

# An Automated Segmentation Framework for Nasal Computational Fluid Dynamics Analysis in Computed Tomography

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## ABSTRACT

28 The use of computational fluid dynamics (CFD) to model and predict surgical outcomes in the nasal cavity  
29 is becoming increasingly popular. Despite a number of well-known nasal segmentation methods being  
30 available, there is currently a lack of an automated, CFD targeted segmentation framework to reliably  
31 compute accurate patient-specific nasal models. This paper demonstrates the potential of a robust nasal  
32 cavity segmentation framework to automatically segment and produce nasal models for CFD. The  
33 framework was evaluated on a clinical dataset of 30 head Computer Tomography (CT) scans, and the  
34 outputs of the segmented nasal models were further compared with ground truth models in CFD simulations  
35 on pressure drop and particle deposition efficiency. The developed framework achieved a segmentation  
36 accuracy of 90.9 DSC, and an average distance error of 0.3 mm. Preliminary CFD simulations revealed  
37 similar outcomes between using ground truth and segmented models. Additional analysis still needs to be  
38 conducted to verify the accuracy of using segmented models for CFD purposes.

39 *Keywords–nasal cavity; image segmentation; computational fluid dynamics; computed tomography*

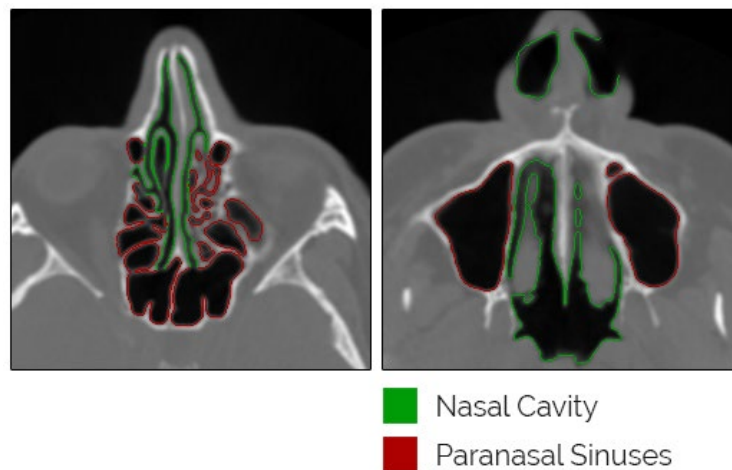
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## 41 **1. Introduction**

42 The nasal cavity’s primary role is to provide humidified, warmed, filtered air before entering the lungs.  
43 Secondary functions include facilitating olfaction, along with ventilation of the paranasal sinuses. To  
44 achieve these specific but varied functions, the nasal cavity has a complex anatomical structure.  
45 Physiological and anatomical conditions in the nasal cavities can result in significant sequelae in the lower  
46 respiratory system. In addition, nasal airway disorders can also disturb other homeostatic systems, such as  
47 sleep and cardiovascular health [1, 2]. Each year, more than 340,000 patients in North America undergo  
48 surgery to correct nasal airway obstruction [2]. However, up to 37% of patients report unsatisfactory or no  
49 improvement after such surgery [3-5]. One of the key reasons for this high failure rate is the lack of objective  
50 methods to predict surgical outcomes [5].

51 In recent years, computational fluid dynamics (CFD) has been proposed as a potential tool for modelling  
52 and predicting surgical outcomes using patient-specific nasal models [5-7]. Through the use of computer-  
53 assisted numerical analysis and simulations, it is possible for CFD to accurately model and derive key  
54 metrics, such as nasal resistance, airflow rate, wall shear stress and heat fluxes. While previous studies have  
55 demonstrated the validity and effectiveness of CFD in modelling the nasal airway [6, 8-10], the process of  
56 patient-specific model creation remains time and labour intensive. There is a critical need for an efficient,  
57 reliable and automated framework to generate patient-specific nasal models to conduct CFD.

58 CFD outcomes are heavily reliant on the accuracy of the patient-specific model used. In order to produce  
59 an anatomically accurate model of a patient's nasal cavity, high quality segmentation is essential. However,  
60 due to the complexity of the nasal cavity anatomy and its connectivity to other airway components of similar  
61 intensity values, such as the paranasal sinuses, it is often difficult to differentiate and segment just the nasal  
62 cavity alone, without performing manual delineation [11]. Figure 1 exemplifies the close proximity of the  
63 paranasal sinuses to the nasal cavity and highlights the connectivity between the two regions. The majority  
64 of existing nasal segmentation methods have either included nearby airway components as part of the  
65 segmentation [12-18], or require some form of manual intervention in order to derive results [17-22].



**Figure 1:** Examples illustrating the issues involved in segmenting the nasal cavity due to the lack of boundary distinction to other airway regions.

69 In the work of Bui et al. [12], a multi-step level-set segmentation procedure was utilised to automatically  
70 segment the nasal cavity and the surrounding paranasal sinuses from cone-beam computed tomography  
71 (CBCT). Whereas Last et al. [13] utilised a level-set based deformable model to segment the nasal cavity  
72 and paranasal sinuses. Other works have used more traditional methods such as thresholding [15, 16] or  
73 region growing [14, 17, 18], where they rely on the similar voxel intensity range, as well as the  
74 interconnectivity of the upper airways to make the segmentation. While these methods were effective at  
75 segmenting airway regions within the human body, they were not able to identify individual components  
76 nor separate them based on anatomical information. Currently, it is difficult to accurately label the  
77 boundaries without manual intervention. This is evident among studies that focused on segmenting just the  
78 nasal cavity itself, such as in the works of Kimura et al. [20] and Alsufyani et al. [22] where thresholding  
79 was used to segment the upper airway, and large amounts of manual delineation was required in order to  
80 separate the nasal cavity from the rest of the airway regions.

81 Keustermans et al. [21] made use of an active shape model (ASM) to semi-automatically segment the nasal  
82 cavity. An ASM is a landmark based statistical shape model (SSM) segmentation method which makes use  
83 of anatomical knowledge derived from a set of training data to segment a particular organ or structure [23,  
84 24]. By modelling the shape variability of the nasal cavity beforehand, they were able to use that  
85 information to significantly reduce the amount of manual intervention required during segmentation when  
86 compared with other literature works, demonstrating the powerful capability of an SSM.

87 In order to achieve a faster and more efficient way of segmenting the nasal cavity, an SSM is essential for  
88 determining the boundaries between the nasal cavity and other airway components. Although [21] made  
89 use of a landmark based SSM for segmentation, it is limited due to the reliance on point correspondence  
90 for shape model representation. As ASMs require every training shape to be constructed with the same  
91 number of points (landmarks) that corresponds, it is difficult to include and model features or shapes that  
92 exist outside the norm. For our goal of establishing patient-specific nasal models for CFD, where a large  
93 majority could likely to be those with diseased or obstructed airways, we require a method which can

94 reliably segment all kinds of cases. Level-set approaches, which are based on evolving contours do not  
95 require point correspondence and can still incorporate statistical and anatomical knowledge as shape priors  
96 into its energy formulation [25, 26]. Although there are known weaknesses for level-set methods during the  
97 segmentation process such as becoming trapped inside a local minimum, these weaknesses can be avoid  
98 when combined with other segmentation methods [27]. As region-based segmentation methods like region  
99 growing have been shown to be effective at segmenting airway regions, combining shape priors from a  
100 level-set SSM with a more advanced region-based method would be the most logical approach for nasal  
101 cavity segmentation.

102 In this study, we demonstrate the efficiency of an automated segmentation framework at optimizing the  
103 nasal model creation process for CFD. Compared with our previous work [11], our proposed framework  
104 makes the following distinctions: (i) the initialization process through the use of superpixels has been  
105 improved [28], in combination with a multi-atlas for seed derivation; and (ii) the framework has been  
106 optimized to reduce the number of manual steps needed for the CFD, by incorporating post-smoothing and  
107 cleaning algorithms, as well as automating the process of pressure inlet and outlet creation using spatial and  
108 anatomical information. The framework was evaluated on a clinical dataset of 30 head CT scans, and the  
109 outputs of the nasal models generated using the segmentation framework were further tested against ground  
110 truth models in CFD simulations. Statistical tests were performed to assess the outcome of the segmented  
111 models against ground truth models for pressure drop and particle deposition.

## 112 **2. Methods**

113 Our algorithm requires the use of a statistical shape model (SSM) which needs to be constructed prior to  
114 segmentation. The segmentation framework can be divided into three main sections: initialization,  
115 segmentation, and post-processing.

## 116 2.1 SSM Construction

117 SSM construction was based on the method of Leventon et al. [25], where a mean offset matrix of the nasal  
118 training data, denoted as  $\{x_1 - \bar{x}, x_2 - \bar{x}, \dots, x_n - \bar{x}\}$  is constructed, with  $x_1$  to  $x_n$  being the signed distance  
119 representations of the training shapes and  $\bar{x}$  being the mean denoted as  $\bar{x} = \frac{1}{n} \sum_{i=1}^n x_i$ . The resulting  
120 eigenvector  $\mathbf{U}$  and eigenvalues obtained from the singular value decomposition (SVD) of the mean offset  
121 matrix holds the decomposed features of the nasal cavity shape. An estimate of a novel nasal shape  $S_{est}$ ,  
122 can be represented by  $k$  principal components in a  $k$ -dimensional vector of coefficients,  $\alpha$ :

$$123 \quad S_{est} = U_k \alpha + \bar{x}. \quad (1)$$

124 For the selection of the training data, we assigned a weight  $w$  to each nasal shape  $t$ , where  $w = \frac{|P \cap t|}{|P| + |t| - |P \cap t|}$ .  
125 Nasal shapes that scored below the third quartile were removed from the training dataset.

## 126 2.2 Initialization

127 An atlas consists of a Computed Tomography (CT) image and a corresponding ground truth segmentation.  
128 Affine and B-spline registration using Elastix [29] was first applied to align  $n$  atlas CT images to the input  
129 target image. The transformation parameters of the registration were then applied on the atlas segmentations  
130 to warp them to the same reference frame as the target image. A probabilistic multi-atlas  $A$  was constructed  
131 as the average of the registered segmentations  $\{G_1, G_2, \dots, G_n\}$  over the total number of atlases  $n$ , denoted  
132 as  $A = \frac{1}{n} \sum_{j=1}^n G_j$ .

133 Thresholding was applied on the input image to extract the position of the airway voxels. By overlaying  $A$   
134 on top of the thresholded image  $T$ , an estimate  $P$  of the nasal cavity was obtained from the union of the  
135 thresholded image and the atlas, defined as:  $P = A \cup T$ . The input image was further cropped in order to  
136 better localize the nasal cavity and to reduce the computation time. Smaller and detached airway regions  
137 captured by  $P$  were removed to ensure accurate seed derivation. We applied the SLIC superpixel [28]  
138 algorithm on the cropped image. Foreground seeds are derived from the cluster centres of the superpixels

139 on the largest airway region that lay within  $P$  at a slice by slice level. Background seeds were derived from  
 140 the cluster centres of tissue regions and airway regions a distance  $\sigma$  away from  $P$ , with  $\sigma$  being a numerical  
 141 parameter specified during initialization. Once the required seeds were derived and an estimate of the nasal  
 142 cavity  $P$  was obtained, the shape priors to capture the statistical variances of the nasal cavity were  
 143 constructed.

### 144 2.3 Segmentation

145 The constructed SSM was embedded in a graph-based segmentation framework and an image was  
 146 formulated as a graph  $G = (V, E)$ , where each vertex  $v \in V$  corresponds to an image voxel and each edge  
 147  $e \in E$  connects two vertices in  $V$ . We utilized the base method of random walk (RW) and formulated the  
 148 Dirichlet energy as  $E_{rw} = z^T LZ$ , where  $L$  is the Laplacian matrix defined in [30] and denotes the pairwise  
 149 affinities among the vertices in  $V$ , and  $z \in R^{|V| \times 2}$  is a labeling vector indicating voxel foreground  
 150 (background) probabilities. In our nasal cavity segmentation problem, we defined a new energy term which  
 151 holds the captured shape variances from the nasal SSM to the labeling vector of image voxels. The labeling  
 152 vector can be optimized by solving a graph Dirichlet problem to produce the final probabilistic labeling.  
 153 The proposed energy term was defined as:

$$154 \quad E_{priors} = (z - (U_k \alpha + \bar{x}_{prob}))^T (z - (U_k \alpha + \bar{x}_{prob})) \quad (2)$$

155 where  $\bar{x}_{prob} = \frac{1}{1 + \exp(\bar{x})}$  and  $z = \begin{bmatrix} z_M \\ z_N \end{bmatrix}$ , where  $z_M$  denotes the predefined labels, i.e. foreground and  
 156 background seeds, and  $z_N$  denotes other labels. Given the definition of  $E_{priors}$ , the complete energy  
 157 function is formulated as  $E_{total} = E_{rw} + E_{priors}$ .

158 An estimation of the nasal cavity shape was obtained by minimizing the proposed functional  $E_{total}(z_N, \alpha)$ ,  
 159 iteratively, with respect to each of its variables  $z_N$  and  $\alpha$ . Starting from  $\alpha = 0$ , the mean shape was  
 160 initialized over the input image. Since  $E_{total}$  is convex, we differentiate  $E_{total}$  with respect to  $z_N$  and find  
 161 the critical point yielding:

162 
$$z_N = (L_N + I)^{-1}(2(U_k \alpha + \bar{x}_{prob}) - B^T z_M) \quad (3)$$

163 where  $I$  is an identity matrix,  $L$  is the Laplacian matrix of the image and  $B$  is the matrix partitioned from  $L$   
 164 which correlates the labeled set to the unlabeled set. Secondly, we use the updated  $z_N$  to differentiate  $E_{total}$   
 165 once more with respect to  $\alpha$ , which yields the following:

166 
$$\alpha = (U_k^T U_k)^{-1} U_k^T (z_N - \bar{x}). \quad (4)$$

167 In order to reduce the amount of over-segmentation while still maintaining the effects of our shape priors,  
 168 the output of  $E_{total}$  was constrained to remain within the boundaries of the nasal airway by computing a  
 169 probability of the estimated foreground voxels and removing those that were located in the tissue regions  
 170 based on their intensity value at each step of the differential iteration.

171 **2.4 Post-Processing and CFD preparation**

172 Post-processing smoothing and cleaning were required in order to prepare the output segmentations for  
 173 CFD modelling. An adaptive smoothing algorithm was applied to smooth the edges that contained rough  
 174 and jagged surfaces. A connected regions tool was applied to filter out displaced segmentation noise to  
 175 ensure a single solid object was outputted to STL which could then be meshed for flow simulation. The  
 176 entrances and exit of the nasal model were automatically detected using spatial and anatomical information,  
 177 and automatically extended by the algorithm in order to clearly establish the inlet and outlet locations  
 178 needed in the CFD model. The final segmentation output was converted into STL ready to be meshed for  
 179 CFD simulation.

180 **2.5 CFD Model Creation**

181 ANSYS Fluent Meshing (version 17.0) was used to create the mesh. The inflow and outflow regions of the  
 182 nasal geometry were first separated from the wall region. Once imported, a wrapping algorithm was applied  
 183 with minimum and maximum surface mesh sizes of 0.1 mm and 2.5 mm, respectively. These parameter  
 184 values were selected empirically after a dimensioned rectangular prism was introduced as a body of



185 influence which acted as a secondary sizing control limiting the size of surface mesh to a maximum of  
186 0.1 mm in regions of the nasal cavity that were separated by narrow gaps. Once the volumetric region was  
187 computed it was automatically meshed using polyhedral elements growing according to the local size field  
188 of the region. Inflation was applied at all walls with the Fluent Meshing default algorithm, which uses a  
189 first aspect ratio of 10, last aspect ratio of 4.8, growth rate of 1.2 and is set to generate five layers at the  
190 walls. Node locations were then automatically adjusted by systematically reducing the threshold for the  
191 maximum skewness to approximately 0.6. The final nasal mesh was imported directly into the ANSYS  
192 Fluent solver ready for simulation.

### 193 **3. Materials and Evaluation Setup**

#### 194 **3.1 Materials**

195 Our dataset which consisted of 30 de-identified head CT scans was acquired from the Department of  
196 Radiology at the Royal Prince Alfred hospital (Camperdown, NSW, Australia), following approval from  
197 the Sydney Local Health District Ethics Committee. The subjects of the 30 CT scans were all males of  
198 Caucasian descent over the age of 40 with different nasal or sinus related complaints. The scans were  
199 acquired using a GE Lightspeed-16 CT Scanner using Helical CT imaging protocols with an average  
200 exposure time of 707 ms. The resulting images maintained a resolution of 220 by 220 mm (512×512 voxels)  
201 and a voxel depth of 1.25 mm. The ground truth data were semi-automatically segmented using Geodesic  
202 Image Segmentation [31] by an experienced operator under the guidance of a nasal surgeon. Each  
203 segmentation took approximately 15 minutes to complete. The segmented data were further re-examined  
204 by a clinical doctor with expertise in CT scan interpretation. The resulting ground truth segmentations were  
205 used in our algorithm as both training data for the level-set SSM and for the construction of the probabilistic  
206 atlas.

### 207 **3.2 Segmentation Evaluation Setup**

208 The leave-one-out cross validation was performed on 30 CT images (30 folds), where 29 ground truth labels  
209 were used each time for the creation of the PA and SSM. The initialization parameter was set to  $\sigma = 5$ ,  
210 which was derived empirically based on experiment validations. We compared our framework with the  
211 locally constrained statistical shape model (LC-SSM) [11] and evaluated our method on both segmentation  
212 accuracy and time. We conducted our experiment on a Windows 8.1 64-bit Desktop PC with i5-3470  
213 3.2 GHz processor and 16 GB DDR3 RAM.

214 The following metrics were used for the evaluation of segmentation results: (i) dice similarity coefficient  
215 (DSC) calculated as the overlap between the two volumes according to:  $DSC = \frac{2|X \cap Y|}{|X|+|Y|}$ , where  $X$  is the  
216 segmentation label and  $Y$  is the ground truth label; (ii) average symmetric surface distance (ASSD in mm);  
217 (iii) average symmetric root mean square surface distance (ASRSD in mm); (iv) maximum surface distance  
218 (MSD in mm); and (v) volumetric overlap error (VOE in %). Further details on the evaluation metrics can  
219 be found in Heimann et al. [32].

### 220 **3.3 CFD Evaluation Setup**

221 Of the 30 nasal segmentation outputs, the best and worst segmentation case based on DSC along with 8  
222 additional segmentation outputs were selected and CFD simulation constructed using ANSYS Fluent solver  
223 (version 17.0). The outcomes of the CFD simulation using the nasal segmentation models were directly  
224 compared against the corresponding ground truth nasal models. Two-tailed paired t test was utilized to  
225 assess the mean difference between using segmented and ground truth nasal models, and a  $p$  value less than  
226 0.05 was considered statistically significant.

227 We based the parameter setting for CFD on Engelhardt et al. [33] which models airflow and particle  
228 deposition in the nasal cavity and presents calculated Reynolds (Re) numbers for various flow rates. For  
229 the breathing rate of 30 L/min a  $Re > 3000$  was calculated, indicating turbulent flow. A flow rate of  
230 30 L/min was selected to replicate fast nasal inhalation as would practically occur with administration of

231 therapeutic nasal sprays. As such, the flow was modelled using the realizable  $k-\varepsilon$  turbulence model and a  
 232 target mass flow rate of  $6.13 \times 10^{-4}$  kg/s (30 L/min) set at the pressure outlet. The total pressure at the inlet  
 233 was set to 0 Pa (gauge). The coupled solver was used with convergence achieved when the locally scaled  
 234 residuals fell below  $10^{-4}$ , which typically required 200 iterations.

235 Once the simulation was converged, Lagrangian particle tracking with a turbulent dispersion model was  
 236 applied. The particle diameter size distribution was described using the Rosin-Rammler distribution, with  
 237 the distribution parameters determined from laser diffraction experimental data obtained by analyzing water  
 238 plumes from a spray bottle using a Malvern Spraytec®. The minimum and maximum diameters were set to  
 239  $0.12 \mu\text{m}$  and  $1000 \mu\text{m}$ , respectively, with a mean diameter of  $85.8 \mu\text{m}$  and a spread parameter of 1.92. After  
 240 the flow had converged, water droplets were injected from each inlet and the simulation completed when  
 241 all the particles had either escaped from the outlet or collided with the rigid walls of the nasal cavity, which  
 242 was set to trap particles upon contact.

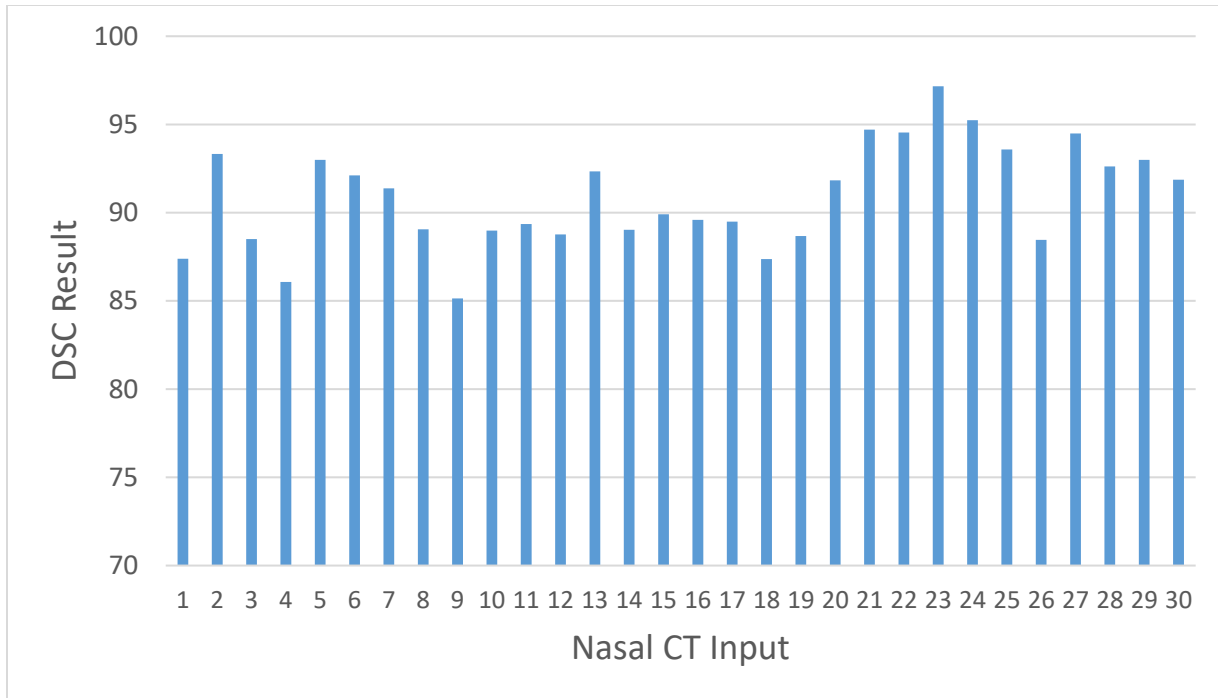
## 243 4. Results

### 244 4.1 Segmentation

245 Table 1 presents the segmentation results of our framework compared with the LC-SSM, evaluated based  
 246 on the metrics described in section 3.2. Our method achieved an averaged Dice Similarity Coefficient (DSC)  
 247 of 90.9%, an averaged Surface Distance error of 0.3 mm, and an averaged Volumetric Overlap Error (VOE)  
 248 of 16.6%. Figure 2 contains the DSC for each individual segmentation case, with case 23 achieving the best  
 249 result (97.1%) and case 9 the worst (85.1%).

Segmentation Accuracy	<i>DSC</i>	<i>ASSD</i>	<i>ASRSD</i>	<i>MSD</i>	<i>VOE</i>
Our method	<b><math>90.9 \pm 2.9</math></b>	<b><math>0.30 \pm 0.1</math></b>	<b><math>0.86 \pm 0.3</math></b>	<b><math>7.5 \pm 1.7</math></b>	<b><math>16.6 \pm 4.8</math></b>
LC-SSM	$90.4 \pm 3.1$	$0.31 \pm 0.1$	$0.86 \pm 0.3$	$7.8 \pm 2.0$	$17.2 \pm 5.4$

250 **Table 1:** Comparative evaluation of our segmentation framework with the LC-SSM.



251

252

**Figure 2:** DSC results of each nasal segmentation

253

Table 2 illustrates the average segmentation speed of our framework. The initialization process took on

254

average 328 seconds to complete, where most of the time was spent on image registration. An average of

255

11.5 seconds was required for the segmentation process to complete. When combining initialization,

256

segmentation, and post-processing, the total average time was 339.6 seconds.

<b>Segmentation Speed</b>	<i>Initialization</i>	<i>Segmentation</i>	<i>Post-processing</i>	<i>Total</i>
Our method	328s ± 3.1s	11.5s ± 1.8s	<1s ± 0.05s	339.6s ± 2.2s

257

**Table 2:** Speed of our segmentation framework

## 258 4.2 CFD

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Case 23 and case 9 were selected as the best and worse segmentation cases based on the DSC result. Cases

260

6, 10, 12, 15, 17, 18, 21, 25 were selected based on their combined average score (90.6 DSC) which was

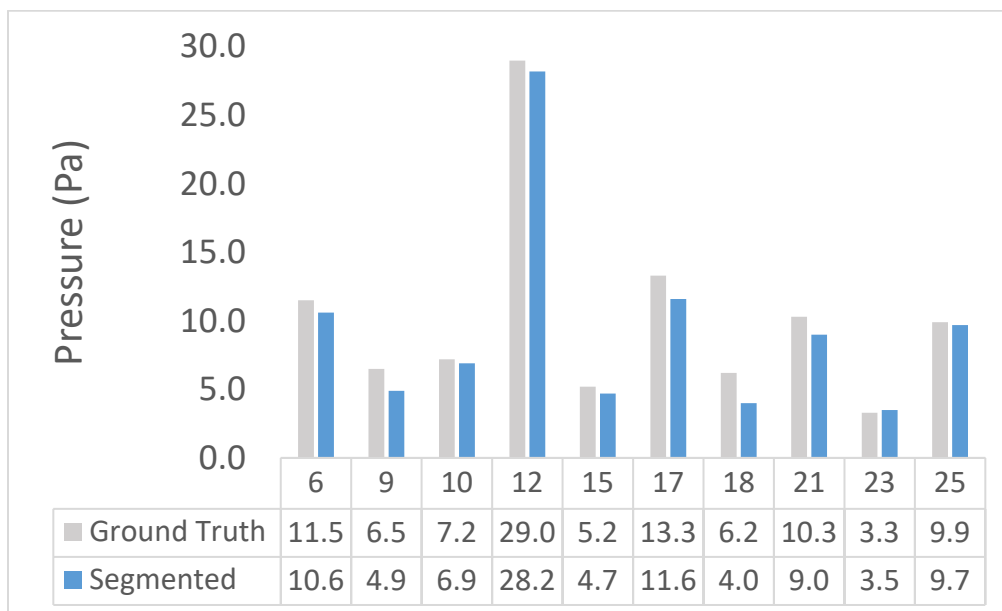
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the nearest to the reported mean DSC average. Figure 3 presents the pressure drop (Pa) for each nasal

262

segmentation case compared against their respective ground truth model. The pressure drops ranged from

263 3.5 Pa (lowest) to 29 Pa (highest), and the results were relatively consistent between the segmented and  
 264 ground truth models with the exceptions of cases 9 and 18 where the differences in pressure drop were  
 265 1.6 Pa (25%) and 2.2 Pa (35%), respectively.

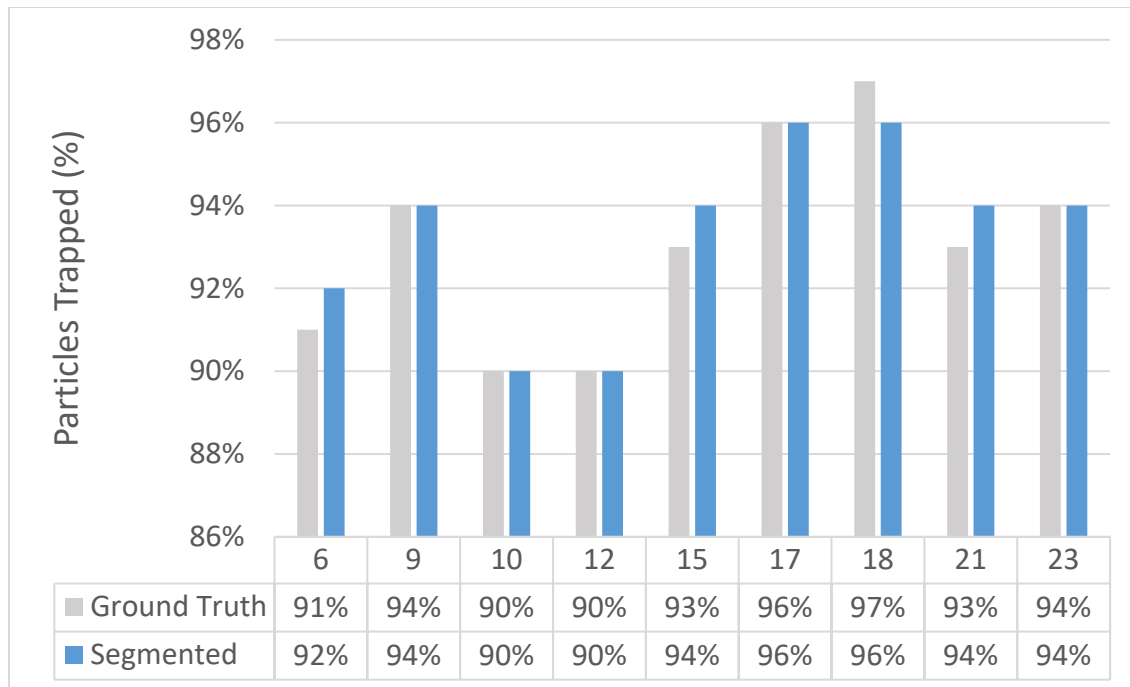


266

267 **Figure 3:** A summary of the pressure drop calculated for the different cases. Results shown in grey are for the ground truth  
 268 models whereas those in blue are for the segmented models.

269  
 270 Figure 4 presents the particle deposition efficiency across the 10 pairs of nasal models calculated as the  
 271 percentage of the input mass flow of particles that were trapped on the walls. The percentage trapped ranged  
 272 from 90% to 97%. Overall, there were very few differences found between the ground truth and segmented  
 273 models. Even for cases that demonstrated greater variation in airflow between the ground truth and  
 274 segmented models (case 9 and 18), the particle deposition remained consistent.

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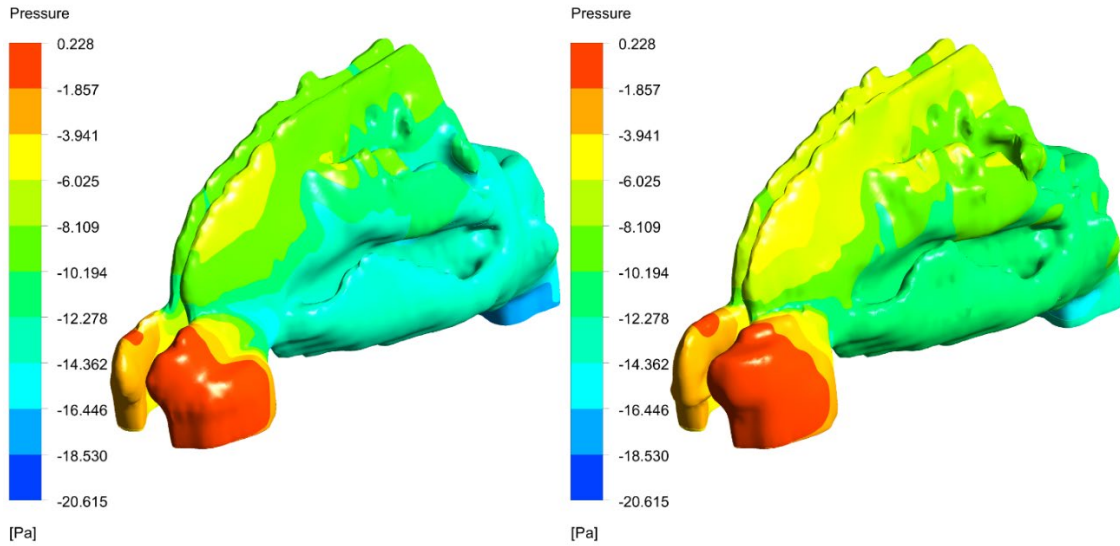


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277 **Figure 4:** The percentage of particle deposited on the walls of the nasal geometry for each case: ground truth model (grey) and  
 278 segmented models (blue).

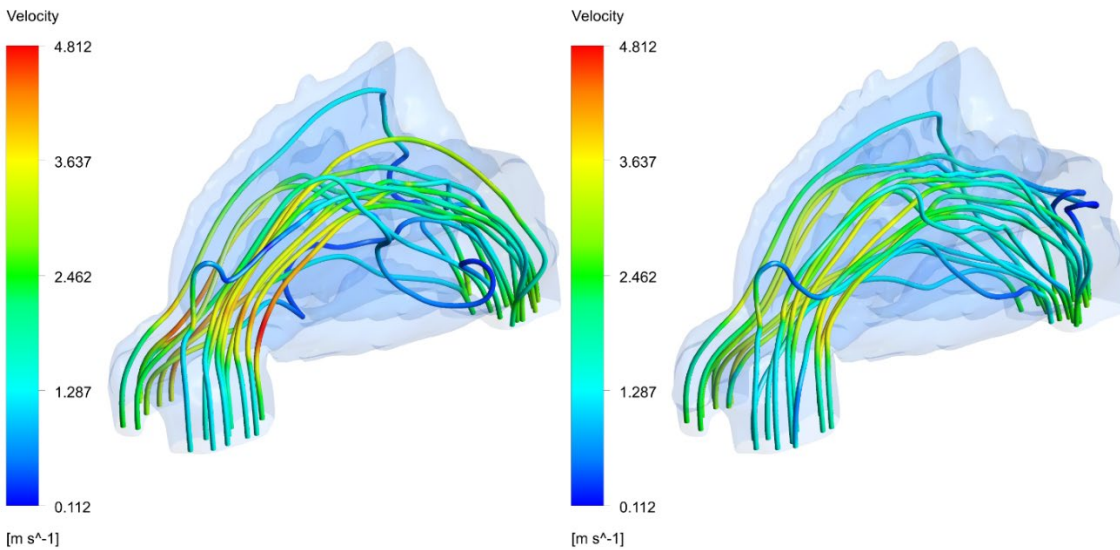
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280 When examining individual cases, both the ground truth and segmented models revealed similar pressure  
 281 distributions across all 10 nasal pairs. Overall, the region comprising of the nasal vestibule and the nasal  
 282 valve were observed to contain the highest pressure drop and velocity magnitude. Figure 5 illustrates the  
 283 wall pressure and the streamline plots of airflow for case 10, where regions of higher pressure in the left  
 284 nostril, as well as in the middle to upper region of the nasal cavities were observed. The outer (inferior  
 285 turbinate) regions displayed less pressure drop when compared with the others, indicating that most of the  
 286 airflow was centred primarily on the middle turbinate region and further disperses to the upper regions.



287

288 **Figure 5a.** Wall pressure plots of case 10 with the ground truth model (left) and segmented model (right) indicating the overall  
 289 change in pressure across the mode from approximately  $-21$  Pa to  $0.2$  Pa.



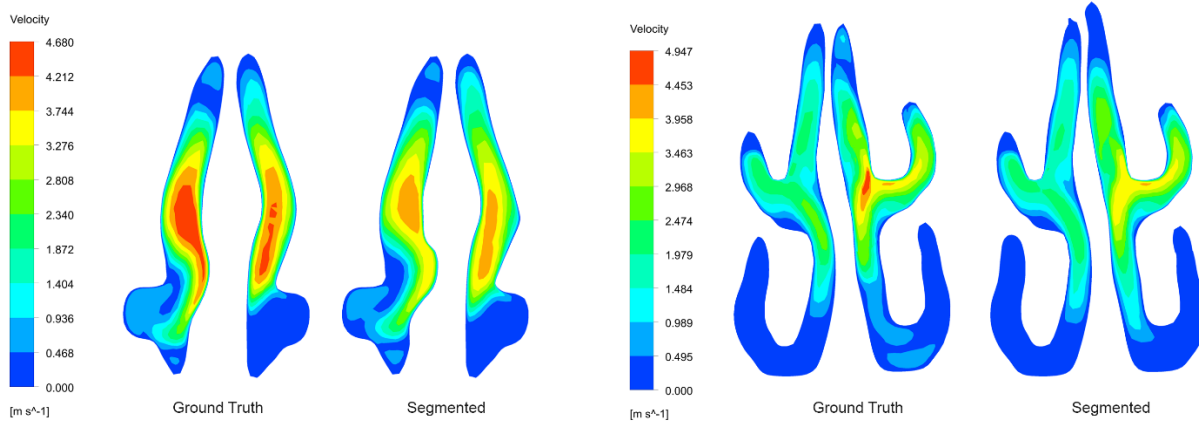
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291 **Figure 5b:** Streamline plots, coloured by velocity magnitude, of case 10 originating from the inlets for the ground truth (left) and  
 292 the segmented model (right) colored by velocity magnitude.

293

294 When looking at the airflow velocities, the middle regions experienced a higher velocity ( $3.6$  m/s) on  
 295 average when compared with the upper regions ( $1.29$  m/s). Figure 6 illustrates the velocity magnitudes  
 296 shown in cross sectional view, where higher velocities were observed in the regions close to the centre of  
 297 each passage, with the highest velocity magnitude ( $4.7$  m/s) occurring in the region adjacent and superior  
 298 to the inferior turbinate of the ground truth model. While the segmented model revealed a similar albeit

302 slightly lower velocity magnitude (4.2 m/s). Despite the similar flow behavior exhibited between the ground  
 303 truth and segmented models, ground truth models were observed on average to have higher velocity  
 304 magnitudes than their counterpart.



305 **Figure 6:** Examples of cross sectional view comparison on the local velocity magnitude. The left example shows the cross section  
 306 at 20mm from the front of the model, and the right example at 40mm from the front of the model.

307 Table 3 presents the overall CFD outcomes between using ground truth and segmented nasal models. No  
 308 significant difference was observed between ground truth and segmented nasal models for pressure drop ( $p$   
 309 = 0.061) and particle deposition ( $p = 0.279$ )

Overall Outcomes	Ground Truth	Segmented	$p$ value
Pressure Drop (Pa)	10.2	9.3	0.061
Particle Deposition (%)	93	93	0.278

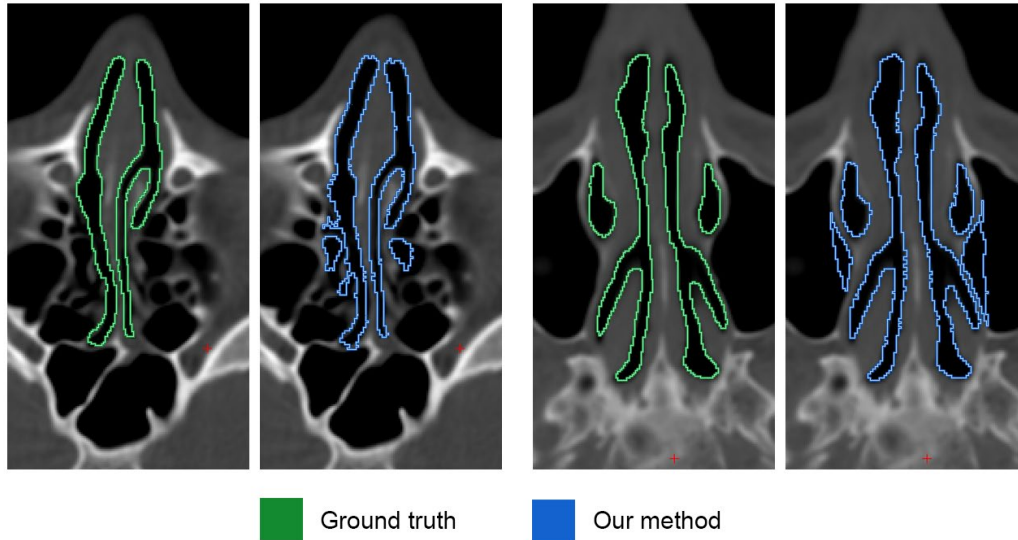
310 **Table 3:** Mean pressure drop (Pa) and percentage of particles trapped for the ground truth and segmented nasal  
 311 models. Two-tailed paired t test was conducted to derive the  $p$  values.

## 312 5. Discussion

313 Overall, our framework was able to automatically segment the nasal cavity from CT images at a relatively  
 314 fast and efficient pace. When compared with our previous method, we were able to improve the  
 315 segmentation accuracy due to the changes made to the initialization process. As a large portion of the nasal  
 cavity consists of thin and narrow pathways that are often located in very close proximity to each other,  
 where sometimes the distance is as thin as two voxels apart, it is especially important to be able to clearly



316 define the foreground and background seeds during initialization. With the use of superpixel clusters, we  
317 were able to minimize over-extension and ensure a more robust selection of foreground seeds.



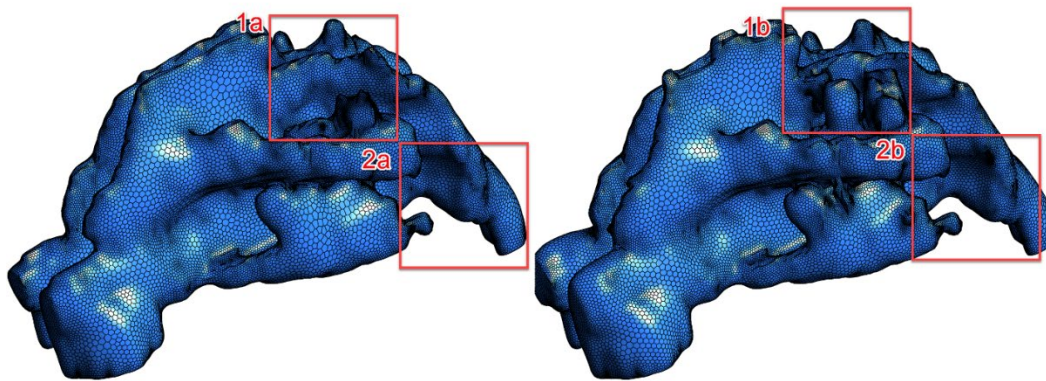
318

319 **Figure 7:** Examples of some of the worst segmentation errors with ground truth labels for comparison. The left side (case 1) depicts  
320 a commonly encountered error of leakage into ethmoid sinuses, while the right side (case 9) depicts a rare over segmentation error  
321 that expanded into the maxillary sinuses.

322 When comparing our results with ground truth segmentation, we noticed that despite having leveraged the  
323 use of a level set SSM, it was still difficult for our method to accurately label some of the boundaries,  
324 especially at the regions surrounding the superior meatus. Based on CT observations, these regions of the  
325 nasal cavity are often in very close proximity to the ethmoid sinuses. Entrances connecting the two airway  
326 components are not fixed, which makes it difficult for an algorithm to correctly judge when to cut the  
327 segmentation. The majority of our results have experienced some form of leakage at this region of the nasal  
328 cavity. The left side of figure 7 shows an example of this problem where parts of the segmentation leaks  
329 into the ethmoid sinuses. The example shown on the right side highlights the worst case of over-  
330 segmentation where due to similar issues, would sometimes cause leakage into the maxillary sinuses.  
331 However, even for our worst case, the size of the leakage was kept relatively contained due to the effect of  
332 our SSM.

333 As the primary objective of our study was centred on creating patient-specific nasal models to conduct CFD  
334 simulations, it was important for our framework to be able to reduce as many of the steps needed prior to  
335 conducting the simulations. Prior to adding the post-processing method component to our framework, we  
336 analysed the key differences between our segmented results and the ground truth models. The first apparent  
337 difference discovered was in the cell count of the mesh. As a result of the sensitivity of the segmentation  
338 process, a large majority of the segmented models had artefactual regions in their morphology, which do  
339 not appear in the ground truth models. Most of these artefacts appear to represent paranasal sinus cells,  
340 particularly ethmoid cells above the middle turbinate, such as in case 12. In many cases, these cells are  
341 connected to the main body of the nasal cavity by narrow channels which when meshed produce large cell  
342 counts because many small-sized cells are required to resolve these regions. During manual segmentation,  
343 these cells are typically excluded from segmentation. We were able to reduce this issue by adding an extra  
344 post-processing cleaning step where algorithms such as 6-way connected regions were applied on the  
345 segmentation outputs in order to remove those that were in the neighbourhood of almost touching or  
346 clipping the central superior nasal airway.

347

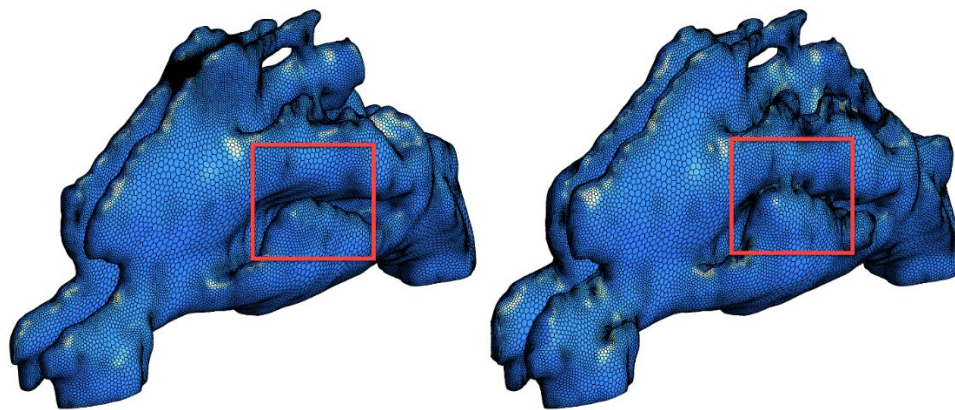


348

349 **Figure 8:** The geometry of the nasal cavity for case 12. The model on the left is the ground truth model and that on the right is the  
350 segmented model. Region 1a and 1b demonstrate the ethmoid sinus region on the model and region 2a and 2b demonstrate lateral  
351 enlargement of the posterior end of the inferior turbinate restricting airflow to the nasopharynx.

352

353 Nevertheless, it remained difficult to fully remove these artefact cells as it would risk exclusion of the  
354 narrow central superior nasal airway that passes immediately between the ethmoid region and the septum.  
355 Looking at case 12 in particular (Figure 8), it was observed that it contained a number of such artefacts.  
356 Additionally, region 2a and 2b revealed a disconnection between the airflow beneath and around the inferior  
357 turbinate and the nasopharynx which although is consistent in both the segmented and ground truth models,  
358 it is not commonly observed in other nasal pairs. When examining the CT image for case 12 in closer detail,  
359 we noticed lateral enlargement of the posterior end of the inferior turbinate in both nasal cavities, restricting  
360 airflow to the nasopharynx and causing both the ground truth and segmented models to be disconnected in  
361 that particular section. In addition, significant medial obstruction in the right nasal cavity of the patient was  
362 observed, resulting in marked reduction in airflow. These sites of obstruction were the primary reason for  
363 the higher pressure drop and increased particle retention which remained consistent in both segmented and  
364 ground truth models. In all other models, the largest pressure drops were noted in the vestibule and nasal  
365 valve area, as is typically seen in the nasal CFD literature [34, 35]. If all models were chosen to represent  
366 clear noses, case 12 would be omitted.



367  
368 **Figure 9:** The nasal cavity geometry for case 21. The model on the left is the ground truth model and that on the right is the  
369 segmented model. The region highlighted by the red square shows a narrow connection formed between the air passage beneath  
370 and above the inferior turbinate in the segmented model.

371  
372 In addition to incorporating ethmoid cells within the segmented model, another artefact was identified,  
373 whereby the segmented model would rarely connect adjacent passages which would normally be separated

374 by very thin tissue. For example, in case 21, in the segmented model, a narrow connection was formed  
375 between the air passage beneath and above the inferior turbinate (Figure 9) that does not appear in the  
376 ground truth model. Although the artefact captured in this example appears small and creates little impact  
377 on the overall nasal segmentation accuracy, its effect on the internal structure remains significant. Such  
378 artefacts create small holes within the geometry allowing flow to pass through, creating pressure and  
379 velocity differences between the ground truth and segmented models. It is an important issue to address  
380 when designing segmentation algorithms with the intent of performing CFD simulations, as artefacts such  
381 as these would alter the expected physical air flow patterns contrary to the actual nasal structure of the  
382 patients from which the CT scans are obtained.

### 383 **5.1 Other work**

384 While the majority of our discussions have been on the accuracy and limitations of our proposed framework,  
385 it is important to also discuss and compare with other related works in general. In the work presented by  
386 Last et al. [13], they made use of a parametric level-set based deformable model to segment the region of  
387 interest (ROI) containing the nasal cavity and paranasal sinuses. Different to other studies, their intended  
388 segmentation target was not just the airway passages but also included the bones and tissues situated within  
389 the nasal region, as the aim of their segmentation was to identify the critical structures in robot assisted  
390 functional endoscopic sinus surgery. The base component of their method was the same as our segmentation  
391 framework where an SSM was constructed from training data. However, the segmentation approach they  
392 employed was a geodesic contour based method [25] and they segmented the CT image one 2D slice at a  
393 time, breaking down the targeted 3D structure into individual components. Such an approach is capable of  
394 achieving high performance for 2D objects but inherently has a higher probability of sharp local jumps in  
395 the combined 3D result.

396 Keustermans et al. [21] employed an ASM for their nasal cavity segmentation. Different to level-set  
397 methods, ASM makes use of landmarks for SSM construction. This approach was highly popular due to  
398 the memory efficiency of using a fewer number of features called “landmarks” to represent a modelled

399 structure, whereas for level-set SSMs every voxel has a corresponding SDM representation. Both  
400 approaches have their strength and weaknesses. For landmark approaches, it is difficult to model complex  
401 shapes with high degrees of variation such as the nasal cavity. While for level-set approaches, larger  
402 numbers of eigen-modes are required due to the intrinsic nature of modelling the variations on the space of  
403 embedded contours. Nevertheless, the flexibility offered from level-set SSMs are higher as they have higher  
404 degrees of freedom and be combined with a number of other methods for segmentation to overcome some  
405 of its weaknesses.

## 406 **6. Conclusions**

407 This paper presents an efficient automated framework for the 3D segmentation of the nasal cavity,  
408 optimized for CFD modelling. Our framework achieved a segmentation accuracy of 90.9 DSC and an  
409 average distance error of 0.3 mm. The nasal models generated from our framework were further evaluated  
410 by calculating flow and droplet impact behaviour using the CFD model ANSYS Fluent in comparison with  
411 the models constructed from ground truth segmentation. Similar outcomes were observed for pressure drop  
412 and particle deposition efficiency between the two groups. We note that the proposed method still shows  
413 some differences that affect the flow locally and we are currently working on methods to detect these using  
414 the CFD data.

## 415 **Conflict of Interest**

416 Robin Huang, Anthony Nedanoski, David Fletcher, Jerome Schmid, Paul Young, Nicholas Stow, Lei Bi,  
417 Daniela Traini, Eugene Wong, Craig Phillips, Ron Grunstein, Narinder Singh, and Jinman Kim declare  
418 that they have no conflict of interest.

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422 **References**

423 [1] N. Mygind and R. Dahl, "Anatomy, physiology and function of the nasal cavities in health and  
424 disease," *Advanced drug delivery reviews*, vol. 29, no. 1-2, pp. 3-12, 1998.

425 [2] N. Bhattacharyya, "Ambulatory sinus and nasal surgery in the United States: demographics and  
426 perioperative outcomes," *The Laryngoscope*, vol. 120, no. 3, pp. 635-638, 2010.

427 [3] H. Dommerby, O. R. Rasmussen, and J. Rosborg, "Long-term results of septoplasty operations,"  
428 *ORL*, vol. 47, no. 3, pp. 151-157, 1985.

429 [4] A. Singh, N. Patel, G. Kenyon, and G. Donaldson, "Is there objective evidence that septal surgery  
430 improves nasal airflow?," *The Journal of Laryngology & Otolaryngology*, vol. 120, no. 11, pp. 916-920,  
431 2006.

432 [5] A. AT Borojeni, D. O. Frank - Ito, J. S. Kimbell, J. S. Rhee, and G. J. Garcia, "Creation of an  
433 idealized nasopharynx geometry for accurate computational fluid dynamics simulations of nasal  
434 airflow in patient - specific models lacking the nasopharynx anatomy," *International journal for  
435 numerical methods in biomedical engineering*, vol. 33, no. 5, p. e2825, 2017.

436 [6] S. K. Kim, Y. Na, J.-I. Kim, and S.-K. Chung, "Patient specific CFD models of nasal airflow:  
437 overview of methods and challenges," *Journal of biomechanics*, vol. 46, no. 2, pp. 299-306, 2013.

438 [7] J. Cisonni, A. D. Lucey, A. J. King, S. M. S. Islam, R. Lewis, and M. S. Goonewardene, "Numerical  
439 simulation of pharyngeal airflow applied to obstructive sleep apnea: effect of the nasal cavity in  
440 anatomically accurate airway models," *Medical & biological engineering & computing*, pp. 1-11,  
441 2015.

442 [8] G. J. de Moraes Garcia, N. A. Bailie, D. A. Martins, and J. S. Kimbell, "Atrophic rhinitis: a CFD  
443 study of air conditioning in the nasal cavity," *Journal of applied physiology*, 2007.

444 [9] P. W. Longest and S. Vinchurkar, "Validating CFD predictions of respiratory aerosol deposition:  
445 effects of upstream transition and turbulence," *Journal of biomechanics*, vol. 40, no. 2, pp. 305-  
446 316, 2007.

447 [10] J. Wen, K. Inthavong, J. Tu, and S. Wang, "Numerical simulations for detailed airflow dynamics  
448 in a human nasal cavity," *Respiratory physiology & neurobiology*, vol. 161, no. 2, pp. 125-135,  
449 2008.

450 [11] R. Huang *et al.*, "A Locally Constrained Statistical Shape Model for Robust Nasal Cavity  
451 Segmentation in Computed Tomography," in *International Symposium on Biomedical Imaging:  
452 From Nano to Macro (ISBI)*, 2016, pp. 1-4: IEEE.

453 [12] N. L. Bui, S. H. Ong, and K. W. C. Foong, "Automatic segmentation of the nasal cavity and  
454 paranasal sinuses from cone-beam CT images," *International journal of computer assisted  
455 radiology and surgery*, vol. 10, no. 8, pp. 1269-1277, 2015.

456 [13] C. Last, S. Winkelbach, F. M. Wahl, K. W. Eichhorn, and F. Bootz, "A model-based approach to  
457 the segmentation of nasal cavity and paranasal sinus boundaries," in *Pattern Recognition*: Springer,  
458 2010, pp. 333-342.

459 [14] K. Tingelhoff *et al.*, "Comparison between manual and semi-automatic segmentation of nasal  
460 cavity and paranasal sinuses from CT images," in *2007 29th Annual International Conference of  
461 the IEEE Engineering in Medicine and Biology Society*, 2007, pp. 5505-5508: IEEE.

462 [15] A. S. El, H. El, J. M. Palomo, and D. A. Baur, "A 3-dimensional airway analysis of an obstructive  
463 sleep apnea surgical correction with cone beam computed tomography," *Journal of Oral and  
464 Maxillofacial Surgery*, vol. 69, no. 9, pp. 2424-2436, 2011.

465 [16] T. Iwasaki *et al.*, "Evaluation of upper airway obstruction in Class II children with fluid-mechanical  
466 simulation," *American Journal of Orthodontics and Dentofacial Orthopedics*, vol. 139, no. 2, pp.  
467 e135-e145, 2011.

468 [17] A. Seo, S. Chung, J. Lee, J.-I. Kim, and H. Kim, "Semiautomatic segmentation of nasal airway  
469 based on collaborative environment," in *Ubiquitous Virtual Reality (ISUVR), 2010 International  
470 Symposium on*, 2010, pp. 56-59: IEEE.

- 471 [18] P. Dastidar, T. Heinonen, J. Numminen, M. Rautiainen, and E. Laasonen, "Semi-automatic  
472 segmentation of computed tomographic images in volumetric estimation of nasal airway,"  
473 *European archives of oto-rhino-laryngology*, vol. 256, no. 4, pp. 192-198, 1999.
- 474 [19] K. Tingelhoff *et al.*, "Comparison between manual and semi-automatic segmentation of nasal  
475 cavity and paranasal sinuses from CT images," in *Engineering in Medicine and Biology Society,  
476 2007. EMBS 2007. 29th Annual International Conference of the IEEE*, 2007, pp. 5505-5508: IEEE.
- 477 [20] S. Kimura *et al.*, "Voxel-based modeling of airflow in the human nasal cavity," *Computer methods  
478 in biomechanics and biomedical engineering*, pp. 1-9, 2019.
- 479 [21] W. Keustermans, T. Huysmans, B. Schmelzer, J. Sijbers, and J. J. Dirckx, "Matlab® toolbox for  
480 semi-automatic segmentation of the human nasal cavity based on active shape modeling,"  
481 *Computers in biology and medicine*, vol. 105, pp. 27-38, 2019.
- 482 [22] N. A. Alsufyani *et al.*, "New algorithm for semiautomatic segmentation of nasal cavity and  
483 pharyngeal airway in comparison with manual segmentation using cone-beam computed  
484 tomography," *American Journal of Orthodontics and Dentofacial Orthopedics*, vol. 150, no. 4, pp.  
485 703-712, 2016.
- 486 [23] T. F. Cootes, C. J. Taylor, D. H. Cooper, and J. Graham, "Active shape models-their training and  
487 application," *Computer vision and image understanding*, vol. 61, no. 1, pp. 38-59, 1995.
- 488 [24] T. Heimann and H.-P. Meinzer, "Statistical shape models for 3D medical image segmentation: a  
489 review," *Medical image analysis*, vol. 13, no. 4, pp. 543-563, 2009.
- 490 [25] M. E. Leventon, W. E. L. Grimson, and O. Faugeras, "Statistical shape influence in geodesic active  
491 contours," in *Computer Vision and Pattern Recognition, 2000. Proceedings. IEEE Conference on,  
492 2000*, vol. 1, pp. 316-323: IEEE.
- 493 [26] A. Wimmer, G. Soza, and J. Hornegger, "A generic probabilistic active shape model for organ  
494 segmentation," in *International Conference on Medical Image Computing and Computer-Assisted  
495 Intervention*, 2009, pp. 26-33: Springer.
- 496 [27] D. Cremers, M. Rousson, and R. Deriche, "A review of statistical approaches to level set  
497 segmentation: integrating color, texture, motion and shape," *International journal of computer  
498 vision*, vol. 72, no. 2, pp. 195-215, 2007.
- 499 [28] R. Achanta, A. Shaji, K. Smith, A. Lucchi, P. Fua, and S. Süsstrunk, "SLIC superpixels compared  
500 to state-of-the-art superpixel methods," *IEEE transactions on pattern analysis and machine  
501 intelligence*, vol. 34, no. 11, pp. 2274-2282, 2012.
- 502 [29] S. Klein, M. Staring, K. Murphy, M. A. Viergever, and J. P. Pluim, "Elastix: a toolbox for intensity-  
503 based medical image registration," *Medical Imaging, IEEE Transactions on*, vol. 29, no. 1, pp. 196-  
504 205, 2010.
- 505 [30] L. Grady, "Random walks for image segmentation," *Pattern Analysis and Machine Intelligence,  
506 IEEE Transactions on*, vol. 28, no. 11, pp. 1768-1783, 2006.
- 507 [31] A. Criminisi, T. Sharp, and A. Blake, "Geos: Geodesic image segmentation," in *Computer Vision-  
508 ECCV 2008*: Springer, 2008, pp. 99-112.
- 509 [32] T. Heimann *et al.*, "Comparison and evaluation of methods for liver segmentation from CT  
510 datasets," *Medical Imaging, IEEE Transactions on*, vol. 28, no. 8, pp. 1251-1265, 2009.
- 511 [33] L. Engelhardt, M. Röhm, C. Mavoungou, K. Schindowski, A. Schafmeister, and U. Simon, "First  
512 steps to develop and validate a CFPD model in order to support the design of nose-to-brain  
513 delivered biopharmaceuticals," *Pharmaceutical research*, vol. 33, no. 6, pp. 1337-1350, 2016.
- 514 [34] C. Croce *et al.*, "In vitro experiments and numerical simulations of airflow in realistic nasal airway  
515 geometry," *Annals of biomedical engineering*, vol. 34, no. 6, pp. 997-1007, 2006.
- 516 [35] D. Doorly, D. Taylor, and R. Schroter, "Mechanics of airflow in the human nasal airways,"  
517 *Respiratory physiology & neurobiology*, vol. 163, no. 1-3, pp. 100-110, 2008.

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