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# Prevalence of carotid artery calcifications among 2,500 digital panoramic radiographs of an adult Brazilian population

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# Abstract

Background: The aim of the present study was to analyze the epidemiological data of digital panoramic radiographs revealing suggestive images of carotid artery calcifications (CAC) from a Northeast Brazilian population. Material and Methods: A cross-sectional retrospective study was conducted with 2,500 digital panoramic radiographs obtained from a single imaging reference center in Northeast Brazil. Images from individuals of both sexes and older than 18 years were included and those that did not cover the region of cervical vertebrae or presented low radiographic quality were excluded. Data were analyzed regarding prevalence, location (bilateral, right or left), sex, and age using the Chi-square test at the significance level of 5%.

Results: An amount of 96 (4%) patients presented suggestive images of CAC. The female sex (p=0.003) and individuals aged up to 70 years (p=0.002) were statically significant. 40.4% were found bilaterally, 37.6% on the right side (p<0.001) and 22% on the left side.

Conclusions: In conclusion, this study showed a low prevalence of suggestive images of CAC in digital panoramic radiographs from a Northeast Brazilian population. It was observed a higher prevalence of CAC associated with female sex, older patients, and right side location.

Key words: Atherosclerosis, carotid artery, digital panoramic radiograph, prevalence.

# Introduction

Coronary artery disease is considered a major cause of worldwide morbidity and mortality (1), being a multifactorial disorder in which multiple genetic variants are combined with several environmental risk factors and deleterious lifestyles (2). It is characterized by the long-term formation of atheromatous plaques within the arterial walls, culminating in atherothrombotic obstructive lesions leading to local tissue damage, such as ischemic stroke, transient ischemic attack, or amaurosis fugax (3). These aspects make the presence of atherosclerotic plaques a relevant health problem since heart attack is one of the main causes of death in the world, as well as it is closely related to severe impairments for the patients affected by this disease (4).

Carotid artery calcifications (CAC) can be visualized on panoramic radiographs, imaging exams that are routinely used in the evaluation of patients with dental issues (5). The presence of CAC in these exams is an indicator of the risk of developing possible cardiovascular events, which justifies the clinical interest of the dentist in being able to identify these alterations early (6). These alterations appear as single or multiple, noncontinuous nodular radiopaque images, located on the intervertebral junction C3-C4, about 1 to 2.5 cm inferior to posterior to the angle of the mandible, or as vertical radiopaque lines representing thin vascular wall calcifications (7,8).

The differential diagnosis of atherosclerotic plaques includes anatomical structures of the neck region such as hyoid bone, trichoid cartilage, the superior horn of the calcified and epiglottic thyroid cartilage, and lesions such as sialoliths, elongation, and calcifications of the styloid process, calcifications in the stylomandibular and stylohyoid ligaments, phleboliths, among others (5,6,8).

Epidemiological studies have shown that the prevalence of CAC found in panoramic radiographs, performed for dental reasons, is around 2% to 5% (9). However, a limited number of studies regarding CAC prevalence on digital panoramic radiographs from South America have been published to date. Thus, the present study aimed to evaluate the epidemiological data of digital panoramic radiographs showing suggestive images of CAC from a significant Brazilian population.

## **Material and Methods**

-Study design

A retrospective cross-sectional study was performed with a sample of 2500 digital panoramic radiographs obtained between December 2011 to December 2014 from a private dental imaging clinic, which is a reference service in the state of Ceará, Brazil. The present study was approved by the Research Ethics Committee under protocol number 285/11. It was used the Cranex D (Soredex, Tuusula, Finland) digital panoramic and cephalometric imaging system. The images were acquired with Frankfurt's horizontal plane parallel to the ground and they were generated with a resolution of 300 dpi and setting parameters adjusted according to the patient's size.

Images of individuals under the age of 18 years, that did not cover the region of cervical vertebrae, or that presented low radiographic quality, were excluded from this sample (n=111). Two previously calibrated examiners (kappa coefficient = 0.856) analyzed the radiographs in order to detect images suggestive of CAC. The images were analyzed using Adobe Photoshop® CC 2015 software (Adobe Systems Incorporated, California, USA) for saturation and contrast in order to standardize images during data evaluation (Fig. 1). CAC were considered present when it was observed heterogeneous

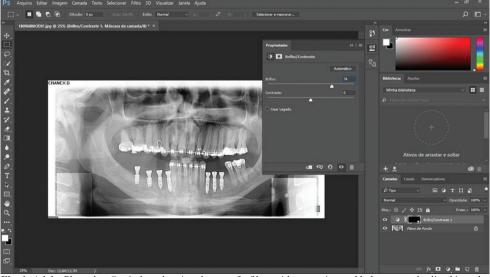


Fig. 1: Adobe Photoshop® window showing the use of a filter with saturation and balance standardized in order to analysis a bilateral CAC on panoramic radiograph.

radiopacities located into the intervertebral space between C3 and C4 (6). Before the radiographic analysis, two examiners were calibrated in order to establish uniform criteria for imaging evaluation. Cohen's Kappa statistic was applied and it was obtained an inter-rater agreement value higher than 0.80. Doubts about the presence of CAC were interpreted by two oral and maxillofacial radiologists. The two radiologists analyzed 118 doubts of the panoramic radiographs being the main differential diagnosis the cricoid cartilage. The studied variables were sex, age, and location (bilateral, right unilateral or left unilateral).

# -Statistical analysis

The data were tabulated in Microsoft Excel 2010 software regarding location (bilateral, right unilateral or left unilateral), gender and age in order to obtain data on the prevalence of calcifications and the results were submitted to statistical treatment by the Chi-square test at the significance level of 5%. Statistical Package for the Social Sciences (SPSS) software 15.0 version for Windows (SPSS Inc.®, Chicago, Illinois, USA) was used for all analysis.

#### Results

In this study, CAC prevalence was 4% (n=96) considering a sample with an estimated power of 99.6%. Table 1 showed a statically significant difference (p=0.003) between females (n=68; 70.8%) and males (n=28; 29.2%). Radiographs suggesting CAC were observed in individuals with age ranging from 25 to 89 years (mean age of  $54\pm13$  years). There was no statically significant difference (p=0.968) between male ( $54\pm15$  years) and female ( $54\pm12$  years) mean age. In addition, sex and age did not show a statically significant association with the right, left or both sides (p>0.05). The most prevalent and statically significant (p=0.002) age group was that one between 50 and 70 years (n=46; 47.9%), followed by individuals aged up to 50 years (n=38; 39.6%).

There was a statically significant amount of patients presenting suggestive images of CACs on the right side (p<0.001). However, there was no difference regarding the occurrence even in the left side (p=0.147) or in a bilateral presentation (p=0.110). According to Table 2, there was no statically significant association of sex with age groups (p=0.443), right side (p=0.635) or left side (p=0.379). Bilateral CAC did not show association with sex (p=0.198; Table 1) or age (p=0.598; Table 3).

# **Discussion**

Atherosclerotic plaques in carotid arteries have been extensively studied in the field of Medicine and Dentistry. Indeed, atherosclerotic plaques have been considered important predictors of cardiovascular disorders and, therefore, their identification could be useful as a tool for medical stratification and health education strategies

Table 1: Characterization of the sample according to sex, age and location

	n	%	<i>p</i> -Value
Sex			
Female	68*	70.8	0.003
Male	28	29.2	
Age (years)			
Up to 50	38*	39.6	0.002
51-70	46*	47.9	
> 70	12	12.5	
Right side			
No	21	21.9	< 0.001
Yes	75*	78.1	
Left side			
No	38	39.6	0.147
Yes	58	60.4	
Laterality			
Unilateral	59	61.5	0.110
Bilateral	37	38.5	

\*p<0.05, chi-square test. Data expressed as absolute and percentage frequency.

according to Gepner *et al.* (10). In a systematic review with a meta-analysis published in 2017 by Gorgui *et al.* (3), the prevalence of atheromatous plaques differed according to geographic regions: South Africa (38.1%), United States (17.2-43%), Japan (29.8%), Europe (5.4%), Spain (60%), and China (44.4-51%).

In dentistry literature, Friedlander and Lander (1981) (7) were the first researchers to report CAC visualized on panoramic radiographs, emphasizing the importance of this exam since it is a routine radiograph required in dental practice and, in many cases, the identification of CAC is an accidental finding (5). These aspects motivated the conduction of the present research, also reinforced by the retrospective collection of a substantial number of digital panoramic radiographs from a single reference center for maxillomandibular imaging exams in Northeast Brazil.

Almog *et al.* reported that the prevalence of CAC observed by panoramic radiographs has been usually described up to 5% with higher occurrence in patients with clinical conditions associated with atherosclerosis (9). The present study showed a percentage of 4% in an adult Northeast Brazilian population and since it had a non-intentional sample recruitment, clinical data were not obtained, which may explain the different prevalence in comparison with other investigations. Regarding epidemiological data on the worldwide prevalence of CAC in panoramic radiographs, there have been reported similar results to the present study in Germany, Saudi Arabia, and Japan (11-13). In Brazil, the largest

Table 2: Characterization of sex according to age, side and laterality.

	Se		
	Female (%)	Male (%)	<i>p</i> -Value
Age (years)			
Up to 50	26	12	0.443
	(38.2%)	(42.9%)	
51-70	35	11	
	(51.5%)	(39.3%)	
> 70	7	5	
	(10.3%)	(17.9%)	
Right side			
No	14	7	0.635
	(20.6%)	(25%)	
Yes	54	21	
	(79.4%)	(75%)	
Left side			
No	25	13	0.379
	(36.8%)	(46.4%)	
Yes	43	15	
	(63.2%)	(53.6%)	
Laterality			
Unilateral	39	20	0.198
	(57.4%)	(71.4%)	
Bilateral	29	8	
	(42.6%)	(28.6%)	

<sup>\*</sup>p<0.05, chi-square test. Data expressed as absolute and percentage frequency.

study that evaluated 8,338 panoramic radiographs was performed in the South region and it showed a CAC prevalence of 6.9% (14). Thus, the present study is the second one in Brazil that presented a significant sample size and it is the largest one from the Brazilian Northeast region to date.

In the present cross-sectional study, the age group between 50 and 70 years showed a statistical significance prevalence. A similar result was observed by Lee *et al.* (13) since the authors found a higher prevalence of images suggestive of atheromatous plaques in patients aged 40 to 70 years. These results reinforce the age as an important risk factor for CAC. In younger patients, only a small number of cases of carotid artery calcifications has been found. In spite of this finding, atheromatous plaques should be suspected at any age, justifying the evaluation of panoramic radiographs in younger patients (15).

This study showed a statistically significant prevalence of CAC in women, which has been commonly found in the literature. Friedlander and Altman (1981) (7) stated that this association may occur due to a decline in the estrogen level after menopause. Since it is a hormone that acts on lipoproteins metabolism by preventing the formation of atheromatous plaques, its reduced level may precipitate the formation of atheromas. Friedland-

Table 3: Characterization of the sample according to age versus sex, side and laterality.

	Up to 50	Age (years) 51-70	> 70	<i>p</i> -Value
Sex				
Female	26	35	7	0.443
	68.4%	76.1%	58.3%	
Male	12	11	5	
	31.6%	23.9%	41.7%	
Right side				
No	7	13	1	0.266
	18.4%	28.3%	8.3%	
Yes	31	33	11	
	81.6%	71.7%	91.7%	
Left side				
No	14	17	7	0.365
	36.8%	37.0%	58.3%	
Yes	24	29	5	
	63.2%	63.0%	41.7%	
Laterality				
Unilateral	21	30	8	0.598
	55.3%	65.2%	66.7%	
Bilateral	17	16	4	
	44.7%	34.8%	33.3%	1

<sup>\*</sup>p<0.05, chi-square test. Data expressed as absolute and percentage frequency.

er *et al.* (16) evaluated the panoramic radiographs of women with a history of amenorrhea greater than 12 months and images suggestive of CAC appeared in 31% of the radiographs. Patil *et al.* (17) reported a 22.9% prevalence of CAC in women aged over 50 years among a sample of 1,214 panoramic radiographs. Regarding occurrence of CAC in men, it has been considered that the smoking habit plays an important risk factor for its development, which differs in relation to women as described by Cohen *et al.* (18).

The right side showed a statistically significant prevalence in the present study. Similar findings were found by Ohba *et al.* (19), which revealed a prevalence of 74% of CAC on the right side. This result is reasonable according to aforementioned authors, which cited that the level of bifurcation of the right carotid artery is most often located between C3 and C4, whereas the left carotid artery shows a bifurcation mostly observed between C4 and C5.

In the present investigation, the differential diagnosis included radiopaque structures resembling atheromatous plaques on digital panoramic radiographs. CAC were considered in the presence of one or more non-continuous adjacent radiopaque nodular images or vertical lines at the intervertebral space between C3 and C4, and located posteriorly and lower than the mandibular angle (20,21). In addition, the present study included as potential differential diagnosis other conditions, including sialoliths, tonsilloliths, as well as stylohyoid ligament and triticeous cartilage calcifications, and the hyoid bone. Sialoliths are usually located in the submandibular gland or in its duct and have a radiographic pattern as a single or multiple calcifications. On panoramic radiographs, they can be seen superimposed on or below the body or ramus of the mandible, which is a most anterior location than observed in CAC. Tonsilloliths are small calcifications developed into tonsillar crypts and are described as radiopaque images superimposed on the mandibular branch on panoramic radiographs. The styloid process has been radiographically described as a cylindrical and radiopaque image projecting forward and down between the mandible ramus and the mastoid process. The hyoid bone is presented as a bilateral and horizontal well-defined radiopaque image located below the mandible. Calcified triticeous cartilage presents as an ovoid radiopaque image, about 2 to 4 mm wide and 7 to 9 mm long, normally found within the air space adjacent to the pharynx (21).

Regarding the study design adopted in this research, it was not possible to use the Doppler ultrasonography in order to confirm the diagnosis of CAC. However, in the field of Oral and Maxillofacial Radiology, we believe that is important to add literature concerning epidemiological data from different geographic locations.

In summary, the present study showed a low prevalence

of suggestive images of CACs on digital panoramic radiographs from an adult South American subpopulation. It was observed a higher prevalence in women and older patients, and the right side was the main location for CAC.

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# Conflict of interest

The authors have declared that no conflict of interest exist.