity, Diabetes mellitus, Hypertension, Body mass index, Metabolic syndrome.

Resumen

Introducción: el comportamiento sedentario (CS) se ha asociado de forma independiente con resultados perjudiciales para la salud en diferentes regiones del mundo. El objetivo de esta revisión sistemática fue examinar si el CS específico de cada dominio se asocia a no a enfermedades cardiometabólicas (ECM) y sus biomarcadores relacionados en adultos sudamericanos.

Métodos: se realizaron búsquedas en nueve bases de datos electrónicas para identificar todos los estudios que habían analizado la asociación entre CS y ECM –por ejemplo, obesidad, diabetes, hipertensión, síndrome metabólico y agrupación de enfermedades crónicas (AEC) – y sus biomarcadores relacionados en adultos sudamericanos. Dos revisores independientes realizaron evaluaciones de los resúmenes/textos completos, el resumen de los datos y evaluaciones de calidad. El protocolo de revisión está registrado en la base de datos PROSPERO (CRD42018099319).

Resultados: de los 1262 artículos identificados en la búsqueda, 262 se revisaron en su totalidad y 20 se utilizaron en el análisis de acuerdo con los criterios de inclusión. El gran CS (principalmente, tiempo sentado y de televisión) se asoció a una mayor probabilidad de obesidad (n = 8), diabetes (n = 6) y AEC (n = 3), así como a valores altos de IMC (n = 8), WC (n = 7), % BF (n = 4), lípidos plasmáticos (n = 4) y glucemia (n = 5). Once de los 20 estudios fueron de alta calidad.

Conclusión: la gran cantidad de tiempo invertido en el CS, principalmente el tiempo sentado y de televisión, se asoció positivamente con la aparición de ECM y sus biomarcadores relacionados en adultos de América del Sur.
INTRODUCTION

South America is comprised of nations and territories containing different environments and many complex and heterogeneous ethnicities, societies, and cultures in a population estimated at over 430 million. In South America upper-middle-income economies are predominant, and major demographic shifts like population growth, urbanization, technological advancements, and ageing are in course. Thus, behavior and environment factors, such as smoking, unhealthy diets (i.e., high energy-rich foods and low fruit and vegetables consumption), and physical inactivity are relevant modifiable risk factor for cardiometabolic diseases (CMD) (2-6).

Concerning physical inactivity, sedentarism is known to bring about serious health consequences and associations with all-cause mortality and other outcomes worldwide (2,7-10). Despite the fact that one in four adults worldwide does not meet the World Health Organization recommendations on physical activity to benefit from a reduced risk of common chronic diseases (11), failing to achieve the public health goals on physical activity is not the same as being sedentary. In this sense, sedentary behavior (SB) refers to activities that do not require significant energy expenditure – i.e., 1.5 METs or lower (12) – and is usually expressed as sitting time (ST) (13) such as in television (TV) viewing, computer use, transport, driving, reading or playing video games whilst in a sitting or reclining posture (14), in different everyday life domains (i.e., home, workplace, commuting, leisure time) (13,15). Sedentary behavior has been independently associated with detrimental health outcomes, including CMD, in adults from different regions worldwide (16-19). Moreover, SB presents high values in populations of different ethnic background worldwide (20) – among adults, the proportion of individuals spending 4 or more hours a day in a sitting position varies from 23.8 % in Southeast Asia, 37.8 % in Africa, 39.8 % in the western Pacific, 41.4 % in the eastern Mediterranean, and 55.2 % in the Americas to 64.1 % in Europe (21).

In South America’s adult population physical inactivity reaches levels of over 40 % (21), and SB has been assessed as almost 6 h/day (22). In addition, the prevalence of CMD and its risk factors in South America is high (4,6,23,24). Nevertheless, the association between SB and CMD and its risk factors in South Americans is not well known. Identifying the association of SB in different domains with CMD is not only important for public health interventions, but also for occupational health, urban planning, and transport-related initiatives. Therefore, the aim of this study was to systematically review the literature to examine whether domain-specific SB is associated with CMD and related biomarkers in South American adults.

METHODS

SEARCH STRATEGY

A broad and specialized search was performed. Studies reporting on the possible association between SB and/or domain-specific SB – e.g., ST, TV viewing time or frequency and overall SB (sedentary activity, screen time, computer time, reading time, passive transport, and sedentary work) – with CMD diseases (primary outcomes) and or related biomarkers (secondary outcomes) in South American adults were examined. The reporting guidelines of the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) statement were followed, and the criteria outlined in A Measurement Tool to Assess Systematic Reviews (AMSTAR) checklist (25,26) were observed. The methodology of this systematic review was prospectively registered with PROSPERO (CRD42018099319; http://www.crd.york.ac.uk/PROSPERO) under the title “Is sedentary behavior associated with risk for cardiometabolic diseases in South American adults? A systematic review”.

This review examined studies completed from January 2010 to December 2018, written in English, Spanish or Portuguese or with translations into these languages. Relevant studies were identified using the following electronic databases: Medline (Medical Literature Analysis and Retrieval System Online /PubMed); Embase (Elsevier); Cochrane Central (The Cochrane Central Register of Controlled Trials The Cochrane Library); Lilacs (Scientific and technical literature of Latin America and Caribe); Science Direct; Bireme (Latin American and Caribe center of information in health science); Scielo.org; Scopus (Elsevier); and SPORTDiscus.

The exposure descriptors used were: a) “sedentary behavior or time or lifestyle”; b) “sitting or reclining or leisure time”; and c) “television or TV view or watch or time”. The outcome descriptors used were: a) “cardiometabolic or metabolic health or markers or risk or disease or syndrome”; and b) “cardiovascular or CVD markers or risk or disease or syndrome”. The population descriptors used were: a) “South America or South American countries and territories”. Appropriate combinations of these descriptors were used to ensure quality, transparency and maximum sensitivity during article retrieval. The search strategies adapted to the indexing systems are available from the authors upon request.

STUDY SELECTION

This review included studies that addressed the amount of time spent in SB or domains of SB as risk factors for CMD, either self-reported or objectively measured, reported on a continuous scale (e.g., minutes/day) or divided into categories (e.g., > 3 h of TV time vs. ≤ 3 h), and time spent in specific domains of SB (i.e., ST, TV time or frequency and overall SB). The presence of CMD such as obesity, diabetes, hypertension, metabolic syndrome (MetS), and a clustering of chronic diseases (CDD - Heart attack, heart failure, angina, hypertension, DM, or arthritis) were the primary outcomes of interest in this review. Related biomarkers like total cholesterol (TC), high density lipoprotein (HDL) and low density lipoprotein (LDL) cholesterol, triglycerides (TG), blood glucose and insulin, glycated hemoglobin (HbA1c), insulin resistance (HOMA-IR), hypercholesterolemia, dyslipidemia and anthropomorphic measures such as body mass index (BMI), waist circumference (WC) and percentage of body fat (% BF) were the secondary outcomes of interest. Articles excluded were those on non-South American populations and/or non-adult populations (mean age < 18 years), those that did not report exposure to SB or the association between SB and health conditions.
outcomes, those that were duplicated, impossible to locate or obtain, and reviews or meta-analyses.

The screening of titles and abstracts for all studies was independently performed by two authors (KAM and AJN) in order to identify potential relevant articles. Likewise, these two authors (KAM and AJN) performed full-text screenings, and with mutual consensus confirmed that studies met the study’s inclusion and exclusion criteria. The authors resolved discrepancies after discussion.

DATA EXTRACTION AND ANALYSIS

Data and informations of interest were extracted by KAM or AJN. The main information obtained was on study characteristics, population characteristics, country of study, sample size analyzed, SB exposure and methods, outcome measurement, and measure of effect or correlation. Study quality was assessed using an adapted 20-item checklist (originally 27 items) of the Downs and Black checklist (27). Good quality was determined by using the overall numeric score of quality out of 20 possible points. Studies showing 16/20 median split or higher were considered of high quality. KAM independently assessed quality using the checklist, and AJN reviewed all scores. Disagreements were resolved upon consensus or were referred by a third researcher.

The study-specific ORs for obesity or diabetes were combined as a generic inverse variance to estimate the pooled OR with 95 % CI by using the inverse variance statistical method with a random effects model. The pooled OR was calculated from a natural logarithm of OR [ln(OR)] and the standard error of ln(OR) was obtained for the 95 % CI. A two-sided p-value lower than 0.05 was considered significant for all analyses. Studies were not included in the meta-analysis if the summary statistics of OR and 95 % CI were not available (31), or when the subjects evaluated were not sedentary (29). This meta-analysis was conducted using the RevMan 5.3 (The Nordic Cochrane Centre, The Cochrane Collaboration, Copenhagen, 2012), a free download at http://tech.cochrane.org/revman/download.

RESULTS

The search identified 1,262 articles, and the screening of their primary titles and abstracts generated 262 articles. Then these 262 articles were reviewed in full and 20 articles met the inclusion and exclusion criteria. Figure 1 shows the flow diagram and the reasons for exclusion. In all, 55 % of the analyzed studies were considered to be high-quality. Five out of 20 studies scored 17 points; six scored 16 points, eight scored 15 points, and one scored 14 points.

Figure 1.
Flow diagram for literature search, January 2010 – December 2018. (SB: sedentary behaviour. Source: prepared by the authors from the study results).
In this review eleven out of the 20 analyzed studies were carried out on Brazilian, seven on Chilean, one on Colombian, and one on Peruvian adults, and were published prior to and including December 2018, specifically from 2010 to 2018. We observed that most studies were published from 2017 on (n = 11). The selected studies analyzed adults (mean age: 30 to 70 years) from urban (n = 20/20) and rural (n = 8/20) areas.

The qualitative analysis revealed that 13 out of 20 studies reported on CMD (obesity, diabetes, hypertension, CCD, and MetS) (Table I), and 16 studies reported results related to CMD biomarkers. Overall, the results were consistent in showing that an increase in time spent in SB was associated with an increase in obesity (n = 8/8), diabetes (n = 6/7), CCD (3/4), hypertension (n = 3/4), and MetS (n = 2/2). Along with these diseases, SB was associated with higher values of the following related biomarkers (Table II): BMI (n = 8/9), waist circumference (n = 7/7), % BF (4/4), HOMA-IR (n = 4/4), glucose (n = 5/6), insulin (n = 4/4), and TG (n = 4/6), and low levels of HDL-c (n = 6/7). Dyslipidemia (1/1) and hypercholesterolemia (1/1) were also positively associated with SB. Despite that, TC (n = 3/6), and LDL-c (n = 3/5) were associated with SB in only three out of six and five studies, respectively. Regarding HbA1c (1/2), this glycemic control biomarker was associated with SB in only one study.

CARDIOMETABOLIC DISEASES (PRIMARY OUTCOMES)

The association between SB and CMD as primary outcome was observed in different studies (Table I). For example, high SB (> 4 h/day) was associated with central obesity in Chilean adults (28); and those categorized as either low SB (i.e., ST of 1.56 ± 0.7 h/day) or high SB (i.e., ST of 5.31 ± 2.2 h/day) and physically active were less likely to have obesity or central obesity when compared to those in the high SB (i.e., ST of 6.14 ± 2.6 h/day) and inactive category (29). In addition, Chilean men exhibiting a higher ST were more likely obese (30). A ST of over 8 h/day during the week discriminated the presence of abdominal obesity, but not of obesity in Brazilian women (31); and obesity was associated with SB in Brazilian adults (32). Furthermore, a higher TV time was associated with greater abdominal obesity (33) and obesity (34) in Brazilian and Peruvian (35) women. The pooled odds ratio for obesity in association with sedentary behavior – 1.16 (1.07-1.25 – is presented in figure 3.

Concerning MetS, Chilean adults in the low or high SB and physically active group were less likely to have MetS when compared to those in the high SB time and inactive category (29). In addition, high ST (> 4 h/day) was positively associated with MetS among Chilean adults (28).

The association between SB and CMD-related biomarkers was also reported in the reviewed studies (Table II). Regarding BMI, its positive association with SB was found in Chilean adults (29,41-43) and in Colombian men (44). Such association was also reported in Chilean men only (2), whereas a positive association of TV frequency with BMI was observed in Brazilian (34) and in Peruvian women only (35). However, SB (> 4 h/day) was negatively associated with BMI in Chilean women, but positively in Chilean men (28). In addition, ST (> 4 h/day) was not associated with BMI in Brazilian adults (45). Concerning % BF, it was positively associated with sedentary behavior and ST in Chilean adults (41-43) and in Colombian men (44).

With regard to WC, its positive association with SB was reported in Brazilian (33) and Chilean (2) women. Chilean adults categorized as ‘high SB-active’ or ‘low SB-active’ showed significant negative associations with WC when compared to those in the ‘high SB-inactive’ category (29). Moreover, in Chilean adults a high ST was positively associated with WC (41). Furthermore, per one-hour decrease in sedentary time, there was a significant decrease in WC among Chilean adults (42). SB was also positively associated with WC in Colombian men (44), whereas SB, but not ST, was positively associated with WC in Chilean adults (43).

With respect to blood glucose, a positive association between ST and fasting glycemia was found in Chilean adults (29,41-43), whereas such association was not observed by others among Chilean adults (2) and in Colombian men (44).
## Table I. Cardiometabolic diseases (primary outcomes) reported in the included studies looking at its association with sedentary behavior in South American adults, January 2010 to December 2018

| First author (year) Reference | Population (age) Sample | SB Measure Domain | Outcome measurement | Measure of effect (primary outcome) | Interpretation | QA - Score (/20) |
|------------------------------|-------------------------|------------------|---------------------|-----------------------------------|----------------|------------------|
| Fernandes (2010) 34          | Brazilians (≥ 30 years) | Self-reported TV time | Self-report          | aPR for obesity                   | TV time was associated with obesity in women | 16               |
|                              | Men: 925                |                  |                     | Men: 0.98 (0.84-1.15)              |                |                  |
|                              | Women: 1,061            |                  |                     | Women: 1.42 (1.09-1.83)*           |                |                  |
| Poterico (2012) 35           | Peruvians (30.6 ± 10.1 years) | Self-reported TV time | Self-report          | aOR (95 % CI) to be obese: 2.6 (2.0-3.5)* | TV viewing frequency was associated with obesity | 16               |
| Cortes (2013) 33             | Brazilians (18-49 years) | Self-reported TV time | Objective           | aPR (CI 95 %) ≥ 5 times/week: Smokers: 1.7 (1.1-2.5)*. Non-smokers: 1.1 (0.9-1.2) | Watching TV ≥ 5 times per week was associated with greater prevalence of AO in smoker women | 16               |
| García (2014) 32             | Brazilians (30-40 years) | Self-reported SA time | Self-report          | aOR (95 % CI) for obesity: Men: 1.27 (1.15-1.41)*  Women: 1.24 (1.04-1.48)* | SB was associated with obesity | 17               |
| Crist-Montero (2017) 29      | Chileans (46.4 ± 18.6 years) | Self-reported ST | Objective            | Reference (high SB/inactive) aOR to have CO: high-SB/active: 0.79 (0.65-0.96)* low-SB/active: 0.71 (0.59-0.84)* | Less sitting time was associated with less chance to have CO | 17               |
| Mussi (2017) 31              | Brazilians (18-25 years) | Self-reported ST | Objective            | Area under ROC curve (95 % CI): Obesity = 0.50 (0.30-0.71) AO = 0.66 (0.57-0.75)* | ST during the week was powerful enough to discriminate AO, but not obesity | 15               |
| Petermann (2017) 30          | Chileans (45.5 ± 18.2 years) | Self-reported ST | Objective            | aOR (95 % CI) to be obese in the highest tercile of ST: Men: [1.97 (1.45-2.68]* Women: [0.88 (0.69-1.12)] | ST was a risk factor for CO among men | 16               |
| Martínez (2018) 28           | Chilean (47 years)      | Self-reported ST | Objective            | aOR (95 % CI) for CO: ST ≥ 4 h/day vs. < 4 h/day: 1.15 (1.02-1.30)* | High ST was associated with central obesity | 15               |
Table I (Cont.). Cardiometabolic diseases (primary outcomes) reported in the included studies looking at its association with sedentary behavior in South American adults, January 2010 to December 2018

| First author (year) Reference | Population (age) Sample | SB Measure Domain | Outcome measurement | Results measure of effect (primary outcome) | Interpretation | QA - Score (/20) |
|------------------------------|--------------------------|-------------------|--------------------|-------------------------------------------|---------------|-----------------|
| **Diabetes**                 |                          |                   |                    |                                           |               |                 |
| Garcia (2014) 32             | Brazilians (30-40 years) Men: 33,139; Women: 14,338 | Self-reported SA time | Self-report         | OR (CI 95 %): for DM: Men: 1.06 (0.88-1.27) Women: 0.86 (0.65-1.13) | SB was not associated with DM | 17 |
| Silva (2015) 38              | Brazilians (60-80 years) Men: 1,256 Women: 2,009 | Self-reported ST | Self-report         | aOR (CI 95 %) to have DM: 1.351 (1.057-1.729)* A daily ST of > 5.5 h was associated with high chances to have DM | 16 |
| Cristi-Montero (2017) 29     | Chileans (46.4 ± 18.6 years) Men: 2,036 Women: 3,004 | Self-reported ST | Objective          | Reference (high SB/inactive) OR to have DM: high-SB/active: 0.45 (0.35-0.59)* low-SB/active: 0.44 (0.34-0.56) | Less ST was associated with less chance to have DM | 17 |
| Diaz-Martinez (2017) 2       | Chileans (41.6 ± 18.6 years) Men: 1,783 Women: 2,674 | Self-reported ST | Objective          | OR (95 % CI) for DM: increase of 1h of ST as independent factor: Men: 1.10 (1.04 – 1.16)* Women: 1.08 (1.02 – 1.14)† | SB was associated with DM and cardiometabolic risk factors | 16 |
| Lemes (2018) 36              | Brazilians (68.9 ± 8.4 years) Men: 147 Women: 410 | Self-reported SA time TV time | Objective          | OR (95 % CI) for DM: overall SA ≥ 3 h/day vs. < 3 h/day: 1.60 (1.09-2.36)* TV time ≥ 3 h/day vs. < 3 h/day: 1.61 (1.11–2.33)* | TV time and overall SA was associated with higher prevalence of DM | 17 |
| Martinez (2018) 28           | Chileans (47 years) Men: 2,047 Women: 2,993 | Self-reported ST | Objective          | OR (95 % CI) for DM: ST ≥ 4 h/day vs. < 4 h/day: 1.27 (1.06-1.52)* | High sitting time was associated with DM | 15 |
| Werneck (2018) 37            | Brazilians (≥18 years) Men: 25,920 Women: 34,282 | Self-reported TV time | Self-report         | aOR (95 % CI) for DM: men [1.64 (1.23-2.67)*] women [1.33 (1.09-1.63)*] | High levels of TV viewing were associated with greater odds of DM | 17 |

(Continuation in the next page)
Table I (Cont.). Cardiometabolic diseases (primary outcomes) reported in the included studies looking at its association with sedentary behavior in South American adults, January 2010 to December 2018

| First author (year) | Reference | Population (age) | Sample | SB Measure | Domain | Outcome measurement | Measure of effect (primary outcome) | Interpretation | QA - Score (20) | Results |
|---------------------|-----------|------------------|--------|------------|--------|---------------------|------------------------------------|----------------|----------------|---------|
| Garcia (2014)       | 32        | Brazilians (30-40 years) | Men: 33,139 Women: 14,338 | Self-reported SA time | Self-report | OR (CI 95 %) for ≥ 2 CD: | Men: one CD = 1.17 (1.09-1.26)*; and ≥ 2 CD = 1.61(1.46-1.78)* | SB was associated with ≥ 2 CD | 17 |
| Sebastião (2017)    | 40        | Brazilians (70.2 ± 7.5 years) | Men: 134 Women: 221 | Self-reported ST | Self-report | ST was not different in individuals presenting none, one or ≥ 2 CD (DM, hypertension or arthritis) | | |
| Ribeiro (2018)      | 39        | Brazilians (70.0 ± 7.6 years) | Men: 373 Women: 447 | Self-reported ST | Self-report | aOR (95 % CI) to have ≥ 2 CD: | Men: 1.76 (1.10-2.80)*; Women: 1.92 (1.15-3.28)* | ST was associated to a higher risk of having ≥ 2 CD (DM, hypertension or arthritis) | 15 |
| Werneck (2018)      | 37        | Brazilians (≥ 18 years) | Men: 25,920 Women: 34,282 | Self-reported TV time | Self-report | OR (95 % CI) for HD: | men [1.96 (1.43-2.69)*], women [1.30 (1.00-1.68)*] | High levels of TV viewing were associated with greater odds of HD | 17 |
| Cristi-Montero (2017) | 29       | Chileans (46.4 ± 18.6 years) | Men: 2,036 Women: 3,094 | Self-reported ST | Objective | Reference (high SB/inactive); OR to have MetS: | high-SB/inactive: 0.63 (0.49-0.82)*; low-SB/inactive: 0.72 (0.59-0.87)* | Less ST was associated with less chance to have MetS | 17 |
| Martínez (2018)     | 28        | Chileans (47 years) | Men: 2,047 Women: 2,993 | Self-reported ST | Objective | OR for MetS: | ST ≥ 4 h/day vs. < 4 h/day: 1.15 | High sitting time was associated with MetS | 15 |
| Werneck (2018)      | 37        | Brazilians (≥ 18 years) | Men: 25,920 Women: 34,282 | Self-reported TV time | Self-report | OR for hypertension: | OR for hypertension: | High levels of TV viewing were associated with greater odds of hypertension | 17 |
| First author (year) Reference | Population (age) Sample | SB Measure Domain | Outcome measurement | Measure of effect (primary outcome) | Results | Interpretation | QA - Score (/20) |
|-------------------------------|-------------------------|-------------------|---------------------|-----------------------------------|---------|----------------|------------------|
| **Hypertension**              |                         |                   |                     |                                   |         |                |                  |
| Garcia (2014) 32 |
| Brazilianis (30-40 years)  |
| Men: 33,139                 |
| Women: 14,338               |
| Self-reported SA time       |
| Self-report                 |
| OR (CI 95 %) for hypertension: |
| Men: 1.00 (0.82-1.22)       |
| Women: 1.04 (0.89-1.22)     |
| SB was not associated with hypertension 17 |
| Cristi-Montero (2017) 29 |
| Chileans (46.4 ± 18.6 years) |
| Men: 2,036                  |
| Women: 3,004                |
| Self-reported ST            |
| Objective                   |
| Reference (high SB/inactive) |
| OR to have hypertension: |
| high-SB/active: 0.52 (0.43-0.63)† |
| low-SB/active: 0.60 (0.50-0.72)† |
| Less ST was associated with less chance to have hypertension 17 |
| Martinez (2018) 28 |
| Chileans (47 years)         |
| Men: 2,047                  |
| Women: 2,993                |
| Self-reported ST            |
| Objective                   |
| OR (95 % CI) for hypertension: |
| ST ≥ 4 h/day vs. < 4 h/day: |
| Men: 1.00 (0.82-1.22)       |
| Women: 1.04 (0.89-1.22)     |
| SB was not associated with hypertension 15 |
| Werneck (2018) 37 |
| Brazilianis (≥ 18 years)    |
| Men: 25,920                 |
| Women: 34,282               |
| Self-reported TV time       |
| Self-report                 |
| aOR (95 % CI) for hypertension: |
| Men [1.36 (1.14-1.63)],   |
| Women [1.20 (1.05-1.37)]   |
| High levels of TV viewing were associated with greater odds of hypertension 17 |

*p < 0.05; †p < 0.01. Source: prepared by the authors from the study results. AO: abdominal obesity; aOR: adjusted odds ratio; aPR: adjusted prevalence ratio; BMI: body mass index; CI: confidence interval; CO: central obesity; CD: chronic disease; DM: diabetes mellitus; HC: hypercholesterolemia; Li: lower limit; MetS: metabolic syndrome; NR: not reported; OR: odds ratio; PA: physical activity; QA: quality assessment; ROC: receiver operating characteristic; SA: sedentary activity; SB: sedentary behavior; ST: sitting time; TV: television; TC: total cholesterol; WC: waist circumference.
A positive association of ST with fasting insulinemia was observed in Chilean adults (41-43) and in Colombian men (44). Furthermore, SB was positively associated with HOMA-IR in Chilean adults (41-43) and in Colombian men (44). Regarding HbA1c, Chilean ‘high SB-active’ adults showed lower values of HbA1c when compared to those in the ‘low SB-active’ category (29). However, no association of SB with HbA1c was reported for Chilean adults (2). Moreover, Chilean adults exhibited a positive association of ST with serum lipids (41-43). Likewise, ST was positively associated with serum TG and HDL-c levels among Colombian men (44). Notwithstanding this, ST during the week or weekend did not discriminate any serum lipid abnormalities in Brazilian women (31). Among Brazilians, BMI and obesity were positively associated with SB in women only (34). Moreover, SB at work was positively associated with chronic diseases (e.g., obesity, hypertension, CCD, and hypercholesterolemia) especially in men as compared to women (32). Finally, a higher adjusted prevalence ratio of dyslipidemia was reported for Brazilian women but was not reported for men (46).

**SUBGROUP ANALYSES**

Subgroup analyses for sex and domain-specific sedentary behavior were performed. Between-sex differences were found in some studies. For instance, obesity (30) and BMI (2) were found to be positively associated with ST in Chilean men only, whereas it was negative in women (28). Waist circumference was found to be positively associated with SB in Chilean women only (2). Among Brazilians, BMI and obesity were positively associated with SB in women only (34). Moreover, SB at work was positively associated with chronic diseases (e.g., obesity, hypertension, CCD, and hypercholesterolemia) especially in men as compared to women (32). Finally, a higher adjusted prevalence ratio of dyslipidemia was reported for Brazilian women but was not reported for men (46).
Table II. Related biomarkers (secondary outcomes) of included studies looking at the association of sedentary behavior and cardiometabolic diseases in South American adults, January 2010 to December 2018

| First author (year) Reference | Population (age) Sample | SB Measure Domain | Outcome measurement | Results Interpretation | QA - Score (/20) |
|------------------------------|-------------------------|-------------------|---------------------|------------------------|------------------|
| Fernandes (2010) 34          | Brazilians (≥ 30 years) Men: 925; Women: 1,061 | Self-reported TV time | Self-report | The highest frequency of TV viewing was significantly associated with BMI in women (aPR: 1.42), but not in men (aPR: 0.96) | 15 |
| Cellis-Morales (2012) 43      | Chileans (37.5 ± 12.8 years) Men: 149; Women: 177 | Accelerometer ST Self-reported SB time | Objective | Significant trends to increase BMI and % BF with increasing time spent in SB (accelerometer) and ST (IPAQ) | 17 |
| Suárez (2012) 44             | Colombians (46.7 ± 6.6 years) Men: 137; Women: 177 | Self-reported ST | Objective | Higher ST was significantly correlated with BMI (rho: 0.225) and % BF (rho: 0.249) in men | 14 |
| Salas (2016) 42              | Chileans (36.6 ± 12.5 years) Men: 137; Women: 177 | Accelerometer SB time | Objective | Per one hour decrease in SB, there were significant improvements in BMI (-0.93 and -0.62 kg/m²) in physically active and inactive and in % BF (-1.04 %) in active men and women | 15 |
| Cristi-Montero (2017) 29     | Chileans (46.4 ± 18.6 years) Men: 2,036; Women: 3,004 | Self-reported ST | Objective | Compared to ‘High-SB & Inactive’ group, those individuals in ‘High-SB & Active’ and ‘Low-SB & Active’ groups showed significant negative associations with BMI (≥ 30.0 kg/m²) (standardized β: -0.182 and -0.156, respectively) | 17 |
| Diaz-Martinez (2017) 2       | Chileans (41.6 ± 18.6 years) Men: 1,783; Women: 2,674 | Self-reported ST | Objective | BMI was positive and significantly associated with ST (> 8 h/day) in men but not women | 16 |
| Gonçalves (2017) 45          | Brazilians (18-65 years) Men: 521; Women: 890 | Self-reported ST | Self-report | Sitting time (>4 h/day) was not significantly associated with BMI in men and women with overweight classified as sedentary and not active (> 10 min PA/week - OR: 1.03, 95 % CI); > 150 min PA/week - OR: 1.11, 95 % CI) | 15 |
| Leiva (2017) 41              | Chileans (37.6 ± 12.9 years) Men: 141; Women: 181 | Accelerometer ST | Objective | Per one hour increase in ST, there were significant adverse changes in BMI (0.69 kg/m²) and % BF (1.03 %) in men and women | 15 |
| Martínez (2018) 28           | Chileans (47 years) Men: 2,047; Women: 2,993 | Self-reported ST | Objective | Sitting time was negatively (OR: 0.69: 95 % CI: 0.59-0.81) associated with BMI (> 25 kg/m²) in women, but positively in men (OR: 1.21: 95 % CI: 1.00-1.49) | 15 |

(Continuation in the next page)
Table II (Cont.). Related biomarkers (secondary outcomes) of included studies looking at the association of sedentary behavior and cardiometabolic diseases in South American adults, January 2010 to December 2018

| First author reference | Population (age) | SB Measure Domain | Outcome measurement | Results | Interpretation | QA - Score (/20) |
|------------------------|------------------|-------------------|---------------------|---------|----------------|-----------------|
| **Waist circumference** |                  |                   |                     |         |                |                 |
| Cels-Morales (2012) 43 | Chileans (37.5 ± 12.8 years) Men: 149; Women: 177 | Accelerometer ST Self-reported SB time | Objective | Significant trends to increase WC with increasing time spent in sedentary time (accelerometer), but not with increasing ST (IPAQ) | 17 |
| Suárez (2012) 44      | Colombians (46.7 ± 6.6 years) Women: 69 | Self-reported ST | Objective | Sitting time was positively significantly correlated with WC (rho 0.266) in men | 14 |
| Cortes (2013) 33      | Brazilians (18-49 years) Women: 13,262 | Self-reported TV time | Objective | WC (≥ 80.0 cm) was positively associated with TV time (≥ 5 times/week) | 16 |
| Salas (2016) 42       | Chileans (36.6 ± 12.5 years) Men: 137; Women: 177 | Accelerometer SB time | Objective | Per one hour decrease in SB, there were significant improvements in WC (-2.32 and -1.65 cm) in physically active and inactive men and women | 15 |
| Cristi-Montero (2017) 29 | Chileans (46.4 ± 18.6 years) Men: 2,036; Women: 3,004 | Self-reported ST | Objective | Adults categorized as ‘High-SB & Active’ or ‘Low-SB & Active’ showed significant negative associations with WC (standardized β: -0.258 and -0.233, respectively), compared to adults categorized as ‘High-SB & Inactive’ | 17 |
| Díaz-Martinez (2017) 2 | Chileans (41.6 ± 18.6 years) Men: 1,783; Women: 2,674 | Self-reported ST | Objective | WC was significantly positively associated with ST (> 8 h/day) in women but not men | 16 |
| Leiva (2017) 41       | Chileans (37.6 ± 12.9 years) Men: 141; Women: 181 | Accelerometer SB time | Objective | Per one hour increase in ST there was significant adverse change in WC (1.95 cm) in men and women | 15 |
| **Glucose**           |                  |                   |                     |         |                |                 |
| Cels-Morales (2012) 43 | Chileans (37.5 ± 12.8 years) Men: 149; Women: 177 | Accelerometer ST Self-reported SB time | Objective | Significant trends to increase fasting glucose with increasing time spent in SB (accelerometer) and ST (IPAQ) | 17 |
| Suárez (2012) 44      | Colombians (46.7 ± 6.6 years) Women: 69 | Self-reported ST | Objective | Sitting time was not significantly correlated with fasting glucose (rho: 0.069) in men | 14 |
| Salas (2016) 42       | Chileans (36.6 ± 12.5 years) Men: 137; Women: 177 | Accelerometer SB time | Objective | Per one hour decrease in SB, there was a significant improvement in glucose (-8.46 and -4.68 mg.dL-1) in physically active and inactive men and women. | 15 |
| Cristi-Montero (2017) 29 | Chileans (46.4 ± 18.6 years) Men: 2,036; Women: 3,004 | Self-reported ST | Objective | Adults categorized as ‘High-SB & Active’ or ‘Low-SB & Active’ showed significant negative associations with fasting glucose (standardized β: -0.238 and -0.174, respectively), compared to adults categorized as ‘High-SB & Inactive’ | 17 |
| Díaz-Martinez (2017) 2 | Chileans (41.6 ± 18.6 years) Men: 1,783; Women: 2,674 | Self-reported ST | Objective | Sitting time (> 8 h/day) was not associated with fasting glucose levels in men and women | 16 |
| Leiva (2017) 41       | Chileans (37.6 ± 12.9 years) Men: 141; Women: 181 | Accelerometer ST | Objective | Per one hour increase in ST there was significant adverse change in glucose (4.79 mg/dL) in men and women | 15 |
| First author | Reference | Population | Sample | SB Measure | Domain | Outcome measurement | Interpretation | QA - Score (/20) |
|-------------|-----------|------------|--------|------------|--------|---------------------|----------------|------------------|
| Celis-Morales | (2012) 43 | Chileans (37.5 ± 12.8 years) | Men: 49; Women: 17 | Accelerometer ST | Self-reported SB time | Lipid profile, hypercholesterolemia, dyslipidemia | Significant trends to increase TG and reduce HDL-c levels with increasing time spent in SB (accelerometer) and ST (IPAQ). | 17 |
| Suarez | (2012) 44 | Colombians (46.7 ± 6.6 years) | Women: 69 | Self-reported ST | | | Sitting time significantly correlated with serum TG (rho: 0.258) and HDL-c (rho: -0.27), but not with TC (rho: 0.07) and LDL-c (rho: 0.082) levels in men. | 14 |
| Moraes | (2013) 46 | Brazilians – urban areas 30-65 years Men: 437; Women: 376 | | Self-reported ST | | | Higher time spent in SA (> 4 h/day) at work (i.e. TV viewing and sedentary work) was associated with serum lipid profile (e.g., TC, TG, HDL-c, LDL-c, and VLDL-c) and insulin resistance (HOMA-IR) in men and women. | 15 |
| Garcia | (2014) 33 | Brazilians (30-40 years) | Men: 33,139; Women: 14,338 | Self-reported SA time | | | Higher time spent in SA (> 4 h/day) at work was significantly associated with hypercholesterolemia. | 17 |
| Salas | (2016) 42 | Chileans (36.6 ± 12.5 years) | Men: 137; Women: 177 | Accelerometer SB time | | | Per one hour decrease in SB, there were significant improvements in insulin (-2.12 and -1.77 pmol/L) and HOMA-IR (-0.81 and 0.56) in physically active and inactive men and women. | 15 |
| Cristi-Montero | (2017) 29 | Chileans (37.5 ± 12.8 years) | Men: 49; Women: 17 | Self-reported SB time | | | Adults categorized as 'High-SB & Active' or 'Low-SB & Active' showed significant negative associations with HbA1c (standardized β: -0.286 and -0.183, respectively) in men and women. | 17 |
| Leiva | (2017) 41 | Chileans (37.6 ± 12.9 years) | Men: 141; Women: 181 | Accelerometer ST | | | Per one hour increase in ST there were significant adverse changes in TC (9.73 mg/dL), HDL-c (-3.50 mg/dL), LDL-c (10.7 mg/dL), and TG in men and women. | 15 |
| Mussi | (2017) 31 | Brazilians (18-25 years) | Men: 26; Women: 161 | Self-reported ST | | | Sitting time during the week or weekend did not discriminate serum lipid (e.g., TC, TG, HDL-c) abnormality among women. | 15 |
| Diaz-Martinez | (2017) 2 | Chileans (41.6 ± 18.6 years) | Men: 1,783; Women: 2,184 | Self-reported ST | | | The time spent sitting (> 8h/day) was not significantly associated with HbA1c levels in men and women. | 16 |
| Leiva | (2017) 42 | Chileans (37.6 ± 12.9 years) | Men: 141; Women: 181 | Accelerometer ST | | | The time spent sitting (> 8h/day) was not significantly associated with HbA1c levels in men and women. | 15 |

*SB: sedentary behavior; BMI: body mass index; HbA1c: glycated hemoglobin; HOMA-IR: homeostasis model assessment-insulin resistance; HDL-c: high density lipoprotein-cholesterol; LDL-c: low density lipoprotein-cholesterol; IPAQ: International Physical Activity Questionnaire; WC: waist circumference. Source: prepared by the authors from the study results.*
The investigation of the association of domain-specific SB with CMD and related biomarkers found that 12 studies reported on ST (2, 28-31, 38-41, 44-46) and four on TV time or TV watching frequency (33-35, 37). The remaining four studies looked at general measures of SB (32, 36, 42, 43). The results demonstrate equivalent trends for each SB domain.

**DISCUSSION**

In this review we thought to systematically review the literature to examine whether domain-specific SB is associated with CMD and its risk factors in South American adults. This is the first review to explore the association of SB with CMD and its risk factors in South American adults. This study summarized the published evidence over the review period (i.e., January 2010 to December 2018), and observed that longer SB (i.e., sitting, TV watching, overall SB time) was associated with CMD (i.e., obesity, DM, and CCD) as well as with its anthropometric (i.e., BMI, WC, and % BF) and metabolic biomarkers (i.e., lipid profile and glucose). Nevertheless, evidence is limited when studies reported the relationship between SB and CMD (i.e., hypertension and MetS) as well as its biomarkers (i.e., HOMA-IR, BI, dyslipidemia, and hypercholesterolemia). It has to be taken into consideration that reports on the associations of SB with CMD and related biomarkers (i.e., CMD and its risk factors) are scarce.

Our results complement the previous evidence from systematic reviews (17, 18, 47) and meta-analyses (16, 19, 48), which demonstrate that high SB is associated with CMD and risk factors in different regions of the world. Based on the studies reviewed here, South American adults are likely at larger risk for developing obesity, diabetes, and CCD when incurring in prolonged SB time. Such findings are consistent with those reported about different populations (18, 19, 48). A significant association of longer time spent in SB was found with obesity (8 studies), DM (6 but 1 study), and CCD (4 studies) in the 20 studies reviewed. For instance, South American (i.e., Chilean) adult men in the highest tertile of ST had a 97 % higher risk for obesity (30). In addition, Lemes et al. (36) reported a 60 % higher risk for diabetes in South American (i.e., Chilean) adults who spent ≥ 3 h/day in sedentary activities, whereas those South American (i.e., Brazilian) men and women who reported a daily ST of 4 h had a 76 % and 82 % higher risk of presenting ≥ 2 CCDs, respectively (39). However, the magnitude of the risk of developing DM with longer sedentary time is approximately twice as high as in the general population and, in general, longer sedentary time is associated with a 14 % greater risk of cardiovascular disease (19). A significant relationship between increased SB and higher BMI (8 but 1 study), WC (7 studies), and % BF (4 studies) was also reported in the reviewed studies. These results are similar to those previously reported in different populations (49, 50) and, more importantly, are consistent with the association of longer time spent in SB with obesity as observed in the present review.

Concerning sex differences, only seven studies reported distinct associations of SB with CMD and its biomarkers. Based on those studies, it appears that South American men are at higher risk for obesity and high BMI when spending a lot of time in SB, as compared to women. Such finding is contradictory to the report that women are more inactive (33.9 %) than are men (27.9 %) worldwide (21). Regarding domain-specific SB in South American adults, most studies associated a higher risk for CMD with longer periods of ST (13 studies), which was followed by the extent of TV (6 studies) and SB (4 studies) times. Whether there are distinct health damages for the different specific domains rather than length of exposure is not known, and thus demands further investigation.

Overall, this systematic review presented a positive association between SB and CMD in South American adults. Whether these risks are truly greater in this population group further studies are needed to elucidate, inasmuch as the insufficiency of available direct evidence on this sub-population limits the ability to ascertain such findings. Despite this, the findings presented here are of relevance for South American governments, since they reinforce the need for public policies to face the deleterious consequences of a sedentary lifestyle.

**STUDY LIMITATIONS**

This review has some limitations. First, there is a lack of studies examining the relationship of SB with CMD and related biomarkers in South American adults, which make it difficult to generalize our findings. Eleven out of 20 analyzed studies were carried out in Brazilian, seven in Chilean, one in Colombian and one in Peruvian adult men and women. Although the population of these four countries accounts for 73 % (~ 316 million people) of the South American population, no data on adults from the other South American countries were found. Moreover, the selective reporting of positive outcomes in individual studies could potentially reduce the generalizability of our findings. Secondly, the predominant use of self-reported data in individual studies is another limitation, since self-reported data are highly susceptible to biases that might have affected the magnitudes of our results. Only three studies in the present review reported on the use of objective measures (accelerometers) of SB time in Chileans (Leiva et al., 2017; Salas et al., 2016; Coles-Morales et al., 2012), but did not assess the association of SB with CMD as their primary outcome. Finally, although the quality of the selected studies, all 20 studies were cross-sectional in design, and as such they may include selection and reverse causality biases, as well as residual confounders. Therefore, since there is no temporality described between the sedentary behavior and the outcomes, interpretations should be cautious.

**CONCLUSIONS**

In conclusion, spending a long time in an overall SB, mainly in a sitting position or watching TV, was positively associated with the occurrence of CMD (i.e., obesity, diabetes, and ≥ 2 CCDs) and related biomarker (i.e., BMI, WC, % BF, lipid profile, and blood glucose) elevations in South American adult men and women.
ACKNOWLEDGEMENTS

The authors thank Dr. Osvaldo Costa Moreira for his technical support.

REFERENCES

1. The World Bank. World Bank Country and Lending Groups. Available at: https://datahelpdesk.worldbank.org/knowledgebase/articles/906519-world-bank-country-and-lending-groups. [Accessed 26 March 2019].

2. Díaz-Martínez X, Stekel L, Martínez MA, Leiva AM, Salas-Bravo C, Labrada AM, et al. Higher levels of self-reported sitting time is associated with higher risk of type 2 diabetes independent of physical activity in Chile. J Public Health 2017;1-7. DOI: 10.1093/aje/kip029.

3. Rubinstein AL, Izarola VE, Calandrini M, Elleriga M, Gutierrez L, Lanas F, et al. Multiple Cardiovascular Risk Factors in the Southern Cone of Latin America: A Population-based Study in Argentina, Chile, and Uruguay. Int J Cardiol 2015;183:82-8. DOI: 10.1016/j.ijcard.2015.01.062.

4. Lanas F, Sérón P, Lanas A. Cardiovascular Disease in Latin America: The Growing Epidemic. Prog Cardiovasc Dis 2014;57:262-7. DOI: 10.1016/j.pcd.2014.07.007.

5. López-Jaramillo P, Sánchez RA, Díaz M, Cobos L, Bryce A, Parra Carrillo JZ, et al. Latin American consensus on hypertension in patients with diabetes type 2 and metabolic syndrome. J Hypertens 2013;31:223-8. DOI: 10.1097/HJH.0b013e32835c4444.

6. Avezum A, Braga J, Santos I, Guimarães HP, Marín-Neto JA, Piegas LS. Cardiovascular disease in South America: current status and opportunities for prevention. Heart 2009;95:1475-82. DOI: 10.1136/hrt.2008.156331.

7. Biddle SJH, Bennie JA, Bauman AE, Chau JY, Dunstan D, Owen N, et al. Too much sitting and all-cause mortality: is there a causal link? BMC Public Health 2016;16:635. DOI: 10.1186/s12889-016-1307-3.

8. Rezende LF, Sá TH, Mielke GI, Viscondi JYK, Rey-Lopez JP, Garcia LMT. All cause mortality attributable to sitting time: analysis of 54 countries worldwide. Am J Prev Med 2016;51:253-63. DOI: 10.1016/j.amepre.2016.01.022.

9. Young D, Hivet M-F, Alhassan S, Camhi SM, Ferguson JF, Katzmarzyk PT, et al. Sedentary behavior and cardiovascular morbidity and mortality: a science advisory from the American Heart Association. Circulation 2016;134:e262-79. DOI: 10.1161/CIR.0000000000000440.

10. Chau JY, Grunseit AC, Choy T. Daily sitting time and all-cause mortality: a meta-analysis. PLoS One 2013;8:e68000. DOI: 10.1371/journal.pone.0086000.

11. Guthold R, Stevens GA, Riley LM, Bull FC, Guthold R, Haskell W, Ekelund U, et al. Factores asociados al desarrollo de obesidad en Chile: resultados de la Encuesta Nacional de Salud 2009-2010. Rev Med Chil 2016;144:262-79. DOI: 10.4067/S0034-98872016000300016.

12. García LM, da Silva KS, Del Duca GF, da Costa FF, Nahas MV. Sedentary behaviour and cardiovascular risk factors in Chilean adults. J Public Health 2017;40:485-92. DOI: 10.1093/p,mid/mfx134.

13. Petermann F, Eliana D, Labrada AM, Martínez SA, Leiva AN, Garrido-Mendez CA, et al. Factors associated with obesity in Chile: results of the Encuesta Nacional de Salud 2009-2010. Rev Med Chil 2016;144:716-22. DOI: 10.4067/S0034-98872017000100013.

14. Biddle SJH, camel C, Stekel L, Petermann F, Garrido-Mendez A, Díaz-Martínez X, Salas-Bravo C, et al. Joint effect of physical activity and sedentary behaviour on cardiovascular risk factors in Chilean adults. J Public Health 2017;40:485-92. DOI: 10.1093/pmid/mfx134.

15. Bottino MC, Pitaña FJG, Pires CGS. Cumulative sitting time as discriminator of overweight, obesity, abdominal obesity and lipid disorders in nursing university. Rev Bras Cirurgia de Veneros 2017;19:40-9. DOI: 10.5380/0990-0337.2017v19n14p40.

16. García LM, da Silva KS, Del Duca GF, da Costa FF, Nahas MV. Sedentary behaviors, leisure-time physical inactivity, and chronic diseases in Brazilian workers: a cross sectional study. J Phys Act Health 2014;11:1622-34. DOI: 10.1123/jpah.2012-0423.

17. Cortes TR, Schlussler mm, Franco-Sena AB, Rebolo F, Kac G. Television viewing and abdominal obesity in women according to smoking status: results from a large cross-sectional population-based study in Brazil. Rev Bras Epidemiol 2013;16:137-45. DOI: 10.1590/S1135-79032013000100013.

18. Fernandes RA, Christofaro DG, Casonatto J, Costa Rosa CS, Costa FF, Freitas IF, et al. Leisure time behaviors: prevalence, correlates and associations with overweight in Brazilian adults: a cross-sectional analysis. Rev Med Chil 2010;138:1-29.S5. DOI: 10.4067/S0034-98872010000100004.

19. Potencio JC, Bernabé-Ortiz A, Mola CL, Miranda JA. Association between television viewing and obesity in Peruvian women. Rev Saúde Pública 2012;46:610-16. DOI: 10.1590/S0034-89102012000400004.

20. Lemos R, Sul T, Turl-Lynch BC, Lee DC, Blair SN, Fernandes RA, et al. Sedentary behaviour is associated with diabetes mellitus in adults: findings of a cross-sectional analysis from the Brazilian National Health System. J Public Health 2017;1-8. DOI: 10.1093/pmid/mfx169.

21. Wurmack AO, Cyriac ES, Collins PJ, Ronque ERV, Szwarcowicz CL, Sandinb LA, et al. TV Viewing in 60,202 Adults From the National Brazilian Health Survey: Prevalence, Correlates, and Associations With Chronic Diseases. J Phys Act Health 2018;15:510-15. DOI: 10.1123/jpah.2017-0317.

22. Silva RC, Meneguci J, Martins TI, Santos AS, Sasaki JE, Tribesa S, et al. Association between time spent sitting and diabetes mellitus in older adults: a population-based study. Rev Bras Cirurgia de Veneros 2015;17:379-88. DOI: 10.5380/0990-0337.2015v17n4p379.
A SYSTEMATIC REVIEW OF CROSS-SECTIONAL STUDIES ON THE ASSOCIATION OF SEDENTARY BEHAVIOR WITH CARDIOMETABOLIC DISEASES AND RELATED BIOMARKERS IN SOUTH AMERICAN ADULTS

39. Ribeiro AS, Pereira LC, Silva DPP, Santos LD, Schoenfeld BJ, Teixeira DC, et al. Physical Activity and Sitting Time are Specifically Associated With Multiple Chronic Diseases and Medicine Intake in Brazilian Older Adults. J Aging Phys Act 2018;16:608-13. DOI: 10.1123/japa.2017-0271

40. Sebastião E, Galvez PAE, Nakamura PM, Papini CB, Kokubun E, Gobbi S. Activity behavior, nutritional status and perceived health in older Brazilian adults: Does the number of chronic diseases matter? Geriatr Gerontol Int 2017;17:2376-82. DOI: 10.1111/ggi.13078

41. Leiva AM, Martínez MA, Cristi-Montero C, Salas C, Ramírez-Campillo R, Díaz-Martínez X, et al. El sedentarismo se asocia a un incremento de factores de riesgo cardiovascular y metabólicos independiente de los niveles de actividad física. Rev Med Chile 2017;145:458-67. DOI: 10.4067/S0034-98872017000400006

42. Salas C, Cristi-Montero C, Fan Y, Durán E, Labarría AM, Martínez MA, et al. Being physically active modifies the detrimental effect of sedentary behavior on obesity and cardiometabolic markers in adults. Rev Med Chile 2016;144:1400-09. DOI: 10.4067/S0034-98872016001100005

43. Celis-Morales CA, Perez-Bravo F, Ibanez L, Salas C, Bailey ME, Gill JM. Objective vs. Self-Reported Physical Activity and Sedentary Time: Effects of Measurement Method on Relationships with Risk Biomarkers. PLoS ONE 2012;7:e36345. DOI: 10.1371/journal.pone.0036345

44. Suárez MF, Arbeláez A, Mosquera M, Ramírez-Vélez R, Pita ACA. Los niveles de ferritina y los marcadores de riesgo cardiovascular se correlacionan con mayor tiempo sedentario auto-reportado en hombres aparentemente sanos. Rev Colom Cardiol 2012;19:4-10. DOI: 10.1016/S0120-5633(12)70097-9

45. Gonçalves PB, Lopes AAS, Silva AAP, Silva JSB, Silva AT, Becker LA, et al. Combined physical activity and sitting time and their contribution to body mass index in adults. Rev Bras Cineantropom Hum 2017;19:174-84. DOI: 10.5007/1980-0037.2017v19n2p174

46. Moraes SA, Checchio MV, Freitas IC, Dampeia e fatores associados em adultos residentes em Ribeirão Preto, SP: resultados do Projeto EPIDCV. Arq Bras Endocrinol Metab 2013;57:691-701. DOI: 10.1590/0004-27302013000900004

47. Wirth K, Klenk J, Brefka S, Dallmeier D, Faehling K, Roqué F, Figuls M, et al. Biomarkers associated with sedentary behaviour in older adults: A systematic review. Ageing Res Rev 2017;35:87-111. DOI: 10.1016/j.arr.2016.12.002

48. Wilmot EG, Edwardson CL, Achana FA, Davies MJ, Gorely T, Gray LJ, et al. Sedentary time in adults and the association with diabetes, cardiovascular disease and death: systematic review and meta-analysis. Diabetologia 2012;55:2895-905. DOI: 10.1007/s00125-012-2677-z

49. Belletiere J, Winkler EAH, Chastin SFM, Kerr J, Owen N, Dunstan DW, et al. Associations of sitting accumulation patterns with cardio-metabolic risk biomarkers in Australian adults. PLoS ONE 2012;7:e36345. DOI: 10.1371/journal.pone.01371

50. Thorp AA, Owen N, Neuneha M, Dunstan DW. Sedentary behaviors and subsequent health outcomes in adults: a systematic review of longitudinal studies, 1996-2011. Am J Prev Med 2011;41:207-15. DOI: 10.1016/j.amepre.2011.05.004