Usefulness of measuring bone density of mandibular condyle in patients at risk of osteoporosis: A cone beam computed tomography study

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ABSTRACT

Objective: Using cone beam computed tomography (CBCT) images, the aim of the study was to evaluate the usefulness of measuring bone density of mandibular condyle (BDMC) in patients at risk of osteoporosis. **Materials and Methods:** Two hundred and four mandibular condyles (46 condyles from males and 158 condyles from females) were examined by CBCT. Using inVivo software (Anatomage, San Jose, Calif), BDMC and mental index (MI) were measured by two observers and correlated using Pearson's correlation coefficients. Patients were divided into high risk and low risk of osteoporosis using 3.1 mm of MI as a cutoff value, and the mean value of BDMC was compared and correlated using independent samples t-test and regression analysis. Receiver-operating characteristic (ROC) curve analyses were also used to examine the predictive power of BDMC. **Results:** The mean value of BDMC was moderately correlated with MI (r = 0.38), and in female patients at low risk of osteoporosis, the mean value of BDMC was significantly higher than in patients at high risk of osteoporosis. In multivariate linear regression analysis, being at risk of osteoporosis is significantly affected by BDMC, age, and gender. ROC analysis showed that the mean value of BDMC had a moderate predictive power for predicting patients at risk of osteoporosis (area under the curve = 0.621 for females and 0.649 for males). **Conclusions:** Measuring BDMC is considered useful in predicting patients at risk of osteoporosis.

Key words: Bone density, cone beam computed tomography, mandibular condyle, osteoporosis

INTRODUCTION

Osteoporosis defined as "a disease characterized by low bone mass and microarchitectural deterioration of bone tissue, leading to enhanced bone fragility and a consequent increase in fracture risk." [1] Complications

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of osteoporosis such as osteoporotic fracture are associated with high morbidity and medical costs. The risk for such complications can be reduced if the

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How to cite this article: Alkhader M, Aldawoodyeh A, Abdo N. Usefulness of measuring bone density of mandibular condyle in patients at risk of osteoporosis: A cone beam computed tomography study. Eur J Dent 2018;12:363-8.

DOI: 10.4103/ejd.ejd_272_17

disease is diagnosed and treated earlier, and this is where the importance of finding new screening tools for osteoporosis comes from.

The current gold standard for diagnosing osteoporosis is dual-energy X-ray absorptiometry (DEXA) or DEXA, which is a test that is used to measure bone mineral density (BMD). [2] Although DEXA is used extensively around the world because of its reliability and low radiation dose, it is not always available and is not considered as cost-effective tool for screening in low-risk patients, such as younger individuals. [2] Quantitative computed tomography (CT) can be used instead of DEXA to diagnose osteoporosis, [3] but it is also expensive and associated with higher radiation dose, thus would not be appropriate as screening tool too.

In an effort to increase dentists role in screening potential osteoporosis patients, and to increase the likelihood that these patient are discovered earlier in life, many indices related to panoramic radiograph have been proposed. These indices can be used to analyze mandibular bone by two ways; either by objective analysis, in which linear measurements are calculated, or by subjective analysis, in which the appearance of the bone is described according to a classification.

The commonly used method for subjective analysis is Klemetti index (KI), which classifies the inferior mandibular cortex appearance to three categories: C1 - the endosteal margin of the cortex is even and sharp on both sides; C2 - the endosteal margin shows semilunar defects and/or seems to form endosteal cortical residues on one or both sides; and C3 - the cortical layer forms heavy endosteal cortical residues and is clearly porous.^[5] Although many studies were conducted and suggested that this index can be used as a screening tool, Calciolari et al.[4] reported that the wide range of outcomes is a sign of high heterogeneity between these studies, this may be attributed to the subjectivity of this index. In fact, in a more recent study, this index was reported to be of low sensitivity and specificity, and concluded that it is of limited use in clinical practice.[6]

On the other hand, other morphometric analysis dependent indices have been described; the most famous of them are panoramic mandibular index (PMI) and mental index (MI). The PMI index is defined as the ratio of the thickness of the inferior mandibular cortex in the mental region over the distance between the lower border of the mandible and either the

inferior or the superior border of the mental foramen.^[4] Some studies showed significant correlation between PMI index and osteoporosis, but others did not.^[7-10] Furthermore, Horner and Devlin showed that the PMI has no significant advantage over the MI.^[11]

The MI (also known as mandibular cortical width or mandibular cortical thickness) is measured at the mental foramen region, along a line passing through the middle of the mental foramen and perpendicular to the tangent to the lower border of the mandible. [4] MI is considered the most established among mandibular cortical indices. [12] Other indices, such as the KI, the gonial, and antegonial indices, were less frequently used and are less established.

Another more recent imaging modality, cone beam CT (CBCT), is being widely used in dentistry. Koh and Kim^[13] were the first to examine the potential use of CT indices on CBCT images for assessment of BMD. Mostafa *et al.*^[14] found a significant correlation between CT mental index (CTMI) scores and osteoporosis.

CBCT is considered the standard of care for imaging osseous structures of the temporomandibular joint (TMJ), mainly because it has better imaging quality and less radiation dose compared to helical CT.^[15] More recent studies suggest using an alternative approach that utilizes two CBCT acquisitions of the right and left TMJs, as this results in >10-fold reduction in the radiation do when compared to full field CBCT.^[16]

As mentioned above, for osseous assessment of TMJ, CBCT images from the small field of view are recommended. However, in such case, the mandible will not be present, and it will not be possible to predict osteoporosis using mandibular cortical indices. Therefore, to enhance dentists' role in screening patients for osteoporosis, and to maximize patients' benefits from getting such images, we find it necessary to know if these images can be used to screen patients for osteoporosis or not.

Kosugi *et al.*^[17] were the first to study the effect of osteoporosis on bone density of mandibular condyle (BDMC) in rats, and they found that the BDMC is less in osteoporotic rats compared to normal rats when measured using morphometric analysis indices for micro-CT scans. To the best of our knowledge, no one has ever studied the relationship between BDMC and osteoporosis in humans; therefore this study was conducted.

MATERIALS AND METHODS

Patients

In our retrospective study, CBCT images for all patients aged 50 years and more were retrieved and evaluated from our CBCT database. All patients underwent CBCT examination for dental implant treatment and other dental purposes at our dental radiology clinic between January 2011 and January 2017. After excluding cases with artifacts or pathology affecting measurements of BDMC or MI, the number of cases included was 204 mandibular condyles (46 condyles from males and 158 condyles from females). Patients aged between 50 and 78 years (mean age 59.75 ± 7.01). The present study is part of a protocol which was approved by the Local Ethical Committee with the protocol number 447/2016.

Cone beam computed tomography examination

As a CBCT apparatus, KODAK 9500 Cone Beam 3D System (Carestream, Rochester, NY, USA) with flat panel detector was used. The imaging area of CBCT is a cylinder with a height of 15–20.6 cm and a diameter of 9–18 cm providing isotropic cubic voxels with sides approximating 0.2–0.3 mm. Only cases examined with 0.2 mm of voxel size were included in the study. The exposure parameters were: 90 kV as a tube voltage, 10 mA as a tube current, and 10.8 s as an exposure time.

Examinations were performed by 360° rotation in the occlusal position with the patient standing and closing their teeth.

Images

Two calibrated observers used in Vivo software (Anatomage, San Jose, Calif) for measuring BDMC and MI.

For measuring BDMC, a coronal section in the middle of each mandibular condyle was used in arch section module. To obtain the coronal section in the middle of mandibular condyle, the middle of mandibular condyle was localized on reconstructed panoramic image using the associated vertical colored bar. A circle touching inner cortical surfaces was drawn using circle notation tool, and then using the distance measurement tool, 6 mm × 6 mm square area of interest was drawn inside the circle starting from top to the bottom. Finally, Hounsfield Units [HUs] measurement tool was used to generate density value for the corresponding condyle. HU measurement tool showed 3 density values at each condyle (minimum, mean, and maximum).

However, we only considered the mean density value for analysis [Figure 1a].

For measuring MI (inferior cortical width of the mandible at mental foramen), arch section module was utilized, and a cross-sectional image in the middle of each mental foramen was used. To obtain a cross-sectional image at the middle of mental foramen, the middle of mental foramen was localized on corresponding panoramic image using associated vertical colored bar, then using the distance measurement tool, the inferior cortical width of the mandible was measured at each mental foramen following the example of Koh and Kim^[13] [Figure 1b].

All images were evaluated on a high definition LCD display, and window settings were fixed for all cases.

Statistical analysis

Data were analyzed using SAS 9.2 (SAS Institute, Cary, NC). Data were described using means and percentages. Pearson's correlation coefficient was used to determine inter-rater correlation between first and second observer measurements of BDMC and MI, and the correlation between BDMC and MI. According to a multi-center study with large sample size, [18] patients with the thinnest MI (\leq 3 mm) have the highest likelihood of osteoporosis. Therefore, using 3.1 mm of MI as a cutoff value, patients were divided into low and high risk of osteoporosis. Independent *t*-test was used to compare means of continuous variables between those with low and high risk of osteoporosis. The association between BDMC and

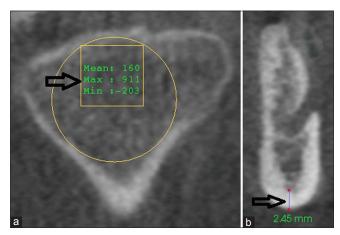


Figure 1: A 68-year-old female patient who visited our dental clinic for implant placement (a) Cross-sectional cone beam computed tomography image at the middle of right mandibular condyle showing bone density values of the square area of interest (arrow). (b) Cross-sectional cone beam computed tomography image at the middle of right mental foramen showing inferior cortical width (mental index) (arrow)

MI after adjusting for age and gender were tested in linear regression analysis when MI is continuous and binary logistic regression when MI is dichotomous. Receiver operating characteristics (ROCs) analysis was used to test the ability of BDMC to predict the risk of osteoporosis. P < 0.05 was considered statistically significant.

RESULTS

This study included a total of 204 mandibular condyles (46 condyles from males and 158 condyles from females). The intra-rater correlations for BDMC (r = 0.99) and MI (r = 0.98) were very strong. Therefore, the two readings for each measurement were averaged to be used for further analysis. The correlation between average BDMC and average MI was moderate (r = 0.32).

Males had significantly higher mean of MI [Table 1]. Although males had higher BDMC mean compared to females, it was not statistically significant [Table 1]. Patients were divided into high risk and low risk of osteoporosis using 3.1 mm of MI as a cutoff value, and females had significantly higher risk than males for osteoporosis [Table 2]. For females only, the mean age was significantly higher for females at high risk of osteoporosis compared to females at low risk of osteoporosis (61.99 [7.6] year vs. 57.38 [4.6] year; P < 0.0001) [Table 3]. Furthermore, females at high risk of osteoporosis have lower BDMC compared to females at low risk of osteoporosis (206 [102.14] vs. 236.21 [83.72] year; P < 0.046) [Table 3]. After adjusting for the effects of age and gender, the mean value of BDMC (F = 10.87, P < 0.0001) remained significantly higher in patients at

Table 1: Gender differences for mental index and density							
	Males (n=46)	Females (n=158)	P				
	Mean (range)±SD	Mean (range)±SD					
Mental index	3.51 (1.65-5.30) 0.91	3.12 (1.29-7.25) 1.01	0.0197*				
Density	245.16 (83.50-512.0)	221.16 (37.5-470.5)	0.1439				

*P values were determined by t-test. SD: Standard deviation

Table 2: Relationship between osteoporosis risk with gender							
Osteoporosis risk	High risk (%)	Low risk (%)	P				
Gender							
Male	13 (28.3)	33 (71.7)	0.0059*				
Female	81 (51.3)	77 (48.7)					
*Females have higher risk than males for osteoporosis; <i>P</i> values were determined by Chi-square test							

low risk of osteoporosis. ROC analysis showed that the mean value of BDMC (area under the curve = 0.621 for females and 0.649 for males) had a moderate predictive power to predict osteoporosis.

DISCUSSION

In this study, we evaluated the usefulness of BDMC in predicting osteoporosis in patients aged 50 years old and more. According to Jonasson *et al.*,^[19] mandibular cortical thickness decreased significantly after the age of 50, and more fractures were shown. Therefore, patients above 50 years old are at risk of osteoporosis and possibility to have low MI is high as shown in similar studies.^[6,20]

As we mentioned in the introduction, a relationship between the density of the condyle in mico-CT images and osteoporosis was established in *in vitro* studies. ^[17] On the other hand, Güngör *et al.* ^[21] showed that a positive correlation is also present between femoral head BMD and left condyle density value in CBCT images. According to our results, the difference in BDMC between high-risk and low-risk osteoporosis group is statistically significant in females. This suggests that the osteoporosis can affect the BDMC in human too.

It is worth mentioning that CT and CBCT images of other parts of the mandible have already been found to be useful for the same purpose. For example, Barngkgei *et al.*^[22] concluded in their study that the radiographic density of the body of the mandible in CBCT images can be used to accurately predict lumber vertebrate and femoral neck osteoporosis. Others, such as Mostafa *et al.*^[14] found that CT cortical index scores, CTMI, and computed tomography mandibular index (CTI) can be used to differentiate between the control and the osteoporotic groups in postmenopausal women.

In the previous studies, which measured MI on cross-sectional CBCT images,^[14,21] MI was significantly higher in patients at low risk of osteoporosis. Our results are also in agreement with these studies.

In accordance with our results, several studies showed that MI has higher values in males and inversely related to age. [20,23-25] Nevertheless, in our study the mean age was significantly higher for females at high risk of osteoporosis compared to females at low risk of osteoporosis; this could be due to the possibility of having more females at menopause or postmenopause at older age.

Table 3: Descriptive statistics for gender, age, bone density of mandibular condyle, and mental index according to the risk of osteoporosis

Gender	Parameter (n)	High risk of osteop	High risk of osteoporosis		Low risk of osteoporosis	
		Mean±SD	n	Mean±SD	n	
Females	MI	2.37 (0.73)	79	3.88 (0.61)	79	<0.0001
	Density	206.12 (102.14)	79	236.21 (83.72)	79	0.0446
	Age	61.99 (7.57)	79	57.38 (4.85)	79	<0.0001
Males	MI	2.42 (0.56)	14	3.99 (0.53)	32	<0.0001
	Density	219.86 (82.22)	14	256.23 (117.64)	32	0.3007
	Age	61.36 (9.60)	14	59.34 (7.16)	32	0.4340

MI: Mental index, SD: Standard deviation

With regard to the dentition status, both of Xu et al.[26] and Aggarwal et al.[27] found significant difference between right and left condyle bone density in patients with unilateral loss of teeth, with the density being lower in the edentulous half. Choi et al. [28] reported on the same issue, but his results specified the presence of molars to be the important factor affecting condyle density. Despite being significant factor, the dentition status remained outside the scope of our study. In the same study of Choi et al., [28] they found that age was affecting factor when it comes to condyle density in both males and females as measured from micro-CT scan. In our study, the age was a significant factor for females only. This may be explained by the fact that the age range in our study included only old patients, while wider age range including younger individuals was selected for the previously mentioned study.

In a study done by Yamada *et al.*,^[29] a correlation between bone density of lumbar spine (L1–L3) and BDMC was present, and both decrease with age in a similar fashion to each other. Their data were obtained using quantitative CT; similarly, our data obtained using CBCT, which is being used by dentists for different dental applications.

Both of the studies support the idea that the BDMC can be used to predict patients at risk of osteoporosis. Although dual X-ray absorptiometry (DXA) as the gold standard for diagnosis of osteoporosis was not utilized in both of the studies, DXA has several limitations especially in obese patients and/or patients with degenerative changes in the spine.^[3] Moreover, DXA is not available in routine dental practice as mentioned previously.

With regards to the usefulness of BDMC in predicting patients at high risk of osteoporosis, the area under the ROC curve was 0.621 for females and 0.649 for males. This indicates moderate predictive power of BDMC. Similar results were obtained by using

MI measurements for predicting osteoporosis. [12,20] Therefore, BDMC might be recommended to be used instead of MI when mandibular bone is not shown in patient images for any reason.

In this study, all patients with MI ≤3 mm were considered at high risk of osteoporosis as shown in previous studies, ^[6,18,30] however, the cutoff value of MI might be related to race or ethnic group. ^[20] We could not test the accuracy of the previous cutoff value; this is considered a limitation of this study. Moreover, the dimensions of area of interest used in this study might not be applicable on images of patients from different ethnic group.

CONCLUSIONS

Taking into account the limitations of the present study, our results indicate that measuring BDMC is considered useful in predicting patients at risk of osteoporosis.

Financial support and sponsorship Nil.

Conflicts of interest

There are no conflicts of interest.

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