

Open access segmentations of intraoperative brain tumor ultrasound images

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▶ To cite this version:

Bahareh Behboodi, Francois-xavier Carton, Matthieu Chabanas, Sandrine de Ribaupierre, Ole Solheim, et al.. Open access segmentations of intraoperative brain tumor ultrasound images. Medical Physics, 2024, $10.1002/{\rm mp.17317}$. hal-04661871

HAL Id: hal-04661871 https://hal.science/hal-04661871v1

Submitted on 25 Jul 2024

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¹ Open access segmentations of intra-operative brain tumor ² ultrasound images

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23	Abstract
24	Purpose: Registration and segmentation of magnetic resonance (MR) and ultrasound
25	(US) images could play an essential role in surgical planning and resectioning brain
26	tumors. However, validating these techniques is challenging due to the scarcity of pub-
27	licly accessible sources with high-quality ground truth information. To this end, we
28	propose a unique set of segmentations (RESECT-SEG) of cerebral structures from the
29	previously published RESECT dataset to encourage a more rigorous development and
30	assessment of image processing techniques for neurosurgery.
31	Acquisition and Validation Methods: The RESECT database consists of MR and
32	intra-operative US (iUS) images of 23 patients who underwent brain tumor resection
33	surgeries. The proposed RESECT-SEG dataset contains segmentations of tumor tis-
34	sues, sulci, falx cerebri, and resection cavity of the RESECT iUS images. Two highly
35	experienced neurosurgeons validated the quality of the segmentations.
36	Data Format and Usage Notes: Segmentations are provided in 3D NIFTI format
37	in the OSF open-science platform: https://osf.io/jv8bk.
38	Potential Applications: The proposed RESECT-SEG dataset includes segmenta-
39	tions of real-world clinical US brain images that could be used to develop and evaluate
40	segmentation and registration methods. Eventually, this dataset could further improve
41	the quality of image guidance in neurosurgery.

42 I. Introduction

Gliomas are the most common malignant primary brain tumors originating from glial cells 43 and are classified into grades 1-4 by the World Health Organization (WHO)^{1,2}. Grades 1-44 are low-grade, while grades 3-4 are high-grade tumors³. Surgical resection is a standard 2 45 treatment for gliomas, and pre-operative magnetic resonance (MR) imaging is used for tumor 46 characterization. However, brain tissue deforms during surgery due to factors like edema and 47 gravity (i.e. brain shift)⁴, rendering pre-operative MR images inaccurate. Acquiring data at 48 different stages during surgery helps the surgeon better monitor the progress of the tumor 49 resection and, consequently, operate more precisely. Intra-operative imaging, particularly 50 intra-operative MR (iMR) and intra-operative ultrasound (iUS) aids surgeons by providing 51 updated guidance^{5,6,7}. While iMR offers superior image quality, it is costly, adds a long 52 time to the operation, and requires dedicated operating rooms^{8,9}. In contrast, iUS is a cost-53 effective, flexible, and versatile modality that presents real-time scanning without altering 54 the surgical workflow^{10,11,12}. Due to the easy procedure of acquiring iUS rather than iMR, 55 several recent studies have demonstrated the use and interest in iUS in neurosurgery^{8,11,12,13}. 56

While iUS presents several advantages in the context of brain tumor resection, ultra-57 sound (US) images can be difficult to interpret. Non-standard imaging planes and unfamiliar 58 contrast are major factors limiting the efficient and widespread use of US in neurosurgery. 59 To mitigate such limitations and fully leverage the advantages of iUS, automated image seg-60 mentation of structures such as tumors within iUS images can provide valuable assistance to 61 neurosurgeons during procedures. Recent automated image segmentation algorithms, such 62 as deep learning (DL) algorithms have made advancements in brain tumor segmentation 63 from both US^{14,15,16,17} and MR^{18,19,20,21} images. However, access to high-quality datasets ex-64 pertly annotated is essential for the development and validation of DL algorithms²². In the 65 context of medical imaging. MR images have seen more readily available datasets compared 66 to other modalities, making them the primary focus for DL algorithm development. The 67 BRATS challenge, among others, stands out as a prominent dataset with valuable annota-68 tions that have significantly contributed to the evolution and refinement of DL algorithms²³. 69 Acquiring such data, especially for iUS images, is expensive and rare. 70

Currently, there are only three publicly available datasets that provide iUS brain im-71 ages, the BITE dataset²⁴, the RESECT database²⁵, and ReMIND²⁶. The BITE dataset 72 contains pre- and post-operative MR scans as well as multiple iUS images of 14 patients. 73 The RESECT database contains pre-operative MR scans and iUS images from 23 patients 74 with low-grade gliomas. The ReMIND dataset contains 369 pre-operative MR scans, 320 3D 75 iUS scans, 301 iMR scans, and 356 pre-operative MR segmentations of 114 patients. None of 76 the abovementioned datasets contain segmentation of anatomical structures in the iUS im-77 ages, thereby hindering the development and validation of iUS processing methods. For the 78 RESECT database, a few research groups have previously conducted segmentations of iUS 79 images^{14,18,27}, However, a portion of these annotations remained inaccessible to the public, 80 and in some instances, only a small subset of cases was segmented, with limited validation 81 procedures in place. 82

In this work, we present the most comprehensive and validated expert segmentations of 83 cerebral structures in iUS images from the RESECT database. The focus of the RESECT-84 SEG dataset is on delineating the tumor in pre-resection iUS 3D volumes and identifying the 85 resection cavity during and after the surgery. To enhance the surgeon's ability to achieve 86 more precise tumor resection, *sulci* and the *falx cerebri*, whenever they were within the 87 field of view, were also delineated. These structures commonly serve as crucial anatomical 88 references for surgeons, given their clear visibility in iUS images. The following sections 89 detail the segmentation and validation protocols for all the mentioned structures in the iUS 90 images and brain tumor segmentation in pre-operative MR images. 91

⁹² II. Acquisition and Validation Methods

In this section, a comprehensive overview of the dataset is presented, along with detailed an-93 notations for various anatomical structures, including tumors, resection cavities, falx cerebri, 94 and *sulci*. The annotation process involved the utilization of two primary tools: ITK-SNAP²⁸ 95 and 3D Slicer²⁹, chosen based on individual preference and familiarity. Furthermore, specific 96 built-in features of these software platforms were leveraged as necessary, such as smoothing 97 and interpolation functionalities, to enhance the accuracy and completeness of the annota-98 tions. Further elaboration on these tools and their respective functionalities is provided in 99 the subsequent paragraphs. Given that both ITK-SNAP and 3D Slicer are widely used tools 100 in the field, the decision to select one over the other was solely based on our inter-group 101 preferences. It is important to highlight that manual segmentation entails human judgment 102 and expertise, enabling nuanced interpretation and adjustments tailored to the anatomical 103 complexities of each case. Consequently, the choice of segmentation software or algorithm 104 does not introduce bias, as it depends on the proficiency and diligence of the annotators. 105

¹⁰⁶ II.A. RESECT Database

The RESECT database²⁵ comprises pre-operative contrast-enhanced T1-weighted and T2 107 FLAIR MR scans alongside three 3D volumes of iUS scans from 23 patients with low-grade 108 gliomas (grade 2) who underwent surgeries between 2011 and 2016 at St. Olavs University 109 Hospital, Trondheim, Norway. The iUS scans were acquired at three different stages of 110 the procedure: before resection, during resection, and after resection for control. These US 111 images were captured by an expert surgeon, and the database includes manual neuroanatomy 112 landmarks, facilitating MR-to-US volume registration and inter-US volume alignment. The 113 details of the image acquisition procedure can be summarized as follows: 114

Pre-operative MR scans: T1-weighted and T2 FLAIR sequences were acquired on 3T
 Magnetom Skyra MR scanners, both with 1 mm isotropic voxel size, except three
 patients who underwent the MR imaging on a 1.5T Magnetom Avanto MR scanner
 with 1 mm slice thickness.

• Intra-operative US scans: 3D US images collected using a 12FLA-L linear probe of Sonowand Invite neuronavigation system with a frequency range of 6-12 MHz integrated with the NDI Polaris optical tracking system.

¹²² II.B. iUS Tumor Segmentation Protocol

Tumoral tissue in US images is typically identified through abnormal echogenicity or tex-123 ture variations compared to healthy tissue. Echogenicity refers to the level of reflectivity 124 or brightness of tissue on a US image. In this context, variations in echogenicity indicate 125 potential areas of malignancy, allowing medical professionals to identify and examine po-126 tential cancerous lesions. In the study, 19 out of 23 cases' iUS images (cases 1 to 23) were 127 segmented, initially following Munkvold et al.'s method²⁷. In these cases, initial US volume 128 segmentations were already available. Four cases (cases 24 to 27) without prior iUS segmen-129 tations relied on MR segmentations to define the tumor region of interest in iUS images. 130 To be more specific, for these four cases, the MR segmentation served as a starting point to 131 define the region of interest (ROI) for the tumor in the iUS images. Further details of the 132 MR segmentation protocol can be found in the work of Munkvold et al.²⁷. 133

In the iUS segmentation protocol, 3D Slicer²⁹, a free and open-source medical image 134 analysis software, was employed to perform ground truth segmentations on the acquired 135 iUS images. For cases 24 to 27 that were initiated with MR segmentations due to brain 136 shift during resection surgery, the boundaries of the tumor in the US images and MR tumor 137 segmentations did not align²⁵. This discrepancy necessitated the registration of MR tumor 138 segmentations to iUS images using available landmarks from the RESECT database. This 139 registration process effectively mitigated the misalignment of tumor borders between the iUS 140 images and their corresponding MR tumor segmentations. Therefore, for cases 24 to 27, the 141 MR segmentations were imported into the 3D Slicer scene and after the registration step. 142 they were utilized as the initial delineation for iUS. 143

In all cases, to refine the initial 3D US tumor segmentations, the Label Map Smoothing module, an existing feature in 3D Slicer, was employed. Subsequently, the smoothed tumor segmentations were manually fine-tuned to ensure accurate coverage of the tumor region in the iUS image. To facilitate further correction, FLAIR MR images were registered to iUS images since MR images were unaffected by brain-shift effects. These complementary overlays served as guidance for generating more precise iUS tumor segmentations. An illustrative example of tumor segmentation is provided in Fig. 3 (a)-(d).

¹⁵¹ II.C. iUS Resection Cavity Segmentation Protocol

Resection cavity segmentation in iUS images encompasses the volume where tissue has been resected or retracted during image acquisition. Resection, a surgical procedure characterized by the complete removal of tissue or tumors, contributes to the formation of a distinct three-dimensional space within the brain known as the resection cavity. On the other hand, retraction, another surgical maneuver, entails cutting tissue and displacing it to the side



Figure 1: Ultrasound image of a resection cavity (a) with and (b) without segmentation.

without complete removal. Hyperechoic signals surrounding cavities in iUS images result 157 from sound attenuation differences between brain tissue and saline water, as well as the 158 presence of blood (see Fig. 1)³⁰. To prevent false positives, only homogeneous, dark signals 159 were considered as cavities, potentially leading to slight underestimation in cases involving 160 blood-filled cavities. In challenging cases with small, entirely blood-filled cavities, segmen-161 tation was not possible due to indistinguishable borders. For example, in three exceptional 162 instances (Case 11 during and after resection, and Case 15 during resection), the cavities 163 appeared notably small and completely inundated with blood, without any noticeable dark 164 signals. 165

It is worth noting that, due to the inherent variability in surgical procedures, determin-166 ing the precise timing of image capture was challenging since it could occur at different stages 167 of the resection process. The crucial factor was ensuring that US images were taken before 168 the surgeon completed the resection entirely, even if residual tumors remained. The segmen-169 tation of resection cavities was conducted using ITK-SNAP²⁸. Initially, regularly spaced 170 slices, approximately one in every five slices, were manually delineated. Subsequently, ITK's 171 morphological interpolation, facilitated by ITK-SNAP's Convert3D command-line tool, was 172 utilized to fill in the remaining slices. Convert3D is one of the companion tools of ITK-SNAP 173 that provides additional features. It is a command-line tool that enables the combination 174 of multiple image processing tasks into efficient mini-programs, making it an integral tool 175 in studies involving hundreds of 3D images. In cases where necessary, additional slices were 176 manually segmented to optimize the interpolation outcome. The segmentation process for 177 most RESECT cases was originally carried out by two raters as part of a previous study¹⁴. 178 Following an assessment of intra- and inter-rater variability, these segmentations were re-179 viewed by a neurosurgeon and were then modified accordingly. Subsequently, for this study, 180 the remaining RESECT cases were segmented, and all cases underwent refinement during 181 the validation protocol detailed in section II.G.. 182



Figure 2: T1 MR (a), (b), and US (c) images of the falx cerebri.

¹⁸³ II.D. iUS *Falx Cerebri* Segmentation Protocol

The *falx cerebri* is the membrane that separates the left and right hemispheres of the brain. 184 This cerebral falx has a hyperechoic signal in iUS images. It presents a characteristic quasi-185 planar shape that appears as a straight line in coronal and axial slices. The falx is also visible 186 in the MR images, especially T1-weighted (Fig. 2). This structure is thus a convenient 187 landmark that can be particularly useful to anchor registration. The falx is not always 188 within the iUS volume due to the limited field of view but can be visible depending on the 189 tumor location. The falx segmentation is, therefore, present for some volumes only. Since 190 the falx' bright signal is similar to *sulci* it is difficult to localize the inferior border of the 191 membrane. We, therefore, used the registered T1-weighted MR images to adjust the falx 192 segmentation in height. Regularly spaced slices were manually delineated using ITK-SNAP 193 and then interpolated with Convert3D. 194

¹⁹⁵ II.E. iUS *Sulci* Segmentation Protocol

For segmentation purposes, *sulci* were defined as folds filled with cerebrospinal fluid (CSF) 196 between brain tissue sections. CSF surrounding the brain, such as between the tissue and the 197 dura mater or tentorium, was not labeled, although it had a similar iUS signal. Sulci were 198 initially segmented with manual delineation every five slices and morphological interpolation 199 using Convert3D when moving through the volume in a single direction (e.g., axial slices). 200 Unlike volumetric structures, *sulci* are thin, complex, folded surfaces. To capture their 201 irregular shapes, each volume was annotated in three viewing directions (axial, sagittal, and 202 coronal), and these segmentations were first interpolated separately and later combined into 203 a union. While this process resulted in slight over-segmentation, it significantly improved 204 sulci delineation, according to annotators and neurosurgeons. In each case, the volume 205 preceding resection underwent segmentation aided by registered MR images to accurately 206 delineate sulci structures. This initial segmentation served as a reference for segmenting 207 the volume during and after resection. The manual segmentation process was facilitated by 208 ITK-SNAP, employing a Wacom One pen tablet, which demonstrated superior speed and 209 precision compared to a conventional computer mouse. 210

²¹¹ II.F. Pre-operative MR Tumor Segmentation

For completeness, we provide segmentations of the tumors in the pre-operative T2 FLAIR images. As the cases in the database are lower-grade gliomas, there is no contrast uptake in the T1 weighted images and the T2 FLAIR images are used to define the tumor boundaries. The tumors were semi-automatically segmented in 3DSlicer using the *GrowCut* algorithm³¹. The resulting segmentations were manually corrected when needed and smoothed with a 2×2 mm median filter.

218 II.G. Data Validation

All segmentations presented in this work were validated by two experienced neurosurgeons (S.D.R., O.S.). The segmentations were presented to the specialists through a case-bycase 3D Slicer scene, including the original iUS image, segmentation masks, and MR images. Figure 3 represents an example of such a scene. We asked specialists to grade all segmentation masks based on five criteria for three types of structures:

- Quality of tumor: smoothness of the boundaries (SMT), identification of tumoral tissues (IdT), exclusion of non-cancerous tissues (ExT)
- Quality of resection cavity: identification of resection cavity (IdR)
- Quality of *sulci* and falx: identification of sulci and falx (IdS)

The grading scheme for each criterion was on a scale of 1 to 5 defined as major improvement 228 needed, minor improvement needed, acceptable, good quality, and excellent, respectively. 229 For each criterion, a score of 3 from both surgeons was needed to pass the quality control 230 of segmentation masks. Otherwise, the masks were revised according to the surgeons' com-231 ments. In determining the choice of a passing score of 3, it is important to clarify that this 232 decision was rooted in the specific criteria established for surgeons evaluating the dataset. 233 The selection of 3 as the pass score was deliberate, as it represented the midpoint on the 234 validation scale of 1 to 5. This choice was informed by the summary of evaluation forms, 235 aiming to identify a common rating among inter-rater assessments. Given the intricate and 236 complex nature of brain structures, achieving consensus on a moderate score like 3 ensured 237 a balanced assessment that accounted for the variability inherent in such evaluations. The 238 final grades for each patient are presented in Table 1. 239



Figure 3: An example of segmentations overlaid with intra-operative ultrasound (iUS) and MR images (green: tumor; yellow: sulci; red: cerebral falx; blue: resection cavity). iUS volume before resection: (a)-(d); iUS volume during resection: (e)-(h).

²⁴⁰ III. Data Format and Usage Notes

The proposed RESECT-SEG segmentations are distributed in the NIFTI format. Upon the acceptance of this paper, they will be available via the OSF open-science https://osf.io/ jv8bk for public viewing and downloading and can be freely used by research laboratories as well as clinical institutes. However, gaining any financial benefits from the distribution of the proposed segmentation dataset is prohibited. The database is under the CC BY-NC-SA 4.0 License.

²⁴⁷ IV. Discussion

The border and shape of brain tumors have long been established as important diagnostic 248 markers in resection surgeries. Several image processing techniques have been adopted to 249 segment tumors which rely on the creation of mathematical descriptions of the tumor border. 250 Similarly, identifying resection cavity contours has been used to evaluate the completeness 251 of tumor resection. Finally, segmenting surrounding cerebral structures can greatly bene-252 fit image analysis during the surgeries. However, validation of image processing techniques 253 needs to be investigated in the case of new data. It is important to highlight that there are 254 few brain datasets accessible to the public, and even those available, such as the $BITE^{24}$. 255 RESECT²⁵, and ReMIND²⁶ datasets, do not include iUS segmentation of brain anatomies. 256 The absence of US segmentation datasets for the brain is largely attributed to the complexity 257 of the task. To this end, we have provided the manual segmentations of cerebral structures 258 in iUS images of the 23-patient RESECT dataset, verified by two expert surgeons through 259 detailed evaluation criteria per brain anatomy. Our proposed expert-annotated RESECT-260

SEG dataset comprises the segmentation masks of brain tumors, resection cavities, the *falx cerebri*, and *sulci*. Tumor segmentations of the pre-operative MR images are also provided as a reference. To the best of our knowledge, this is the first study that publicly provides a comprehensive expert-annotation segmentation of iUS images. The challenging procedure of delineating brain iUS images has impeded the publications of such segmentations. This validated dataset serves as a crucial asset for evaluating and benchmarking various segmentation methods, thereby driving advancements in brain imaging research.

Table 1: Quality control grade chart for segmentation masks before, during, and after resection. Grades of the two neurosurgeons are given side-by-side in each cell. For Case 11 (during and after resection) and Case 15 (during resection), the resection cavity was not labeled (see section II.C.).

Validation Scores										
	Before Resection				During Resection		After Resection			
Patient ID	SMT	IdT	ExT	IdS	IdR	IdS	IdR	IdS		
1	44	45	43	34	$3 \ 3$	44	3 4	$4 \ 4$		
2	44	$4\ 3$	$3 \ 5$	44	3 4	4 4	35	45		
3	44	$3 \ 3$	$4\ 3$	33	3 4	33	44	43		
4	34	35	45	34	3 4	4 4	4 5	$3 \ 4$		
5	34	$4 \ 4$	44	34	3 5	$3\ 4$	4 5	4 3		
6	34	45	$3 \ 5$	34	45	4 4	4 5	4 4		
7	34	$3 \ 5$	44	34	$3 \ 3$	4 4	4 5	4 4		
8	43	$4 \ 4$	45	34	3 5	$3\ 4$	3 5	4 4		
11	44	$4 \ 4$	$3 \ 5$	44		$4 \ 3$		4 3		
12	34	$4\ 3$	$4\ 3$	44	$3\ 4$	$3\ 4$	4 5	45		
13	44	$4 \ 4$	$3\ 4$	43	$3 \ 5$	45	3 5	43		
14	44	$3\ 4$	$4 \ 4$	44	$4 \ 3$	4 4	44	$3 \ 4$		
15	34	$3 \ 3$	$4\ 3$	44		4 4	44	4 4		
16	4 4	$4 \ 4$	$4 \ 4$	44	$3 \ 4$	4 4	34	$3 \ 4$		
17	34	45	45	44	4 5	4 4	34	4 4		
18	44	$3\ 4$	$3\ 4$	44	$3\ 4$	$3\ 4$	3 5	35		
19	34	$3 \ 5$	$3 \ 5$	43	$3 \ 5$	35	3 5	$4 \ 4$		
21	3 4	$3\ 4$	44	3 4	4 5	4 4	4 5	$4 \ 4$		
23	34	$3 \ 5$	$3 \ 5$	34	$3 \ 3$	4 4	3 5	4 4		
24	44	$4 \ 4$	$3 \ 5$	35	$3 \ 4$	4 4	3 4	4 4		
25	44	$3\ 4$	3 4	34	$4 \ 3$	$3\ 4$	3 4	$3 \ 3$		
26	4 5	45	$3 \ 5$	35	$3 \ 5$	35	35	$3 \ 3$		
27	4 5	33	35	44	33	$3\ 4$	35	3 4		

SMT: smoothness of the boundaries, IdT: identification of tumoral tissues, ExT: exclusion of non-cancerous tissues, IdS: identification of sulci and falx. IdR: identification of resection cavity

The proposed RESECT-SEG dataset offers substantial utility, focusing primarily on two pivotal applications that have the potential to revolutionize brain tumor diagnosis and

treatment. As the first application, it accelerates the development of advanced image analy-270 sis algorithms for brain tumor detection and segmentation, whether based on deep learning 271 or energy minimization. With the growing number of segmentation algorithms, there is 272 a need for comprehensive evaluation, and this dataset provides a standardized metric for 273 rigorous testing, propelling advancements in brain cancer treatment. Therefore, the pro-274 posed dataset offers an opportunity for both technical and clinical communities to rigor-275 ously test their algorithms. Additionally, it offers a unique opportunity for algorithms to 276 excel in multi-instance detection and segmentation, enhancing performance beyond binary 277 segmentation tasks. Multiple methodologies are available for binary segmentation problems; 278 however, recent studies suggest that integrating instances into deep learning algorithms not 279 only enhances performance through parallel multi-instance segmentation but also achieves 280 a comprehensive improvement overall³². Consequently, this dataset acts as a catalyst in 281 refining computer-aided diagnosis (CAD) systems, enabling multi-instance and multi-organ 282 analyses, thereby revolutionizing brain tumor diagnosis and treatment. 283

The second application is transformative, focusing on developing and validating 284 segmentation-based registration algorithms to address brain shift challenges during surgery. 285 Brain shift, involving tissue deformation and displacement, poses precision hurdles in surgery. 286 Integrating this dataset into registration algorithms provides them with expert-annotated 287 tumor segmentations as a foundation. These resources empower algorithms to dynamically 288 recalibrate pre-operative images in real-time, aligning them with evolving intraoperative con-289 ditions. The result is an advanced neuronavigation system, offering surgeons accurate, real-290 time patient anatomical visualization. This leads to enhanced resection control, minimizing 291 structural damage risk and optimizing tumor removal. The synergy between segmentation 292 and registration algorithms has the potential to redefine neurosurgery, equipping surgeons 293 with a powerful tool to navigate brain shift complexities, ultimately ensuring safer surgeries, 294 better patient outcomes, and improved resection control. This advancement holds promise 295 for revolutionizing the field of neurosurgery. 296

The proposed RESECT-SEG dataset stands as a pivotal resource for advancing image 297 processing techniques in neurosurgery. While it offers valuable segmentations of cerebral 298 structures, it's essential to acknowledge its limitations. One such concern is the potential 299 lack of representation of diverse clinical scenarios. Ensuring the dataset encapsulates a broad 300 spectrum of anatomical variations, tumor types, and imaging modalities is crucial for its 301 relevance and applicability in broader contexts. Evaluating the generalizability of proposed 302 techniques beyond the confines of the RESECT-SEG dataset is imperative. Nevertheless, 303 by adhering to rigorous evaluation protocols, conducting thorough validation processes, and 304 maintaining transparent reporting standards, the RESECT-SEG dataset can significantly 305 improve its credibility and impact in advancing neurosurgical image processing techniques. 306

³⁰⁷ V. Conclusion

In this study, the most comprehensive and validated expert delineations of cerebral structures within iUS images from the RESECT database were proposed. The primary focus lay in outlining tumor boundaries within pre-resection iUS 3D volumes and tracking the resection cavity both during and post-surgery. Additionally, delineated *sulci* and the *falx cerebri* were further provided to enhance surgical precision. This dataset presents an invaluable resource for both the training and evaluation of DL-based segmentation algorithms and registration methodologies, thereby presenting a rigorous challenge to their capabilities. This collective effort is poised to catalyze advancements in brain tumor treatment and surgical interventions, ultimately benefiting patients and furthering the realm of medical science.

317 Acknowledgements

This work has been supported by the Natural Sciences and Engineering Research Council of Canada (NSERC), the French National Research Agency through the frameworks Computer Assisted Medical Interventions (ANR-11-LABX-0004 CAMI Labex) and the Multidisciplinary Institute in Artificial Intelligence (ANR-19-P3IA-0003 MIAI@Grenoble Alpes).

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