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RESEARCH ARTICLE 3D dento-maxillary osteolytic lesion and active contour segmentation pilot study in CBCT: semi-automatic vs manual methods

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Objectives: This study was designed to evaluate the reliability of a semi-automatic segmentation tool for dento-maxillary osteolytic image analysis compared with manually defined segmentation in CBCT scans.

Methods: Five CBCT scans were selected from patients for whom periapical radiolucency images were available. All images were obtained using a ProMax® 3D Mid Planmeca (Planmeca Oy, Helsinki, Finland) and were acquired with $200-\mu m$ voxel size. Two clinicians performed the manual segmentations. Four operators applied three different semi-automatic procedures. The volumes of the lesions were measured. An analysis of dispersion was made for each procedure and each case. An ANOVA was used to evaluate the operator effect. Non-paired *t*-tests were used to compare semi-automatic procedures with the manual procedure. Statistical significance was set at $\alpha = 0.01$. Results: The coefficients of variation for the manual procedure were 2.5–3.5% on average. There was no statistical difference between the two operators. The results of manual procedures can be used as a reference. For the semi-automatic procedures, the dispersion around the mean can be elevated depending on the operator and case. ANOVA revealed significant differences between the operators for the three techniques according to cases.

Conclusions: Region-based segmentation was only comparable with the manual procedure for delineating a circumscribed osteolytic dento-maxillary lesion. The semi-automatic segmentations tested are interesting but are limited to complex surface structures. A methodology that combines the strengths of both methods could be of interest and should be tested. The improvement in the image analysis that is possible through the segmentation procedure and CBCT image quality could be of value.

Dentomaxillofacial Radiology (2015) 44, 20150079. [doi: 10.1259/dmfr.20150079](http://dx.doi.org/10.1259/dmfr.20150079)

Cite this article as: Vallaeys K, Kacem A, Legoux H, Le Tenier M, Hamitouche C, Arbab-Chirani R. 3D dento-maxillary osteolytic lesion and active contour segmentation pilot study in CBCT: semi-automatic vs manual methods. Dentomaxillofac Radiol 2015; 44: 20150079.

Keywords: CBCT; dento-maxillary osteolytic image; jaw cysts; 3D dento-maxillary imaging; image segmentation

Introduction

The use of CBCT is increasing in dento-maxillofacial practice. CBCT can be beneficial in dento-maxillofacial imaging a[nd](#page-7-0) can be considered as the present reference technique. $1-3$ CBCT has a high resolution and causes lower radiation exposure compared with multislice CT (MSCT). CBCT enables the clinician to obtain linear

Correspondence to: Dr Karen Vallaeys. E-mail: karen.vallaeys@univ-brest.fr The present study has contributed to testing some semi-automatic segmentation protocols on dento-maxillary imaging. To our knowledge, no studies have yet been carried out on CBCT and odontogenic cysts. Region-based active contour segmentation was comparable with the manual procedure for delineating circumscribed osteolytic dento-maxillary lesions. However, region-based and boundary-based segmentation are not suitable for all cases. These approaches are limited when segmenting objects with complex surface structures. A hybrid methodology, combining the strength of both manual and semi-automatic protocols, needs to be developed in order to provide a suitable method for the majority of cases, regardless of clinical situation, extent and position of the lesion. The improvement of the image analysis must be considered through the segmentation procedure and CBCT image quality. This work was partly supported by the French ANR within the Investissements d'Avenir programme (LabEx CAMI) under reference ANR-11-LABX-004. Received 7 March 2015; revised 15 May 2015; accepted 20 May 2015

measurements of anatomic structures and lesions in the maxillary and mandibular bones on three planes of spac[e an](#page-7-0)d is considered to be accurate in vivo and in $v_{itro.}^{4–6}$ It can give accurate volumetric and linear measurements that are of interest for monitoring treatment outcomes and patient follow-up during the healing phase. Several studies have been published confirming the high accuracy of CBCT for diagnosing periapical lesions and radiolucency.⁴

In the field of dento-maxillary osteolytic images, radicular cysts are the main diagnosis for jaw cysts and cyst-like lesions of the mandible, accounting for $>50\%$ of cases.[8](#page-7-0) Most non-healing lesions submitted for biopsy are classified as granulomas or cysts (73% of cases).[9](#page-7-0) Radicular cysts are inflammatory disorders of the periradicular tissues, caused by persistent microbial infection of the root canal system of the affected teeth. The prevalence of periapical radiolucency is very high, broadly equivalent to one radiolucency per patient.^{[10](#page-7-0)} These pathoses cause bone resorption and may generate extensive tissue destruction that requires surgical intervention. Maxillary approaches require surgeons to drill the bone to expose target lesions while avoiding the critical structures within it, such as the neurovascular bundle, maxillary sinus or nasal fossa. Indeed, a misguided action may create damage causing permanent injury, paraesthesia etc. An understanding and an evaluation of the anatomical specificities of individual patients are important to develop a case-specific risk assessment. In complex cases when surgery is planned, a planning and guidance support

Figure 1 Example of manual segmentation result. A, anterior; I, inferior; L, left; P, posterior; R, right; S, superior.

Table 2 Operator effect, ANOVA $\alpha = 0.01$

Case	MAN	EB	RC	<i>EBDA</i>
	0.189	0.0119	0.126	$\leq 0.0001*$
	0.373	$< 0.0001*$	0.771	$0.0007*$
3	0.668	0.063	$\leq 0.0001*$	0.0226
$\overline{4}$	0.772	$\leq 0.0001*$	$\leq 0.0001*$	$0.0031*$
	0.028	0.599	0.335	$0.0094*$

EB, semi-automatic procedure with edge-based contour evolution; EBDA, semi-automatic procedure with edge-based contour evolution by adjunction of an anisotropic diffusion filter; MAN, manual procedure; RC, semi-automatic procedure with region competition contour evolution.

* Statistically significant difference.

system would be beneficial. Three-dimensional (3D) representation of pathological structures must be very accurate to be suitable for virtual planning, highlighting its relation with the surrounding anatomical structures, especially with the critical areas of the inferior alveolar dental nerve in the mandible, the maxillary sinus and nasal cavi-ties.^{[11](#page-7-0)} Some studies have proposed and validated a method for estimating preoperative and postoperative bone volume using CBCT imaging in vitro.^{[12](#page-7-0)} Presently, there is no standardized method clinically usable in vivo.

The first step of preparation for surgical planning and bone volume evaluation is tissue segmentation. This needs to be performed as quickly as possible with as few user interactions as possible and an accurate result. Voxels are gathered according to certain criteria that make it possible to isolate images within the structures that need to be analysed. Accurate segmentation of the CBCT image is an essential step for generating 3D models for the diagnosis and treatment planning of patients with voluminous osteolytic disease. This is a critical step because it affects the quality of subsequent measurement and analyses. Inaccurate information may lead to mistakes of measurement and treatment simulation. The difficulty of segmentation lies in the variation of the target structure, which differs according to the clinical case. Several segmentation techniques exist: manual, semi-automatic and fully automatic.

The most straightforward approach to segmentation is the manual method, where the user outlines the structures slice by slice. By defining a region of interest (ROI) in each slice, a volume of interest can be composed after combining all successive ROIs. Manual segmentation is often used as a reference compared with all automated segmentation procedures, but manual segmentation is very time consuming and user dependent.¹

In complex images, automatic methods lead to inaccuracies that often require the operator to correct the

results, 13 which is one reason why semi-automatic methods are of interest. They combine the efficiency and repeatability of automatic segmentation with the sound judgment that can only come from human expertise.[14](#page-8-0) Many semi-automatic segmentation protocols have been described in the literature. Among these, two active contour segmentation methods have become well-established theoretical approaches: Geodesic Ac-tive Contours and Region Competition.^{[15,16](#page-8-0)} The human expert must specify the initial contour, balance the various forces that act upon it and monitor the evolution of the segmentation. Semi-automatic segmentation methods greatly assist the clinician in this task.

ITK-SNAP (Penn Image Computing and Science Laboratory, Philadelphia, PA), an open source medical image processing application, implements these two methods. This software application provides a combination of manual and semi-automatic tools for extracting structures in 3D image data of different modalities and from different anatomical regions.¹⁴

The purpose of this study was to compare semiautomatic segmentation methods for CBCT scans with manual segmentation for dento-maxillary osteolytic 3D image reconstruction and analyses.

Methods and materials

Study sample

For this study, a total of five CBCT data sets of patients, presenting a periapical radiolucent image with a CBCT periapical index score from 4 to 5 ED (expansion and destruction of periapical bone; see [Table 1](#page-1-0)), were selected. This index was developed on the basis of criteria established from measurements corresponding to periapical radiolucency interpreted on CBCT scans $(Table 1)$ $(Table 1)$. All the images have been anonymized.

CBCT imaging protocol, data acquisition and data export

All the CBCT image data sets were obtained using ProMax® 3D Mid Planmeca X-ray source equipment (Planmeca OY, Helsinki, Finland). The data sets were obtained for diagnosis and before treatment of the patients. All data sets were acquired with a 200 - μ m isotropic voxel size, *i.e.* with a $0.2 \times 0.2 \times 0.2$ mm spatial resolution, two fields of view (7×5 or 4×5 cm), a tube (anode) voltage of 90 kV and a tube (anode) current of 11–13 mA. The image detector was a flat panel.

Table 3 Manual procedure, statistical data

Case												
<i>Operator</i>												
Mean volume $(mm3)$	428.95	437.68	1651.90	1620.04	871.00	877.79	406.40	408.94	116.49	135.33		
Median volume (mm ³)	427.78	438.66	1630.17	1619.36	881.79	879.15	404.57	405.16	112.94	134.55		
Standard deviation	9.30	2.27	53.46	3.43	19.83	15.99	1.12	8.85	6.20	7.47		
CV(%)	2.17	0.52	3.24	0.83	2.28	l.82	2.74	2.16	5.32	5.52		

CV, coefficient of variation.

Image acquisition was performed in a standing position by a single 360° rotation with a variable scanning time (18–26 s pulsed X-ray). The number of slices (images) in each set was 251. The acquisition time ranged from 12.252 to 12.312 ms. All available data sets were exported in digital imaging and communications in medicine format.

Segmentation protocol

The method selected for the segmentation procedures used ITK-SNAP software (Penn Image Computing and Science Laboratory).^{[11,14,17](#page-8-0)} This software is an open source medical image processing application that provides a combination of manual and semi-automatic tools for extracting structures in 3D image data of different modalities and from different anatomical regions[.14](#page-8-0) It was developed based on the library of image analysis algorithms, Insight ToolKit, and the library of visualization algorithms and advanced modelling techniques, Visualization ToolKit.

Manual volumetric segmentation

Two clinicians experienced in endodontics and dental imaging performed the manual segmentations (MAN, Operators A and B). The data were segmented twice to verify the variability of the manual segmentation.

Manual segmentation was performed with the open source software ITK-SNAP v. 2.4.0.^{[18](#page-8-0)} Every case $(n = 5)$ was segmented manually using slice-by-slice boundary drawing in all three orthogonal views. This procedure was performed three times for each case at a minimum time interval of 2 weeks. The time spent by the two operators ranged from 1 to 3 h depending on the case. The manual segmentations performed served as the gold standard [\(Figure 1\)](#page-1-0).

Semi-automated segmentation

Semi-automatic segmentations were performed using the same software mentioned above. Three different procedures were applied. The first two methods were developed and proposed by ITK-SNAP. The procedures use 3D active contour methods to compute feature images based on the CBCT image grey level intensity and boundaries. The principle of active contours is to evolve an initialized curve towards the outlined object of interest. The first method forces the active contour to slow down near edges, or discontinuities, of intensity (edge-based contour evolution or EB). The second method causes the active contour to be attracted to boundaries of regions of uniform intensity and to reach equilibrium (region competition contour evolution or $R\ddot{C}$.¹⁴ The third method was developed from edge-based contour evolution by adjunction of an anisotropic diffusion filter (EBDA).

Four operators applied the protocol: three dental clinicians experienced in endodontics and dental imaging and an engineer experienced in image analysis. It was performed five times for each case with an interval of 2 weeks between the same procedures. Two operators performed both manual and semi-automatic procedures.

The operators selected the best parameters according to the case and to the method.

The segmentation results were visualized by a 3D display of the segmented structures. Once the procedure had been finished, the software provided the volume of the segmented structure (mm³). The volumes of the lesion 3D reconstructions were used to provide statistics. Time spent was less than 10 min for all operators. All the results were classified according to the case, the technique and the operator.

Statistical methods

All data were analysed using StatView v. 5.0 (SAS Institute Inc., Cary, NC). The accuracy of different approaches was calculated by comparing the volumes (mm³) obtained by semi-automatic with manual segmentation. For each protocol and each case, an analysis of dispersion among-operator and within-operator sum of squares (mean, standard deviation and coefficient of variation) was made. An ANOVA test was used to evaluate the operator effect. An a posteriori Bonferroni–Dunn test was also applied if a statistically significant difference was previously noticed, to determine which means differ. Nonpaired t-tests were used to compare semi-automatic with manual procedures. Statistical significance was set at $\alpha =$ 0.01. The null hypothesis was that there was no significant difference in lesion volume measurement between the semi-automatic methods and the manual delineation.

Results

Manual protocol

The coefficient of variation for the manual protocol ranged from 0.5% to 5.5% with a mean of 2.5–3.5% within operator. The correlation coefficient between the operators was 0.9997. There was no statistically significant difference between the two operators, for each pair, by ANOVA 1%. The results of the manual segmentation protocol can therefore be used as a reference [\(Tables 2](#page-2-0) and [3](#page-2-0)).

Semi-automatic protocol

The coefficients of variation for the semi-automatic protocol were quite high for Case 4 and especially for Case 5, meaning that the dispersion around the mean can be high, depending on the operator ([Tables 4](#page-3-0)–[6\)](#page-5-0).

Operator effect

The ANOVA analysis showed significant differences among the operators, for the edge-based technique, regarding Case 2 and 4 segmentations. The post hoc test (Bonferroni –Dunn) showed a significant difference between Operators A and D, B and C, B and D and C and D for Case 2; and A and C, A and D, B and C and B and D for Case 4.

There were significant differences among operators for the region competition procedure in Cases 3 and 4 (ANOVA). With the post hoc test, there were significant

differences between Operators A and C, A and D and B and D for CBCT Case 3; and for all the operators except A and C for CBCT Case 4.

There were significant differences between the operators in all cases except Case 3 for the edge-based with anisotropic filter procedure (ANOVA). The post hoc test showed significant differences between A and D, B and D and C and D for Case 2; B and C and B and D for Case 4; A and C and B and C for Case 5; and between all the operators except between A and D for Case 1 ([Table 2\)](#page-2-0).

Comparison between protocols

In cases where the operators agreed with each other, a comparison was made with an unpaired t-test. In the other cases, the difference between operators forcibly led to a difference of segmentation between manual and semi-automatic protocols.

Cases 1 and 5 had comparable results between the manual and edge-based procedures (non-significant difference). In Case 5, the result could be explained by the existence of an important dispersion in the measures, making this particular result unreliable.

The comparison between the manual and region competition procedures showed non-significant differences in three cases (1, 2 and 5). For Case 5, the conclusion is the same as before. Cases 1 and 2 had no cortical bone destruction, unlike the three other cases.

For all cases, results of the EBDA procedure were significantly different from those of manual segmentation. This technique does not allow accurate semiautomatic segmentation as it did not provide reproducible results.

Discussion

The aim of this study was to test and compare a manual segmentation protocol with three active contour segmentations for the delineation of dento-maxillary osteolytic lesions. The manual protocol was validated as a reference in our study. The objective was to develop a semi-automatic protocol comparable to the manual procedure with an acceptable coefficient of variation and no statistically significant difference regarding lesion volume for all cases tested. The present investigation produced mixed results. For the semi-automatic procedures, region competition segmentation had the least operator effect and gave the best results. This procedure was comparable to manual segmentation for the lesion without destruction of the cortical bone. The EBDA procedure had the largest operator effect and did not provide reproducible results. The semi-automatic protocol tested here was interesting but was not usable in all cases. These methods were not repeatable and remained unreliable in complex cases. All parameters need to be improved. A post-treatment would be necessary in most of our test cases. Edge-based segmentation methods, which use local edge information to evolve the contour of edges, fail at boundaries where the edges between the

ROI and background are not sharp or clear. Regionbased segmentation methods use different intensity distributions of the ROI and background to separate them. These methods fail in areas where the region inside the ROI has similar intensity levels to those of the background.¹⁹ In these periradicular lesions, bone resorption was not uniform. There were many variations in their shape, their size, their extension and their intensity distribution. The boundaries of such lesions can be circumscribed, sharp (Cases 1 and 2) or, conversely, anarchical and non-homogeneous (Case 5). The cortical bone can be blown or broken over a portion of the lesion (Cases 3 and 4).

The tested procedures are a good basis for work, but it is essential to develop a tool more suited to this type of pathology. Hybrid segmentation protocols could overcome these limitations and problems.[19,20](#page-8-0) Despite this, it is likely that a manual post-treatment will always be necessary, in cases where cortical bone is broken or there are few sharp or clear boundaries, to remove the overflow generated by the semi-automatic protocol, even when optimized.

One recent study examined an innovative tool, a 3D smart brush, which seems to provide a reliable and fast method of segmentation in odontogenic cyst and tumour surgery, although this study was performed on MSCT rather than CBCT.^{[21](#page-8-0)} Several studies have been performed on CBCT in order to evaluate and optimize the segmentation step for the head and neck region, including condyle, 22 22 22 and teeth and jaws. 23,24 23,24 23,24 However, to our knowledge, no studies have yet been performed on CBCT and odontogenic cysts.

In the present investigation, five CBCT image data sets were used. The selected scans had a 200 - μ m resolution, which was the common resolution proposed by our equipment for 4×5 or 7×5 cm fields of view with normal settings. This is a compromise between the noise and possibilities of image processing. The high-resolution settings, proposed by our CBCT device, lead to high noise in the images that, at present, prevent accurate segmentation. CBCT has been described to visualize high contrast structures such as bone in a comparable way to conventional CT. The level of radiation exposure is lower, and the metal artefacts are reduced.^{[25](#page-8-0)} CBCT with a limited volume is well suited for the diagnosis of periapical pathology, as used in planning for endodontic surgery. $3,2$ Limited volume CBCT provides images at high spatial resolution, with low radiation doses and displaying accurate measurements. $26,2$

Its disadvantage is the lack of tissue information.^{[25](#page-8-0)} Segmentation in CBCT is often more complex because the image is noisier than in MSCT, owing to lower bone signal-to-noise ratio. The image contrast between the tooth root and the alveolar bone is lower in CBCT. The tooth boundary is too ambiguous to be exactly defined.¹⁹

The contrast, signal-to-noise ratio and sharpness of the images obtained from CBCT acquisitions still need to be improved to make the segmentation easier. 29 29 29 The initial image quality is very important. Parameter variations lead to significant differences in image quality. This influences the potential to visualize the structures that need to be analysed. The voxel size, acquisition quality, accurate resolution and limited noise need to be adapted to the clinical situation. The image must provide information in adequate quantity to allow a sufficient degree of certainty when making a decision. In vivo conditions impose some settings that affect image quality. Indeed, acquisition time must be as short as possible to limit radiation exposure. However, it is CBCT scan time and voxel size that are the main factors influencing jawbone model accuracy.^{[30](#page-8-0)} Scans of real patients contain more artefacts and reduction in spatial resolution owing to natural micro-movements of the jaw.

CBCT has somewhat lower segmentation accuracy than MSCT; but, considering the results obtained with low radiation, the short scanning time and good image quality, CBCT could be helpful for surgery in the orofacial region. 31

ITK-SNAP software was chosen because it was developed for the treatment of medical images. Therefore, active contours ("snakes") were adapted for our investigation. It was validated for segmentation of brain MRI images.¹⁴ It is also a free open source software: anybody can use it and can modify the source code to adapt and optimize the segmentation algorithm to their needs.

Semi-automatic segmentation methods are user-guided. The work requires a manual element and guidance, consisting of fast and accurate refinement techniques to assist the human operator. 13 Computation time for the semiautomatic processing was less than 10 min compared with the manual procedure, which took >1 h.

Image segmentation is a fundamental problem in image analysis. The segmentation procedure gives the surgeon information about the total volume and dimensions of a lesion and the spatial relationship between the lesion and any high-risk structure in its proximity. It is a helpful additional process in preoperative diagnosis.⁸

The accuracy of segmentation relies on the grey value and threshold value defined by the operator. Periradicular cyst variability and the frequent absence of a strong grey gradient near their boundaries make the segmentation step difficult, which leads to an overextension of the segmentation beyond the limits of the lesion with the semi-automatic procedure. Owing to the lack of distinctive features that can describe the boundary of the lesions, it is difficult to achieve segmentation with a high degree of accuracy. This leads to underestimation or overestimation of size. It is hard to segment accurately, even using the manual procedure, when visual features (boundaries, regions etc.) are not sufficiently dense, robust or discriminating. Each procedure requires some degree of compromise.

To limit these problems, segmentation requires a thorough knowledge of the images to be analysed and of the information to be extracted subsequently. A good knowledge of the situation enables the user to separate artefacts and noise from what is of interest. The subjectivity involved in this method can lead to different results being found by two users or by a single user between two segmentations of the same tooth. Prior training of users in displaying the segmentation of the images before starting is interesting 29 and should be considered. In our study, half of the operators were not highly trained for the segmentation protocols to be tested. The aim was to determine if these protocols can be applied by a novice or a non-expert. In any case, subjective ability of the observers must also be taken into account.

In the medical field, it seems to be impossible for a segmentation algorithm to meet all the requirements owing to the large numbers of characteristic differences between organs and tissues.^{[32](#page-8-0)}

Manual post-processing, which benefits from the skilled eye and judgment of a clinician, is still required to rectify the semi-automatic procedure. The problem this approach poses is that it usually takes longer (up to several hours) and is therefore not practical.

Segmentation is not easy and its complexity is not often taken into account.^{[33](#page-8-0)} Nevertheless, high-quality segmentation within CBCT images of patients requiring oral surgery is a prerequisite for surgical planning, simulation and guidance. Likewise, image quality of CBCT is a prerequisite for accurate segmentation. The CT scanners have excellent spatial accuracy, but their quality needs to be improved in terms of signal-to-noise ratio, contrast and measurement of the radiodensity. Acquisition parameters have to be correctly chosen by the clinician. The CBCT manufacturers also should optimize the technical characteristics of their products to obtain better quality CT scans.

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Research on advanced segmentation methods is essential to obtain a semi-automatic accurate individual segmentation of dento-maxillary osteolytic lesions and their environments.¹¹

The present study has contributed to testing some semi-automatic segmentation protocols on dentomaxillary imaging. Further investigations need to be performed in order to improve segmentation quality.

Conclusion

Region-based active contour segmentation was comparable to the manual procedure for delineating circumscribed osteolytic dento-maxillary lesions. However, region-based and boundary-based segmentations are not suitable for all cases. These approaches are limited when segmenting objects with complex surface structures. A hybrid methodology, combining both manual and semi-automatic protocols, needs to be developed in order to provide a suitable method for the majority of cases, regardless of clinical situation, extent and position of the lesion.

The improvement of the image analysis must be considered both through the segmentation procedure and through CBCT image quality (limited noise, augmented contrast etc.).

Acknowledgments

The authors would like to thank Dr Vincent Morin, Associate Professor, for his statistical help.

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