

## **Reliability of anatomic structures as landmarks in three-dimensional cephalometric analysis using CBCT**

**Pegah Naji<sup>a</sup>; Noura A. Alsufyani<sup>b</sup>; Manuel O. Lagravère<sup>c</sup>**

### **ABSTRACT**

**Objective:** To identify anatomic structures in three dimensions and examine their reliability to be used as landmarks in a three-dimensional coordinate cephalometric analysis, using cone-beam computerized tomography (CBCT).

**Materials and Methods:** Thirty CBCT images were randomly selected for landmark location. Forty-two anatomic landmarks, which are not included in the traditional cephalometric landmarks, were chosen based on radiographic characteristics that make them pragmatic to mark in the CBCT image slices. The principal investigator marked the full set of landmarks on the software by navigating in the X, Y, and Z axes for every image three times, with each measurement trial being at least 1 week apart. One other investigator also located the landmarks once for each image for reliability purposes. Intraclass correlation coefficients (ICCs) were used to analyze the mean differences in landmark location in all axes.

**Results:** Intra- and interexaminer reliability for x, y, and z coordinates for all landmarks had ICC greater than 0.95 with confidence interval of 0.88–0.99. Mean measurement differences found were <1.4 mm for all landmarks in all three coordinates. Mean measurement error differences obtained in the principal investigator's trials were primarily <0.5 mm.

**Conclusion:** The most reliable and reproducible landmarks tested for use in CBCT are mental foramina, infraorbital foramina, inferior hamulus, dens axis, foramina transversarium of atlas, medial and lateral condyles of the mandible, superior clinoid processes, and mid-clinoid. (*Angle Orthod.* 2014;84:762–772.)

**KEY WORDS:** Cone-beam computerized tomography; Three-dimensional cephalometrics; X-rays; Orthodontics

### **INTRODUCTION**

For several decades, two-dimensional (2D) cephalometric radiographs have been used effectively for orthodontic diagnosis and treatment planning. Although 2D cephalometric radiography has been effective, it does present some deficiencies and limitations. The information in these radiographs is

subject to projection errors as well as landmark identification problems and measurement errors because they can only produce 2D data from a three-dimensional (3D) object.<sup>1,2</sup> In comparison, 3D volumetric imaging, such as cone-beam computerized tomography (CBCT) presents a better geometric accuracy and spatial resolution than regular computerized tomography with relatively low radiation dosages.<sup>2–5</sup>

In the past decade, the introduction of CBCT in dental radiology and its wide adoption for many clinical applications in dentistry has revived interest in using 3D cephalometric analysis for routine orthodontic cases.<sup>6</sup> Compared with the multislice computerized tomography, CBCT is less expensive, is more accessible, and offers a lower dose of radiation.<sup>7,8</sup> Furthermore, CBCT measurements are not significantly influenced by variation in skull orientation or head position during image acquisition.<sup>9,10</sup> The challenge for clinicians is to understand and interpret 3D imaging.<sup>11</sup> Currently, there are no specific guidelines about how to analyze this type of 3D image, especially in the

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orthodontic field. For this reason, new standards are required and clinicians need special training when dealing with 3D craniofacial images.

Linear measurements made on CBCT-derived 3D images are sufficiently clinically accurate and reliable.<sup>12</sup> Some studies have attempted to verify whether landmarks used in conventional imaging can be extrapolated to CBCT.<sup>1,13–18</sup> Some landmarks that are used in the 2D cephalometric analysis are challenging to locate in the CBCTs.<sup>15</sup> Many landmarks have demonstrated low reliability in CBCT, and therefore, are not suitable for use in 3D imaging. In these studies some of the least reliable conventional landmarks in CBCT were gonion, orbitale, condylion, A point, ANS, PNS, and pogonion.<sup>15–18</sup> Using conventional landmarks in the 3D analysis showed that 3D landmark identification using CBCT can offer consistent and reproducible data if a protocol for operator training and calibration is followed.<sup>14</sup> Some studies<sup>19,20</sup> have explained the need for a 3D-based measurement analysis when using a 3D Cartesian system. The reliability and repeatability of the identification landmarks are very high if the 3D cephalometric landmarks are defined correctly.<sup>21</sup> The purpose of this study is to identify several anatomic landmarks and test their reliability in CBCT images for use in 3D cephalometric analysis.

## MATERIALS AND METHODS

Institutional Ethical Review Board approval was obtained for the study. Thirty CBCT images of subjects (age range = 12–17 years) were randomly selected from the University of Alberta Orthodontic Graduate Program pool. All scans were taken for orthodontic diagnosis and treatment planning. Any CBCT scans demonstrating motion artifact were replaced. All CBCT scans were taken using the I-CAT machine (Imaging Sciences International, Hatfield, Pa) with a collimation height scan of 13 cm, scan time of 20 seconds, and resolution of 0.3 mm voxel size. Each image was converted to a Digital Imaging and Communications in Medicine (DICOM) format and analyzed using the Avizo software AVIZO 7.0, (Visualization Sciences Group, Burlington, Mass). The DICOM format images were rendered into a volumetric image.

Forty-two anatomic landmarks that are not included in the traditional cephalometric landmarks were chosen based on the radiographic characteristic of head and neck hard tissue structures, which are feasible to be localized in the CBCT images. The radiographic characteristics for choosing the landmarks were being presented as a small radiolucency, on a short curve, at the tip of a spine, and with a distinct contrast difference of the adjacent structures. The x, y, and z coordinates

of each landmark were defined based on the Cartesian system used by Lagravere et al.<sup>2</sup> (Table 1; Figures 1 through 12) to standardize the anatomic identification in the three planes of space and to guide the selection of the most precise location in the sagittal, axial, and coronal views.

The principal investigator marked the landmarks in the software using the virtual spherical markers of 0.5 mm diameter in the x, y, and z axes; the center of each marker represented the exact location of the landmark. The examiners marked a landmark in the first slice in which it was visible, moving mesiodistally, superiorly-inferiorly or anteriorly-posteriorly, and then adjusting the marker in other coordinates. For example, for marking the superior left clinoid process, the left clinoid process was marked when it was visible first on the axial slice moving inferiorly; the position was then adjusted in the coronal and sagittal views if needed.

The landmarks were marked three times for every image, and each measurement trial was at least 1 week apart. One other investigator also located the landmarks once for each image for reliability purposes. To reduce exhaustion effect, both investigators would discontinue the process of landmark identification once they felt tired and then resumed the process another day.

Mean differences and descriptive statistics were calculated for all landmarks and different axes. Intraclass correlation coefficients (ICCs) were calculated to determine the intra- and interexaminer reliability of each landmark. The mean differences of all the trials of the 3D cephalometric measurements of the 42 landmarks and their standard deviations are summarized in Table 2 for all three axes. These were obtained by averaging the differences between the four total trials of landmark location of the principal investigator and the second examiner for every axis. A landmark was considered clinically reliable if it presented <1.5 mm mean difference.

## RESULTS

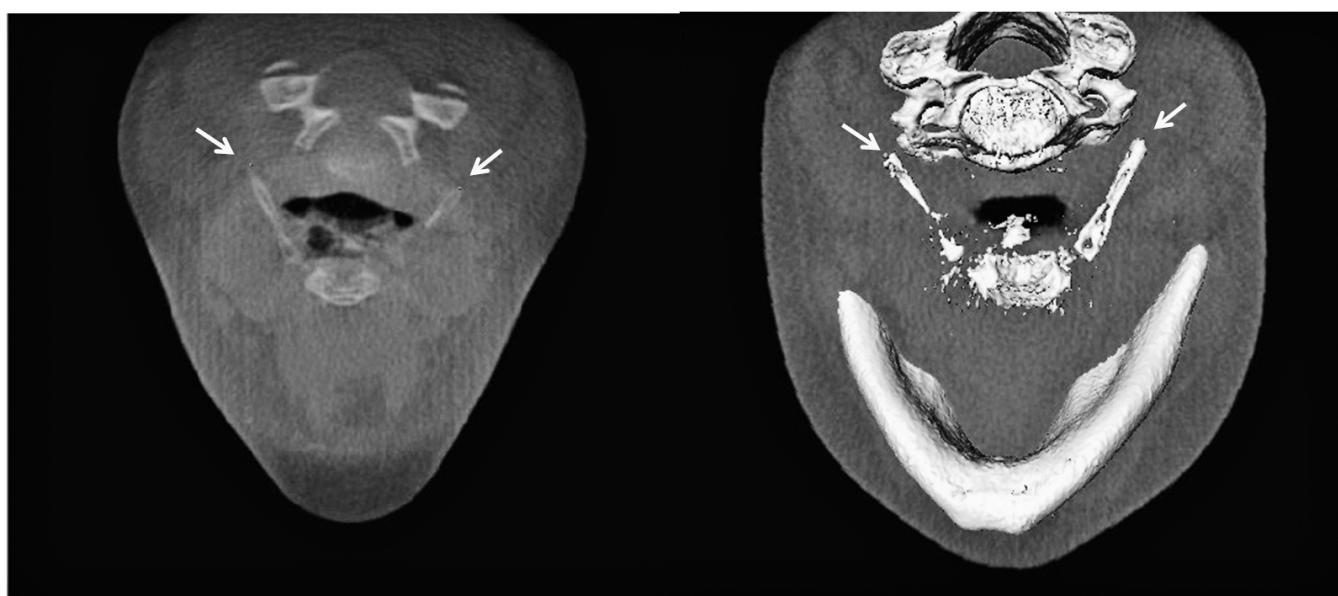
Intraexaminer reliability for x, y, and z coordinates for all landmarks had ICC greater than 0.98 with 95% confidence interval (CI) = -0.97, 1.00. Mean measurement error differences obtained in the principal investigator's trials were primarily <0.5 mm. Interexaminer reliability for x, y, and z coordinates for all landmarks has ICC >0.95 (CI = -0.88, 0.99). Mean differences of the intraexaminer and interexaminer measurements were all <1.4 mm. The only landmarks with mean values >1 mm in any of the x, y, and z coordinates were right and left posterior-inferior concha (1.39 mm and 1.09 mm, respectively), oral

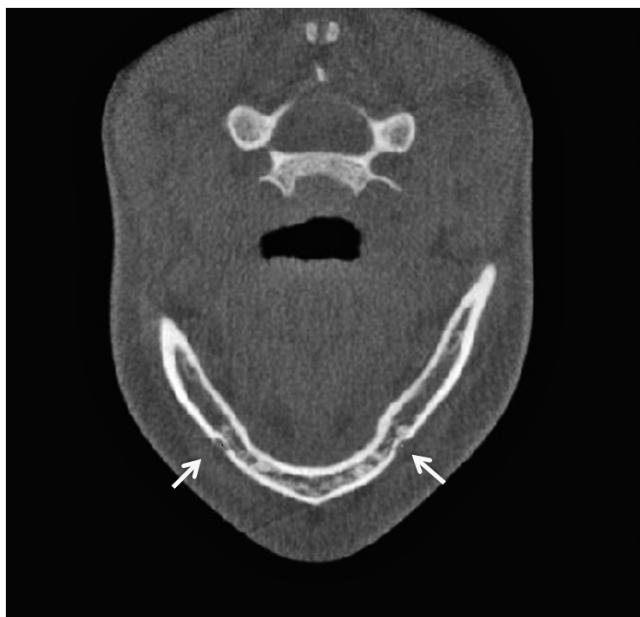
**Table 1.** Three-Dimensional Cephalometric Landmarks

Three-Dimensional Landmark	Definition	Sagittal View	Axial View	Coronal View
Posterior hyoid right (Figure 1)	Most middle-posterior point of the hyoid right greater cornu	Posterior-most point	Posterior-most point	Middle-posterior-most point
Posterior hyoid left (Figure 1)	Most middle-posterior point of the hyoid left greater cornu	Posterior-most point	Posterior-most point	Middle-posterior-most point
Mental foramen right (Figure 2)	Most middle-lateral point of the right mental foramen	Middle-most point	Middle-lateral-most point	Middle-lateral-most point
Mental foramen left (Figure 2)	Most middle-lateral point of the left mental foramen	Middle-most point	Middle-lateral-most point	Middle-lateral-most point
Lingula right (Figure 3)	Most posterior-medial point of mandible's right lingula sharp spine	Middle-posterior-medial-most point	Posterior-medial-most point	Middle-medial-most point
Lingula left (Figure 3)	Most posterior-medial point of mandible's left lingula sharp spine	Middle-posterior-medial-most point	Posterior-medial-most point	Middle-medial-most point
Dens axis (Figure 5)	Most middle-superior point of odontoid process of axis/C2	Superior-most point	Middle-superior-most point	Superior-most point
Right transversarium atlas (Figures 3 and 4)	Center of the right foramen transversarium of atlas/C1	Middle-most point	Middle-most point	Middle-most point
Left transversarium atlas (Figures 3 and 4)	Center of the right foramen transversarium of atlas/C1	Middle-most point	Middle-most point	Middle-most point
Right transverse process atlas (Figure 4)	Most middle-lateral point of the right transverse process of atlas/C1	Middle-lateral-most point	Lateral-most point	Middle-lateral-most point
Left transverse process atlas (Figure 4)	Most middle-lateral point of the left transverse process of atlas /C1	Middle-lateral-most point	Lateral-most point	Middle-lateral-most point
Inferior right hamulus (Figure 4)	Most middle-inferior point of right medial pterygoid process	Inferior-most point	Middle-inferior-most point	Inferior-most point
Inferior left hamulus (Figure 4)	Most middle-inferior point of left medial pterygoid process	Inferior-most point	Middle-inferior-most point	Inferior-most point
Inferior right lateral pterygoid plate (Figure 4)	Most middle-inferior point of right lateral pterygoid process of sphenoid	Inferior-most point	Middle-inferior-most point	Inferior-most point
Inferior left lateral pterygoid plate (Figure 4)	Most middle-inferior point of left lateral pterygoid process of sphenoid	Inferior-most point	Middle-inferior-most point	Inferior-most point
Right infraorbital (Figure 6)	Center of the right infraorbital foramen	External-most point	Middle-external-most point	Middle-most point
Left infraorbital (Figure 6)	Center of the left infraorbital foramen	External-most point	Middle-external-most point	Middle-most point
Superior right condyle (Figure 8)	Most middle-superior point of right condyle of the mandible	Superior-most point	Middle-superior-most point	Superior-most point
Posterior right condyle (Figure 8)	Most middle-posterior point of right condyle of the mandible	Posterior-most point	Posterior-most point	Middle-posterior-most point
Anterior right condyle (Figure 8)	Most middle-anterior point of right condyle of the mandible	Anterior-most point	Anterior-most point	Middle-anterior-most point
Superior left condyle (Figure 8)	Most middle-superior point of left condyle of the mandible	Superior-most point	Middle-superior-most point	Superior-most point
Posterior left condyle	Most middle-posterior point of left condyle of the mandible	Posterior-most point	Posterior-most point	Middle-posterior-most point
Anterior left condyle	Most middle-anterior point of left condyle of the mandible	Anterior-most point	Anterior-most point	Middle-anterior-most point
Superior right clinoid process (Figure 9)	Most middle-superior point of right posterior clinoid process of the sphenoid bone	Superior-most point	Superior-most point	Middle-superior-most point
Superior left clinoid process (Figure 9)	Most middle-superior point of left posterior clinoid process of the sphenoid bone	Superior-most point	Superior-most point	Middle-superior-most point
Mid-clinoid (Figure 9)	Midpoint of the line between the most superior points of two posterior clinoid processes of the sphenoid bone	Midpoint of right and left clinoid processes	Midpoint of right and left clinoid processes	Midpoint of right and left clinoid processes
Lateral right condyle	Most middle-lateral point of right condyle of the mandible	Middle-lateral-most point	Lateral-most point	Lateral-most point
Lateral left condyle	Most middle-lateral point of left condyle of the mandible	Middle-lateral-most point	Lateral-most point	Lateral-most point

**Table 1.** Continued

Three-Dimensional Landmark	Definition	Sagittal View	Axial View	Coronal View
Medial right condyle (Figure 5)	Most middle-medial point of right condyle of the mandible	Middle-medial-most point	Medial-most point	Medial-most point
Medial left condyle (Figure 5)	Most middle-medial point of left condyle of the mandible	Middle-medial-most point	Medial-most point	Medial-most point
Inferior right styloid process (Figure 3)	Most middle-inferior point of right styloid process of the temporal bone	Inferior-most point	Middle-inferior-most point	Inferior-most point
Inferior left styloid process (Figure 3)	Most middle-inferior point of left styloid process of the temporal bone	Inferior-most point	Middle-inferior-most point	Inferior-most point
Lower right first molar furca	Center of the furcation of mandibular right first molar	Center of the furcation	Center of the furcation	Center of the furcation
Lower left first molar furca (Figure 11)	Center of the furcation of mandibular left first molar	Center of the furcation	Center of the furcation	Center of the furcation
Upper right first molar furca (Figure 12)	Center of the furcation of mesial, distal, and palatal roots of maxillary right first molar	Center of the furcation	Center of the furcation	Center of the furcation
Upper left first molar furca (Figures 11 and 12)	Center of the furcation of mesial, distal, and palatal roots of maxillary left first molar	Center of the furcation	Center of the furcation	Center of the furcation
Posterior right inferior concha (Figure 7)	Most middle-posterior point of the right inferior nasal concha	Posterior-most point	Middle-posterior-most point	Middle-posterior-most point
Posterior left inferior concha (Figure 7)	Most middle-posterior point of the left inferior nasal concha	Posterior-most point	Middle-posterior-most point	Middle-posterior-most point
Oral incisive foramen (Figures 4 and 10)	Middle-inferior point of oral orifice of the nasopalatine canal	Inferior-most point	Middle-most point	Middle-inferior-most point
Nasal incisive orifice (Figure 10)	Middle-superior point of nasal orifice of nasopalatine canal (or the midpoint of the two orifices, if two nasal orifices are present)	Superior-most point	Middle-superior-most point	Superior-most point
Mandibular canal at lower right first molar furca (Figure 12)	Center of alveolar canal at the level of lower right first molar furca	—	—	Center of alveolar canal at the level of lower right first molar furca
Mandibular canal at lower left first molar furca (Figure 12)	Center of alveolar canal at the level of lower left first molar furca	—	—	Center of alveolar canal at the level of lower left first molar furca

**Figure 1.** Posterior hyoid right, posterior hyoid left in the axial view and three-dimensional reconstruction.



**Figure 2.** Mental foramen right, mental foramen left in the axial view.

incisive foramen (1.25 mm), inferior left styloid (1.07 mm), right lingula (1.03 mm), and inferior right and left lateral pterygoid (1.16 mm and 1.10 mm, respectively); the highest amount was 1.39 mm for the right posterior-inferior concha. The posterior hyoid left was not present in two CBCTs and was only marked in 28 samples.

Landmarks that had mean differences  $\leq 0.5$  mm in all coordinates were right and left mental foramina,

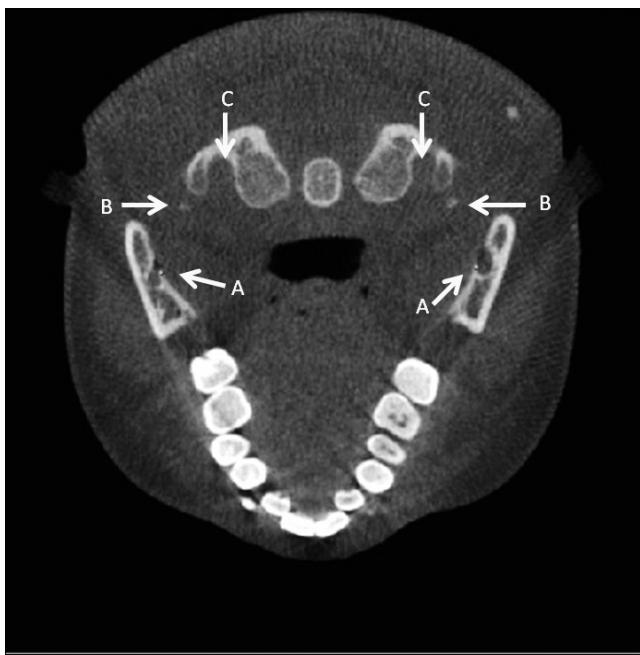


**Figure 4.** (A) Right and left inferior medial pterygoid plate. (B) Right and left transverse process atlas. (C) Right and left transversarium atlas. (D) Right and left inferior lateral pterygoid plate. (E) Oral incisive foramen in the axial view.

dens axis, right and left transversarium atlas, right and left inferior hamulus, right infraorbital foramen, medial right condyle, and lateral left condyle.

## DISCUSSION

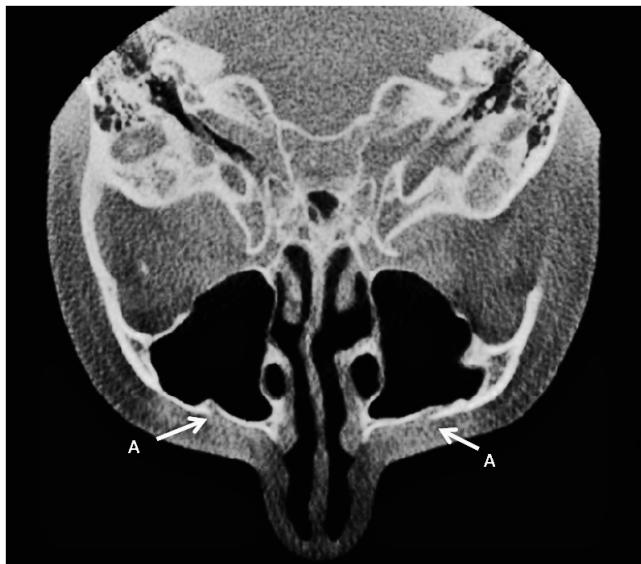
Studies show that CBCT craniometric measurements are more accurate than traditional measurements and can potentially be used as a quantitative orthodontic diagnostic tool.<sup>13,22</sup> The main advantage in tomographic scans is the possibility of interacting with



**Figure 3.** (A) Lingula right and left. (B) Right and left transversarium atlas. (C) Right and left inferior styloid process in the axial view.



**Figure 5.** (A) Right and left medial condyle. (B) Dens axis in the coronal view.



**Figure 6.** Right and left infraorbital foramina in the axial view.

previously stored data, which in turn generates new images and allows the practitioner to reconstruct synthesized cephalograms that are comparable with conventional radiographs, based on a single 3D measurement.<sup>23,24</sup> However, the use of lateral cephalograms might be limited in some cases; for example, lateral cephalograms are difficult to accurately superimpose because of the difference between the right and left sides, such as differences in scaling ratios, variations in head positioning, and overlapping of various cranial structures.<sup>25</sup>

Some studies have analyzed the reliability of taking landmarks used in traditional cephalometry and

locating them in CBCTs.<sup>1,7</sup> Some studies have used CBCT-synthesized cephalograms and showed that the reproducibility of exams obtained from conventional cephalograms and CBCT-synthesized cephalograms was similar.<sup>24</sup> Traditional cephalometric landmarks have been defined using a 2D x-ray, giving them just two axes to worry about.

With CBCT, visualization of structures can be done using all three axes, thus opening the possibility of analyzing new 3D-defined landmarks to use with 3D cephalometric analysis. However, the development of 3D landmark-based cephalometric analysis requires definition of 3D landmarks on complex curving structures.<sup>14</sup> Practical considerations of identification errors, coupled with an essential need for biological relevance and a balanced representation of components of the craniofacial form, limit the number and nature of landmarks available for analysis.<sup>14</sup> Historically, landmarks such as articulare were used because of the ease in landmark location on 2D cephalometric projections, but these projected superimposed structures do not exist in the actual 3D facial structure. For these reasons, the development of 3D landmark-based cephalometric analysis demands suitable operational definitions of the landmark location in each of the three planes of space.<sup>14,26</sup>

In this study, the goal was to analyze the ease of locating and the reliability of anatomic structures or sites to be used in the development of a 3D analysis. Clinical significance of variation in repeated landmark location is difficult to define and depends on the purpose of analysis. In traditional, clinical 2D cephalometrics, a landmark error <1 mm is considered a



**Figure 7.** Posterior right and left inferior concha in the coronal view and the posterior right inferior concha in the sagittal view.



**Figure 8.** (A) Anterior right condyle in the axial and sagittal views. (B) Posterior right condyle in the axial and sagittal views. (C) Superior right condyle in the sagittal view.

precise measurement. Variations  $<1.0$  mm are unlikely to have clinical significance if the cranial base landmarks are used for linear or angular measurements.<sup>27</sup> Variations  $<0.5$  mm are probably not clinically significant, and variations between 0.5 mm and 1.0 mm might be clinically relevant.<sup>22</sup> Whether this measure is applicable in 3D cephalometrics might be questionable because a third axis is introduced in 3D cephalometrics that may add errors to the overall error.<sup>7,25</sup>

All landmarks studied had mean differences  $<1.4$  mm in all three coordinates. However, seven axis coordinates of all the landmarks identified had a value  $\geq 1$  mm. Several landmarks showed mean

differences close to 0.5 mm in all three coordinates, which makes them reliable to be used in 3D analysis. During this study, the examiners faced some difficulties in locating the landmarks according to the 3D definitions, which could have affected the mean values measured. The difficulties and possible causes that make these landmarks hard to locate are listed in Table 3.

In this study, the oral incisive foramen showed 1.25 mm of mean value measurement in the z coordinate. The size of the foramen, location in the palate curvature, and anatomic variation of this structure could have caused the wide range in



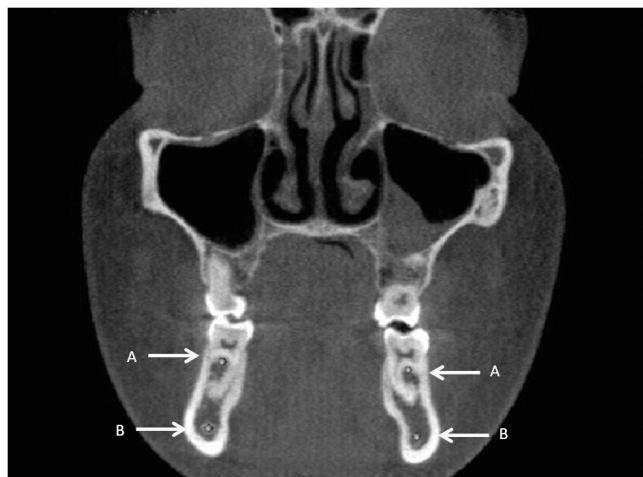
**Figure 9.** (A) Superior right and left clinoid processes in the axial and sagittal views. (B) Mid- clinoid in the axial view.



**Figure 10.** (A) Nasal incisive orifice. (B) Oral incisive foramen in the sagittal view.

localizing the landmark. De Oliveira-Santos and colleagues<sup>28</sup> demonstrated that additional palatine foramina at least 1 mm in diameter may be found in around 16% of patients of both genders and different age groups. Also, the oral incisive foramen could have been identified in different coronal planes because of its large dimension. Furthermore, it is located on the hard palate curvature, which makes it difficult to unquestionably mark its most inferior point.

The size and shape, contrast, and location of the anatomic structure can influence its accurate localizing. The precise 3D definition of the structure in three dimensions also influences localizing the landmarks.<sup>14</sup> For example, the midpoint of upper first molar furca could be marked close to the buccal, mesial, or distal



**Figure 12.** (A) Right and left upper first molar furca. (B) Right and left mandibular canal at first lower molar furca in the coronal view.

furca, within a wide area that might be more than the favorable range of 1 mm. In some of the studied CBCT slices, the posterior point of inferior concha was more like a flat plane than a pointy or curved structure, which makes it hard to define as a single point. In this study, the inferior left styloid showed a mean 1.07 mm of mean measurement value in the z coordinate. Stylohyoid ligament, which connects the styloid process to the lesser horn of the hyoid bone, can be subject to calcification.<sup>29</sup> In a study, Alpoz and colleagues<sup>30</sup> showed ratios of different patterns of the styloid process from the normal styloid process (68.3%), elongated styloid process (27.1%), calcified stylohyoid ligament (1.7%), and absent stylohyoid chain (2.5%). These variations might affect the examiner's judgment in the precise localizing of the landmarks.



**Figure 11.** (A) Upper left first molar furca. (B) Lower left first molar furca in the sagittal view.

**Table 2.** Absolute Mean Value Measurement and Standard Deviation (mm) in Coordinates of Landmarks Based on All Trials (Intra- and Interexaminer)

Landmarks	N	X Axis		Y Axis		Z Axis	
		Mean	SD	Mean	SD	Mean	SD
Posterior hyoid right	30	0.59	0.37	0.82	0.58	0.71	0.51
Posterior hyoid left	28	0.56	0.50	0.71	0.49	0.88	0.63
Mental foramen right	30	0.48	0.32	0.39	0.26	0.31	0.27
Mental foramen left	30	0.40	0.20	0.44	0.28	0.40	0.36
Lingula right	30	0.43	0.26	0.94	0.61	1.03	0.78
Lingula left	30	0.41	0.25	0.70	0.49	0.79	0.60
Dens axis	30	0.46	0.28	0.42	0.20	0.13	0.18
Right transversarium atlas	30	0.34	0.19	0.47	0.27	0.50	0.37
Left transversarium atlas	30	0.41	0.20	0.38	0.26	0.39	0.34
Right transverse process atlas	30	0.47	0.33	0.65	0.52	0.56	0.40
Left transverse process atlas	30	0.48	0.33	0.65	0.39	0.50	0.35
Inferior right hamulus	30	0.47	0.35	0.50	0.28	0.37	0.41
Inferior left hamulus	30	0.54	0.37	0.45	0.28	0.36	0.35
Inferior right lateral pterygoid plate	30	1.16	0.72	0.91	0.61	0.48	0.41
Inferior left lateral pterygoid plate	30	1.10	0.58	0.92	0.59	0.54	0.55
Right infraorbital	30	0.51	0.47	0.51	0.20	0.57	0.54
Left infraorbital	30	0.51	0.31	0.54	0.37	0.66	0.43
Superior right condyle	30	0.63	0.36	0.41	0.25	0.12	0.13
Posterior right condyle	30	0.73	0.43	0.50	0.31	0.62	0.47
Anterior right condyle	30	0.86	0.67	0.38	0.29	0.47	0.32
Superior left condyle	30	0.65	0.50	0.43	0.24	0.17	0.19
Posterior left condyle	30	0.96	0.74	0.46	0.24	0.56	0.36
Anterior left condyle	30	1.00	0.79	0.37	0.23	0.61	0.42
Superior right clinoid process	30	0.60	0.51	0.33	0.18	0.17	0.23
Superior left clinoid process	30	0.64	0.68	0.41	0.20	0.16	0.23
Mid-clinoid	30	0.56	0.45	0.43	0.25	0.56	0.92
Lateral right condyle	30	0.22	0.22	0.53	0.27	0.59	0.51
Medial right condyle	30	0.25	0.24	0.52	0.30	0.51	0.36
Inferior right styloid process	30	0.80	0.92	0.55	0.54	0.73	0.55
Lower right first molar furca	30	0.66	0.34	0.49	0.26	0.40	0.19
Upper right first molar furca	30	0.74	0.35	0.46	0.31	0.59	0.33
Posterior right inferior concha	30	0.93	0.64	1.39	0.82	0.54	0.30
Posterior left inferior concha	30	0.88	0.49	1.09	0.72	0.45	0.24
Lower left first molar furca	30	0.71	0.47	0.55	0.40	0.55	0.29
Upper left first molar furca	30	0.80	0.66	0.48	0.24	0.77	0.48
Inferior left styloid process	30	0.96	0.70	0.77	0.68	1.07	0.99
Medial left condyle	30	0.32	0.19	0.55	0.29	0.71	0.35
Lateral left condyle	30	0.21	0.17	0.54	0.32	0.54	0.35
Oral incisive foramen	30	0.40	0.30	0.97	0.70	1.25	0.71
Nasal incisive orifice	30	0.42	0.26	0.55	0.39	0.62	0.33
Mandibular canal at lower right first molar furca	30	0.77	0.36	0.62	0.36	0.65	0.49
Mandibular canal at lower left first molar furca	30	0.88	0.60	0.77	0.63	0.87	0.66

Bone thickness, bone mineralization, and bone density can contribute to the identification of the bone boundaries.<sup>31</sup> The quality of the scan is a crucial factor in landmark identification error. Locating the inferior alveolar nerve canal at the level of mandibular first molar furca can be influenced by the contrast and quality of the radiograph. It can be well-defined in sharp and high-quality scans and challenging to locate using scans with low signal to noise ratio. This can also be attributed to the low-density contrast with adjacent landmarks due to bone quality; however, this correlation needs to be confirmed with further study.

Locating the most inferior point of the medial pterygoid, the pterygoid hamulus was of high accuracy

and reliable; however, locating the most inferior point of the lateral pterygoid process can be challenging because of the flat edge of the inferior border of the lateral pterygoid plate. Foramen transversarium of the atlas showed high reproducibility while locating. Even though it is relatively large, foramen requires searching through three or four slices. Gupta and colleagues<sup>31</sup> reported incomplete foramen transversarium of the atlas in 8.57% of specimens. However, the center of this foramen can be estimated based on the boundaries of the transverse processes. It is suggested that the size and reliability of foramina transversarium of the axis and C3 should be studied for inclusion in cephalometric landmarks.

**Table 3.** Challenges Presented When Trying to Identify Each Landmark

Landmark	Reason
Posterior hyoid right and left	Landmark can be missed in some scans, that is, is not included in the field of view
Inferior right and left lateral pterygoid plate	Some plates might have an inferior flat border, which makes it a broad surface for localization
Infraorbital foramen	Obliquity of the foramen renders localization of the center difficult
Points along condyle	Condylar surfaces can be flat, which makes localization of a single point difficult
Inferior right and left styloid processes	Eagle syndrome and calcification/ossification of the stylohyoid ligament could occur, thereby losing this landmark, which means it will be indistinguishable from the calcified/ossified ligament
Posterior right and left inferior concha	In some cone-beam computerized tomographic scans the inferior concha had a flat posterior surface, making it hard for precise localizing
Molar furca	Presence of restorations could cause beam-hardening artifact and make it difficult to estimate the location
Nasal incisive orifice	Presence of multiple nasal incisive foramina renders their localization difficult
Oral incisive foramen	Broad obliquity of the foramen renders localization of the center difficult

Overall, all landmarks showed a mean difference <1.4 mm, which makes them reliable for use in 3D cephalometric analysis; however, some specific landmarks showed a much lower mean difference in locating and can be used for 3D cephalometric analysis. Future studies are recommended to assess the deviation of the marked landmarks from the skull model using radiopaque markers and to evaluate changes in these landmarks as subjects age.

## CONCLUSION

- The most reliable and reproducible landmarks tested for use in CBCT are mental foramina, infraorbital foramina, inferior hamulus, dens axis, foramina transversarium of atlas, medial and lateral condyles of the mandible, superior clinoid processes, and mid-clinoid.

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