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Comparison of Trabecular Bone Mineral Density Measurement Using Hounsfield Unit and Trabecular Microstructure in Orthodontic Patients Using Cone-Beam Computed Tomography

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Abstract: The aim of this study was to measure the bone mineral density of specific regions of maxilla, mandible, and first cervical vertebra using the Hounsfield unit and trabecular microstructure pattern analysis and to compare the two methods. In this study, cone-beam computed tomography (CBCT) images were obtained from 58 patients. Trabecular thickness, trabecular number, trabecular separation, and bone volume fraction were measured in 484 regions for trabecular microstructure parameters and Hounsfield unit was measured for the grayscale value. There was no difference in bone mineral density between the right and left side in every site and between males and females. Trabecular thickness and trabecular number were high in the order of anterior base of the maxilla, mandibular body, first cervical vertebra, and mandibular condyle. Bone volume fraction and Hounsfield unit were high in the order of anterior base of the maxilla, mandibular body, mandibular condyle, and first cervical vertebra ($p < 0.05$). Trabecular thickness, trabecular number, and bone volume fraction was positively correlated to the Hounsfield unit, and trabecular separation was negatively correlated to the Hounsfield unit ($p < 0.005$). This study suggests that it is possible to compare the bone mineral density of trabecular bone in various sites using the Hounsfield unit and trabecular microstructure pattern analysis.

Keywords: bone mineral density; trabecular microstructure; Hounsfield unit; cone-beam computed tomography

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1. Introduction

Changes in the density of trabecular bones are often associated with local or systemic diseases. Changes in the trabecular bone density also affect the corrective movement of teeth associated with the orthodontic treatment [1,2]. Therefore, when evaluating the trabecular bone density, it is important to make a qualitative evaluation as well as a quantitative evaluation of the bone.

Bone mineral density (BMD) can be measured based on radiodensity in computed tomography (CT) or assessed by analyzing the microstructure of the trabecular bone. Hounsfield unit (HU) is mainly used to measure the radiographic density value [3]. Misch's bone density classification was classified from D1 to D5 based on HU values [4]. The mandible showed the BMD from D1 to D3 from the anterior to the posterior, and the maxilla showed the BMD from D2 to D4 from the anterior to the posterior region. Re-

cently, the introduction of high-resolution micro-CT shows the microstructure of the trabecular bone as an objective indicator and can be used for trabecular bone density analysis [5].

Cone-beam computed tomography (CBCT) became popular in the dental field for the three-dimensional evaluation of structures [6]. CBCT has the advantage of obtaining high resolution images with less radiation dose and cost than other conventional CT. Ibrahim et al. reported that the micro structural evaluation of trabecular bone using CBCT correlated with that of micro-CT [5].

A representative case of measuring trabecular bone density is for the diagnosis of osteoporosis, which measures the density of the lumbar spine or femur [7]. Guerra et al. reported that the trabecular bone density measurement using the mandible or cervical vertebra in dental CT showed a high correlation with the conventional trabecular bone density measurement through a systemic review [8]. The analysis of trabecular bone density using CBCT will be a useful reference for the diagnosis and treatment of various diseases.

In this study, the anterior base of the maxilla, mandibular body, mandibular condyle, and the first cervical vertebra were selected to measure the trabecular bone density. The aim of this study was to evaluate the bone mineral density measurement methods in various regions of the CBCT image using the Hounsfield unit and trabecular microstructure parameters in patients undergoing an orthodontic treatment, and to compare and find out the correlation between the two methods.

2. Materials and Methods

This study was conducted on 484 regions measured in 58 patients who visited the department of orthodontics, Korea University Guro Hospital between 2014 and 2019, and obtained diagnostic records including CBCT images. Inclusion criteria were as follows.

- (1) Patients without defects in the trabecular bone density measurement area.
- (2) Patients with no symptoms of temporomandibular disorder (TMD).
- (3) Patients without a history of bone diseases such as osteoporosis.
- (4) CBCT images having a quality suitable for research.

Table 1 showed the number of samples, mean age, and standard deviation. This study was approved by the Institutional Review Board (IRB) of Korea University Guro Hospital (no. 2019GR0189).

Table 1. Characteristics of the samples.

	Male	Female	Total
Number of regions measured	272	192	464
Number of patients	34	24	58
Mean age (SD)	23.8 (6.5)	19.5 (7.0)	22.0 (7.0)

SD: Standard deviation.

The CBCT equipment used in this study was PaX-Reve3D (Vatech Korea, Seoul, Korea), the tube voltage was 80 kV, the tube current was 3.3 mA, and the voxel size was 250 μm . Before the trabecular bone density measurement, the head position was reoriented with the Frankfort Horizontal plane (right and left orbitale, right portion).

The trabecular bone density was measured using the Hounsfield unit and trabecular microstructure parameters in the anterior basal bone beside the incisive foramen of the maxilla, the mandibular body; medial side of mental foramen, the mandibular condyle; center of medial and lateral pole; and the lateral mass of first cervical vertebra, at the right and left region, respectively (Figure 1).

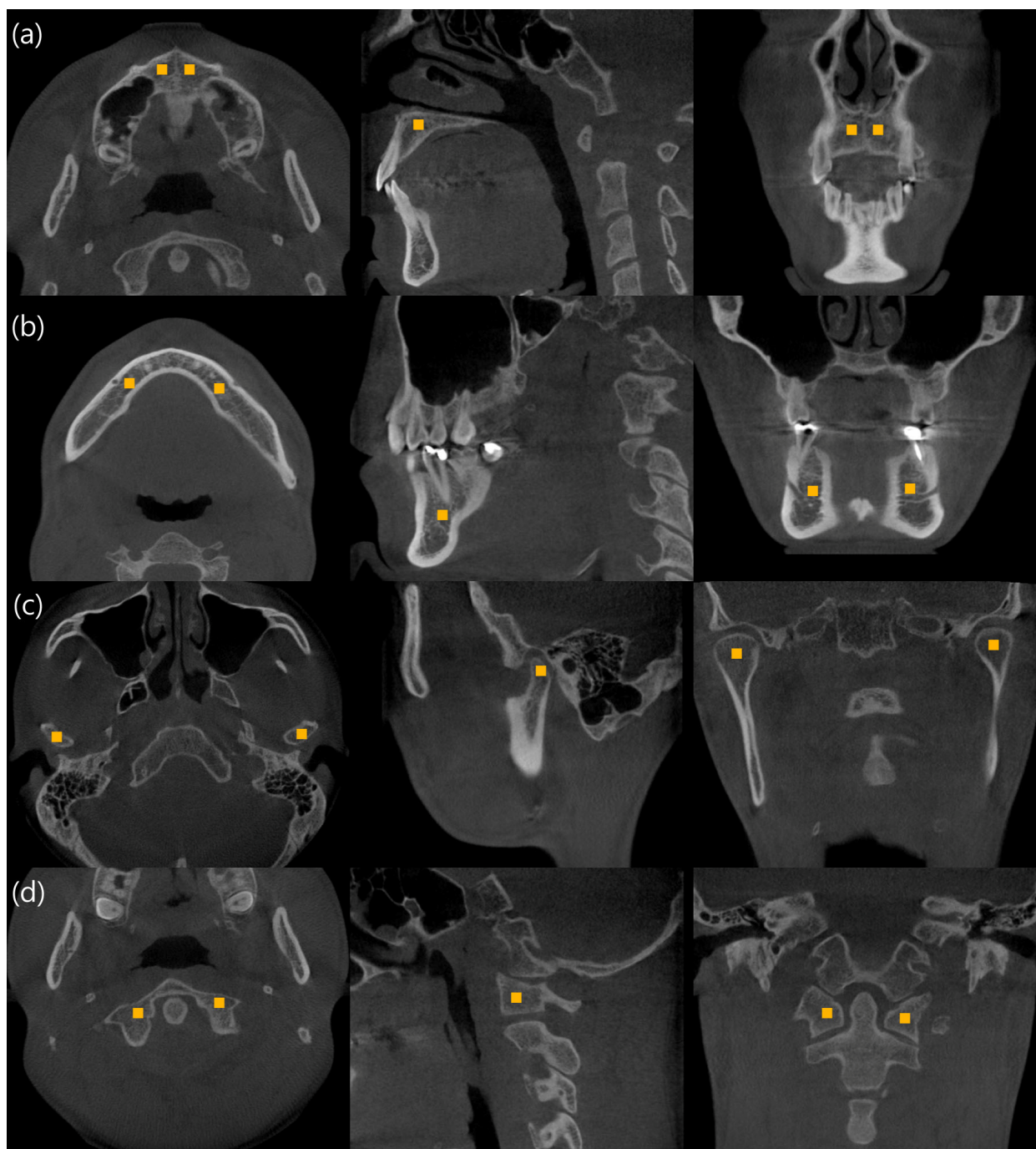


Figure 1. Regions of trabecular bone density measurement (yellow square points): (a) Anterior basal bone beside the incisive foramen in maxilla (Mx); (b) body of mandible, medial side of mental foramen (Mn); (c) condyles of mandible, center of medial, and lateral pole (Co); (d) lateral mass of the first cervical vertebra (Cv).

The Invivo5 (Anatomage, San Jose, CA, USA) software was used for the Hounsfield unit measurement, and the CTan (Bruker micro-CT, Kontich, Belgium) software was used for the trabecular microstructure analysis (Figure 2).

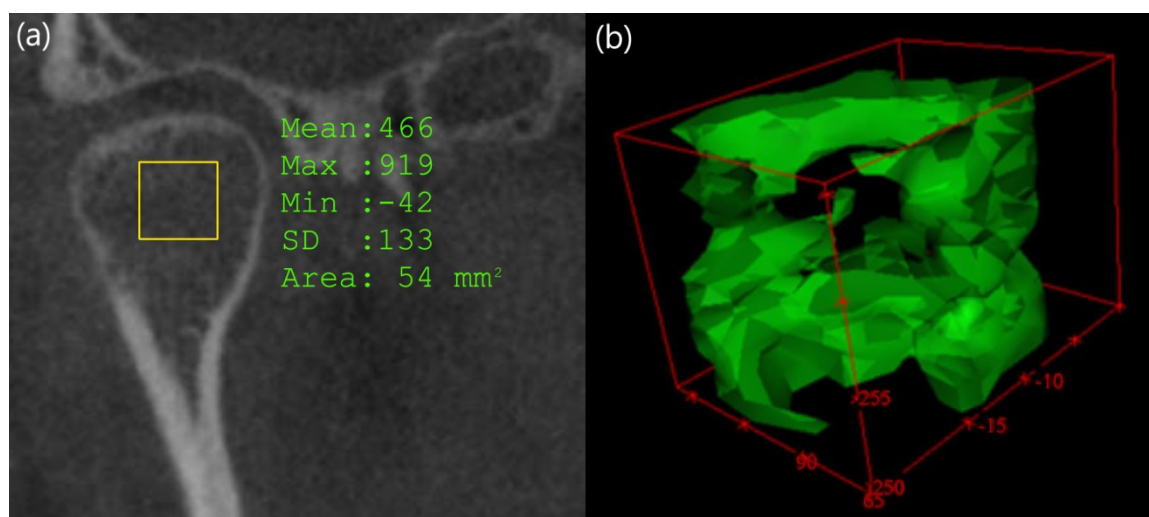


Figure 2. Two methods of trabecular bone density measurement: (a) Hounsfield unit, (b) trabecular bone microstructure parameters from reconstructed three-dimensional (3D) images.

Each region was evaluated in three-dimensions (3D) with a size of $4 \times 4 \times 4 \text{ mm}^3$ cube. The trabecular microstructure parameters used in this study were described in Table 2.

Table 2. Description of trabecular microstructure parameters.

	Description	Unit
Tb.th	Trabecular thickness: Mean thickness of trabeculae	mm
Tb.n	Trabecular number: Measure of the average number of trabeculae per unit length	1/mm
Tb.sp	Trabecular separation: Mean distance between trabeculae	mm
BV/TV	Bone volume fraction: Ratio of the segmented bone volume to the total volume of the region of interest	%

The Wilcoxon signed-rank test was performed to determine whether there was a difference in the measured values of left and right of each region and a difference in the measured trabecular bone density for each region. The Mann-Whitney U test was used to identify differences in the measured trabecular bone density between genders. Spearman correlation was used to check the correlation coefficient between two measurement methods of the measured trabecular bone density. The SPSS software (ver 25.0; IBM Corp., Armonk, NY, USA) was used for all the analysis.

3. Results

3.1. Bone Mineral Density Measurement in the Right and Left Side and in Males and Females

The bone mineral density measurement of the left and right measurements for each region and the difference between the left and right side were obtained. The Wilcoxon signed-rank test showed no significant difference in left and right measurements of all the areas (Table 3).

Table 3. Bone mineral density measurement in the right and left side and the differences between males and females.

Region		Right		Left		Difference		<i>p</i> -Value
		Median	IQR	Median	IQR	Median	IQR	
Mx	Tb.th	3.79	1.11	3.71	0.79	0.10	0.94	0.568
	Tb.n	0.14	0.04	0.14	0.03	0.00	0.03	0.298
	Tb.sp	3.57	0.82	3.52	0.85	−0.02	0.81	0.640
	BV/TV	54.52	10.23	53.10	11.81	0.49	9.34	0.502
	HU	1441.41	332.27	1505.36	268.68	−74.48	331.90	0.268
Mn	Tb.th	3.45	0.67	3.42	0.68	0.04	0.69	0.497
	Tb.n	0.14	0.06	0.14	0.07	0.02	0.10	0.572
	Tb.sp	3.62	1.17	3.71	0.91	−0.08	0.98	0.354
	BV/TV	49.86	13.25	48.03	11.85	1.79	11.21	0.374
	HU	1401.38	304.89	1416.11	283.95	−13.08	290.25	0.248
Co	Tb.th	3.17	0.51	3.24	0.43	−0.06	0.58	0.349
	Tb.n	0.14	0.04	0.13	0.03	0.01	0.03	0.130
	Tb.sp	4.46	3.96	4.55	3.75	−0.31	4.73	0.811
	BV/TV	48.07	21.83	44.58	23.50	−0.75	33.20	0.868
	HU	653.02	315.91	715.49	378.01	−49.77	383.40	0.402
Cv	Tb.th	3.22	0.60	3.35	0.47	−0.14	0.65	0.201
	Tb.n	0.13	0.09	0.14	0.05	−0.01	0.07	0.366
	Tb.sp	4.50	3.79	3.96	2.48	0.79	2.89	0.659
	BV/TV	43.86	28.81	49.26	20.35	−6.51	25.54	0.383
	HU	511.09	236.07	575.71	279.92	−45.18	235.75	0.186

IQR: Interquartile range; Tb.th: Trabecular thickness; Tb.n: Trabecular number; Tb.sp: Trabecular separation; BV/TV: Bone volume fraction; HU: Hounsfield unit; Mx: Anterior base of the maxilla; Mn: Mandibular body; Co: Mandibular condyle; Cv: First cervical vertebra.

In addition, the bone mineral density measurement and differences between males and females were measured. The Mann-Whitney U test showed that there was no significant difference between genders in the measured values of all the areas (Table 4).

Table 4. Bone mineral density measurement in males and females and the differences.

Region		Male		Female		Difference		<i>p</i> -Value
		Median	IQR	Median	IQR	Median	IQR	
Mx	Tb.th	3.92	0.90	3.71	0.59	−0.21	0.31	0.647
	Tb.n	0.14	0.02	0.15	0.04	0.01	0.02	0.776
	Tb.sp	3.51	0.64	3.50	0.74	−0.01	0.10	0.825
	BV/TV	54.36	10.58	54.20	9.80	−0.16	0.78	0.991
	HU	1482.81	269.65	1456.87	213.89	−25.94	55.76	0.752
Mn	Tb.th	3.42	0.46	3.50	0.69	0.08	0.23	0.412
	Tb.n	0.14	0.04	0.14	0.06	0.00	0.02	0.328
	Tb.sp	3.68	0.94	3.75	0.87	0.07	0.07	0.290
	BV/TV	51.41	9.97	47.01	12.97	−4.40	3.00	0.602
	HU	1441.03	234.83	1346.83	304.60	−94.20	69.77	0.097
Co	Tb.th	3.19	0.30	3.23	0.39	0.04	0.09	0.468
	Tb.n	0.14	0.03	0.14	0.03	0.00	0.01	0.412
	Tb.sp	4.38	2.26	4.31	3.53	−0.07	1.27	0.962
	BV/TV	45.61	13.95	49.63	20.43	4.02	6.48	0.122
	HU	656.03	253.57	739.79	343.88	83.76	90.31	0.067
Cv	Tb.th	3.23	0.48	3.37	0.43	0.14	0.05	0.190
	Tb.n	0.13	0.07	0.14	0.05	0.01	0.02	0.728
	Tb.sp	4.48	3.23	4.10	1.86	−0.38	1.37	0.776
	BV/TV	45.92	24.52	52.51	19.55	6.59	4.97	0.507
	HU	522.27	232.53	549.13	255.92	26.86	23.39	0.764

IQR: Interquartile range; Tb.th: Trabecular thickness; Tb.n: Trabecular number; Tb.sp: Trabecular separation; BV/TV: Bone volume fraction; HU: Hounsfield unit; Mx: Anterior base of the maxilla; Mn: Mandibular body; C: Mandibular condyle; Cv: First cervical vertebra.

3.2. Comparison between the Bone Mineral Density Parameters among Regions

3.2.1. Trabecular Thickness (Tb.th)

Results for the comparison among regions of the Tb.th values were shown in Table 5. The value at the anterior base of the maxilla was the highest among all regions and was statistically significant. The Tb.th value in the mandibular body was larger than in the first cervical vertebra, and was significantly higher than in the mandibular condyle. There was no statistically significant difference between the first cervical vertebra and the mandibular condyle.

Table 5. Comparison of the bone mineral density parameters among regions.

	Mx	Mn	Co	Cv	<i>p</i> -Value
	Median(IQR)	Median(IQR)	Median(IQR)	Median(IQR)	
Tb.th	3.84(0.85) ^a	3.49(0.53) ^b	3.21(0.36) ^c	3.25(0.50) ^{bc}	0.025
Tb.n	0.14(0.034) ^a	0.14(0.05) ^{ab}	0.13(0.03) ^b	0.14(0.05) ^{ab}	0.032
Tb.sp	3.51(0.73) ^c	3.74(1.05) ^{bc}	4.51(3.21) ^a	4.27(2.79) ^{ab}	0.015
BV/TV	53.42(9.78) ^a	50.29(12.35) ^b	47.50(16.72) ^{bc}	46.58(24.06) ^c	0.008
HU	1458.69(227.17) ^a	1414.19(265.78) ^b	697.14(264.81) ^c	540.51(226.40) ^d	<0.001

IQR: Interquartile range; Tb.th: Trabecular thickness; Tb.n: Trabecular number; Tb.sp: Trabecular separation; BV/TV: Bone volume fraction; HU: Hounsfield unit; Mx: Anterior base of the maxilla; Mn: Mandibular body; Co: Mandibular condyle; Cv: First cervical vertebra. ^{a,b,c,d} comparison of the bone mineral density parameters among regions.

3.2.2. Trabecular Number (Tb.n)

Results for the comparison among regions of the Tb.n values were shown in Table 5. The value at the anterior base of the maxilla was the highest among all regions. The value

of the maxilla did not show a statistically significant difference from the mandible and the first cervical vertebra, but showed a significant difference from the mandibular condyle. There was no statistically significant difference between the mandibular body, the first cervical vertebra, and the mandibular condyle.

3.2.3. Trabecular Separation (Tb.sp)

Results for the comparison among regions of the Tb.sp values were shown in Table 5. The value in the mandibular condyle was the highest among all regions. The value of the mandibular condyle did not show a statistically significant difference from the values of the first cervical vertebra, but showed significant differences from those of the mandibular body and the anterior base of the maxilla. Tb.sp in the first cervical vertebra was significantly greater than that in the anterior base of the maxilla. There was no statistically significant difference in Tb.sp between the mandible and the anterior base of the maxilla.

3.2.4. Bone Volume Fraction (BV/TV)

Results for the comparison among regions of the BV/TV values were shown in Table 5. The value at the anterior base of the maxilla was significantly higher than all the other regions. BV/TV in the mandibular body was significantly higher than in the first cervical vertebra and did not show a significant difference from the mandibular condyle. The measured values in the first cervical vertebra and the mandibular condyle showed no significant difference.

3.2.5. Hounsfield Unit (HU)

Results for the comparison among regions of the HU values were shown in Table 5. The value was highest in the order of the anterior base of the maxilla, the mandibular body, the mandibular condyle, and the first cervical vertebra.

3.3. Comparison Between the Trabecular Bone Density Measurement Method Using the Hounsfield Unit and Trabecular Microstructure Parameters

Spearman correlation was used to find the correlation between two measurement methods (Figure 3). Correlation coefficients were obtained from trabecular microstructure parameters and Hounsfield unit values in the first cervical vertebra, which are known to have a similar accuracy to the general bone density. The Tb.th, Tb.n, and BV/TV of the trabecular pattern showed a significant positive correlation with the HU value, and the Tb.sp showed a significant negative correlation with the HU value.

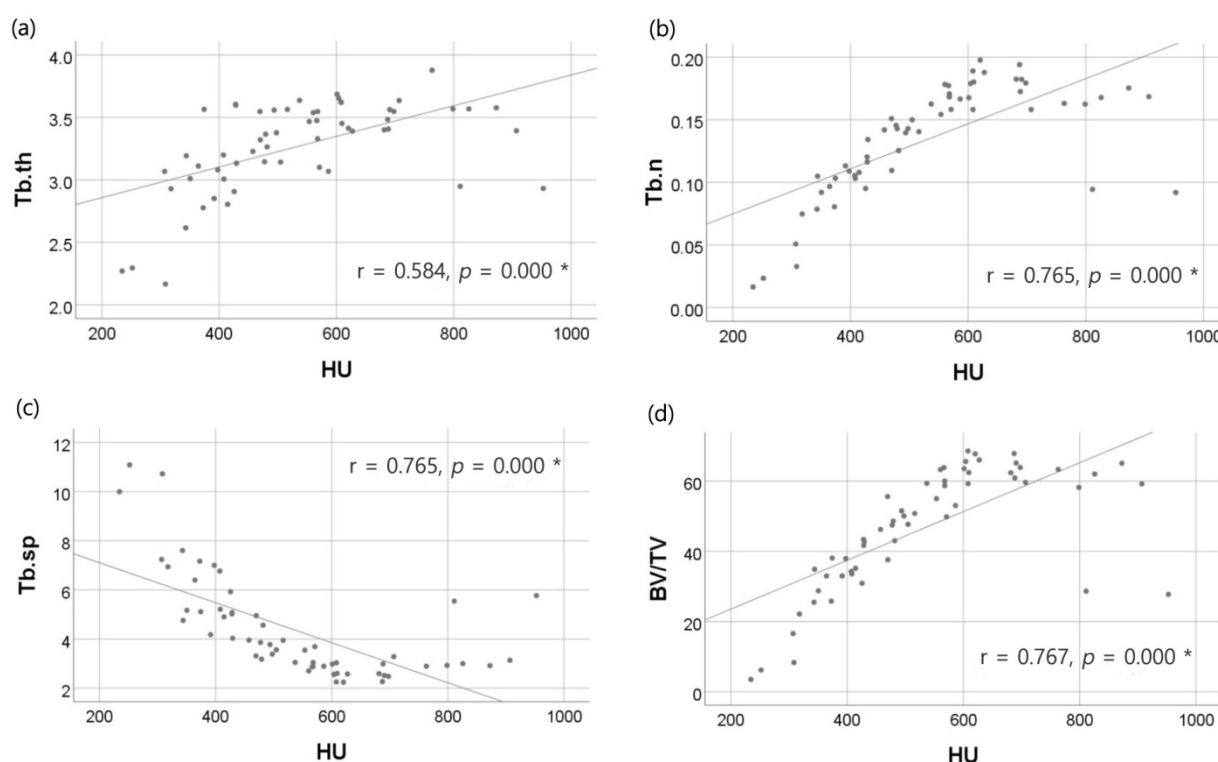


Figure 3. Correlations between the Hounsfield unit measurement and trabecular microstructure parameters: (a) Positive correlation between Hounsfield unit (HU) and trabecular thickness (Tb.th); (b) positive correlation between HU and trabecular number (Tb.n); (c) negative correlation between HU and trabecular separation (Tb.sp); (d) positive correlation between HU and bone volume fraction (BV/TV). * $p < 0.001$

4. Discussion

Since bone-related diseases such as osteoporosis increase the probability of the bone fracture, several methods have been proposed to accurately measure the trabecular bone density [9]. With regard to an orthodontic treatment, the trabecular bone density also affects the speed of the tooth movement. As the trabecular bone density increases, the rate of the tooth movement tends to decrease, which is associated with a greater resistance to the tooth movement in the mandibular molar [10]. As a result, the mandibular molar serves as a better anchor.

The cortical bone and the trabecular bone are divided based on the degree of porosity. The cortical bone has a porosity of 0–30%, whereas the trabecular bone has a porosity of 30–90% [11]. For this reason, in the cortical bone, the bone density can be represented to some extent by only the opacity of radiographs, but since the trabecular bone is a mixture of plate-like and columnar structures, it is difficult to accurately measure the density of the trabecular bone without considering three-dimensional structural characteristics [12,13].

In the past, two-dimensional methods such as histological analysis and serial sectioning methods have been used to measure the three-dimensional structure of bones. This not only shows the limitations of the two-dimensional structure, but also has the disadvantage of being more invasive since tissue specimens should be produced. However, the analysis using CT is not only non-destructive compared to the histological analysis, but also has the advantage of realizing a more accurate three-dimensional image at high resolution [14,15].

The Hounsfield unit defines the radiation density of distilled water as 0 HU and the radiation density of air as 1000 HU at a standard pressure and temperature, and determines the radiation density of the measurement object based on these two radiation densities. In this study, the effect of trabecular bone density according to age and gender was

not significant in both the trabecular morphometry analysis and Hounsfield unit analysis. In the maxilla and mandible, males showed slightly higher Hounsfield unit than females, but were not statistically significant. Dutra et al. reported that bones that constantly undergo reconstruction were affected by the gender and dental condition [16]. Some previous studies reported that the bone density reduction in the mandible is apparent in women [17,18]. Pavlova and Peliakov reported that the difference between males and females was the delayed calcification in females, resulting in the less calcified bone [19]. However, Yong et al. reported that the difference between trabecular bones by age and gender was still controversial [20]. Klemetti et al. reported that the bone mineral density of the mandibular alveolar bone was influenced by the activity of the masticatory muscle than on the difference in gender [21]. In this study, it was judged that there was no difference according to the age or gender since the study subjects were relatively young patients undergoing an orthodontic treatment.

Misch reported that the mandible had a higher trabecular bone density than the maxilla, as a result of measuring the trabecular bone density based on Hounsfield unit measurements [4]. He also reported that bone densities tended to decrease from the anterior to posterior in both the maxillary and mandibular trabecular bones when analyzed by the Hounsfield unit. In this study, the Tb.th, BV/TV, and HU showed higher values in the anterior base of the maxilla than in the mandibular body. The reason why the value in the maxilla was higher than that in the mandible seems to be that the measurement region in the maxilla was located in the anterior position relative to the mandible, and the measurement of the maxilla was made in the basal bone. In previous studies, there have been studies that use CT to analyze the structure of the second cervical vertebra [22,23]. Montemurro et al. identified a new anatomical entity, named the Y-shaped trabecular structure of the odontoid process, on axial CT scans [23].

Tb.sp at the anterior base of the maxilla was the lowest among all regions in this study. Tb.sp is the average distance between trabeculae, and since it is related to the mean diameter of the cavities containing the bone marrow, not the bone, the low Tb.sp value tends to have a high trabecular bone density. The Hounsfield unit of this study was higher in the order of anterior base of the maxilla, mandibular body, mandibular condyle, and first cervical vertebra. In a medical CT, these scales are calibrated to express absolute values, but since the CBCT equipment may differ calibration, they are expressed as relative values rather than absolute values. If the imaging equipment is different or the settings are different, HU values may be different [24]. However, in this study, since the same region was measured with the same CBCT equipment and setting, the reliability of the comparison was secured even if the absolute value of HU was not measured. Among the trabecular microstructure parameters, bone volume fraction (BV/TV), a three-dimensional quantitative analysis, showed the most similar results to HU. When analyzing the structural characteristics of the trabecular bone, it is desirable to use Tb.th, Tb.n, Tb.sp, and it is considered that the method of BV/TV or HU is for the evaluation of the overall bone.

The limitation of this study was that the resolution of the used CBCT equipment was 250 μm , and some structural traits of the trabecular bone in the maxilla and mandible may be in fact below this threshold, so that cannot be completely measured [14]. If a higher-resolution CT is used, a more accurate three-dimensional structure of the trabecular bone and measurement of the Hounsfield unit will be possible. However, in that case, there is a disadvantage that more radiation doses are required than CBCT. The second was that the number of samples was small, so it is necessary to conduct the study with a large sample in the further research. Finally, this study was conducted on orthodontic patients, but the age distribution of orthodontic patients is often young and healthy, so the results may differ from those of elderly patients. Future studies on samples for various age groups and various systemic medical histories are needed. Koc, A. et al. reported that no significant difference was found when comparing the trabecular bone structures of edentulous patients and fully dentate patients [25]. In addition to age and gender, it may be

necessary to control variables for other factors that may affect the trabecular bone density, such as the characteristics of skeletal patterns.

5. Conclusions

Both trabecular bone density measurement methods using the Hounsfield unit and trabecular microstructure patterns can be compared between regions in CBCT images. In addition, the measurement method using the Hounsfield unit showed a significant correlation with the method represented by trabecular microstructure parameters.

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Institutional Review Board Statement: The study was conducted according to the guidelines of the Declaration of Helsinki, and approved by the Institutional Review Board of Korea University Guro Hospital (No. 2019GR0189).

Informed Consent Statement: Patient consent was waived because the x-ray image was taken for treatment use and there is no identifiable patient information.

Data Availability Statement: The data underlying this study will be available on reasonable request to the corresponding author.

Conflicts of Interest: The authors declare no conflict of interest.

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