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COMPARATIVE LINEAR ACCURACY AND RELIABILITY OF CONE BEAM CT
DERIVED 2-DIMENSIONAL AND 3-DIMENSIONAL IMAGES CONSTRUCTED USING
AN ORTHODONTIC VOLUMETRIC RENDERING PROGRAM.

By

Danielle R. Periago
B.A., Oral Roberts University, 2000
D.M.D., University of Louisville, 2005

A Thesis
Submitted to the Faculty of the
Graduate School of the University of Louisville
in Partial Fulfillment of the Requirements
for the Degree of

Masters of Science in Oral Biology

Department of Orthodontics and Pediatric Dentistry
University of Louisville
Louisville, KY

August, 2007

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A Thesis Approved on

June 12, 2007

by the following Thesis Committee:

Thesis Director

DEDICATION

This thesis is dedicated to my husband

Juan, who always gives me

unconditional love and support in all of my endeavors, and to my daughter Hope,

who gives me the hope and strength to persevere.

ACKNOWLEDGEMENTS

I would like to sincerely thank my thesis director, Dr. William C. Scarfe, for his guidance and patience in this project. A special thanks needs to be given to the Department of Radiology, Dr. William C. Scarfe and Dr. Allan G. Farman, for providing me with the opportunity to access and use the Cone Beam CT. I would also like to thank my other thesis committee members, Dr. Anibal Silveira and Dr. James Scheetz for their guidance and support. I would finally like to express a special thanks to my co-resident, Dr. Mazyar Moshiri, for his help and collaboration throughout this project.

ABSTRACT

COMPARATIVE LINEAR ACCURACY AND RELIABILITY OF CONE BEAM CT DERIVED 2-DIMENSIONAL AND 3-DIMENSIONAL IMAGES CONSTRUCTED USING AN ORTHODONTIC VOLUMETRIC RENDERING PROGRAM.

Danielle R. Periago, D.M.D.

June 12, 2007

The purpose of this project was to compare the accuracy and reliability of linear measurements made on 2D projections and 3D reconstructions using Dolphin 3D software (Chatsworth, CA) as compared to direct measurements made on human skulls.

The linear dimensions between 6 bilateral and 8 mid-sagittal anatomical landmarks on 23 dentate dry human skulls were measured three times by multiple observers using a digital caliper to provide twenty orthodontic linear measurements.. The skulls were stabilized and imaged via PSP digital cephalometry as well as CBCT. The PSP cephalograms were imported into Dolphin (Chatsworth, CA, USA) and the 3D volumetric data set was imported into Dolphin 3D (Version 2.3, Chatsworth, CA, USA). Using Dolphin 3D, planar cephalograms as well as 3D volumetric surface reconstructions were (3D CBCT) generated. The linear measurements between landmarks of each three modalities were then computed by a single observer three times. For 2D measurements, a one way ANOVA for each measurement dimension was calculated as well as a post hoc Scheffe multiple comparison test with the anatomic distance as the control group. 3D measurements were compared to anatomic truth using Student's t test ($P \leq 0.05$). The intraclass correlation coefficient (ICC) and absolute linear and percentage error were determined as indices of intraobserver reliability.

Our results show that for 2D mid sagittal measurements that Simulated LC images are accurate and similar to those from PSP images (except for Ba-Na), and for bilateral measurements simulated LC measurements were similar to PSP but less accurate, underestimating dimensions by between 4.7% to 17%. For 3D volumetric renderings, 2/3rd of CBCT measurements are statistically different from actual measurements, however this possibly is not clinically relevant

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CHAPTER I

INTRODUCTION

Radiographic imaging is an important diagnostic adjunct in the assessment of skeletal and dental relationships for the orthodontic patient. Historically, cephalometric analysis of the maxillofacial complex for orthodontic diagnosis and treatment planning has been determined from linear and angular measurements made on film or digital two dimensional (2D) cephalograms. Over the past decade, cone beam computed tomography (CBCT) specifically for imaging the maxillofacial region has been developed. CBCT is capable of providing sub-millimeter spatial resolution for images of the craniofacial complex with relatively short scanning times (10-70 sec.) and generally lower radiation dosages than ascribed to fan-beam or helical CT imaging methods.[1] Time and dose requirements for CBCT have been suggested to be a similar order of magnitude to other dental radiographic modalities.[2-4]

While CBCT images provide useful information for the orthodontist in regard to the position and location of impacted teeth and other pathologies, datasets can be used to generate both two dimensional (2D) planar projection and three dimensional (3D) surface or volume rendered images for use in orthodontic assessment and treatment planning. CBCT has a number of advantages compared to conventional CT imaging for cephalometric imaging including sub-millimeter resolution and reduced radiation exposure. Perhaps the most important clinical advantage is that CBCT volumetric datasets can be exported as DICOM files, imported into personal computers and third party software used to provide 3D reconstruction of the craniofacial skeleton. This possibility, and the increasing access of CBCT imaging in orthodontics, is a

component of the paradigm that is directing imaging analysis from 2D cephalometry to 3D visualization of craniofacial morphology.[5] The availability of fast scan CBCT now provides multi-planar reformatted (MPR) imaging and the possibility of 3D image reconstruction of the maxillofacial complex with minimal distortion.

The linear accuracy of CBCT derived 2D and 3D reconstructions have not been previously reported for orthodontic assessment. Therefore this study was made to compare the accuracy of linear measurements made on 2D planar projections and 3D surface reconstructions generated using Dolphin 3D software (Chatsworth, CA) applied to CBCT DICOM datasets with direct measurements made on human skulls.

The aims of this study were to compare the:

- 1) reliability of linear measurements made from lateral cephalometric plain projection digital images and ray sum 2D simulated cephalometric projections derived from CBCT volumetric datasets using Dolphin 3D software (Chatsworth, CA) to direct measurements made on a sample of 25 human skulls.
- 2) accuracy of linear measurements made from lateral cephalometric plain projection digital images and ray sum 2D simulated cephalometric projections derived from CBCT volumetric datasets using Dolphin 3D software (Chatsworth, CA) to direct measurements made on a sample of 25 human skulls.
- 3) reliability of linear measurements made on CBCT derived 3D surface rendered volumetric images generated using Dolphin 3D software (Chatsworth, CA) to direct measurements made on a sample of 25 human skulls.
- 4) accuracy of linear measurements made on CBCT derived 3D surface rendered volumetric images generated using Dolphin 3D software (Chatsworth, CA) to direct measurements made on a sample of 25 human skulls.

The Limitations of Conventional Film Based Cephalometric Analysis

Since 1931, 2D transmission X-ray images have been used to identify specific skull landmarks from which vertical and antero-posterior skeletal and dental dimensions are derived. These lateral skull radiographs, made under standard projection conditions, are currently the image format used in the analysis of both bony and soft tissue landmarks for orthodontic diagnostic purposes as well as for growth evaluation. Post-treatment cephalograms may also serve to evaluate orthodontic treatment outcome and success. Traditionally, cephalograms have been utilized for their cost and radiation efficiency as well as their ease of use. However, characteristics related to projection geometry such as inherent magnification, superimposition of bilateral anatomic structures and distortion as well as the nature of the detector system can diminish accuracy and reliability in evaluation of craniofacial structures and anomalies.

Digital Cephalometrics

Many conventional film based cephalostats are being replaced by digital systems. The advantages of digital cephalometric imaging versus conventional film based modalities include instantaneous imaging, lack of user and performance sensitive chemical developing processes, facilitated patient communication, ease of storage and retrieval, and the ability to enhance images for size or contrast.[6-8] Currently, three methods are available to produce digital images: digitization of film radiographs, solid state systems (charge-coupled device – CCD; complementary metal oxide semiconductor – CMOS; thin film transistor - TFT), and photostimulable phosphor systems (PSP).[6-10] Secondary capture through digitization of film radiographs can be achieved using a scanner with a radiograph/transparency adaptor. This method allows for digitization of all film radiographs, however, it is important to note that the quality of scanned images cannot exceed the quality of the original radiograph.[8] CCD detectors are sometimes incorrectly listed in the dental literature as direct digital imaging modalities, because the output is transferred via cables to a computer system and digitized by the frame grabber.[10]

They are in fact usually indirect imaging devices as they employ a scintillator in most cases, similar to that used with indirect screen film. CCD is the more costly option for cephalometry in orthodontics. Photostimulable phosphor systems (PSP) are reusable and use an imaging plate that superficially resembles scintillating screens used for traditional extra-oral radiography.[8] These phosphor plates are illuminated by a solid state laser beam to release photoluminescence. The released light is photomultiplied and collected by a digital imaging chip and the signals are then analyzed by the image processor.[6-10]

Image quality in cephalometrics either analog or digital, is determined by two parameters: image accuracy and image quality

Image Accuracy

Cephalometric radiography is based on use of a standardized, reproducible head position in relation to the X-ray source and detector. Ear rods are used to prevent the head from rotating about the vertical, sagittal and transverse axes. A third reference, a nasal positioner, may be used to prevent the nose from rotating about the transverse axis. However, when the device is used to contact the external auditory meatus and soft tissues of the patient, the head can be incorrectly positioned sagittally, antero-posteriorly, or vertically, as the head can be slightly rotated within the head-holding device.

Due to these errors caused by different positioning of the head, cephalometric linear and angular measurements can vary depending on the different locations of anatomic structures against the central ray. Malkoc *et al.* found that horizontal linear and angular measurements between the horizontal planes on lateral cephalograms were subject to changes from 16.1% to 44.7% with a 14° rotation of the head position. For PA cephalograms, they reported horizontal linear measurements, particularly mandibular length, were subject to a projection error of up to 34.9% with head rotation.[11]

2D transmission cephalometric radiography is subject to inherent geometric differential magnification. All resulting images are magnified, because X-rays do not radiate parallel to the whole part of the projected object. The ratio of magnification varies in the different planes, and hence the image is distorted. In cephalometric radiography, each landmark is not located at the same distance from the focal area of the anode. As a result, changes may be caused in the relationship of the landmarks to one and another on the cephalogram.[12,13]

Image Clarity

Clarity is the term used to describe the visibility of diagnostically important detail in an image. It is determined by two factors; radiographic contrast and image quality. Radiographic contrast is the ability to determine the difference in density between areas of the image. For both analog radiographic film and digital detectors contrast depends on radiation energy, subject contrast and scatter; however, a fourth element, detector contrast, is also a factor due to inherent dissimilarity between detection systems.

Image quality is defined as the ability to record each point in an object as a point on the detector. For film imaging it is partly determined by radiographic mottle (a feature of the film screen system and film graininess), sharpness and resolution. For digital detectors, seven essential characteristics should be considered: size of active area, signal-to-noise ratio, contrast resolution, spatial resolution, modulation transfer function, quantum efficiency and detective quantum efficiency.[8,15,16]

1. **Active Area:** No standard active areas have been specified for digital imaging systems comparable to the ISO/ANSI standards for the conventional X-ray film. For solid-state extra-oral systems, a narrower receptor is sometimes used for detecting the image and the image is formed via virtual movement. The plates used in storage phosphor systems can be cut to exactly replicate the size of their film counterparts and exposure is similar to cassette motion.

2. **Signal-to-Noise Ratio:** For any imaging system, the useful signal must be compared with background noise which, in analog film, is comparable to the base density plus fog. The base plus fog density for conventional processed film is about 1/20 of the signal density. Both newer CCD and PSP systems outperform film in signal-to-noise ratios (SNS) if base plus fog is considered to be equivalent to SNR. Newer CCD systems exhibit a SNR of approximately 50:1. No matter what the system, all SNRs improve with increased radiation dose.
3. **Contrast Resolution:** In imaging, the ability to separate and distinguish depends upon contrast between adjacent structures. Using current display monitors, working on the WINDOWS system, the maximum number of gray levels is 242 because the operating system in the past has been reported to use 14 shades and the total supported shades is 256 for an 8-bit display. This is usually the maximum contrast resolution available.
4. **Spatial Resolution:** Resolutions comparable to those of conventional cephalometric radiographs are readily obtained using digital systems/detectors. Table 1 compares detector resolution for a number of currently available conventional film, CCD systems and PSP systems.[14]
5. **Modulation Transfer Function:** MTF is the ability of the detector to transfer the modulation of the input signal at a certain frequency to its output and deals with the display of contrast and object size. MTF is responsible for converting contrast values of different sized objects into contrast intensity levels within the image. Therefore, modulation transfer function (MTF) is a useful measure of true or effective resolution, because it accounts for the amount of contrast and blur over a range of spatial frequencies.[15]

6. **Quantum Efficiency:** The average number of electrons photoelectrically emitted from a photocathode per incident photon of a given wavelength in a phototube.
Quantum efficiency (QE) is a quantity defined for a photosensitive device such as photographic film or a charge-coupled device (CCD) as the percentage of photons hitting the photoreactive surface that will produce an electron–hole pair. It is an accurate measurement of the device's sensitivity. It is often measured over a range of different wavelengths to characterize a device's efficiency at each energy.
Photographic film typically has a QE of much less than 10%, while CCDs can have a QE of well over 90% at some wavelengths.[16]
7. **Detective Quantum Efficiency:** Detective quantum efficiency (DQE) refers to the efficiency of a detector in converting incident x-ray energy into an image signal, and is calculated by comparing the signal-to-noise ratio at the detector output with that at the detector input as a function of spatial frequency. It is dependent upon radiation exposure, spatial frequency, MTF, and detector material as well as the quality of the radiation applied. High DQE levels indicate that less radiation is needed to achieve identical image quality, therefore, improved image quality can be obtained by increasing DQE and leaving radiation exposure constant. An ideal detector would have a DQE of 1, indicating that all radiation energy is absorbed and converted into image information. However, in clinical practice the DQE of digital detectors is limited to roughly 0.45 at 0.5 cycles/mm.[15]

Table 1. Comparison of maximum resolution of Imaging Modalities

	<i>Analog Film (T-Mat G)</i>	<i>Storage Phosphor</i>	<i>CCD-Based</i>	
<i>Maximum Resolution</i>	<i>OP 100</i>	<i>OP 100 DenOptix</i>	<i>OP 100 DigiPan</i>	<i>Prototype OP 100 D</i>
lp/mm	>5; <6	>5; <6	>4.47; <4.86	>5; <6

Digital cephalometric images have been reported to be clinically and diagnostically acceptable for orthodontic treatment planning purposes;[7-10] however, there is a need to further compare various radiographic modes of image capture for cephalometry such as conventional vs. digital radiographs and scanned conventional films vs. digital radiographs.[9]

Advanced Imaging Modalities in Orthodontics

Advanced technologies are those that acquire images using a digital receptor and that provide the possibility of multiple planar reformatting (MPR). In these modalities, multiple images become truly inter-relational in that direct comparisons in multiple planes can be made. Some advanced technologies that are available to image the maxillofacial complex include magnetic resonance imaging (MRI), fan-beam computerized tomography (CT), and Cone Beam Computed Tomography (CBCT). The basis of advanced imaging is the recording of transmitted, attenuated x-rays of an object by a digital receptor to produce a digital image. Digital images are composed of pixels, or picture elements, arranged in a 2-dimensional rectangular grid. Each pixel has a specific size, color, intensity value, and location within an image and is the smallest element of the digitized image. In general, radiographic images use gray color with an intensity value between 8 bits (256 shades of gray) and 12 bits (4096 shades of gray). The number of pixels per given length of an image (pixels/mm), the number of gray levels per pixel (bits), and the management of the gray levels determine image resolution or the degree of sharpness of the image. A voxel is a three-dimensional stack of bitmapped images, (each voxel having a height, width, and thickness) and is the smallest element of a three-dimensional image.[18]

Computed Tomography

In addition to utilizing images that are digital, technological advancements now allow dentistry to create images of the maxillofacial region in 3-dimensions. The first 3D imaging technique used in dentistry was computerized tomography (CT). CT units can be divided into two

groups based on the acquisition X-ray geometry: fan beam and cone beam (Figure 1). Essentially, the latter method for capturing an image differs from the traditional CT in that it does so by cone beam volumetric tomography. A three-dimensional X-ray beam passes through the object volume investigated. Simultaneously, the beam hits a two-dimensional extended detector and forms a true volumetric acquisition in a single scan (Figure 1).

Figure 1. X-ray beam projection scheme comparing conventional or "fan beam" CT (a.) and cone beam CT (b.) geometry (Images courtesy Predag Sukovic, Xoran Technologies, Ann Arbor, MI USA)

Fan beam acquisition

CT scanners consist of an X-ray source and detector mounted on a rotating gantry (Figure 1a). During one rotation of the gantry, the detector detects the flux (I) of x-rays that have passed through the patient. These integrals constitute so-called "raw data" that are then fed into an image reconstruction method that generates cross-sectional images whose pixel values correspond to linear attenuation coefficients. Such machines acquire image data through a thin, broad, fan shaped X-ray beam which is transmitted through the patient. These scanners use a large, arc-shaped detector that acquires an entire projection without the need for translation. This rotate-only design, frequently referred to as "fan-beam", utilizes the power of the X-ray tube much more efficiently than the previous generations. Recent advances in CT include multi-row detectors and spiral scanning. Multi-row scanning allows for the acquisition of several cross-sectional slices at the same time, reducing scanning times. Today's state-of-the-art scanners have 64 rows of detectors. Spiral (helical) scanning incorporates a moving table with the rotating X-ray tube, with the net effect that the X-ray tube describes a helical path around the patient.

Cone Beam Acquisition

CBCT scanners often utilize a 2D flat panel detector (Figure 1b), which allows for a rotation of the gantry to generate a scan of the entire region of interest using a 180 degree or greater rotation (up to two 360 degree rotations), as compared to conventional CT scanners whose multiple "slices" must be stacked to obtain a complete image. In comparison with conventional fan-beam or spiral-scan geometries, cone-beam geometry has higher efficiency in X-ray use, inherent quickness in volumetric data acquisition, and potential for reducing the cost of CT. Conventional fan-beam scans are obtained by illuminating an object with a narrow, fan-shaped, beam of X-rays. The X-ray beam generated by the tube is focused to a fan-shaped beam by rejecting the photons outside the fan, resulting in a highly inefficient use of the X-ray photons. Further, the fan-beam approach requires reconstructing the object slice-by-slice and then stacking

the slices to obtain a 3D representation of the object. Each individual slice requires a separate scan and separate 2D reconstruction. The cone beam technique, on the other hand, requires only a single scan to capture the entire object with a cone of X-rays. Thus, the time required to acquire a single cone-beam projection is the same as that required by a single fan-beam projection.

However, since it takes several fan beam scans to complete the imaging of a single object, the acquisition time for the fan beam tends to be much longer than with the cone beam. Although it may be possible to reduce the acquisition time of the fan beam method by using a higher power X-ray tube, this increases the cost and size of the scanner as well as the electric power consumption, thus making the design unsuitable for a compact scanner.

Although CBCT equipment has existed for over two decades, only recently has it become possible to develop clinical systems that are both inexpensive and small enough to be used in operating room, medical offices, emergency rooms, and intensive care. Four technological and application-specific factors have converged to make this possible. First, compact and high-quality flat-panel detector arrays were developed. Second, the computer power necessary for cone-beam image reconstruction has become widely available and is relatively inexpensive. Third, x-ray tubes necessary for cone-beam scanning are orders-of-magnitude less expensive than those required for conventional CT. Fourth, by focusing on head/neck scanning only, one can eliminate the need for sub-second gantry rotation speeds that are needed for cardiac and thoracic imaging. This significantly reduces the complexity and cost of the gantry.

CBCT in Oral and Maxillofacial Imaging

Currently available CBCT units in the United States are the NewTom QR DVT 3G (Quantitative Radiology s.r.l., Verona, Italy), CB MercuRay (Hitachi Medical Corp., Chiba-ken, Japan), i-CAT (Danaher/Imaging Sciences International, Hatfield, PA), Iluma, (Kodak Dental Imaging, Atlanta, GA/Imtec Imaging, Ardmore, OK, USA), Galileos, (Sirona Dental Systems, Charlotte NC), and 3D Accu-i-tomo – XYZ Slice View Tomograph, (J. Morita Mfg.

Corp., Kyoto, Japan), Promax (Planmeca Oy, Helsinki, Finland). All but the two are capable of imaging the skull to include most anthropometric landmarks used in cephalometric analysis (Figure 2)(Table 2).

a.

b.

c.

d.

Figure 2: Examples of current commercially available CBCT units for dento-maxillofacial radiology. a. Newtom 9000G (Quantitative Radiology, Verona, Italy) b. CB MercuRay (Hitachi, Medical Corp., Kashiwa-shi, Chiba-ken, Japan) c. 3D Accuitomo – XYZ Slice View Tomograph, (J. Morita Mfg. Corp., Kyoto, Japan) d. i-CAT (Danaher/Imaging Sciences International, Hatfield, PA)

Table 2. Comparative Specifications of FDA-Approved CBCT Systems (Modified from: [19])

<i>Vendor</i>	<i>AFP-Dent-X</i>	<i>J. Morita Mnfr. Corp.</i>	<i>Imaging Sciences Intl.</i>	<i>Hitachi Medical Systems</i>	<i>Kodak Dental Systems</i>	<i>Sirona Dental Systems</i>
CBCT Name	NewTom 3G	3D Accu-i-tomo	iCAT	CB MercuRay	ILUMA Ultra Cone Beam CT Scanner	Galileos
Headquarters	Elmsford, NY	Kyoto, Japan	Hatfield, PA	Tokyo, Japan	Ardmore, OK	Charlotte, NC
Initial FDA/CDRH Approval	March 2001	May 2003	October 2003	October 2003	November 2005	July 2006
Grayscale	12 Bit	12 Bit	12 Bit	12 Bit	14 Bit	12 Bit-sw 16 bit
Foot Print (H x W x D) (meters)	2 x 2 x 0.74	2.08 x 1.62 x 1.2	1.83 x 1.12 x 1.49	2.25 x 1.96 x 1.9	1.06 x 1.42 x 2.15	2 x 1.60 x 1.60
Image Detector	Image intensifier/CCD	Cesium iodide CsI/amorphous silicon flat panel	Cesium iodide CsI/amorphous silicon flat panel	Image intensifier/CCD	127- micron amorphous silicon flat panel	Proprietary Siemens Technology
Rotation per scan	1	1	1 or 2	1	Single 360° Rotation	210° 200 single shots
Patient Positioning	Supine	Seated	Seated	Seated	Seated with rear-head stabilization	Standing/sitting
Pre-Installed Software	NewTom 3G	i-Dixel	Xoran Cat	CBWorks	ILLUMINAVISION3D	SIDEXIS/GALAXIS
Scan time (s)	5.6-36	17	10-4-	9.6	20-40	14
mA	15 max	1-10	3-5	2-15	4-7	5-7
Kv	110 max	60-80	120	120	120	85
Scan diameter (cm)	25	4-6	17	25	17-19	15
Scan height (cm)	15-30	4-6	6-27.4	15-30	10-19	15
Slice width (mm)	0.1-0.5	0.125-2.0	0.2-0.4	0.1-0.5	0.0936-0.4	Voxel size: 150/300 microns

The cone-beamed technique uses a single scan in which the x-ray source and a reciprocating x-ray detector are attached by a “U-“ or C-arm and rotate around the patient’s head acquiring multiple projection scan images. The field of view (FOV) or area of interest able to be covered is primarily dependent on the detector size (Image intensifier/CCD, CMOS or a:SiTFT field dimensions) and beam projection geometry. While the FOV can be varied by the application of zoomed image reconstruction (e.g. MercuRay [Hitachi, Medical Corp., Kashiwa-shi, Chiba-ken, Japan]) this is usually done at the loss of image resolution.

Data is obtained from a series of multiple single projection scan images as the x-ray source rotates around the patient’s head. The number of projection scans comprising the data set is variable, depending on the system, and is referred to as the frame rate. With a higher the frame rate, more information is available to reconstruct the image: however, the signal-to-noise of individual MPR slices is also higher. The advantage of a higher frame rate is that it reduces metallic artifact, but this is usually accomplished with a longer scan time and therefore radiation dosage. A number of units have variable exposure cycles. For example, the i-CAT has a choice of 10 second, 20 second (standard) and 40 second scans. The number of basis images produced is roughly proportional to the exposure time reflecting a relatively constant frame rate.

CBCT Advantages

Because CBCT provides images of high contrasting structures well, it is extremely useful for evaluating osseous structures. Combined with the limitation of FOV, CBCT is therefore well suited towards the imaging of the craniofacial area. Currently, limitations exist in the application of this technology for soft tissue,[23, 24] but efforts are being directed towards the development of software algorithms to improve signal-to-noise and optimize available contrast.

The utilization of CBCT technology in clinical practice provides a number of

potential advantages compared with conventional CT related to the beam limitation, scan time reduction, and image display. Specifically the advantages of CBCT are as follows [19]:

- 1) **Variable FOV.** Collimation of the CBCT primary x-ray beam enables limitation of the X-radiation to the area of interest. For most (but not all) CBCT systems an optimal FOV (field of view) can be selected for each patient based on suspected disease presentation and the region to be imaged. For example, radiographic investigation of the mandible can be performed by selection of an appropriate FOV. This functionality provides additional dose savings by limiting the irradiation field to fit the FOV, with a resulting exposure reduction to the patient.
- 2) **Sub-millimeter resolution.** Maxillofacial diagnostic CBCT units all use mega-pixel solid state devices for x-ray detection providing a minimal voxel resolution of < 0.25mm isotropically, exceeding the specifications of commonly used multi-slice CT systems in terms of spatial resolution.
- 3) **High speed scanning.** Because CBCT acquires all projection images in a single rotation, scan time can be reduced enormously. In the fan-beam CT system, particularly in high resolution, each thin slice thickness can take up to several tens of seconds. However, various CBCT systems can scan an entire head in 10 seconds or less. While faster scanning times usually mean less number of projections from which to reconstruct the MPR images, motion artifact due to subject movement is reduced. Reconstruction times vary depending on FOV and scanning speed.
- 4) **Dose reduction.** Preliminary reports indicate that CBCT patient absorbed dose can be significantly reduced when compared to conventional CT used with manufacturer recommended sequences.[25] The Newtom 9000 system (Quantitative Radiology, Verona, Italy) also has an automatic exposure control device which selects the starting intensity of the x-ray beam, depending on the size of the patient, and modifies the anodic current according to the density of the transversed tissues (maximum value

15mA). This reduces the patient absorbed dose to approximately that of a film-based periapical survey of the dentition [26-28] or 1-7 times that of a single panoramic image (varying with the panoramic system used).[29, 30] Depending on bone density, a traditional CT exposes the patient to approximately 6-8 times that amount when evaluating either the maxilla or mandible [29] and 15 times the amount of CBCT exposure when imaging both the maxilla and mandible.[31] Table 3 compares radiation exposures from CBCT and other imaging modalities.

Table 3. Radiation Exposures from CBCT and other Imaging Modalities (Modified from: [19])

<i>Machine & Technique</i>	<i>Effective Dose (μSv) using 1990 ICRP</i>	<i>Dose in single panoramic multiples</i>	<i>Dose in days per capita background</i>	<i>Dose in % medical CT equivalent</i>	<i>Dose % annual per capita background</i>
NewTom 3G full (12") FOV	45	7	4	2.1	1.2
NewTom 3G w/ chin tilt & thyroid shield	28	4	3	1.3	0.8
CB MercuRay full (12") FOV 10 mA-100kV	477	74	48	22.7	13.2
CB MercuRay P (9") FOV	289	45	29	13.8	8
CB MercuRay I (6") FOV 125 (maxillary)	169	26	17	12	4.7
CB MercuRay I (6") FOV w/ chin tilt	125	19	12	5.9	3.5
iCAT full (12") FOV	135	21	13	6.4	3.7
iCAT w/ chin tilt & thyroid shield	57	9	6	2.7	1.6
Panoramic (OrthoPhos Plus DS)	6	1	1	0.3	0.3
CT maxilla & mandible	2100	385	243	100	58.3
CT maxilla	1400	164	103	100	38.9
Galileos	Pending	Pending	Pending	Pending	Pending

5) **Voxel isotropy.** The smallest element of a volumetric dataset is the voxel. Voxels have a dimension of thickness as well as the height and width of a 2-dimensional pixel. Voxel representation and therefore resolution are dependent on lateral slice thickness, determined principally by the matrix size of the detector and longitudinal slice thickness (body axis), which in conventional CT is determined by slice pitch, a function of gantry motion. Therefore, conventional CT data is obtained anisotropically, where axial voxel dimensions are equal, but where coronal dimensions are greater and are determined by slice pitch, usually a 1mm minimum (Figure 3a). Therefore, spatial resolution in the longitudinal slice (body axis direction) is poorer than that of lateral slice. On the other hand, the CBCT uses a 2D detector and the same high resolution is obtained in the longitudinal slice (body axis direction) and lateral slice (transverse direction). This voxel representation is known as *isotropic* (Figure 3b). Because of this characteristic, coronal multi-planar reformatting (MPR) of CBCT data has the same resolution as axial data.

a. Anisotropic Voxel

b. Isotropic Voxel

Figure 3: Comparison of voxel acquisition features on conventional “fan beam” CT (a.) and “cone beam” CT (b.)

- 6) ***Real time analysis and manipulation.*** Although conventional CT data is inherently digital, images are supplied to referring clinicians as fixed format, hard copies on film transparencies. CT image algorithms necessary to reformat the data require the computing power of workstations. While such data can be “converted” and imported into proprietary programs for use on personal computers (e.g. Simplant and Simplant CMF: Materialise, Ann Arbor, MI, USA; Procera: Nobel Biopharma, Sweden)) this process is expensive and requires an intermediary stage that potentially extends the diagnostic phase. Reconstruction of CBCT data is performed natively by a personal computer. In addition, availability of software to the user, not just the radiologist, is available either via direct purchase or innovative “per use” license from the various vendors (e.g. Danaher/Imaging Sciences International). Further, because the original data is isotropic, it can theoretically be re-orientated such that the patient’s anatomic features are re-aligned. At least one manufacturer has incorporated this capability into both their acquisition and viewer software (Imaging Sciences International). Finally, the availability of cursor-driven measurement algorithms provides the clinician with an interactive capability for real-time dimensional assessment.
- 7) ***Display modes unique to maxillofacial imaging.*** CBCT software can reconstruct the projection data to provide as many as 512 coronal, sagittal and axial MPR frames. Common to all standard viewing layouts are usually preset options providing display of coronal, sagittal and axial MPR frames. Basic manipulations include zoom or magnification, window/level, the capability to add annotation and measurement algorithms. Some proprietary software is capable of advanced imaging processing functions including:
- a. Oblique MPR such as linear oblique MRP (useful for TMJ assessment) or curved oblique MPR providing a “panoramic” image.
 - b. Cross-sectional imaging provides sequential multi-slice images usually

- perpendicular to the “panoramic” MPR, useful in implant site assessment or lateral oblique MPR which has application in the assessment of the TMJ.
- c. Variable slice thickness adjustments for oblique MPR images provide the clinician with the possibility of producing undistorted plain radiograph projection-like images. One example is the creation of a cephalometric plane projection, either sagittally or coronally. This is developed by increasing the slice thickness of a mid sagittal MPR plane to the width of the head (130-150mm) to produce an image composed of the summed voxels, an image which has been referred to as “Ray Sum”. This image can be exported and analyzed using third party proprietary cephalometric analysis software. This functionality may potentially reduce the need for additional radiographic exposure. Oblique MPR images along the curve of the dental arch with slice thickness comparable to the in-focus image layer of panoramic radiographs (25-35mm) can also be individually created to provide a “panoramic” radiograph customized for each patient. However, unlike conventional panoramic radiographs, these MPR images are undistorted and are free from projection artifacts.
 - d. Maximum intensity projection (MIP). This is a three dimensional volume rendering technique which is used to visualize high-intensity structures within volumetric data. At each pixel, the highest data value encountered along a corresponding viewing ray is depicted. In combination with oblique MPR and selection of wide slice thickness, this technique is capable of providing 3D surface images. This is particularly useful in cephalometric radiography.
 - e. Surface and volume rendering algorithms are available with some software which provides three-dimensional reconstruction and presentation of data that can be interactively adjusted.
 - f. Previously unavailable for viewer use, numerous image enhancement algorithms

are now able to optimize image presentation. While the diagnostic efficacy of the application of these algorithms is yet to be studied, preliminary investigations indicate that sharpening and edge filters show the greatest potential in refining anatomic structures for interpretation.

- 8) ***Variable acquisition modes.*** Many, but not all, units are capable of variable scanning fields of view (FOV) from large FOV capable of imaging the entire craniofacial complex (currently up to 13.2cm with i-CAT and 19cm with CB MercuRay) to limited FOV for specific diagnostic tasks. The Iluma at the time of this research was limited to one full FOV.

CBCT Applications

The advent of CBCT technology has paved the way for the development of relatively small and inexpensive CT scanners dedicated for use in dento-maxillofacial imaging. Manufacturers' web sites provide numerous examples illustrating the value of CBCT in evaluating the position of impacted teeth, supernumerary teeth, maxillary sinus position (in reference to maxillary molars), mandibular canals, and lingual nerves. Maxillofacial applications of CBCT imaging have also been reported for oral and maxillofacial surgery,[34-38] implantology, [39-42] and craniofacial assessment in orthodontics.[43-48] A number of researchers have reported high dimensional accuracy of maxillofacial CBCT in measurement of facial structures.[42,49] Other examples of this modality's uses include surgical assessment of pathology, and preoperative/postoperative assessment of craniofacial fractures.[24,28,33]

Figure 4. Surface-rendering reconstruction of i-CAT CBCT data set (3DVR, Allovision, Geenville, SC) produces interactive volumetric image that can be manipulated to display bony surfaces of maxillofacial complex from various standard orientations.

Applications in Orthodontics

In orthodontics CBCT imaging has current and potential applications in the diagnosis, assessment and analysis of patients with maxillofacial orthodontic and orthopedic anomalies.

In diagnosis, CBCT provides numerous display modalities that can assist the assesment of numerous dental conditions of concern in orthodontics including impacted and supernumerary teeth. The exact position of impacted teeth and their relationships to adjacent roots or other anatomical structures (eg, the mandibular canal) can be comprehended, so that surgical exposure and subsequent movement can be planned. Some of the most significant potential gains from the introduction of CBCT in orthodontics are the ability of integration of information. Instead of

looking at individual diagnostic records—the panoramic radiograph, the cephalogram and its concurrent analysis, the dental models and the patient photographs—a single volume that contains all of this information is now available allowing for a unique appreciation of the inter-correlations between all planes and structures. Image integration, particularly three dimensional imaging, may help to overcome a number of inherent deficiencies in orthodontic treatment planning by providing adequate visualization of anatomical structures.[51, 53] These include assessment of:

- 1) Temporomandibular joint condition prior to treatment particularly if related to condylar trauma and structural development during growth
- 2) Osseous structural conditions in the sagittal, vertical and transverse plane
- 3) Alveolar bone width of available bone for buccolingual movement of teeth (i.e. arch expansion or labial movement of incisors) and evaluation of fenestrations and dehiscence on the buccal and lingual surfaces.
- 4) Tooth inclination and torque: 3D evaluation of the axial inclination of teeth might provide information to supplement that obtained from models.
- 5) Root resorption: Current CT machines could have too low resolution to detect early stages of root resorption due to orthodontic movement, but advances in technology might permit this in the future.
- 6) Soft tissue relationships: Lip length is currently measured on lateral radiographs, but mouth width is not. Three-dimensional data could provide information on the relationship of the corners of the mouth to the underlying dentition. Also, cheek thickness and cheek prominence are soft tissue variables that could be investigated in relation to dental arch width and facial esthetics.
- 7) Tongue size and posture: Volume measurements of the tongue could provide a more objective assessment of size, to aid in the diagnosis of open bites and arch-width discrepancies.

- 8) Airway assessment: Volume measurements of the airway could assess patency, especially in patients who are suspected of mouth breathing, adenoid hypertrophy, or sleep apnea. Nasal morphology and turbinates can be clearly seen in CT scans.
- 9) Patients requiring surgery and those with syndromes and clefts: Surgical planning for such patients can benefit from 3D imaging. 3D data are especially helpful in patients with asymmetry, where true dimensions can be measured, without the problems of magnification or distortion, from which our customary 2D projections suffer. In patients with clefts, bone and soft-tissue defects can be understood much better.

There is an increasing desire in orthodontics to integrate the images of all functional elements, both hard and soft tissue, in the assessment of patients with maxillofacial anomalies. Currently, this is performed using a combination of photographic and radiographic images and study models. Due to the fact that orthodontics involves assessment of hard tissue and soft tissue interactions, such as the effects of tooth movement on esthetics and on functional elements such as occlusion and TMJ, it is highly desirable to have one imaging modality that provides images of all existing elements therefore leading to a better assessment of the interactions present.

Traditionally, conventional cephalometric projections such as the lateral cephalogram, posterior anterior, and submentovertex were used individually or in combination to provide two dimensional representations of structures in three planes of space. There was no single imaging technique readily available to the orthodontist that provided accurate representation of all osseous aspects of the TMJ complex and associated structures until the recent commercialization of CBCT. Hilgers *et al.* studied CBCT multi-planar reformatted projections for TMJ examination to compare the accuracy of linear measurements of the TMJ and related structures with similar measurements made using conventional cephalograms and with the anatomic truth. Using a digital caliper, the investigator measured linear dimensions between 11 anatomic sites to assess the anatomic truth for 25 dry human skulls. All skulls were imaged using i-CAT CBCT and digital cephalograms (PSP) were made in all three orthogonal planes (lateral cephalometric,

posterior anterior, and submentovertex). Linear measurements were made on seven custom CBCT reconstructions and the digital cephalograms. Results showed that all CBCT measurements were accurate; however, three of five lateral cephalometric (LC) measurements, four of five posterior anterior (PA) measurements, and four of six submentovertex (SMV) measurements varied significantly from the truth. Intra-observer CBCT measurements were highly reliable compared to the anatomic truth, and significantly more reliable than measurements made from LC, PA, and SMV images. The authors conclude that custom oblique MPR reconstructions using CBCT provides accurate and reliable linear measurements of mandibular and TMJ dimensions.[54]

Three Dimensional Cephalometrics

Since cephalometric radiology was developed, numerous analyses have been proposed to facilitate communication between practitioners and to describe how individual patients vary from norms derived from other studies. None the less, current cephalometric analyses are two dimensional diagnostic renderings derived from a three dimensional structure. Cephalometric measurements made on 2D radiographs are subject to projection, landmark-identification, and measurement errors.[56-58] The major source of cephalometric error is landmark-identification, which is influenced by many factors such as the quality of the radiographic image, the precision of landmark definition, the reproducibility of the landmark location, the operator, and the registration procedure. Although some cephalometric landmarks are located in the midsagittal plane, many are located at different depth fields leading to increased distortion errors.[56-58] In addition, in lateral cephalometry, it is difficult to determine the difference between right and left sides for superimposition of images, and the sides have different enlargement ratios. It is also difficult to detect deformities in the midfacial area and reading films is difficult due to the superimposition of cranial structures.[59] Despite the potential errors innate to this technique,

cephalometric radiographs are still widely used and, in many cases are essential in the diagnosis and treatment of the patient.

To compensate for the drawbacks of 2D measurements, many techniques have been developed. These techniques include the orientator, [60] the coplanar stereometric system, [61] the multiplane cephalometric analysis, [62] the basilar multiplane cephalometric analysis, [63] and the bilanar cephalometric stereoradiography[64].

Since the mid 1970s, 3D analyses and related procedures in orthodontics have been attempted through several different approaches.[56] There have been three dimensional cephalometrics proposed that use a combination of lateral and frontal cephalograms. These methods rely on the identification of the same point on both radiographs and the implementation of geometry to calculate the point three dimensionally. These approaches, however, are not truly three dimensional and have obvious limitations in that the accuracy depends on a correct correspondence between the landmark locations on the two radiographs, and points not visible on both radiographs cannot be used.[51] Advances in the use of 3D imaging software have permitted important changes in the perception of 3D craniofacial structures.[56] CBCT produces a lower radiation dose than spiral CT and is comparable to conventional radiographs. Because of its volumetric data, CBCT allows secondary reconstructions, such as sagittal, coronal, and para-axial cuts and 3D reconstructions of various craniofacial structures.[34,43,56] Unlike the traditional cephalometric radiograph, the CBCT produces images that are anatomically true (1:1 in size) 3D representations, from which slices can be displayed from any angle in any part of the skull and provided digitally on paper or film. Other reasons for the implementation of 3D cephalometry include:[59]

- 1) actual measurements can be obtained
- 2) a spatial image of the craniofacial structures can be produced
- 3) the 3D image can be rotated easily by changing the rotational axis
- 4) the inner structures can be observed by removing the outer surfaces

5) various anatomical areas can be observed independently by changing the density

According to Hajeer there are numerous benefits of 3D imaging in orthodontics including: pre- and post-orthodontic assessment of dentoskeletal relationships and facial esthetics, auditing orthodontic outcomes with regard to soft and hard tissues, 3D treatment planning and 3D soft and hard tissue prediction. 3D orthodontics also offers efficiency in archiving 3D facial, skeletal and dental records for treatment planning, research and medico-legal purposes.[55] Some authors indicate that three dimensional CBCT images may be useful in the assessment of growth and development.[18,20,35-37,50]

However, many practitioners are accustomed to working with traditional two dimensional cephalograms and may be hesitant to turn to 3D, however, 2D conventional measurements do not have to be abandoned when moving to 3D implementation. Three dimensional data can be rendered as a 2D projection resembling a radiograph allowing traditional analyses to be completed, and customary cephalometric points can also be digitized in 3D on the volumetric rendering itself.[51] Halazonetis believes that the push at implementation of 3D imaging in cephalometrics will lead to an introduction of new landmarks and new analyses which also incorporate advances from related fields, such as geometric morphometrics.[51]

Several CBCT systems permit reconstructions that are comparable with traditional cephalometric projections. Recently, Farman and Scarfe reported a methodology for generating simulated lateral cephalometric images from CBCT using “ray-sum” multiplanar reformatted (MPR) volume reformation.[65] The authors describe a methodology in which existing CBCT image data sets acquired using a 20-second exposure cycle were used to create two dimensional projection images. The three methods of acquisition involved 1) Scout method: exporting the lateral scout radiograph taken initially to confirm the patient’s position, which only provided a lateral cephalogram, 2) Basis image method: selecting the individual lateral and anteroposterior basis images with the least anatomic discrepancies between the right and left sides corresponding to lateral and posteroanterior cephalometric projections and 3) Ray-sum method: manipulation of

the volumetric data set allowed for the development of cephalometric images in all three orthogonal planes. The ray-sum method includes two dimensional cephalometric reconstructions that were developed by increasing the slice thickness of each plane, hence providing an image composed of the summed voxels, or a ray-sum image. The authors indicate that the major difference between the scout or basis image method, or conventional cephalometric images, and the ray-sum method, is that ray sum image projections are orthogonal and have equal magnification between the beam's entrance and exit sides of the patient. The authors were able to produce slices equal to the dimension of the chosen voxel resolution, thus removing anatomic superimposition of landmarks and allowing for more precise definition of bony landmarks. The authors suggest that the use of 3D surface rendering techniques such as volume rendering (Figure 4) and maximum intensity profile algorithms (Figure 5) will redefine orthodontic treatment planning due to the ability to view 3D volumes of the maxillofacial complex from any plane.

Figure 5. Application of maximum intensity projection algorithms to ray-sum projections show relationships of numerous elements (eg, angulation of tooth roots in alveolar bone)

because of their transparent nature. Ray-sum projections provide surface representation of CBCT volumetric data as posteroanterior, submentovertex, and lateral skull images.[65]

Figure 6: Integration of hard- and soft-tissue volumetric data are achieved through surface- and volume-rendering techniques. Visualization of dental occlusion from different perspectives can be achieved via production of surface images of selected maxillofacial structures.[65]

In a recent study, Moshiri *et al.* showed that data from full field scanners can be used to generate simulated cephalometric images.[66] This observational cross-sectional *in vitro* study was conducted to compare the accuracy of linear measurements made on planar images from photostimulable phosphor based cephalograms and two dimensional (2D) simulated lateral

cephalograms derived from full field cone beam computed tomography (CBCT) with direct measurements made on human skulls. The investigator measured the linear dimensions between 15 anatomical landmarks on 23 dentate dry human skulls using a digital caliper to provide nine orthodontic linear measurements (S-N, Ba-N, M-N, ANS-N, ANS-PNS, Pog-Go, Go-M, Po-Or and Go-Co). The skulls were stabilized and imaged with CBCT with a single 360°, 20s, 0.4mm voxel resolution scan. Three 2D simulated cephalometric projections were created: 1) Scout (S), 2) “ray-sum” reconstructed (RS) and 3) basis projection single frame (F) images. Conventional lateral cephalograms (LC) were acquired using a Quint Sectograph and a storage phosphor imaging plate system. TIFF Images were imported into a cephalometric analysis program (Dolphin Imaging Cephalometric and Tracing Software, Chatsworth, CA, USA) and a single observer computed the linear measurements between landmarks and compared them to the anatomic truth. The results showed that the ICC for LC was significantly less than for skull and all CBCT derived modalities. Statistical differences between modalities were found for all measurements except Po-Or ($p=0.27$). For S-N, Ba-N, ANS-PNS and N-M, values for lateral cephalogram measurements were significantly different from actual dry skull dimensions, whereas CBCT values did not differ from the dry skull measurements. All modalities provided significantly different measurements for Pog-Go and Go-M. For ANS-N and Go-Co all CBCT measurements were significantly less than lateral cephalogram measurements. In addition for Go-Co, measurements from scout images were significantly different from actual dimensions. The study concluded that for most measurements in the sagittal plane, simulated 2D lateral cephalometric projections from CBCT are more accurate than lateral cephalogram images. The authors also add that while cephalometric images generated from single CBCT basis projections provide added accuracy in cephalometric analysis, there was no additional advantage in using ray sum images generated from the CBCT volumetric dataset.

Adams *et al.* conducted a study to evaluate and compare traditional 2D cephalometric analysis to a 3-D imaging system with regard to accuracy in recording the anatomical truth as

defined by physical measurements taken using a calibrated caliper.[67] The study used nine dried human skulls to locate thirteen skeletal landmarks both by traditional 2D cephalometry as well as the three dimensional approach. The high average intra-class correlation (0.995), variance (.054 mm²), and standard deviation (SD $\pm$ 0.237 mm) as averaged over 76 measurements derived from precision calipers, using the predetermined 13 skeletal landmarks, established these physical measurements as the gold standard for comparison of the two radiographic methods. The measurements from the 2D model indicated higher variability, with a larger mean standard deviation (6.94 mm) compared with the 3D measures (0.54 mm). The 2D analysis lacked precision as compared with the 3D analysis (points clustered within 0.5 mm). As compared to the gold standard, the ranges between the two systems demonstrated a much larger magnitude of potential error inherent in the 2D system. According to the study, when comparing the actual distance of anatomical distances as measured on a human skull to the measurements derived from a 2D or 3D model, the 3D method is more accurate and precise than the 2D. According to the authors, “Evaluating distances in 3D space with a 2D image grossly exaggerates the true measure and offers a distorted view of craniofacial growth.”

Chidiac *et al.* compared measurements from human skulls and their images from lateral and PA cephalometric radiographs and CT scanograms on thirteen adult skulls. They were unable to reveal any statistically significant differences between mean angular values on cephalometric radiographs and CT views. For sagittal distances, the highest correlation was between the direct measure of condylion-to-pogonion and its radiographic image ($r = 0.73$). Correlations between radiographic and skull transverse measures were higher ($0.46 < r < 0.80$) than the corresponding skull vs. CT measures ($0.06 < r < 0.38$). CT and CR images are 2D slices and projections, respectively, of 3D structures. They found that radiographic images have a distortion (approximately 8%) that brings Co-Pg closer to its anatomic distance, inadvertently contributing to better clinical planning, particularly in orthognathic surgery. The pattern of distortion of PA images was in opposite directions for CR and CT views. They concluded that cephalograms and

CT scanograms are close in depicting angular relations of structures, but they differ in the accuracy of imaging linear measurements, because the location and size of an object within the imaged 3D structure varies with both records.[68]

Most recently, Chan *et al.* compared eight measurements [(sagittal (Sella-Nasion, ANS-PNS), transverse (biorbital, bicoronoidal, and palatal width) and vertical (upper, lower, and posterior facial height)] between 12 commonly used craniometric landmarks made directly on five dry skulls to traditional cephalometry and CBCT (Hitachi CB MercuRay system) using three fields of view (6", 9", and 12"). Intraoperator analysis for skull, CBCT and cephalometric measurements showed good correlation ($r > 0.93$). Both cephalometric and all CBCT measurements showed high correlation ($r > 0.96$) and no statistical significant difference when compared to skull measurements. The average absolute difference between cephalometric and skull measurements was 3.34 ± 4.55 mm. Comparing skull to CBCT measurements, 6", 9", and 12" FOV images showed differences of 0.53 ± 0.46 mm, 0.48 ± 0.44 mm, and 0.46 ± 0.45 mm respectively. They concluded that CBCT measurements showed reliability and more linear measurement accuracy than cephalometry and that CBCT linear measurement accuracy improved as voxel size decreased.[69]

Although recent studies have shown that CBCT derived images are accurate in regard to linear cephalometric measurements,[66,67] the current challenge for clinicians is to understand and interpret 3D imaging, because there is currently no specific way to analyze these 3D images, and interpretation limitations still exist.[56] Lagraverre *et al.* proposed a reference landmark for use in three dimensional cephalometric analysis with 3-dimensional volumetric images.[56] CBCT scans were obtained on 10 patients, all using the same imaging protocol of having the patient lie down with the Frankfort horizontal plane perpendicular to the floor. Images were converted into DICOM format and then rendered into volumetric images using AMIRA software. The investigators used the sagittal, axial, and coronal slices and the 3D image reconstruction for landmark positioning. A point located equidistant to the points in the centers of each foramen

spinosum (ELSA) was established as the reference point ($x=0$, $y=0$, $z=0$ coordinates).

Traditionally used cephalometric landmarks were located on the volumetric images and coordinates of the different landmarks were determined with respect to that reference.

Coordinates of ELSA were registered in a datasheet in the form of x , y , and z dimensions for the 10 subjects measured at three independent times. Present statistical tests do not consider 3D data values, therefore in order to find the intraexaminer reliability, it was necessary to convert all 3D values (x , y , and z) to a sole value using the Delta E formula obtained from the Commission Internationale de l'Eclairage $L^*a^*b^*$ color systems (Vienna, Austria). This system was applied because both use similar Cartesian coordinate systems. The intraexaminer reliability was determined to be $\kappa = 0.998$. Other cephalometric landmarks were then located in different parts of the images where linear and angular measurements could be determined. ELSA as an $x = 0$, $y = 0$, $z = 0$ reference point in 3D images was used because the location of the foramina spinosum was shown to have a low identification error in both the vertical and horizontal planes. The reason in choosing this landmark was twofold: 1) it is a small circle when viewed axially and is easy to locate by using the condyle and the glenoid fossa as guides, and 2) published literature has demonstrated that most of the cranial base growth (>85%) occurs in a child's first 5 years with only minor changes after that age. The authors state that although 3D imaging is a new type of auxiliary examination in orthodontics, no validated method of describing change exists. Most clinicians analyze these images by visually identifying the structures seen without exact measurements or other quantitative analysis. The authors conclude that because ELSA has high intrareliability that it is an adequate reference point for 3D cephalometric analysis.

Although three-dimensional imaging provides volumetric images that can be compared to reality in a 1 to 1 ratio, there is no validated method to describing change with this modality, because most clinicians simply analyze the images with no exact measurements or quantitative analysis.[70] By establishing a precise and reliable instrument for analyzing images produced by 3-D technology, clinicians may have new possibilities for determining changes produced by

certain types of orthodontic treatment. In a subsequent study, Lagravere *et al.* propose certain landmarks and planes to standardize 3D cephalometric image orientation.[70] CBCT scans were obtained on 10 adolescents free from craniofacial anomalies. Images were converted into DICOM format and then rendered into volumetric images using AMIRA software. The investigators used the sagittal, axial, and coronal slices as well as the 3D reconstruction of the images for landmark positioning. To determine orientation planes, the reference point ELSA from the previous study was located, then points located at the superior-lateral border of the external auditory meatus (SLEAM) bilaterally and on the mid-dorsum of foramen magnum (MDFM) were located. Coordinates (in mm) were established for these three points with respect to ELSA and intrareliability values were determined by using the intraclass correlation coefficient for all four points. The axial-horizontal plane (x-y plane) was then determined by using both superior external auditory meatus and ELSA; the sagittal-vertical plane (z-y plane) was formed by ELSA and mid-dorsum foramen magnum perpendicular to the x-y plane. Because all points are located on structures that are not significantly affected by growth after 5 years of age these planes are adequate for standardizing the orientation of 3D images and eliminating the possibility of different results when using other landmarks or structures that might be influenced by growth or treatment. With these planes, the effect of the patient's head position during image acquisition for analysis would be eliminated. The authors conclude that ELSA, rSLEAM, ISLEAM, and MDFM have high intrareliability when locating them with 3D images. The x-y and z-y planes formed by the respective points are an adequate way to standardize the orientation of 3D images.

Conventional 3D CT Imaging Accuracy

The clinical applicability of 3D CT has been evaluated in many studies, and a number of authors have investigated the accuracy of reconstruction software using conventional fan beam derived data sets.[71-73] Recent studies have indicated that there is a high degree of accuracy of

3D reconstructions.[74-76] with differences between measurements and actual dimensions being 2mm to 3mm.[77,78]

The accuracy of craniometric measurements in 3D surface rendering technique has previously been reported,[75] and recently a new 3D CT volume rendering protocol *in vitro* and *in vivo* was established regarding the mental foramen, testing the accuracy and precision of the system.[79] However, there had previously been no report concerning the validation of the soft tissue and the corresponding bone craniometric measurements using specific computer system tools in association with a 3D-CT volume rendering technique. Therefore, Cavalcanti *et al.*[74] investigated the precision and accuracy of anthropometric measurements using 3D conventional (spiral) CT volume rendering by imaging 13 cadaver heads and compared the dimensional accuracy of 10 linear measurements on 2D and 3D reconstructed images performed by two radiologists with those obtained using a spatial digitizer. They used craniofacial measurements including Al–Al (Nasal breadth), G–Op (Skull length), N–Me (Facial height), N–Ns (Nasal height), Po–Al (Camper’s plane), Po–G (Distance between Po and G), Po–Me (Distance between Po and M), Po–N (Distance between Po and N), Po–Ns (Distance between Po and Ns), and Zy–Zy. They found no statistically significant differences between interobserver and intraobserver measurements or between imaging and physical measurements in both 3D-CT protocols. The standard error was found to be between 0.45% and 1.44% for all the measurements in both protocols, indicating a high level of precision. Furthermore, there was no statistically significant difference between imaging and physical measurements ($P \geq 0.01$). The error between the mean actual and mean 3D-based linear measurements was 0.83% for bone and 1.78% for soft tissue measurements, demonstrating high accuracy of both 3D-CT protocols. The authors concluded that the new methodology allowed for a qualitatively high 3D resolution in both bone and soft tissue parameters. They also express that the anthropometric measurements in 3D-CT were considered to be accurate and precise for craniofacial applications.

Recently, Swennen *et al.* developed a new voxel-based 3D cephalometry method.[80]

From a single computed tomography data set, virtual lateral and frontal cephalograms are computed and linked with both hard and soft tissue 3D surface representations, allowing the setup of a precise and reproducible 3D cephalometric reference system[81,82] and reliable and accurate definition of 3D cephalometric hard and soft tissue landmarks[83,84]. Voxel based 3D cephalometry was developed and validated by using spiral multi-slice CT (MS-CT) data.[85] Statistical analysis showed that MS-CT 3D cephalometry is highly accurate and reliable with intraobserver measurement errors as low as 0.88, 0.76, and 0.84 mm for horizontal, vertical, and transverse orthogonal measurements, respectively. Interobserver measurement error was also low: 0.78, 0.86, and 1.26 mm for horizontal, vertical, and transverse orthogonal measurements respectively. Squared correlation coefficients showed high intraobserver and interobserver reliability.[86,82] The authors state that MS-CT cephalometry is a powerful craniofacial measurement tool with several advantages:[80]

- 1) truly volumetric 3D depiction of hard and soft tissues of the skull
- 2) real size (1:1 scale) and real time 3D cephalometric analysis
- 3) no superimposition of anatomic structures
- 4) high accuracy and reliability
- 5) the setup of a biological meaningful 3D cephalometric reference system for cross-sectional and longitudinal analysis of craniofacial changes.
- 6) MS-CT Cephalometry is a major improvement over conventional 2D cephalometry, however, some drawbacks do exist:[80]
- 7) horizontal positioning of the patient during record taking falsifies the position of the soft tissue facial mask
- 8) lack of a detailed occlusion due to artifacts
- 9) limited access for the routine craniofacial patient because of higher cost
- 10) higher radiation exposure than other craniofacial x-ray acquisition systems

Most recently Park *et al.*[59] have described organized, methodological approaches to cephalometric analysis of 3D CT images. Axial images of 30 subjects were taken using CT Hispeed Advantage (GE Medical System, Milwaukee) and reconstructed into 3D models using Vworks 4.0 (Cybermed, Seoul, Korea). Horizontal, midsagittal, coronal, maxillary, mid-maxillary, mandibular, and mid-mandibular planes were all established. 19 Landmarks were first designated on the 3D surface model, and their positions were verified in multiple planar reformat mode, then the Vworks 4.0 and Vsurgery (Cybermed) programs were used to measure the 3D models. The following measurements were determined:

- 1) Zygoma: facial index, midface angle, and Bc point
- 2) Maxilla: canting, rotation, divergence, A-point, and PNS point
- 3) Mandible: canting, rotation, divergence, body length, ramal height, gonial angle, chin prominence, internal ramal inclination, external ramal inclination, lateral ramal inclination, B-point, Pog point, Me point, and mandibular facial width
- 4) Facial convexity (indicates the protrusive state of Bc, A, B, and Pog to the coronal plane)

The results show that cephalometric measurements of the subjects were comparable with the normal Korean averages (t test, $p \leq .01$) and no statistically significant differences were found. All landmarks were reproducible, and there was no significant intraexaminer error between the 2 sessions ($p \geq .01$). The authors do suggest that there are some limitations when using conventional 3D CT as a diagnostic tool. Relatively large errors in the vertical position (z-coordinate) compared with the anteroposterior (y-coordinate) and transverse (x-coordinate) positions were found. The authors state that these errors can be overcome if thin slices are used during the reconstruction. The authors also express that high cost and radiation dose of conventional CT are major disadvantages, and can be improved upon by using cone beam CT, which offers a dose similar to the range of a conventional dental radiographic examination (40 to 50 μ Sv). In addition, in some craniofacial deformities, Orbitale or Porion are deviated, therefore, points in the

horizontal plane should not be used as the reference plane. This limitation can also be overcome by using CBCT, in that CBCT can take an image in the natural head position, and the horizontal reference plane can be parallel to the floor, which is not influenced by Porion and Orbitale. The authors conclude that valuable information can be obtained from a 3D CT reconstruction, and that good treatment results can be obtained with a more precise diagnosis, and the continuous development of 3D analysis will provide more accurate data on a patient.[59]

Potential of CBCT 3D Cephalometry

The application of CBCT technology has allowed the development of a new generation of commercial volumetric dentofacial imaging acquisition systems.[59] CBCT scanners allow image acquisition of a large part of the craniofacial complex with only a 360° rotational sequence, and with dedicated CB reconstruction algorithms a CT data volume is obtained. [86] These scanners focus mainly on bony imaging, leading to a significant decrease in radiation dose. Interesting advantages of CBCT 3D cephalometry for the future include:[80]

1. Reduced radiation exposure
2. Natural shape of the soft tissue facial mask because of the vertical scanning procedure (i-CAT, CB MercuRay)
3. Reduced artifacts at the level of the occlusion
4. Increased access for the routine dentofacial patient because of in-office imaging
(sufficiently compact to be installed in orthodontic and oral surgery outpatient clinics and private practices)
5. Reduced cost

Current limitations of CBCT 3D cephalometry include the scanning volume and positional dependency of the image value of a structure in the field of view of the scanner.[80] The NewTom 3G, i-CAT, and CB MercuRay CBCT scanners all have a scanned volume that is sufficient enough for the setup of the anatomic Cartesian 3D cephalometric reference system and

3D cephalometric hard and soft tissue analyses that do not involve the calvarium or complete ears. However, the 3D Accu-i-tomo and NewTom 9000 systems are not suitable for 3D cephalometry methods due to scanning volumes that are too small.[80] In CBCT systems, the image value of an organ is dependant upon the position in the image volume. Hence, x-ray attenuation of CBCT acquisition systems currently produces different HU values or radiographic densities for similar bony and soft tissue structures in different areas of the scanned volume. An example of this would be that dense bone has a specific image value at the level of menton, but the same bone has a significantly different image value at the level of the cranial base.[80] Vannier states that when new developments in the synthesis and optimization of CBCT reconstruction algorithms allow the full exploitation of the potential of area detectors in CBCT, that CBCT will provide even more important benefits in craniofacial imaging.[44] Therefore it is suggested that improvements in both CB reconstruction algorithms and post-processing will solve or reduce this problem soon.[80]

In conclusion, CBCT derived 3D cephalometry has a number of potential advantages for cephalometric imaging including sub-millimeter resolution, reduced radiation exposure, and inclusion of soft tissue profile. Perhaps the most important clinical advantage is that CBCT volumetric data can be exported as DICOM files and imported into personal computer based software to provide 3D reconstruction of the craniofacial skeleton. This possibility and the increasing access of CBCT imaging in orthodontics is a component of the paradigm that is directing imaging analysis from 2D cephalometry to 3D visualization of craniofacial morphology.[5] The availability of fast scan CBCT now provides an alternate imaging modality capable of providing a 3D representation of the maxillofacial complex with minimal distortion using multi-planar reformatted (MPR) images.

CHAPTER III

METHODS AND MATERIALS

Overview

This investigation was designed as an observational *ex vivo* comparison of linear cephalometric dimensions measured on a sample of 23 skulls to measurements obtained from three modalities 1) digital 2D lateral cephalograms, 2) CBCT derived simulated lateral cephalometric images generated using Dolphin 3D software (Chatsworth, CA) and, 3) Shaded surface 3D reconstructions generated using Dolphin 3D software. This study did not require approval by the Institutional Human Remains Committee, University of Louisville, Louisville KY. The skulls were imaged using CBCT and extraoral lateral cephalometric skull projections. CBCT volumetric data sets were oriented to represent standard head positioning and simulated lateral cephalograms generated using Dolphin 3D software. Twenty cephalometric dimensions were measured on the skulls with a vernier caliper and compared to magnification corrected measurements obtained from displayed images for each of the three modalities. A single rater (PI) performed the measurements on three independent occasions for each modality and the mean measurements and standard deviations compared with ANOVA with *post hoc* analyses.

Sample

Twenty three intact dentate human skulls (n=23) were obtained through the Department of Anatomical Sciences and Neurobiology at the University of Louisville. No demographic data was available on the available human remains so therefore the sample was

not identified by age, gender or ethnic group. The skulls were selected upon the basis of the following criteria: presence of a full permanent dentition and a reproducible, stable occlusion. A stable mandibular relation to the maxilla was achieved for each skull. Careful consideration was taken to assure that proper positioning of the condyle within the glenoid fossa was achieved and reproducible upon imaging and acquisition of measurements. Occlusal interdigitation was also considered in achieving proper mandibular and maxillary alignment. No skulls presented any damage or abnormalities that would interfere with the cephalometric dimensions of interest. All of the skulls were inspected and deemed eligible for providing measurements in the study by two independent observers, Dr. William C. Scarfe and Dr. Mazyar Moshiri.

To provide soft-tissue equivalent attenuation, two latex balloons filled with water were placed in the cranial vault prior to imaging. To separate the mandibular condyle from the temporal fossa, a 1.5 mm thick Styrofoam wedge was placed in the joint space between the glenoid fossa and the condylar head. For all images, the teeth were placed in centric occlusion (maximum intercuspation) and the jaws were held closed by bilateral metal springs. A custom plastic head holder, with a polyvinyl chloride pipe extension for placing into the foramen magnum, was constructed to support the skulls during imaging (Figure 7).

Figure 7. Materials used for imaging of skulls: gloves filled with water, skull holder and foam wedges, and skull.

Imaging

Two modalities were used to image the 23 skull specimens:

1. ***Lateral Cephalometric Radiography.*** Extraoral lateral cephalometric projection radiography was performed using a Quint Sectograph (Model QS 10-1627W; Denar Corporation) using a 10:1 parallel grid. This device maintains a source-to-mid-sagittal distance of 5 ft. Exposure settings were at 78 kVp, 200 mA, and 2/15s. These were determined by subjective evaluation of image quality of multiple images over a range of exposures taken on a skull with a 1cm thick Perspex attenuation material over the exit beam. Skulls were positioned in the cephalostat in standard lateral cephalometric position and stabilizing by two ear pieces inserted into the external auditory meati of the skull and oriented such that the Frankfort plane was parallel to the floor. The skull was oriented such that the sagittal plane was parallel to the X-ray beam with the facial bones of the left side of the skull of the specimen closest to the

imaging detector. The central ray of the beam was directed at the external auditory meati.

Figure 8. Skull positioning for lateral cephalometric radiograph

2. ***Cone beam computed tomography (CBCT).*** CBCT images were acquired of the entire skull using the i-CAT™ CBCT unit (Imaging Sciences International, Hatfield, PA, USA). The device was operated at 1-3mA and 120 kV using a high frequency, constant potential, fixed anode with a nominal focal spot size of 0.5mm. Each skull was positioned into the device supported by the constructed plastic head holder. The hard tissue chin of each skull was inserted into the chin holder and vertical and horizontal laser lights on the device used to position the head. The head was oriented such that the mid-sagittal was perpendicular to the floor and the horizontal laser reference was along an imaginary line at the intersection of the posterior maxillary teeth and alveolar ridge (Figure 9).

Figure 9. Skull positioning for cone beam computed tomography scan

Lateral scout radiographs were then taken and small adjustments made to the skull orientation such that discrepancies between bilateral structures (e.g. posterior and inferior border of the mandibular ramus, zygomatic arch) were less 5mm (Figure 10).

Figure 10. Scout images of a skull demonstrating initial position of specimen with excessive tilt (a) and after adjustment immediately prior to scanning (b)

This positioned the skull for imaging such that the minimal amount of distortion occurred and that analysis with the proprietary software could be facilitated. A single 360⁰ rotation, 20 s. scan, comprising 306 basis projections was then made for each skull with a “Full” field of view (17.0 cm (diameter) x 13.2 cm (height)) collimation using XoranCat acquisition software (Xoran Technologies, Ann Arbor, MI, version 1.7.7). Exposure parameters were controlled by automatic exposure control.

Primary reconstruction of the data was performed immediately after acquisition. Data sets were stored as DICOM and imported into Dolphin 3D Imaging for orientation and construction of PA cephalogram.

Pre-Processing of PSP Images

Extra-oral cephalometric images were acquired using an extra-oral photostimulable storage phosphor 8” x 10” imaging plate and scanned at 300dpi and 16-bit TIFF using the DenOptix Imaging system (Gendex/Kavo, Lake Zurich, IL, USA). The proprietary software used was VixWin 2000 (Version 1.2) digital imaging software (Gendex/Kavo, Lake Zurich, IL, USA). Images were exported from VixWin as lossless 16-bit TIF format without image enhancement. For display and analysis extraoral images were imported into a commercial photographic imaging software (Adobe Photoshop V7.0, 2002; Adobe, San Jose, CA) and images equalized prior to measurement (Figure 11). This was performed to standardize post processing and image display rather than use the proprietary VixWin software equalization algorithm. Equalization redistributes the brightness values of pixels so that they more evenly represented over the entire range of brightness levels. After detecting the brightest and darkest values in the composite image, they are remapped such that the brightest value

represents white and the darkest represents black. Brightness is then equalized by distributing the intermediate pixel values evenly through the gray scale.

Figure 11 Example of effect of post processing using PhotoShop on a lateral cephalometric image (a.) before and (b.) after image equalization.

CBCT images were acquired with a mega pixel (1024 x 1024) flat-panel hydrogenated amorphous silicon detector with cesium iodide scintillator and secondary reconstructed images reformatted from 306 projections for 20 sec scans each providing a pixel matrix size of 0.4mm.

Use of Dolphin Imaging Program for Analysis

The Dolphin Imaging Cephalometric and Tracing Software (Chatsworth, CA) was used exclusively to analyze both the lateral cephalometric digital and CBCT data.

Lateral Cephalometric PSP Analysis

Equalized lateral cephalometric images were imported into the dolphin program. Because of the differences in image dimensions and dpi, calibration of each image was necessary. Therefore a 100-mm radiographic film calibration ruler (Model PN 130-0168, Dolphin Imaging Cephalometric and Tracing Software, Chatsworth, CA USA) was placed in the mid-sagittal plane of the skull perpendicular to the radiographic beam. This image was used to calculate the image-casting magnification factor for PSP images and calibrate distances measured in the cephalometric analysis program. The calculated calibration magnification factors and resulting dpi used within the software program are shown in Table 4.

Table 4. Features of imaging plate and CBCT digital images

<i>Image</i>		<i>Image Dimensions (Width x Height)</i>		<i>Image</i>			
<i>Modality</i>	<i>Type</i>	<i>Matrix Size (Pixels)</i>	<i>Physical Size (inches)</i>	<i>File size (kB)</i>	<i>dpi</i>	<i>Calibration Factor</i>	<i>Final dpi</i>
Projection radiograph	Lateral Cephalometric	2847 x 2386	9.5 x 7.96	6,804	300	9.92%	327

Images were coded and the PI viewed all images on a 20.1-inch flat panel color active matrix TFT (FlexScan L888, Eizo Nanao Technologies Inc., Cypress, CA) screen with a resolution of 1600 x 1200 at 85 Hz and a 0.255 mm dot pitch, operated at 24 bit. A custom analysis within the program was developed that directed the observer to identify specific anatomic landmarks on the images (Table 5). Landmarks were identified by using a cursor-driven pointer. The resultant analysis provided specific linear measurements (Table 6) which were exported as text data. Some anatomic landmarks required identification of bilateral structures that are inherently difficult to view on conventional cephalometric images. For these points the observer was instructed to attempt to differentiate between left and right

features and construct a point midway between them. If the feature could not be differentiated, both right and left condyle landmarks were considered coincident. Although theoretically this would lead to differential magnification of left- and right-sided condylar images compared with the mid-sagittal correction factor, the minimal influence on absolute dimension in relation to the size of the feature and the summation of these distances should average the effect.

CBCT Reconstruction and Analysis

The CBCT data was exported from the XoranCat software in DICOM multi-file format and imported into Dolphin 3D (Pre-release V.2.3, Dolphin Imaging, Chatsworth, CA) on the same computer. All constructions and measurements were performed on a 20.1-inch flat panel color active matrix TFT (FlexScan L888, Eizo Nanao Technologies Inc., Cypress, CA) screen with a resolution of 1600 x 1200 at 85 Hz and a 0.255 mm dot pitch, operated at 24 bit. Two reconstructions were performed of the volumetric dataset:

- 1) Reconstruction of a planar lateral cephalometric projection from CBCT dataset using Dolphin 3D software
- 2) Reconstruction of three-dimensional volumetric renderings from CBCT dataset using Dolphin 3D software

For both reconstructions, first the 3D surface rendering was generated by manually adjusting the threshold of visible pixel levels. This was the method used for segmentation (Fig. 12).

Figure 12 Image capture from Dolphin 3D program demonstrating the segmentation screen. The hard tissue volume segmentation is selected (upper left) and using the segmentation cursor (lower left) the displayed gray level of the voxels is dynamically altered to provide the most realistic appearance of the skull with minimal loss of cortical bone due to thin structures and minimal superimposition of artifacts and soft tissue.

Reconstruction of a planar lateral cephalometric projection

For the reconstruction of the planar cephalometric images, the “Build Radiographs” icon was selected and the Dolphin 3D program constructed 2-dimensional lateral cephalometric renderings from the acquired data set. These images were copied and subsequently imported into a cephalometric analysis program (Dolphin Imaging Cephalometric and Tracing Software, Chatsworth, CA) and images were then analyzed using the same methodology as described above for the two-dimensional digital lateral

cephalometric images. Current versions of Dolphin 3D generate 7 variations of images (Dolphin 1, Dolphin 2, Dolphin 3, Ray-Sum, Emboss, MIP and Trace Filter) which are automatically copied into the Dolphin Imaging system. The pre-release version used in this study did not incorporate these features and therefore only one image was available for export (referred to as “Dolphin 3”). Dolphin 3 is a modified Ray-Sum image with optimized gamma, sharpness, contrast and brightness

Figure 13 Image capture from Dolphin 3D program demonstrating the “Build X-Rays” screen for the lateral cephalometric projections. The hard tissue volume segmented is displayed (upper left) and using the arrow cursor (left) the direction of the reconstruction (either left or right) and volume to be included in the reconstruction (determined by manually changing the position of the left or right white limits or performed semi-automated by selection of the lateral margins) are determined. The resultant image is shown and exported to Dolphin Imaging.

Reconstruction of a 3D Shaded Surface Volumetric Renderings

For the three-dimensional volumetric renderings, after segmentation of the image, the cephalometric landmarks were located and marked on the surface rendered volumetric image. The Dolphin 3D software allowed 3D CBCT measurements from different views using rotation and translation of the rendered image. Landmarks were identified by using a cursor-driven pointer. This was performed by a sequence of pre-set volumetric orientations (Figure 14-17).

Figure 14. PA 3D Volumetric Orientation allowed identification of Na, A point, ANS, B point, Gn, Pog and Or bilaterally located. (5=Nasion ; 6=ANS , 7=A point , 8=B point, 9=Pogonion , 10=Gnathion , 11=left Orbitale , 12=right Orbitale)

Figure 15. SMV 3D Volumetric Orientation allowed identification of Me, PNS and Ba.

(1=Menton, 2=PNS, 3=Basion)

Figure 16. Left and Right Lateral ((Only Right shown) 3D Volumetric Orientation allowed

identification of Co, Po and, Go. (13= Condylion, 14= Porion, 15= Gonion)

Figure 17. Superior 3D Volumetric Orientation allowed identification of cranial fossa and S.

(4=Sella)

Because Sella is a spatial landmark, for the purposes of this study, Sella was assumed to be on the anterior of the clivus, midway between the anterior clinoid processes. As the volume rendering was reoriented, the positions of the previously positioned landmarks were verified and, if necessary, relocated.

Finally measurements between specific landmarks were made. For the pre-release software version used, points and planes were unnamed. Therefore it was necessary to select points to identify a linear plane. This was performed in a specific sequence such that linear measurement 1 corresponded to Na-A, linear measurement 2 corresponded to Na-B and so forth. In this way the resulting analysis provided specific linear measurements (Figure 18) which could be exported as text data. This procedure was repeated three times by the first author.

Figure 18. CBCT 3D image generated from segmentation demonstrating identification of landmarks and linear dimensions measured in the study.

Measurement Analysis

On each skull, fourteen conventional craniometric anatomic landmarks were identified on using an indelible marker (Table 5). Operational definitions of these landmarks were developed considering the vagaries of established definitions and in regard to minimizing location subjectivity.

Table 5. Definition of anthropometric topographic points Used as Landmarks for the

Measurements

<i>Landmark (abbreviation)</i>	<i>Definition</i>
Sella (S)	Midpoint of rim between the anterior clinoid process in the median plane
Nasion (Na)	Mid-sagittal point at the junction of the frontal and nasal bones at the naso-frontal suture
Basion (Ba)	Most inferior point on the anterior margin of foramen magnum, at the base of the clivus
Anterior Nasal Spine (ANS)	Most anterior limit of the of the floor of the nose, at the tip of the anterior nasal spine
Posterior Nasal Spine (PNS)	Point along palate immediately inferior to the pterygomaxillary fossa
A point (A)	Signifies the apical base, the juncture of maxillary basal bone with alveolar bone. The most posterior point between the innermost curvature of the anterior nasal spine and the crest of the labial alveolar plate
B point (B)	Signifies the apical base, the juncture of mandibular basal bone with alveolar bone. The most posterior point between pogonion and the crest of the labial plate
Menton (Me)	Most inferior point along the curvature of the chin in the mid-sagittal plane
Gnathion (Gn)	Most anterior inferior point along the curvature of the chin in the mid-sagittal plane
Pogonion (Pog)	Most anterior point along the curvature of the chin in the mid-sagittal plane
Gonion* (Go)	Point along the angle of the mandible, midway between the lower border of the mandible and posterior ascending ramus
Porion* (Po)	Most superior point of the anatomic external auditory meatus (anatomic Porion)
Condylion* (Co)	Most posterior superior point of the mid-planed contour of the mandibular condyle
Orbitale* (Or)	Most inferior point on infraorbital rim

* Bilateral landmark

Twenty linear measurements characterizing the horizontal and vertical dimensions of the maxillofacial skeleton were then developed from a consideration of the operational definitions and with regard to sagittal assessments provided by conventional extraoral lateral

cephalometric plane projection imaging (Table 6). To establish the true distances between the selected anatomic points, measurements were made by two research associates three times independently using an electronic digital caliper (27-500-90, GAC, Bohemia, NY). The mean of the measurements was designated as the anatomic truth.

Table 6. Definition of Linear Planes used in the Cephalometric Analysis

<i>Abbreviation</i>	<i>Name</i>	<i>Definition</i>
<i>Mid-sagittal Planes</i>		
S-Na	Sella-Nasion	A line connecting Sella and Nasion used to represent the cranial base in the midsagittal plane
Na-Ba	Nasion-Basion	A line connecting Nasion and Basion used to represent the cranial base in the Ricketts Analysis
ANS-PNS	Anterior Nasal Spine- Posterior Nasal Spine	A line connecting ANS and PNS. Used to represent the palatal plane or angle of the maxilla
Na-A	Nasion-A point	A line connecting Nasion and A Point. A sagittal reference line for the maxilla's anterior-posterior position in the Steiner Analysis
Na-B	Nasion-B point	A line connecting Nasion and B Point. Represents the anterior-posterior position of the mandible in the Steiner Analysis
Na-Me	Nasion-Menton	A line connecting Nasion and Menton. Represents total anterior face height.
Na-ANS	Nasion-Anterior Nasal Spine	A line connecting Nasion and ANS. Represents upper facial Height
Ba-ANS	Basion-Anterior Nasal Spine.	A line connecting Basion and Anterior Nasal Spine
<i>Bilateral Planes</i>		
Pog-Go	Pogonion-Gonion	A line connecting Pogonion and Gonion. Represents mandibular body length
Pog-Co	Pogonion-Condylion	A line connecting Pogonion and Condylion. Represents mandibular unit length
Go-Me	Gonion-Menton	A line connecting Gonion and Menton. Represents the mandibular plane
Go-Co	Gonion-Condylion	A line connecting Gonion and Condylion. Represents posterior facial height
Go-Gn	Gonion-Gnathion	A line connecting Gonion and Gnathion. Used to represent the mandibular plane in the Steiner Analysis

Measurements on digital extraoral images were performed by the author on three separate occasions for each skull in the Dolphin Cephalometric and Tracing Software program (Chatsworth, CA).

A custom lateral cephalometric analysis template was designed within the Dolphin imaging program measuring the linear dimensions of interest to this study. Using the custom analysis the PI identified the selected cephalometric landmarks on each lateral cephalometric cephalogram three times and linear distances were calculated by the measurement algorithm implemented by the Dolphin software.

The same steps were repeated for the simulated lateral cephalometric images constructed from the CBCT volumetric data sets; however no equalization was necessary for these images. One set of measurements was taken for each skull at the 20 s scanning interval. The monitor used was a 17". (Proview, California) flat panel TFT color monitor with a screen resolution of 1280 x 1024, a 0.23mm dot pitch set at an image quality of 32-bit.

While the measurement algorithm of the CBCT software is calibrated with respect to voxel dimensions and provides a true 1:1 image ratio, direct measurements from the digital extraoral projections suffer from differential magnification.

Data Collection

All measurements from the Dolphin custom cephalometric analysis were exported with the "data" export function into a text document. The text documents were entered, rearranged and data subsequently exported into a Microsoft Excel 2003 (Microsoft, Redmond, WA, USA) database. Mean $\pm$ standard deviation of three independent replicates of the 20 measurements were calculated for each skull sample and for the three modalities (digital photostimulable storage phosphor lateral cephalometric (PSP), simulated lateral cephalometric generated from the CBCT volumetric dataset (CBCT LC) and, the 3D shaded surface volumetric rendering (CBCT 3D). The data files were then coded for use with the

Statistical Package for the Social Sciences software (SPSS, Version 12.0, SPSS, Inc., Chicago, IL, USA) which was used to conduct the analyses.

Statistical Analyses

Because of the inherent difference in comparing 2D dimensions to 3D measurements, for the purposes of the study, data was analyzed accordingly.

The first analysis (3D Analysis) comprised comparisons of the anatomic measurements from the skulls to those derived from 3D volumetric renderings (Anatomic measurement vs. CBCT 3D). The second analysis (2D Analysis) comprised comparisons of the anatomic measurements from the skulls to those derived from 2D representations (Anatomic measurement vs. CBCT LC vs. PSP)

The Statistical Package for the Social Sciences version 12.0 software (SPSS, Chicago, IL, USA) was used for data analysis. To determine intra-observer reliability, the average measures intraclass correlation coefficient (ICC) was determined for repeat measurements. In addition, absolute and percentage mean error; standard deviations and 95% confidence levels were calculated. Absolute and percentage linear means of triplicate measurements between specified anatomic landmarks were calculated and standard deviations and 95% confidence levels calculated. For the 3D Analysis, the digital measurements of the skulls were taken as ground truth and compared with the PSP lateral cephalograms as well as the 3D CBCT measurements with a two tail paired Student's *t* test at an *à priori* level of significance of $P \leq .05$. For the 2D Analysis, the digital measurements of the skulls were taken as ground truth and bilateral measurements were pooled to provide a mean. This skull dimension was compared with measurements from the PSP lateral cephalograms as well as the simulated lateral cephalometric from CBCT with a one way ANOVA using Scheffé *post hoc* analysis. The *à priori* level of significance for both analyses was $P \leq .05$.

CHAPTER IV

RESULTS AND CONCLUSIONS

Because of the inherent difference in the measurements obtained from the modalities investigated, the analysis of data was performed in two sections.

1. The first analysis (3D Analysis) comprised comparisons of the anatomic measurements from the skulls to those derived from 3D volumetric renderings (Anatomic measurement vs. CBCT 3D).
2. The second analysis (2D Analysis) comprised comparisons of the anatomic measurements from the skulls to those derived from 2D representations (Anatomic measurement vs. CBCT LC vs. PSP)

For each analysis intra-observer reliability and modality accuracy were calculated and compared to anatomic dimensions obtained from the skull.

Analysis of 3D Measurements

Measurement Reliability

Table 7 shows the average measure ICC of triplicate skull measurements made by multiple raters compared to triplicate linear measurements made by the first author for each linear dimension for 3D CBCT. The mean skull ICC (0.996 ± 0.007 ; range; 0.981 to 0.991) was significantly higher than the mean ICC for 3D CBCT (0.976 ± 0.016 ; range; 0.941 to 0.993) ($t=5.468$, $P<.001$).

Table 7. Average measure intra-class correlation coefficient for triplicate measurements for mid-sagittal and bilateral linear measurements by multiple observers on 23 skulls and for a single observer on CBCT 3D reconstructions.

<i>Plane</i>	<i>Location</i>	<i>Modality</i>					
		<i>Skull</i>			<i>CBCT 3D Reconstructions</i>		
		<i>Mean</i>	<i>CL</i>		<i>Mean</i>	<i>CL</i>	
			<i>Lower</i>	<i>Upper</i>		<i>Lower</i>	<i>Upper</i>
S-N	Mid-sagittal	0.998	0.996	0.999	0.941	0.882	0.973
Ba-Na	Mid-sagittal	0.996	0.994	0.998	0.993	0.987	0.997
ANS-PNS	Mid-sagittal	0.994	0.989	0.997	0.954	0.907	0.979
Na-A	Mid-sagittal	0.999	0.997	0.999	0.964	0.928	0.983
Na-B	Mid-sagittal	0.996	0.994	0.998	0.964	0.927	0.983
Na-Me	Mid-sagittal	0.997	0.995	0.999	0.994	0.988	0.997
Na-ANS	Mid-sagittal	0.997	0.995	0.999	0.958	0.917	0.981
Ba-ANS	Mid-sagittal	0.990	0.981	0.996	0.979	0.957	0.990
Pog-Go	Right	0.999	0.999	0.999	0.987	0.974	0.994
	Left	0.999	0.999	0.999	0.990	0.980	0.995
Pog-Co	Right	0.999	0.999	0.999	0.952	0.904	0.978
	Left	0.969	0.998	0.999	0.972	0.944	0.987
Go-Me	Right	0.999	0.999	0.999	0.982	0.964	0.992
	Left	0.999	0.999	0.999	0.987	0.974	0.994
Go-Co	Right	0.999	0.998	0.999	0.990	0.981	0.996
	Left	0.999	0.999	0.999	0.983	0.966	0.992
Go-Gn	Right	0.997	0.993	0.999	0.965	0.931	0.984
	Left	0.995	0.990	0.998	0.986	0.972	0.994
Po-Or	Right	0.996	0.994	0.998	0.980	0.96	0.991
	Left	0.996	0.992	0.998	0.987	0.974	0.994

CL = 95% confidence interval

Table 8 shows the absolute and mean percentage intrarater measurement error for 3D CBCT and skull measurements. Mean percentage measurement error for 3D CBCT ($2.31 \pm 2.11\%$; Range; $1.07 \pm 0.72\%$ to $3.86 \pm 1.85\%$) was significantly higher than repeated

measurements to directly determine anatomic skull dimensions ($0.63 \pm 0.51\%$; Range; $0.29 \pm 0.13\%$ to $1.18 \pm 0.58\%$) ($t = 16.6$, $P < .001$). For all measurements except Ba-Na, 3D CBCT mean absolute and percentage error was significantly higher than direct skull repeated measurements.

Modality Accuracy

Table 9 provides comparison of mean linear measurements obtained from 3D CBCT reconstructions and actual skull dimensions. 3D CBCT measurements were significantly different to actual skull measurements for 13 of the 20 parameters; seven of the eight mid-sagittal measurements (all except Na-A) and six of the 12 bilateral measurements (Pog-Go_{right}, Go-Me, Go-Gn_{left}, and Po-Or). The greatest mean difference of 3.32 mm (3.56%) was found with Ba-ANS. In eight of 20 measurements (40%), an average difference of less than 1 mm, three of which were significantly different from actual measurements was found. In six measurements (30%) a mean difference of between 1 mm and 1.5 mm was found, and for five skulls the CBCT measurements were significantly different from actual measurements made directly on the anatomic specimens. In four measurements (20%) an average difference of between 1.5 mm and 2 mm was found, all of which were significantly different from the assumed anatomic truth. Finally for two measurements (10%) a mean difference greater than 2mm was found, one of which was significantly different from the mean direct skull measurements. In all comparisons, except Na-B and Go-Gn bilaterally, 3D CBCT measurements were less than skull measurements. The mean percentage difference between the mean “actual” and mean 3D-based linear measurements was -1.13% (s.d. $\pm 1.47\%$) and ranged from -0.27 for Na-A to -3.44% for ANS-PNS.

Table 8. Comparison of absolute and percentage mean error for mid-sagittal and bilateral linear measurements between dimensions of planes on 23 skulls and measurements CBCT 3D reconstructions.

Plane	Location	Skull				CBCT 3D Reconstructions					
		Absolute Error (mm)		Percentage Error (%)		Absolute Error		Percentage Error		Significance	
		Mean $\pm$ s.d.	CL	Mean $\pm$ s.d.	CL	Mean $\pm$ s.d.	CL	Mean $\pm$ s.d.	CL	t	p
S-N	MS	0.41 $\pm$ 0.2	0.09	0.7 $\pm$ 0.34	0.15	1.82 $\pm$ 0.86	0.37	3.86 $\pm$ 1.85	0.80	7.61	< 0.001
Ba-Na	MS	0.69 $\pm$ 0.35	0.15	0.7 $\pm$ 0.36	0.15	0.85 $\pm$ 0.60	0.26	1.07 $\pm$ 0.72	0.31	1.1	0.280
ANS-PNS	MS	0.55 $\pm$ 0.25	0.11	1.18 $\pm$ 0.58	0.25	1.30 $\pm$ 1.46	0.63	3.55 $\pm$ 4.00	1.73	2.43	0.019
Na-A	MS	0.33 $\pm$ 0.16	0.07	0.64 $\pm$ 0.31	0.13	1.44 $\pm$ 1.08	0.47	3.52 $\pm$ 2.62	1.13	4.90	< 0.001
Na-B	MS	0.33 $\pm$ 0.18	0.08	0.38 $\pm$ 0.20	0.09	1.41 $\pm$ 2.02	0.87	1.96 $\pm$ 2.76	1.19	2.53	0.015
Na-Me	MS	0.49 $\pm$ 0.64	0.28	0.45 $\pm$ 0.54	0.23	1.05 $\pm$ 0.55	0.24	1.22 $\pm$ 0.65	0.28	3.16	< 0.01
Na-ANS	MS	0.32 $\pm$ 0.17	0.08	0.7 $\pm$ 0.36	0.16	1.16 $\pm$ 0.86	0.37	3.2 $\pm$ 2.45	1.06	4.54	< 0.001
Ba-ANS	MS	1.07 $\pm$ 0.93	0.40	1.16 $\pm$ 0.99	0.43	2.11 $\pm$ 2.12	0.91	2.98 $\pm$ 3.13	1.35	2.15	0.040
Pog-Go	Rt	0.29 $\pm$ 0.10	0.05	0.35 $\pm$ 0.13	0.06	1.24 $\pm$ 1.12	0.48	1.88 $\pm$ 1.77	0.76	4.04	< 0.001
	Lt	0.3 $\pm$ 0.11	0.05	0.36 $\pm$ 0.13	0.06	1.21 $\pm$ 0.80	0.34	1.83 $\pm$ 1.14	0.49	5.40	< 0.001
Pog-Co	Rt	0.33 $\pm$ 0.15	0.06	0.29 $\pm$ 0.13	0.06	2.27 $\pm$ 1.80	0.78	2.49 $\pm$ 2.05	0.89	5.16	< 0.001
	Lt	0.35 $\pm$ 0.24	0.10	0.31 $\pm$ 0.21	0.09	1.61 $\pm$ 1.65	0.72	1.77 $\pm$ 1.91	0.83	3.60	< 0.001
Go-Me	Rt	0.31 $\pm$ 0.15	0.06	0.38 $\pm$ 0.19	0.08	1.33 $\pm$ 0.86	0.37	2.12 $\pm$ 1.31	0.56	5.66	< 0.001
	Lt	0.40 $\pm$ 0.59	0.25	0.51 $\pm$ 0.75	0.33	1.11 $\pm$ 0.90	0.39	1.77 $\pm$ 1.38	0.60	3.17	< 0.01

CL = 95% Confidence Interval

Table 8 (Continued). Comparison of absolute and percentage mean error for mid-sagittal and bilateral linear measurements between dimensions of planes on 23 skulls and measurements CBCT 3D reconstructions.

		Skull				CBCT 3D Reconstructions					
		Absolute Error (mm)		Percentage Error (%)		Absolute Error		Percentage Error		Significance	
Plane	Location	Mean ± s.d.	CL	Mean ± s.d.	CL	Mean ± s.d.	CL	Mean ± s.d.	CL	t	p
Go-Co	Rt	0.37 ± 0.18	0.08	0.65 ± 0.30	0.13	0.97 ± 0.71	0.31	2.13 ± 1.50	0.65	3.91	< 0.001
	Lt	0.30 ± 0.12	0.05	0.54 ± 0.23	0.1	1.12 ± 0.97	0.42	2.45 ± 2.03	0.78	4.04	< 0.001
Go-Gn	Rt	0.69 ± 0.51	0.22	0.88 ± 0.68	0.29	1.56 ± 1.24	0.54	2.44 ± 1.98	0.86	3.08	< 0.01
	Lt	0.71 ± 0.66	0.29	0.88 ± 0.78	0.34	1.43 ± 0.77	0.33	2.22 ± 1.28	0.55	3.36	< 0.01
Po-Or	Right	0.56 ± 0.23	0.10	0.7 ± 0.28	0.12	1.22 ± 0.90	0.39	1.96 ± 1.45	0.63	3.41	< 0.01
	Left	0.58 ± 0.18	0.08	0.74 ± 0.23	0.10	1.10 ± 0.68	0.29	1.80 ± 1.13	0.49	3.54	< 0.001

* MS=Mid-sagittal; Rt=right, Lt=left CL = 95% Confidence Interval

Table 9. Comparison of Absolute and Percentage differences for mid-sagittal and bilateral linear measurements between dimensions of planes on 23 skulls and measurements CBCT 3D reconstructions.

Plane	Location	Skull	CBCT 3D Reconstructions	Difference					
				Absolute		Percentage		Significance	
				Mean	CL	Mean	CL	t	p
S-N	MS	59.31 ± 3.75	58.56 ± 3.67	-0.75	(-1.25 to -0.24)	-1.26	(-2.11 to -0.4)	-3.059	0.006
Ba-Na	MS	98.81 ± 4.99	97.18 ± 4.92	-1.63	(-2.02 to -1.25)	-1.65	(-2.04 to -1.27)	-8.72	<0.001
ANS-PNS	MS	47.70 ± 3.25	46.06 ± 3.29	-1.64	(-2.32 to -0.95)	-3.44	(-4.86 to -1.99)	-4.94	<0.001
Na-A	MS	51.00 ± 3.36	50.86 ± 3.23	-0.14	(-0.63 to 0.34)	-0.27	(-1.24 to 0.67)	-0.61	0.546
Na-B	MS	88.64 ± 5.71	89.75 ± 6.07	1.11	(0.59 to 1.62)	1.25	(0.67 to 1.83)	4.5	<0.001
Na-Me	MS	108.58 ± 7.06	107.77 ± 7.03	-0.81	(-1.26 to -0.36)	-0.75	(-1.16 to -0.33)	-3.77	0.001
Na-ANS	MS	46.41 ± 2.86	45.58 ± 2.85	-0.83	(-1.10 to -0.55)	-1.79	(-2.37 to -1.19)	-6.17	<0.001
Ba-ANS	MS	93.17 ± 6.31	89.85 ± 7.40	-3.32	(-4.96 to -1.68)	-3.56	(-5.32 to -1.8)	-4.19	<0.001
Pog-Go	Rt	83.33 ± 5.83	81.86 ± 5.65	-1.48	(-2.08 to -0.87)	-1.78	(-2.50 to -1.04)	-5.09	<0.001
	Lt	84.20 ± 9.76	81.78 ± 5.71	-2.42	(-5.31 to 0.47)	-2.87	(-6.31 to 0.56)	-1.73	0.097
Pog-Co	Rt	114.77 ± 5.39	114.20 ± 4.84	-0.57	(-1.32 to 0.17)	-0.50	(-1.15 to 0.15)	-1.59	0.127
	Lt	114.63 ± 5.44	113.88 ± 5.02	-0.75	(-1.67 to 0.16)	-0.65	(-1.46 to 0.14)	-1.69	0.104
Go-Me	Rt	79.89 ± 5.59	78.16 ± 5.19	-1.73	(-2.35 to -1.12)	-2.17	(-2.94 to -1.4)	-5.83	<0.001
	Lt	79.31 ± 5.93	78.21 ± 5.29	-1.09	(-1.64 to -0.55)	-1.37	(-2.07 to -0.69)	-4.14	<0.001

CL = 95% Confidence Limit

Table 9 (Continued). Comparison of Absolute and Percentage differences for mid-sagittal and bilateral linear measurements between dimensions of planes on 23 skulls and measurements CBCT 3D reconstructions.

Plane	Location	Skull	CBCT 3D Reconstructions	Difference				Significance	
				Absolute		Percentage			
				Mean	CL	Mean	CL	t	p
Go-Co	Rt	56.38 ± 4.97	56.22 ± 4.71	-0.16	(-0.74 to 0.43)	-0.28	(-1.31 to 0.76)	-0.547	0.590
	Lt	56.54 ± 5.04	56.00 ± 4.21	-0.53	(-1.07 to 0.002)	-0.94	(-1.89 to 0.0)	-2.067	0.051
Go-Gn	Rt	79.85 ± 6.52	81.18 ± 6.02	1.33	(-0.064 to 2.72)	1.67	(-0.08 to 3.41)	1.98	0.061
	Lt	79.78 ± 6.20	81.04 ± 5.92	1.27	(0.38 to 2.16)	1.59	(0.48 to 2.71)	2.95	0.007
Po-Or	Right	78.97 ± 4.79	77.59 ± 4.43	-1.38	(-2.06 to -0.69)	-1.75	(-2.61 to -0.87)	-4.16	<0.001
	Left	78.00 ± 4.82	76.33 ± 4.83	-1.67	(-2.41 to -0.93)	-2.14	(-3.09 to -1.19)	-4.68	<0.001

* MS = Mid-sagittal; Rt = right; Lt = left CL = 95% Confidence Interval

Analysis of 2D Measurements

Measurement Reliability

Table 10 shows the average measure ICC of triplicate skull measurements made by multiple raters compared to triplicate linear measurements made by the first author for each linear dimension for PSP and CBCT LC. The one-way ANOVA identified a significant difference ($F=3.29$; $p=.048$), however *post hoc* analysis was unable to identify differences between PSP (0.973 ± 0.052 ; range; 0.942 to 1.0), skull (0.997 ± 0.003 ; range; 0.996 to 0.998), or CBCT LC (0.999 ± 0.001 ; range; 0.998 to 0.999) ICC.

Table 11 shows the absolute and mean percentage intrarater measurement error for PSP, CBCT LC and skull measurements. One-way ANOVA analysis demonstrated differences between modalities with absolute ($F=13.71$, $p<.001$) and percentage error measurements ($F=12.52$, $p<.001$). Mean absolute measurement error for skull dimensions ($.49 \pm .22\text{mm}$; Range; .362 - .612) were significantly lower than PSP ($.93 \pm .44\text{mm}$; Range; .68 to 1.18) or CBCT LC ($1.19 \pm .39\text{mm}$; Range; .97 to 1.41). Mean percentage measurement error for skull dimensions ($.66 \pm .27\%$; Range; .50% - .82%) were significantly lower than PSP ($1.39 \pm .77\%$; Range; .94% - 1.83%) or CBCT LC ($1.75 \pm .60\%$; Range; 1.4% - 2.09%). For all measurements, except Po-Or, CBCT LC mean absolute and percentage error was significantly higher than direct skull repeated measurements.

Modality Accuracy

Table 12 provides a comparison of mean linear measurements obtained from PSP, CBCT LC and actual skull dimensions. Overall no differences between modalities were found ($F=.098$, $p=.91$). However for all of the bilateral structures, except Go-Co, skull dimensions were significantly less than either PSP or CBCT LC measurements. This difference varied from 4.7% (Po-Or) to 17% (Go-Me). No differences were found between measurements obtained from PSP or CBCT cephalometric images for all bilateral structures.

For all mid-sagittal measurements, except Ba-Na, no differences between modalities were found. For Ba-NA PSP measurements were significantly larger than skull or CBCT LC dimensions by 4.5%.

Table 10. Average measure intra-class correlation coefficient for multiple measurements of linear dimensions by multiple observers on 23 skulls and for a single observer on photostimulable storage phosphor cephalograms (PSP) and simulated cephalograms generated from CBCT volumetric datasets (CBCT LC).

<i>Plane</i>	<i>Modality</i>								
	<i>Skull</i>			<i>Lateral Cephalometric images</i>					
				<i>PSP</i>			<i>CBCT LC</i>		
	<i>Mean</i> <i>CL</i>			<i>Mean</i> <i>CL</i>			<i>Mean</i> <i>CL</i>		
	<i>Mean</i>	<i>Lower</i>	<i>Upper</i>	<i>Mean</i>	<i>Lower</i>	<i>Upper</i>	<i>Mean</i>	<i>Lower</i>	<i>Upper</i>
S-N	0.998	0.996	0.999	0.794	0.588	0.906	0.997	0.995	0.999
Ba-Na	0.996	0.994	0.998	0.994	0.988	0.997	0.999	0.998	0.999
ANS-PNS	0.994	0.990	0.997	0.972	0.944	0.987	0.999	0.999	0.999
Na-A	0.999	0.997	0.999	0.986	0.972	0.994	0.999	0.999	0.999
Na-B	0.996	0.994	0.998	0.997	0.994	0.999	0.999	0.998	0.999
Na-Me	0.997	0.995	0.999	0.999	0.999	0.999	0.999	0.999	0.999
Na-ANS	0.997	0.995	0.999	0.993	0.986	0.997	0.997	0.995	0.999
Ba-ANS	0.990	0.981	0.996	0.998	0.996	0.999	0.998	0.998	0.999
Pog-Go (Lt.)	0.999	0.999	0.999	0.985	0.970	0.993	0.999	0.998	0.999
Pog-Go (Rt.)	0.999	0.999	0.999	---	---	---	---	---	---
Pog-Co (Lt.)	0.999	0.999	0.999	0.987	0.974	0.994	0.999	0.999	0.999
Pog-Co (Rt.)	0.999	0.999	0.999	---	---	---	---	---	---

CL = 95% Confidence Interval

Table 10 (Continued). Average measure intra-class correlation coefficient for multiple measurements of linear dimensions by multiple observers on 23 skulls and for a single observer on photostimulable storage phosphor cephalograms (PSP) and simulated cephalograms generated from CBCT volumetric datasets (CBCT LC).

<i>Plane</i>	<i>Modality</i>								
	<i>Skull</i>			<i>Lateral Cephalometric images</i>					
				<i>PSP</i>			<i>CBCT LC</i>		
	<i>Mean CL</i>			<i>Mean CL</i>			<i>Mean CL</i>		
	<i>Mean</i>	<i>Lower</i>	<i>Upper</i>	<i>Mean</i>	<i>Lower</i>	<i>Upper</i>	<i>Mean</i>	<i>Lower</i>	<i>Upper</i>
Go-Me (Lt.)	0.997	0.995	0.999	0.979	0.959	0.991	0.999	0.999	0.999
Go-Me (Rt.)	0.999	0.999	0.999	---	---	---			---
Go-Co (Lt.)	0.999	0.999	0.999	0.977	0.953	0.989	0.999	0.999	0.999
Go-Co (Rt0.)	0.999	0.999	0.999	---	---	---	---	---	---
Go-Gn (Lt.)	0.995	0.990	0.998	0.982	0.963	0.992	0.999	0.998	0.999
Go-Gn (Rt.)	0.997	0.993	0.999	---	---	---	---	---	---
Po-Or	0.998	0.997	0.999	0.972	0.944	0.987	0.999	0.998	0.999
	0.998	0.996	0.998	---	---	---	---	---	---

CL = 95% Confidence Limit

Table 11. Comparison of absolute and percentage mean error for mid-sagittal and bilateral linear measurements between dimensions of planes on 23 skulls and measurements made on photostimulable storage phosphor cephalograms (PSP) and simulated cephalograms generated from CBCT volumetric datasets (CBCT LC).

<i>Plane</i>	<i>Location*</i>	<i>Skull</i>		<i>PSP</i>		<i>CBCT LC</i>		<i>Significance</i>	
		<i>Absolute</i>	<i>Percentage</i>	<i>Absolute</i>	<i>Percentage</i>	<i>Absolute</i>	<i>Percentage</i>	<i>F</i>	<i>p</i>
S-N	MS	0.42 ± 0.20	0.70 ± 0.34	0.85 ± 2.12	1.45 ± 3.65	1.91 ± 1.86	2.49 ± 3.40	3.16	0.049
Ba-Na	MS	0.69 ± 0.35	0.71 ± 0.36	0.61 ± 0.75	0.60 ± 0.75	1.25 ± 0.72	1.94 ± 1.63	6.95	0.002
ANS-PNS	MS	0.55 ± 0.25	1.18 ± 0.58	0.93 ± 0.65	1.98 ± 1.41	0.89 ± 0.47	1.29 ± 0.82	4.11	0.021
Na-A	MS	0.33 ± 0.16	0.64 ± 0.31	0.64 ± 0.35	1.25 ± 0.66	0.97 ± 0.82	1.60 ± 1.48	8.63	0.000
Na-B	MS	0.34 ± 0.18	0.38 ± 0.20	0.64 ± 0.36	0.71 ± 0.39	1.01 ± 0.57	1.45 ± 0.97	15.77	0.000
Na-Me	MS	0.49 ± 0.64	0.45 ± 0.55	0.35 ± 0.23	0.32 ± 0.20	1.12 ± 0.69	1.5 ± 0.82	12.38	0.000
Na-ANS	MS	0.33 ± 0.17	0.70 ± 0.36	0.41 ± 0.23	0.90 ± 0.50	2.10 ± 1.93	3.43 ± 3.76	18.04	0.000
Ba-ANS	MS	1.07 ± 0.93	1.16 ± 0.99	0.51 ± 0.42	0.54 ± 0.44	1.37 ± 0.78	1.96 ± 1.35	7.91	0.001
Pog-Go	Ave. Lt/Rt	0.30 ± 0.07	0.35 ± 0.08	1.37 ± 1.01	1.97 ± 1.51	1.01 ± 0.88	1.67 ± 1.81	11.52	0.000
Pog-Co	Ave. Lt/Rt	0.34 ± 0.14	0.30 ± 0.12	0.99 ± 0.90	0.93 ± 0.79	1.09 ± 0.74	1.52 ± 0.99	8.42	0.001
Go-Me	Ave. Lt/Rt	0.35 ± 0.29	0.45 ± 0.38	1.40 ± 1.14	2.18 ± 1.88	0.88 ± 0.74	1.17 ± 0.94	9.90	0.000
Go-Co	Ave. Lt/Rt	0.34 ± 0.10	0.60 ± 0.18	1.52 ± 0.10	2.81 ± 1.90	0.71 ± 0.57	1.04 ± 0.93	19.03	0.000
Go-Gn	Ave. Lt/Rt	0.70 ± 0.37	0.88 ± 0.45	1.72 ± 1.14	2.33 ± 1.62	1.18 ± 0.67	1.72 ± 0.94	9.31	0.000
Po-Or	Ave. Lt/Rt	0.57 ± 0.13	0.72 ± 0.15	1.09 ± 0.85	1.44 ± 1.07	1.15 ± 1.48	1.65 ± 2.42	2.41	0.098

* MS=Mid-sagittal; Rt=right, Lt=left

Table 12. Comparison of Mean Linear measurements ($\pm$ s.d.) between cephalometric landmarks on Storage Phosphor and CBCT Derived Simulated Lateral Cephalometric Images of 23 Skulls.

<i>Plane</i>	<i>Location</i>	<i>Skull</i>	<i>Lateral Cephalometric</i>		<i>Significance</i>	
			<i>Storage phosphor</i>	<i>Simulated from CBCT</i>	<i>F</i>	<i>p</i>
S-N	MS	59.31 $\pm$ 3.75	59.76 $\pm$ 2.41	58.60 $\pm$ 3.65	0.71	0.5
Ba-Na ^a	MS	98.81 $\pm$ 4.99	103.30 $\pm$ 5.63	100.33 $\pm$ 5.02	4.4	.016
ANS-PNS	MS	47.70 $\pm$ 3.25	46.87 $\pm$ 2.94	46.98 $\pm$ 2.87	0.5	0.61
Na-A	MS	51.00 $\pm$ 3.36	50.87 $\pm$ 2.78	50.66 $\pm$ 2.59	0.08	0.93
Na-B	MS	88.64 $\pm$ 5.71	89.64 $\pm$ 5.83	88.82 $\pm$ 5.95	0.19	0.83
Na-Me	MS	108.58 $\pm$ 7.06	109.38 $\pm$ 6.98	107.96 $\pm$ 6.64	0.24	0.79
Na-ANS	MS	46.41 $\pm$ 2.86	46.25 $\pm$ 2.58	45.94 $\pm$ 2.49	0.19	0.83
Ba-ANS	MS	93.17 $\pm$ 6.31	96.11 $\pm$ 6.48	94.80 $\pm$ 5.98	1.23	0.29
Pog-Go ^b	Bilat	83.76 $\pm$ 7.46	70.83 $\pm$ 5.77	71.44 $\pm$ 6.48	28.08	<0.001
Pog-Co ^b	Bilat	114.7 $\pm$ 5.35	105.40 $\pm$ 5.25	105.76 $\pm$ 5.92	21.02	<0.001
Go-Me ^b	Bilat	79.6 $\pm$ 5.72	65.46 $\pm$ 5.27	65.16 $\pm$ 6.06	48.39	<0.001
Go-Co	Bilat	56.46 $\pm$ 4.93	55.06 $\pm$ 5.02	55.72 $\pm$ 4.68	0.47	0.63
Go-Gn ^b	Bilat	79.81 $\pm$ 6.24	74.37 $\pm$ 5.92	75.12 $\pm$ 6.65	5.09	0.009
Po-Or ^b	Bilat	78.49 $\pm$ 4.73	75.02 $\pm$ 3.71	74.31 $\pm$ 4.04	6.57	0.003

^a Significant difference between Truth and the PSP lateral cephalometric ($p=.018$). ^b Significant difference between truth and lateral cephalometrics from both methods (both $p<.0001$)

CHAPTER V

DISCUSSION

No single imaging technique has been readily available that provides an accurate, easily interpreted representation of all osseous aspects of the craniofacial region in one exposure. Previously, a plain lateral cephalogram in combination with a panoramic radiograph and occasionally a posterior anterior or submentovertex projection has been used for standard imaging analysis in orthodontics. The introduction of maxillofacial cone beam imaging equipment provides clinicians with an opportunity to generate 2D simulated projections and 3D volumetric renderings easily using relatively inexpensive third party personal computer based software. The rapidly emerging availability of this technology will undoubtedly expand the use and application of 3D imaging, particularly in the field of orthodontics. The aim of this study was to compare linear dimensions between common cephalometric landmarks on a sample of skulls to both 3D measurements and 2D measurements obtained from reconstructed CBCT images and planar PSP lateral cephalometric radiographs.

For 2D analysis we found that for most mid-sagittal measurements, simulated lateral cephalometric images produced from reconstructed CBCT volumetric dataset were similar to those from PSP images. In addition we found these measurements to be accurate, except for Ba-Na from PSP, which overestimated this dimension. For bilateral measurements, simulated CBCT LC measurements were similar to those obtained from PSP however, as expected because of projection geometry, both modalities were less accurate, underestimating dimensions by between 4.7% to 17%. Our results compare well to those of Chan *et al.*[69] who also found no statistically

significant difference between skull, traditional frontal cephalometric and CBCT measurements. Our results also compare favorably with previous work done by Chidiac *et al.*[68] who were unable to reveal any statistically significant differences between mean angular values on cephalometric radiographs and CT views.

For 3D measurements we found that for two-thirds of dimensions, CBCT measurements were statistically significantly different from actual measurements. However analyzing the absolute and percentage differences, this statistical significance probably does not translate into clinical relevance. Statistical differences most likely resulted from small standard deviations within the measurements. In addition, the greater intraobserver variability demonstrated by the 3D CBCT measurements may have also contributed. These results are very similar in magnitude to those of Cavalcanti *et al.* [74] However; they reported no statistically significant difference between imaging and physical measurements. Their mean difference between actual and 3D-based linear measurements was 0.83% as compared with $-1.13 \pm 1.47\%$ in the present study. In the present study, 40% of measurements had an average difference of less than one millimeter, 70% had an average difference of less than 1.5 mm and 90% had an average difference of less than 2 mm. These absolute differences compare favorably with those reported by additional authors.[48, 49]

For two dimensional analyses, the only sagittal measurement where lateral cephalograms were inaccurate was Na-Ba. This inaccuracy is most likely due to the intrinsic difficulty in locating Basion on a lateral cephalogram. Geelen *et al.* [87] compared the reproducibility of cephalometric landmarks on conventional films and PSP images displayed both as hard copy and on computer screen and found Basion to have the third highest envelope of identification intrarater error (after Porion and Orbitale) with a mean deviation of $1.28 \pm 1.06\text{mm}$, $1.46 \pm 1.54\text{mm}$ and $1.99 \pm 1.67\text{mm}$ respectively. The anatomic location of Basion, which is situated between muscle and parenchyma, is considered to be an area of the image of high anatomic,

structural noise, where there is a low signal to noise ratio. The gray areas of the image are resultantly blurred therefore making anatomic landmark identification more difficult.

We anticipated differences in cephalometric dimensions obtained from bilateral planes compared to skull measurements. This is because of the concept of linear projective transformation.[88] This concept explains that due to the position and orientation of equidistant lines in 3D space, that these lines are projected onto the film plane as different lengths due to inherent geometric projection errors. So any measurement made on the film is not correct when compared to the "Anatomic Truth."

For two dimensional measurements, mean absolute and mean percentage intrarater measurement error for skull dimensions ($.49 \pm .22\text{mm}$) were significantly lower than PSP ($.93 \pm .44\text{mm}$) or CBCT LC ($1.19 \pm .39\text{mm}$). Mean percentage measurement error for skull dimensions ($.66 \pm .27\%$) were significantly lower than PSP ($1.39 \pm .77\%$) or CBCT LC ($1.75 \pm .60\%$). This difference may be due to the fact that all skull landmarks were physically located with an indelible marker, serving as a fiducial reference point for physical measurements. The digitization of landmarks on PSP and CBCT lateral cephalograms was performed three times independently by the PI and therefore it was necessary to re-locate landmarks and re-measure distances with each tracing. The lack of a digital indelible landmark position for PSP and CBCT lateral cephalometric radiographs contributed to the overall increased mean absolute measurement error for these two modalities. This result is also unexpected because of the inherent resolution differences between PSP images and CBCT derived images. PSP cephalometric images have an inherent spatial resolution of 300 dpi whereas CBCT images have a 0.4mm voxel size (63.5 dpi). However Held *et al.* (2001) found that the ability to identify landmarks did not differ significantly between cephalometric images scanned at resolutions ranging from 600 to 75 dpi.

There are numerous factors which should be considered when applying the results of this investigation to clinical situations. First and foremost this study was performed on human skulls.

The accuracy of measurement distances between three dimensional landmarks on actual patients may be affected by a reduction in image quality due to soft-tissue attenuation, metallic artifacts and patient motion. Variation in scanning protocol such as voxel size and number of basis projection images may also influence measurement accuracy. Therefore it would be expected that the dimensional accuracy of all measurements would be somewhat less on patient derived data.

There are also some potential limitations when using 3D images derived from CBCT data. Three dimensional volumetric depictions depend on appropriate segmentation – the thresholding of bone pixel values and suppression of surrounding tissue values to enhance the structure of interest. This process is dependent on the software algorithm, the spatial and contrast resolution of the scan, the thickness and degree of calcification or cortication of the bony structure and the technical skill of the operator. In this study, the Dolphin 3D software provides a semi-manual method of segmentation, dependent on the interaction of the operator with the data to produce a visually acceptable 3D rendering. These limitations result in deficiencies or voids in the surface of the image. These occur in regions that are represented by few voxels or have gray values still representing bone, but outside the threshold. These areas include the posterior and anterior superior walls of the maxillary sinus, bone overlying the dentition and cortical bone of the mandibular condyle. Consequently this may lead to greater landmark identification error and subsequent measurement error. Anatomic landmarks whose accuracy may be affected by poor segmentation include A point, ANS, PNS, Porion, and Condylion.

In addition, the method of establishing dimensional truth could have potentially contributed to bias in the results. While the landmark identification and measurements on the 3D rendered images were repeated three times by a single observer, the landmark identification on the skulls was performed only once and measurements performed three times by two observers independently. This reduced the error of point identification on the skulls; however, the establishment of a consensus landmark location was necessary to provide a fiducial reference to

which we could assess the inherent clinical inaccuracies of both landmark identification and measurement associated with the 3D image rendering.

We were unable to completely simulate soft tissue effects of attenuation on image quality in this study as this would have been problematic in positioning and orientation of the skulls. While the use of water balloons placed within the cranial cavity provided some degree of soft tissue attenuation, the lack of peripheral attenuation material may have allowed easier identification of landmarks on 3D surface rendered images.

While the results of this investigation indicate that it is possible to generate reliable and accurate simulated lateral cephalometric images from CBCT volumetric data sets, before this technique is recommended it will be necessary to compare the relative diagnostic yield from existing imaging series. In addition it is necessary to compare the measurement accuracy of 2D images obtained from CB volumes using dose reduction techniques such as decreasing the number of basis images. It has been reported effective dose $E_{(ICRP1990)}$ and $E_{(ICRP2005 \text{ draft})}$ for the i-CAT is 134.8 μ Sv and 193.4 μ Sv respectively for a 12'' FOV scan and a 20 sec. scan. While it is hoped that a CBCT examination would never be substituted for a panoramic examination alone, in the case of orthodontic diagnosis, performance of a CBCT scan to generate a panoramic, lateral and posterior–anterior (PA) cephalometric images may be considered.[89] Dose calculations using ICRP 1990 tissue weights are 6.2 μ Sv for a direct digital panoramic image[2] and 3.4 μ Sv for 2 cephalometric images.[90] If the calculation of E is modified to include salivary glands then the panoramic and cephalometric doses increase to 22 μ Sv and 6.8 μ Sv, respectively. Therefore a conventional digital orthodontic extraoral series (panoramic, lateral cephalometric and PA) would provide a dose of between 9.61 μ Sv [$E_{(ICRP1990)}$] and 28.8 μ Sv [$E_{(ICRP2005 \text{ draft})}$]. Comparing this radiation detriment from conventional digital to CBCT derived simulated images using the standard number of projections (20 sec., 306 projections), CBCT imaging is between 6 [$E_{(ICRP2005 \text{ draft})}$] and 14 [$E_{(ICRP1990)}$] times these conventional exposures.

CHAPTER VI

SUMMARY AND CONCLUSIONS

The purpose of this project was twofold: 1) to compare the accuracy and reliability of linear measurements on lateral cephalograms made conventionally using photostimulable storage phosphor imaging plates (PSP) and simulated lateral cephalometric images from ray sum reconstructions of cone beam computed tomographic (CBCT) volumetric datasets of the same anatomic specimens and 2) to compare the accuracy and reliability of measurements of 3D shaded surface volumetric reconstructions.

While the ICC was high for both 2D and 3D modalities, the mean absolute and mean percentage intrarater measurement error for skull dimensions were significantly lower than PSP cephalograms, CBCT cephalograms and 3D CBCT images. Measurements on I-CAT CBCT simulated lateral images using Dolphin Software were as reliable and as accurate as PSP cephalometric measurements. We found that for most mid-sagittal measurements, simulated lateral cephalometric images produced from reconstructed CBCT volumetric dataset were similar to those from PSP images. In addition we found that these measurements were accurate, except for Ba-Na from PSP, which overestimated this dimension. For bilateral measurements, simulated CBCT LC measurements were similar to those obtained from PSP however, as expected by considerations of projective geometry, underestimated dimensions by between 4.7% to 17%.

While many linear measurements between cephalometric landmarks on 3D volumetric surface renderings obtained using Dolphin 3D software generated from i-CAT CBCT datasets may be statistically significantly different from anatomic dimensions, most can be considered to be sufficiently clinically accurate ($-1.13 \pm 1.47\%$) for craniofacial analyses.

While CBCT imaging provides reliable and accurate 2D simulation images and accurate and reliable 3D rendering, dose considerations with the adoption of any technique involving ionizing radiation must be considered. Conventional digital orthodontic extraoral series (panoramic, lateral cephalometric and PA) provide a dose of between $9.6\mu\text{Sv}$ [$E_{(\text{ICRP1990})}$] and $28.8\mu\text{Sv}$ [$E_{(\text{ICRP2005 draft})}$]. Comparing this to the theoretical dose from CBCT i-CAT imaging using the standard number of basis projection images (306) ($E_{(\text{ICRP1990})}$ and $E_{(\text{ICRP2005 draft})}$ of $134\mu\text{Sv}$ and $194\mu\text{Sv}$ respectively), CBCT imaging still provides between 7 [$E_{(\text{ICRP2005 draft})}$] and 14 [$E_{(\text{ICRP1990})}$] times more radiation detriment.

While CBCT imaging provides clinically reliable and accurate 3D volume rendered images and 2D simulation images, dose considerations demand that evidence-based selection criteria should be developed for CBCT in orthodontics that take into account the ALARA principle [65].

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CURRICULUM VITAE

Danielle R. Periago

I. Academic Degrees

Orthodontic Certificate	University of Louisville	Louisville, KY	In Progress
MS	University of Louisville	Louisville, KY	In Progress
DMD	University of Louisville	Louisville, KY	2005
BA	Oral Roberts University	Tulsa, OK	2000

II. Awards and Honors

- Summa Cum Laude Graduate: University of Louisville School of Dentistry, 2005
- American Academy of Oral Medicine Award: University of Louisville School of Dentistry, 2005
- American Academy of Oral and Maxillofacial Pathology Award: University of Louisville School of Dentistry, 2005
- Dean's Scholar: University of Louisville School of Dentistry; Fall 2005, Spring 2004, Fall 2003, Spring 2003, Fall 2002, Spring 2002, Fall 2001
- Excellence in General Pathology Award: University of Louisville School of Dentistry, 2003
- Delphi Society: University of Louisville School of Dentistry, 2003
- National Boards Part I Scholarship Winner: University of Louisville School of Dentistry, 2003
- Summa Cum Laude Graduate: Oral Roberts University, 2000
- National Spanish Honor Society: Oral Roberts University, 2000
- Award for Best Senior Thesis Defense: Modern Language Department, Oral Roberts University, 2000
- President's List (4.0 GPA): Oral Roberts University, 1996-2000 (7 semesters)
- Dean's List (3.9 GPA): Oral Roberts University, 1998 (1 semester)
- National Freshman Honor Society: Oral Roberts University, 1996-2000
- Student Association Freshman Council: Oral Roberts University, 1996
- Paul Ogil Foundation Scholarship Winner: 1996

IV. Professional Affiliations

- American Association of Orthodontists (AAO): student member, 2005-present
- American Dental Association (ADA): member, 2005-present
- American Student Dental Association (ASDA): member, 2001-2005
- Louisville American Student Dental Association (LASDA):
 - Member, 2001-present
 - Class Representative, 2002-2003
 - Pre-dental Liaison, 2003-2004
- Delphi Society: member, 2003-present
- National Spanish Honor Society: member, 2000
- Oral Roberts University Student Association:
 - Academic Peer Advisor, 1999-2000
 - Dorm Representative, 1996-1997
 - Freshman Council, 1996
- Tri-Beta Biological Honor Society: member, 1996-1998

V. Extracurricular Activities/Community Service

- Fluent in the Spanish language
- Academic Tutor: University of Louisville, 2003-2004
 - General Pathology
 - National Boards Part I
- Smile Kentucky Program 2003-2004
- Oral Cancer Screening: Highland Community Outreach, 2004
- Kentucky State Fair Sealants Program, 2002-present
- March of Dimes
- University Community Outreach program: Oral Roberts University, 1996-2000
- Inner City Los Angeles Mission Trip: Oral Roberts University, 1998
- Bangkok, Thailand Mission Trip: Teen Mania Missions, 1995.