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## FULL PAPER

# Accuracy and precision of manual segmentation of the maxillary sinus in MR images—a method study

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**Objective:** To assess the accuracy and precision of segmentation of the maxillary sinus in MR images to evaluate the potential usefulness of this modality in longitudinal studies of sinus development.

**Methods:** A total of 15 healthy subjects who had been both craniofacial CT and MR scanned were included and the 30 maxillary sinus volumes were evaluated using segmentation. Two of the authors did segmentation of MRI and one of these authors did double segmentation. Agreement in results between CT and MRI as well as inter- and intraexaminer errors were evaluated by statistical and three-dimensional analysis.

**Results:** The intraclass correlation coefficient for volume measurements for both method error, inter- and intra-examiner agreement were > 0.9 [maximal 95% confidence

interval of 0.989–0.997,  $p < 0.001$ ] and the limit of agreement for all parameters were < 5.1%. Segmentation errors were quantified in terms of overlap [Dice Coefficient (*DICE*) > 0.9 = excellent agreement] and border distance [95% percentile Hausdorff Distance (*HD*) < 2 mm = acceptable agreement]. The results were replicable and not influenced by systematic errors.

**Conclusion:** We found a high accuracy and precision of manual segmentation of the maxillary sinus in MR images. The largest mean errors were found close to the orbit and the teeth.

**Advances in knowledge:** MRI can be used for 3D models of the paranasal sinuses with equally good results as CT and allows longitudinal follow-up of sinus development.

## INTRODUCTION

The use of three-dimensional (3D) software to construct models for representation of body compartments has gradually gained ground and has become useful for diagnostics, surgical planning and simulation.<sup>1–3</sup> Furthermore, 3D models could become useful in following development of sinuses, which is of particular interest in patients with chronic rhinosinusitis (CRS). CRS may slow and alter pneumatization<sup>4</sup> and is a major concern in individuals with Cystic Fibrosis<sup>5,6</sup> and Primary Ciliary Dyskinesia.<sup>7,8</sup> In general, the maxillary sinus is one of the most common sinuses for disease<sup>9</sup> and surgery, which is why it is of particular interest.

In understanding pathology, a complete knowledge of the normal development is necessary and is currently based on cross-sectional studies using CT,<sup>10–13</sup> cone beam CT (CBCT),<sup>14–16</sup> and a few using MRI.<sup>17–19</sup> Hence, longitudinal

studies are clearly called for to elucidate the growth and development of the paranasal sinuses further. In principle, CT scans would be the best modality;<sup>20</sup> however, the increased cancer risk associated with consecutive CT scans during childhood<sup>21</sup> makes such a study unacceptable. Even though lower doses are seen with CBCT,<sup>22</sup> an extensive follow-up would imply a significant amount of radiation, and the Food and Drug Administration still recommend their use only when clinically necessary. An alternative solution would be to employ longitudinal MRI without using ionising radiation.

The primary purpose of the present study was to assess the accuracy and secondary the precision of segmentation of the maxillary sinus in MR images in order to evaluate the potential usefulness of this modality in longitudinal studies of normal individuals and patients with sinus disease.

## METHODS AND MATERIALS

### Subjects

15 subjects (10 females and 5 males; median age: 18 years, range: 14–20 years) without sinonasal disease or history of sinus surgery were included in the study. While one patient underwent MR 28 days after the CT scan, all the others had both scans done within the same day. The majorities of patients were diagnosed with osteoarthritis in the temporomandibular joint and were referred to CT to evaluate bony changes and MR for articular disk condition. Other indications were trigeminal neuralgia, severe trismus, phlegmone (cellulitis) and mandibular protrusion. Since this is a retrospective study, subjects did not receive any additional radiation exposure.

### Scanning procedure

All 15 subjects were scanned with a 1.5T MR scanner (neurovascular head and neck coil; GE Healthcare, Milwaukee, WI). We applied a 3D  $T_1$  weighted image sequence. The imaging protocol consisted of sagittal fast gradient echo sequences (TR/TE/FA/NEX, 10.5 ms/4.8 ms/15 degree/1) 280 × 280 × 224 mm FOV; 512 × 512 × 320 matrix size (voxel size was 0.55 × 0.55 × 0.7 mm).

CT examination was performed by a 64-row multidetector CT scanner (Light Speed VCT; GE Healthcare, Milwaukee, WI) and applied helical (spiral) mode. A dose reduction algorithm [3D mA Modulation (Smart mA), GE] was applied. CT images were taken at 120 kVp and 200 mA. Axial images were obtained with the X-ray beam parallel to the occlusal plane or mandibular plane and displayed with bone window [WW/WL: 4,000/800 Hounsfield units (HU)]. The field of view was 25 × 25 cm. The matrix size was 512 × 512 (voxel size was 0.49 × 0.49 × 0.625 mm). No enhancement methods were used in either CT or MR.

### Image Registration

For each image pair, the CT scan was oriented according to the Frankfort horizontal plane and the mid-sagittal plane by manually placing four anatomical landmarks (the two *orbits*, the deepest point on the infraorbital rim, and the two *porions*, the highest point on the upper margin of the external acoustic meatus bilaterally). Subsequently, the MR image was registered to the CT using a voxel-based rigid image registration algorithm (Image Registration Toolkit, IRTK, IXICO PLC, London, UK) applying normalised mutual information as a similarity measure.<sup>23</sup> Thereafter, fast flickering between the CT and MR images was done for quality control.

### Segmentation

The aligned scans were imported into Analyze™ (v. 12.0, Minneapolis, MN). Segmentation of the CT images was carried out with "Auto Trace", a two-dimensional seeded region growing and level-set method, using a seed pixel and predefined thresholds of -1024 to -318 HU. Every slice in the transversal plane was reviewed and any errors were manually corrected. Segmentation of MRI was manual using the "Spline Edit" tool, a curve interpolated between points manually set by the observer for each slice. In the context of this work, the segmentation performed on CT is considered as the gold standard (GS). While only one author

did segmentation of CT, two authors did segmentation of MRI and one of these did double segmentation with a minimum of 1-month interval between the two. Observer 1 (O1) and observer 2 (O2) were calibrated in terms of segmentation procedure and any uncertainty of the general delineation was discussed with an ENT doctor and an oral maxillofacial radiologist before the study was initiated. Thereafter, both observers carried out a pilot segmentation on three subjects each, which were the last ones to be segmented with a minimum of 1.5-month interval from the training period.

### Analysis

An error analysis was carried out based on segmentation of left and right sinuses from 15 individuals, resulting in a total sample size of 30.

### Volume error

The volume of each sinus was calculated by multiplying the number of voxels with the voxel dimensions. The intraclass correlation coefficient (ICC) was the measure of agreement for volume comparisons. A modified Bland-Altman approach was used to test for random and systematic errors by plotting the relative difference in volume of the sinuses segmented in CT and MRI.<sup>24</sup>

### Magnitude and spatial distribution of error

An investigation of the magnitude of local errors was carried out by comparing pairs of segmented sinus surfaces from the same individual.

Six types of comparisons were carried out:

- (1) O1AvsGS: comparison of observer 1's first segmentation (A) to the gold standard
- (2) O1BvsGS: comparison of observer 1's second segmentation (B) to the gold standard
- (3) O2vsGS: comparison of observer 2's segmentation to the gold standard
- (4) O1AvsO1B: intraexaminer comparison between observer 1's two segmentations
- (5) O2vsO1A: interexaminer comparison of observer 2's segmentation to observer 1's first segmentation
- (6) O2vsO1B: comparison of observer 2's segmentation to observer 1's second segmentation

Dice Coefficient (*DICE*) and Hausdorff Distance (*HD*)<sup>25</sup> were used as comparison metrics. *DICE* provides a measure of the amount of overlap between two segmentations and ranges from 0 (no overlap) to 1 (full overlap). *HD* represents the maximum distance between corresponding segmentation boundary points. A 95% Hausdorff percentile (*HD95*) was also calculated, representing the maximum distance when the largest 5% of distances were discarded (this is the default setting in the Insight Toolkit).<sup>26</sup> Segmentation boundaries were formed as polygonal surfaces using Marching Cubes.<sup>27</sup> Corresponding points on two surfaces were determined as the distance  $d_1$  from a point in one (source) surface to the nearest location in the other (target) surface. Another set of distances,  $d_2$ , were also calculated by switching the role of source and target surfaces, providing, in general, different values than  $d_1$ . *HD* is therefore, in practice, calculated as the

maximum of all distances  $d_1$  and  $d_2$ . Histogram distributions of distances were also plotted, as well as mean histogram distributions (one for each of the six types of comparison).

An investigation of the spatial distribution of the local differences (“errors”) ( $d_1$  or  $d_2$ ) was carried out by color coding the source surface by the distance value, providing a distance or “error” map  $E$ . In the plotted distributions and color coded surfaces the distances were given a sign depending on whether the source surface was inside or outside of the target surface. A mean signed distance ( $\bar{D}$ ) different from zero thus indicates a systematic difference between a pair of segmentations. Mean error maps  $\bar{E}$ , averaging the error maps from all individuals were also constructed, but it required establishment of detailed point correspondence between surfaces from different individuals; this was achieved using a method similar to the method by Szeliski et al<sup>28</sup>.

### Statistics

Calculation of ICC was done in SPSS (v. 23) based on inter- and intraobservations with a fixed number of two examiners. The two-way mixed procedure was used and results for average measures were reported. Bland Altman plots were calculated and illustrated using RStudio (v. 0.99.879). Power analysis was based on a paired t-test and found to be 0.93 in detecting a 2% difference between CT and MRI with a two-sided significance level of 0.05. Normality was checked with a Shapiro-Wilk test and verified by plotting the residuals in a QQ-plot.

## RESULTS

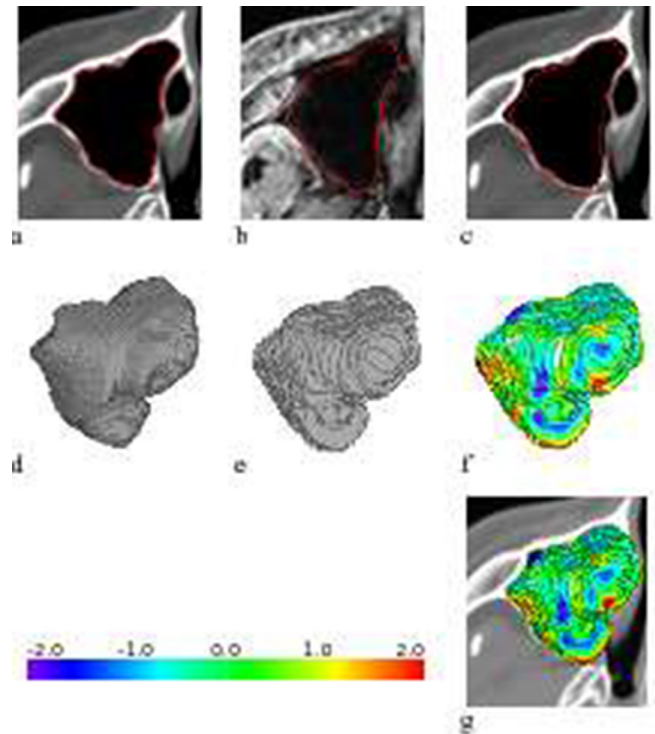
### Registration

The visual quality control of registration results with fast flickering between CT and MRI showed good agreement of the maxillary borders and related anatomical structures (Figure 1).

### Segmentation

Figure 2 shows an example of segmentation of a maxillary sinus. It may be seen how tracing of the sinus’ border differs between MR and CT in some locations. Figure 2 also shows corresponding 3D surface reconstructions, indicating that similar 3D shapes are obtained in the two different modalities. An example error map  $E$  for comparison type O1A vs GS is also shown, indicating local errors up to about  $\pm 2$  mm. Segmentation of a single sinus took between 1 and 3 h depending on the shape and size of the sinus. Overall, O2 used considerably less time on segmentation than O1.

Figure 2. Example of segmentation. Axial slices. (a) Semi-automatic delineation (red) on CT. (b) Manual delineation (red) on MR. (c) Manual delineation (red) on MR shown on corresponding CT slice. (d) CT surface reconstruction from semi-automatic segmentation. (e) MR surface reconstruction from manual segmentation. (f) The MR surface reconstruction colour coded according to an error map  $E$  providing an accuracy of MR surface reconstruction in terms of calculated closest point distance in mm between the surfaces in d) and e). (g) Same as f) but shown with CT image slice for spatial context.



### Volume reliability

The mean volume of assessed sinuses was 21.5 cm<sup>3</sup> (range 12 to 28 cm<sup>3</sup>). In accordance with the classification by Landis and Koch<sup>29</sup>, the intraexaminer and interexaminer ICC (Table 1) indicate an excellent accuracy and precision (ICC range 0.995–0.998,  $p < 0.001$ ). The limits of agreement were less than 5%. The Bland-Altman plots (Figure 3) did not

Figure 1. Example of images included in the study. (a) Axial slice of CT scan. (b) Axial slice of MR scan shown after voxel-based rigid registration to the CT scan. (c) Fusion with 50% CT and 50% MR after registration.

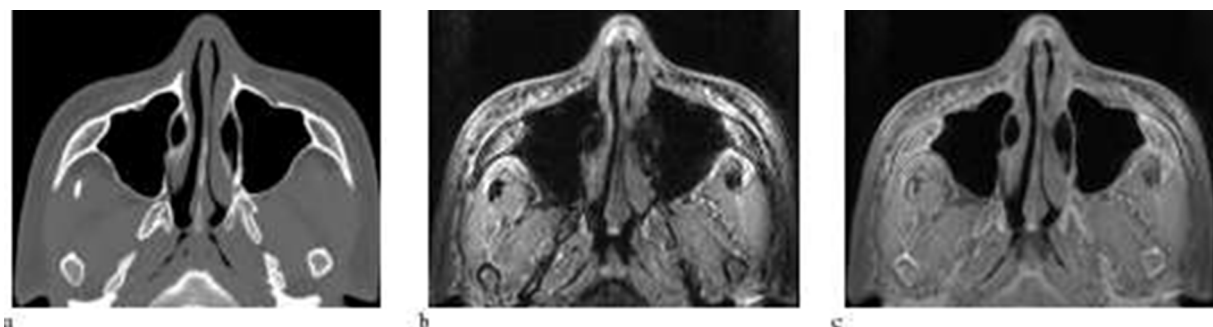


Table 1. Method, intraexaminer and interexaminer reliability for volume measures of the maxillary sinus using CT and MR segmentation ( $n = 30$ ).

Comparison	Intraclass correlation coefficient	95% CI	Mean Diff [%]	SD [%]	Limit of agreement (Mean Diff. $\pm$ 1.96 SD) [%]
O1AvsGS	0.998	0.996–0.999	0.20	1.6	–2.9 to 3.3
O1BvsGS	0.996	0.992–0.998	0.72	2.0	–3.2 to 4.7
O2vsGS	0.997	0.994–0.999	–0.06	1.9	–3.9 to 3.8
O1vsGSa	0.998	0.995–0.999	0.46	1.6	–2.6 to 3.5
O1AvsO1B	0.998	0.995–0.999	0.52	1.8	–3.0 to 4.0
O1AvsO2A	0.995	0.989–0.997	–0.26	2.46	–5.1 to 4.6

CI, confidence interval; GS, gold standard; Mean Diff, mean difference; O1, observer 1; O1A, first segmentation by observer 1; O1B, second segmentation by observer 1; O2, observer 2; O2A, first segmentation by observer 2; SD, standard deviation.

<sup>a</sup>For O1, a mean of volumes for O1A and O1B was used to compare with GS.

show any evidence of systematic errors for any of the observers.

### Comparisons

Table 2 presents results of the six comparisons in terms of Dice Coefficient (*DICE*), Hausdorff Distance (*HD*), 95% percentile Hausdorff Distance (*HD95*) and mean signed distance (*D*). Minimum *DICE* values are  $>0.9$  indicating excellent overlap between segmentations for all six comparisons. The maximum *HD* was large (about 8 mm), while the maximum *HD95* was much smaller ( $<1.7$  mm) and smaller for observer 1 than for observer 2. The mean signed distance was very close to zero, indicating no systematic differences.

Figure 4 presents mean of the distance distributions for the four comparisons O1A vs O1B, O2 vs O1A, O1A vs GS and O2 vs GS (see the section on "Magnitude and spatial distribution of error" for explanation). It may be seen that the values are approximately normally distributed and with mean values close to 0; thus no systematic errors seem to be present. It is also evident that the narrowest distribution is the one describing O1A vs O1B, followed by O2 vs O1A, O1A vs GS and finally O2 vs GS.

Mean error maps  $\bar{E}$  for the comparisons O1A vs GS are shown in Figure 5. In addition, the comparisons O2 vs GS, O1A vs

O1B and O2 vs O1A can be seen in the supplemental material (Supplementary Figure 1). Regions with the largest mean error were seen especially at the superior and inferior parts of the maxillary sinus, and also at the medial part to some extent.

### DISCUSSION

Although 3D models based on MRI would seem to be potentially useful in examining the development of the paranasal sinuses in children and adolescents because of no ionising radiation exposure, nearly all 3D studies of the sinuses to date are based on CT or CBCT data and accuracy or precision of MR segmentation have not been analysed so far.

The present study showed high accuracy and precision of segmentation of the maxillary sinus in MRI compared with CT. The ICC for volume-measurements indicated excellent approximation of volume. In 95% of the cases, it was possible to predict the sinus volume with a maximal error of 5% using MRI. The large *DICE* values found ( $>0.9$ ) confirmed this, indicating that accuracy as well as inter- and intraobserver error were small. The high *HD* values found reflected the presence of noise in the segmentation border. *HD* is a measure of agreement that is known to be sensitive to outliers. Using *HD95* (95th quantile of distances) revealed that 95% of the distances were below 2.5 mm. In comparison, the average maxillary sinus develops approximately 1 cm in all dimensions and triples in volume from 3 to 18 years of age.<sup>17</sup> In general, O1 had higher accuracy compared to O2, which may be explained by considerably more time spent on segmentation by O1. In addition, we also presented analysis of spatial localisation of deviations and found the largest mean errors to be located in the superior and inferior part of the sinus; areas that are in close relation to the orbit and the teeth respectively. This is most likely because of the difficulty to distinguish the paper-thin bone of the orbital floor from air in the sinus, while the teeth constitute an uneven floor of the maxillary sinus and their placement relative to the maxilla varies between individuals.

Both left and right sinuses were included for each of the 15 subjects. Therefore, the method error has contributions from both intersubject and left-right intrasubject variability. This approach is sensible since the segmentation procedure should be carried out on both left and right sinuses.

Figure 3. Bland-Altman plots of precision of volume estimate. GS, gold standard (CT); O1A, first segmentation by observer 1; O1, mean of two segmentations (O1A and O1B, see the text) by observer 1; O1B, second segmentation by observer 1; O2, segmentation by observer 2. Solid line: mean difference between GS- and MR-volume. Dotted lines: two standard deviations of difference between GS- and MR-volume.

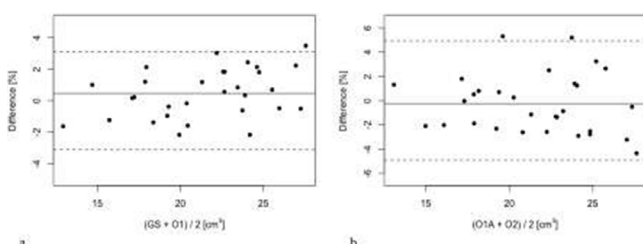


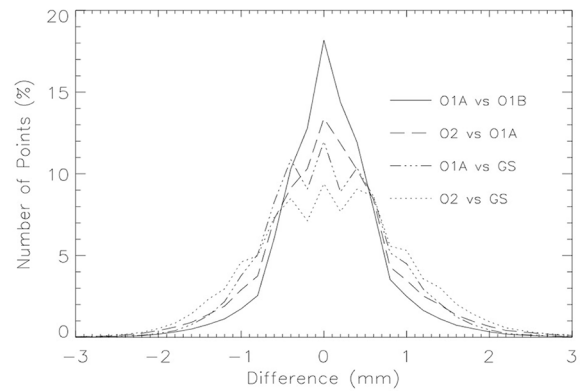


Table 2. Comparison metrics

Comparison	D' (mm)		DICE		HD (mm)		HD95 (mm)		Mean		Max	
	Mean	Min	Mean	Min	Mean	Min	Mean	Min	Mean	Min	Mean	Min
O1A vs GS	0.93	0.92	3.64	2.37	5.49	1.43	1.25	1.91	1.43	1.25	5.49	1.91
O1B vs GS	0.93	0.91	3.37	2.50	5.76	1.44	1.24	1.77	1.44	1.24	5.76	1.77
O2 vs GS	0.92	0.90	4.37	2.61	7.94	1.72	1.25	2.45	1.72	1.25	7.94	2.45
O1A vs O1B	0.94	0.93	3.33	2.31	4.83	1.25	0.92	1.71	1.25	0.92	4.83	1.71
O2 vs O1A	0.92	0.91	4.33	2.63	8.09	1.62	1.25	2.09	1.62	1.25	8.09	2.09
O2 vs O1B	0.92	0.91	4.39	2.72	8.00	1.67	1.31	2.27	1.67	1.31	8.00	2.27

GS, gold standard; O1A, first segmentation by observer 1; O1B, second segmentation by observer 1; O2, observer 2. Values of D' (mean signed distance), mean and range (Min and Max) of Dice Coefficient (DICE), Hausdorff Distance (HD) and 95% Hausdorff percentile (HD95) for six comparisons as indicated. Three uppermost rows pertain to measurement of accuracy; three lowermost to measurement of precision

Figure 4. Distribution of error magnitude. Mean histograms  $\bar{H}$  shown for four types of comparisons as indicated. GS, gold standard; O1A, first segmentation by observer 1; O1B, second segmentation by observer 1; O2, observer 2.

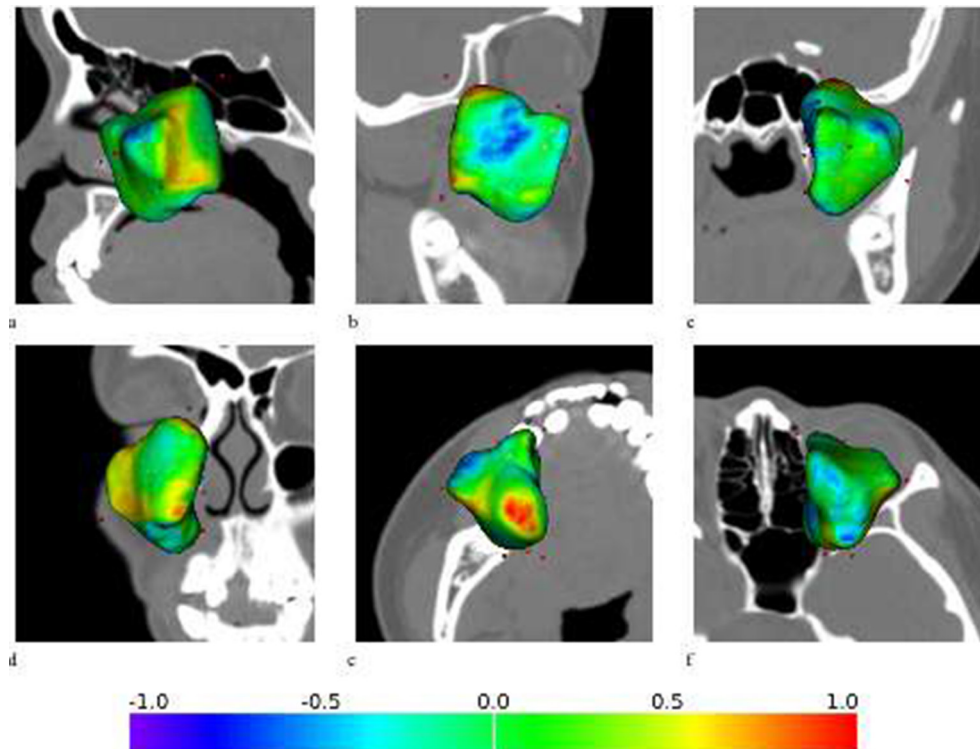


Some previous method studies in the field have used manual segmentation by a single observant as GS,<sup>30,31</sup> but this has shown to have greater than expected intervariability.<sup>32</sup> Furthermore, the precision of manual and semi-automatic segmentation on CT images of the maxillary sinus is previously shown to not be significantly different.<sup>33</sup> The choice of semi-automatic segmentation as GS in this study was based on these considerations as well as the authors' experience with segmentation based on predefined and fixed HU as more reproducible than manual segmentation, since very few or in some cases even no corrections are made by the observer. Nevertheless, this has not been investigated in a study and should therefore be mentioned as a limitation. Another limitation is the limited number of two observers with only one of these doing double segmentation. Therefore, the reader should interpret the results of repeatability in light of this.

Due to ethical considerations, CT data on healthy children is sparse which is why the minimum age in our study was 14 years of age. This may leave a limitation since it can be argued that this study cannot be generalised to children. On the other hand, we included patients with various sinus volumes (range 12–28 cm<sup>3</sup>) and shapes. Furthermore, our study finds that the job of distinguishing paper-thin bone from tissue and air can be overcome with excellent results. This issue does not differ in a paediatric population and the authors argue that the method can be applied in younger individuals with reasonable results. After all, manual segmentation is probably the best approximation of sinus volume based on MRI compared to far simpler stereological methods used in previous MRI studies.<sup>18</sup> Because of its time requirements, however, this method is only useful in research contexts and CT substitutes derived from MRI data could be an interesting and more time efficient alternative.<sup>34</sup>

Previous MRI studies on paranasal sinuses have refused using segmentation for volume assessments<sup>17–19</sup> since the delineation of bony structures on MRI at first seems difficult and imprecise. These results have a particular interest in patients with CRS, but may also be useful in other fields such as obstructive sleep apnoea, where volumetric assessments from MRI have shown to be useful in diagnostics.<sup>35</sup> Finally, this allows us to use manual segmentation for a future project with longitudinal MRI data and

Figure 5. Mean error map  $\bar{E}$  for the comparison O1A (a) vs GS (b). The error maps are shown on a mean maxillary sinus surface (mean of 15 surfaces) and with CT slices shown behind the surface for indication of orientation. Red dots represent landmarks used for alignment. Lateral (a), medial (b), anterior (c), posterior (d), superior (e) and inferior (f) view. Difference between the compared error maps is shown in mm (range  $-1$  to  $1$ ). Negative (bluish) values indicate that the compared segmentation (a) is smaller than the other (b) (a is inside b surface), while positive (red dish) colours indicate that the compared segmentation (a) is larger than the other (b) (a is outside b surface). GS, gold standard; O1A, first segmentation by observer 1.



has a great interest for otolaryngologists in understanding pathological sinus development.

## CONCLUSION

Our results show high accuracy and precision of segmentation of the maxillary sinus in MR images. While maximal errors were found to be relatively large, there were only a few of them. The mean errors were small and not of clinical relevance. The largest mean errors were seen in relation to the orbit and the teeth.

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## CONSENT

Written and informed consent was achieved from each subject, and the “Osaka University Ethical Committee” approved the study (application no. “H21-E16”, “Improvement of accuracy of image diagnosis using CT and MRI”).

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