

Semi-automatic CT Image Segmentation using Random Forests Learned from Partial Annotations

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Abstract: Human tissue segmentation is a critical step not only in the process of their visualization and diagnostics but also for pre-operative planning and custom implants engineering. Manual segmentation of three-dimensional data obtained through CT scanning is very time demanding task for clinical experts and therefore the automation of this process is required. Results of fully automatic approaches often lack the required precision in cases of non-standard treatment, which is often the case when computer planning is important, and thus semi-automatic approaches demanding a certain level of expert interaction are being designed. This work presents a semi-automatic method of 3D segmentation applicable to arbitrary tissue that takes several manually annotated slices as an input. These slices are used for training a random forest classifiers to predict the annotation for the remaining part of the CT scan and final segmentation is obtained using the graph-cut method. Precision of the proposed method is evaluated on various CT datasets using fully expert-annotated segmentations of these tissues. Dice coefficient of overlap is 0.976 ± 0.014 for hard tissue segmentation and 0.978 ± 0.008 for kidney segmentation, achieving competitive results with other task-specific methods.

1 INTRODUCTION

Three-dimensional segmentation of hard tissues in medical Computed Tomography (CT) is a first step required not only for reliable 3D visualization for diagnostic purposes but also for precise presurgical planning of orthopedic surgeries (Jun and Choi, 2010; Wu et al., 2014) or craniofacial surgery (Chim et al., 2014; Parthasarathy, 2014). Recently, number of 3D-printable patient-specific implants (Tetsworth et al., 2017) has also started to grow. Since these applications require very accurate anatomical models, the segmentation is often performed by clinical experts slice-by-slice. As this task is very time demanding and tedious considering the number of slices in CT data, the automation of this process is highly desirable. However, obstacles such as low-contrast tissue boundaries and scatter which are usually present in CT data make the automatic segmentation very difficult. Also, methods need to be robust against various pathologies and anatomical variability.

Although fully automatic approaches often yield acceptable results in standard cases where the patient anatomy doesn't strongly deviate from the norm, the

aforementioned applications often require precision in cases of abnormalities such as fractures, lesions etc. where these automatic methods very often fail. Semi-automatic approaches allow the user to introduce a priori case-specific information regarding the desired result of the segmentation and alter the result after the automatic segmentation, in case adjustment is needed.

In this paper, we propose a new semi-automatic method of 3D segmentation of arbitrary tissue in medical CT scans based on Graph-Cut method and Random Forests. The method searches for globally optimal binary segmentation of the volumetric data with respect to probability fields acquired from Random Forest classifiers trained online on several expert-annotated slices. This method yields satisfactory results while using as few as two orthogonal user-annotated input slices as shown in Figure 1, allowing for segmentation of objects with various deformities or pathologies. Additionally, we introduce a novel way of encoding the voxel-wise classification into both the region-term and boundary-term energies of the graph-cut framework.

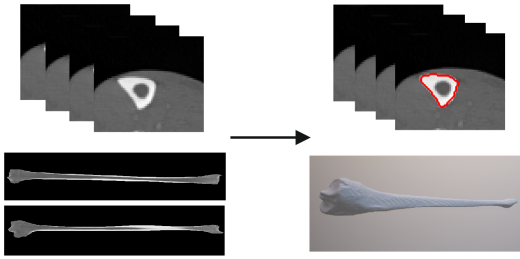


Figure 1: An example of using the method. Input CT data and two orthogonal user-annotated input slices (left) and the segmentation result (right).

2 RELATED WORK

A number of methods of 3D medical data segmentation is available in the literature up to date. The most simple methods are segmentation of hard tissues using adaptive thresholding of acquired Hounsfield units (HU), which was applied to segmentation of long bones (Rathnayaka et al., 2011) as well as cranial bones (Pakdel et al., 2012) and region growing (Xi et al., 2014). All these methods are often used in medical research as well as practice for their simplicity, user-friendliness and easy understanding of the method so behavior of a method can easily be predicted. However, these methods require optimal setting of threshold values and even then they are prone to failure in locations where object boundaries are very thin and low-contrast, as is often the case in CT data due to partial volume artifact.

The more sophisticated approaches based on active contour models such as level-sets (Pinheiro and Alves, 2015) are able to complete the segmented object boundary in these low contrast areas by making assumptions about smoothness and continuity of the boundary, but also tend to fail in areas of low intensity gradient and are strongly dependable on user initialization of the object shape.

Methods utilizing statistical shape models (Yokota et al., 2013) or active shape models (He et al., 2016) form another category of segmentation approaches that include preliminary creation of model of the segmented object based on a training dataset acquired from previous examinations. Such model can then be registered to the current patient data using manually located landmarks (Virzi et al., 2017) or automatically (Chen et al., 2010). Use of shape models however assumes that enough training data of the object is available to sufficiently cover the physiological variability and it usually doesn't anticipate notable deviations in the patient data.

In recent years, graph-cut methods based on (Boykov and Jolly, 2001) gained some popularity as the

method is able to reliably and efficiently find globally optimal segmentation of the object. While these methods often require multiple iterations of user interaction to achieve acceptable results (Shim et al., 2009), another of their advantage is that they can easily utilize additional information such as output of specialized edge detector (Krčah et al., 2011) or shape model (Keustermans et al., 2012).

Convolutional neural networks (Krizhevsky et al., 2012) are currently one of the fastest developing methods in area of image processing. With the advancements in computational power and expansion of various image databases, they became the state of the art in numerous areas such as semantic image segmentation (Zheng et al., 2015) as well as volumetric medical data segmentation (Prasoon et al., 2013). In this area, however, the available datasets still lack the sufficient scale and therefore the applications usually rely heavily on synthetic dataset augmentation (Milletari et al., 2016; Ronneberger et al., 2015). This comes at cost of reduced robustness similar to the model-based methods.

Classification random forests (Loh, 2011) is an ensemble machine learning method which uses random subsets of available training dataset to construct a set of binary decision trees. Each of these trees, given a vector of input data, then outputs histogram of class probabilities and finally either voting or averaging strategy is used to extract the final classification of the ensemble of trees. The method has been widely used in computer vision in recent years. While random forests have lately been outperformed by other machine-learning approaches such as convolutional neural networks in some image classification and segmentation tasks, random forests usually require a much smaller training dataset and shorter training time.

In this paper, we show that random forests learned using a few user-annotated slices only, in matter of minutes, can achieve classification accuracy sufficient to provide local voxel-wise predictions of tissue boundaries for final segmentation step based on global graph-cut optimization.

3 GRAPH-CUT SEGMENTATION WITH VOXEL-WISE PROBABILITIES FROM RANDOM FOREST CLASSIFICATION

The segmentation method proposed in this paper first requires the user to manually denote contour of the

object in several slices, for example 3 orthogonal slices through the middle of the data volume. Next, three voxel-wise random forest classifiers are trained on the user annotated slices. Applying these classifiers to the whole data volume outputs the probability for each voxel to be a part of the object, its inner edge and its outer edge. These three probability fields are then converted into a graph structure where minimum cut/maximum flow algorithm is employed to find the optimal binary segmentation of the object and its background while re-using the user-annotated slices as hard constraints.

3.1 Random Forest Classification

We formulate the segmentation as a binary classification problem where each voxel is assigned probabilities of belonging inside or outside the segmented object based on HU values of its surrounding voxels. A lot of work has been put into choosing image features suitable as input into the methods based on random forests in the available literature. Various features such as first- and second-order statistics of the patch surrounding the voxel (Cuingnet et al., 2012), entropy metrics and Gabor features (Mahapatra, 2014) and Haar-like features (Larios et al., 2010) have been used for various segmentation tasks. However, our results show that using only the intensity values from a small volume patch surrounding the voxel yields sufficient trade-off between the results and training complexity, which is important for semi-automatic method.

As mentioned earlier, we train following three separate random forest classifiers:

- Object detector trained using the user-annotated input binary object mask as the training data.
- Outer edge detector trained using the difference between the user-annotated mask and its corresponding 2D binary dilatation as the training data.
- Inner edge detector trained using the difference between the user-annotated mask and its corresponding 2D binary erosion as the training data.

The examples of detector outputs are shown in Figure 2 (d). The reason for training three separate classifiers is explained in the following chapter.

3.2 Global Segmentation Refinement using Graph-cut

In our method, graph-cut is used to refine the local outputs of random forest classifiers into a compact three-dimensional binary object. In the original work by (Boykov and Jolly, 2001), the region-

and boundary-term energies used to find the optimal boundary are conventionally derived from original image data as follows:

$$R_p(\text{"obj"}) = -\ln[Pr(I_p|O)] \quad (1)$$

$$R_p(\text{"bkg"}) = -\ln[Pr(I_p|B)] \quad (2)$$

$$B_{\{p,q\}} = \exp\left(-\frac{(I_p - I_q)^2}{2\sigma^2}\right) \cdot \frac{1}{\text{dist}(p,q)}, \quad (3)$$

where $Pr(I_p|O)$ and $Pr(I_p|B)$ are probability values corresponding to the intensity value of voxels extracted from histogram of user-annotated areas of image, I_q and I_p are intensity values of the voxels and σ is variance of intensity in data volume. Graph-cut then finds the optimal segmentation by minimizing the region-term energy over every individual voxel and boundary-term energy over every combination of neighboring voxels (6-neighborhood in 3D space).

Intuitively, we can replace the histogram-derived probabilities in region term with the more task-specific output of object detecting random forest classifier from the previous step as it is defined for every individual voxel. The new region-term energy is then defined as

$$R_p(\text{"obj"}) = -\ln[RF_{obj}(p)] \quad (4)$$

$$R_p(\text{"bkg"}) = -\ln[1 - RF_{obj}(p)], \quad (5)$$

where $RF_{obj}(p)$ is output of random forest object classifier for voxel p .

When defining the boundary-term energy for refining the output of voxel-wise classifier, most approaches such as (Mahapatra, 2014) decide to keep the intensity-derived energy values or simply replace the intensity difference with the difference of classifier output. This, however, discards the ability of the classifier to decide whether the object-background boundary is likely between two given voxels. While it is not practical to train and then evaluate a random forest classifier on each possible combination of neighboring voxels, it is possible to train two voxel-wise classifiers and then combine their outputs. Considering a pair of neighboring voxels p and q with corresponding inner edge detector outputs $RF_{in}(p)$ and $RF_{in}(q)$ and outer edge detector outputs $RF_{out}(p)$ and $RF_{out}(q)$, the probability that the object-background boundary lies in between these voxels is:

$$P_{edge}(p,q) = RF_{in}(p) \cdot RF_{out}(q) + RF_{out}(p) \cdot RF_{in}(q). \quad (6)$$

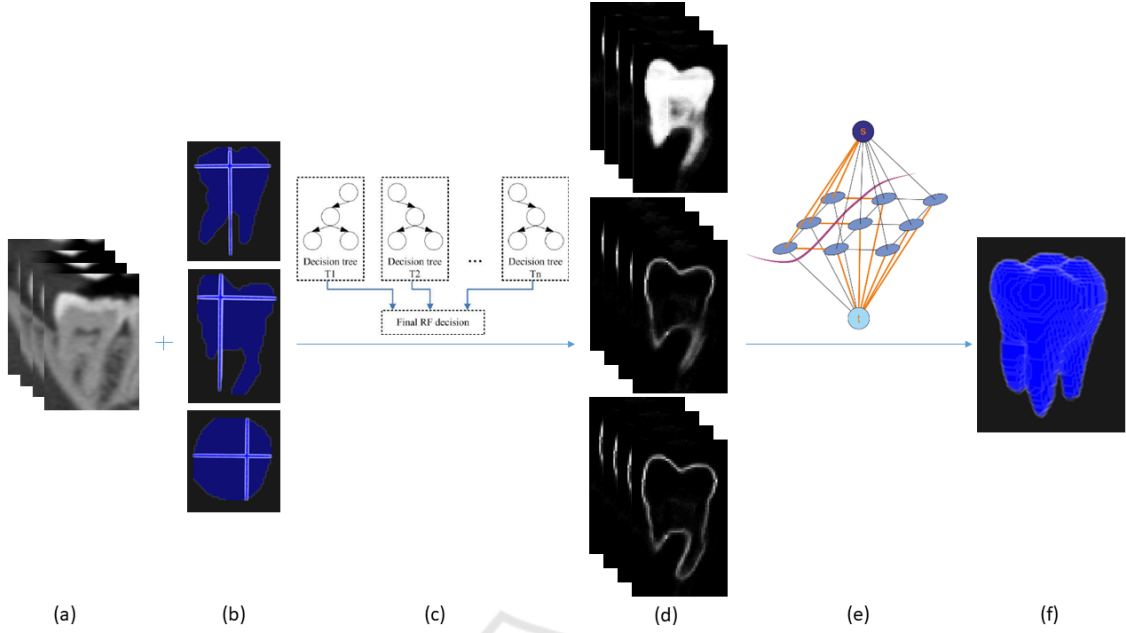


Figure 2: Overview of the segmentation framework. Original CT data (a) serve as an input along with several user-annotated slices (b). Random forests (c) are then used to produce voxel-wise predictions of the object and its outer and inner edge (d). These outputs are then integrated into a single graph structure (e) where min-cut/max-flow algorithm is employed to find final segmentation (f).

Therefore, we can define the new boundary-term energy in following way:

$$B_{\{p,q\}} = -\ln[RF_{in}(p) \cdot RF_{out}(q) + RF_{out}(p) \cdot RF_{in}(q)]. \quad (7)$$

While this approach is similar to (Browet et al., 2016) who also add "border" class to the classifier before graph-cut, they derive the boundary-term energy as maximum of the border penalization of the two voxels while our method computes the penalization for border *between* the two voxels specifically.

Similarly to the original method, we further use the user-annotated slices as hard constraints by setting the region-term energy of corresponding voxels to

$$R_p("obj") = 0 \quad (8)$$

$$R_p("bkg") = \infty \quad (9)$$

on voxels that user marked as object and to

$$R_p("obj") = \infty \quad (10)$$

$$R_p("bkg") = 0 \quad (11)$$

on the remaining voxels of the annotated slice. This forces the final segmentation to correspond to the user's initial shape description as the voxels already

annotated as the object can never be assigned background label and analogically the remaining voxels in the slice can never have the object label. Finally, all segmented objects not directly connected to any user-defined hard-constraints are discarded as false detections.

4 EXPERIMENTS AND RESULTS

We conducted experiments on a medical CT dataset that can be divided into three groups. The first group (A) consisted of 5 standard longitudinal bone scans with 1.27 mm voxel spacing and 0.5 mm slice thickness. These scans were fully manually segmented by medical expert and could therefore be used for quantitative analysis of our method. The dataset included three tibia, one humerus with higher level of noise and one radial bone with higher level of blur. Second group (B) consisted of two high-resolution mandible scans. First scan with 0.4 mm voxel spacing and 0.4 mm slice thickness included a toothless mandible and had expert segmentation made in 11 axial slices. The second mandible with 0.3 mm voxel spacing and 0.2 mm slice thickness had expert segmentation made for individual teeth in 3 orthogonal slices for each tooth. Scans in this group could not be used for quantitative analysis due to lack of ground truth and only illustrate that the method can be used

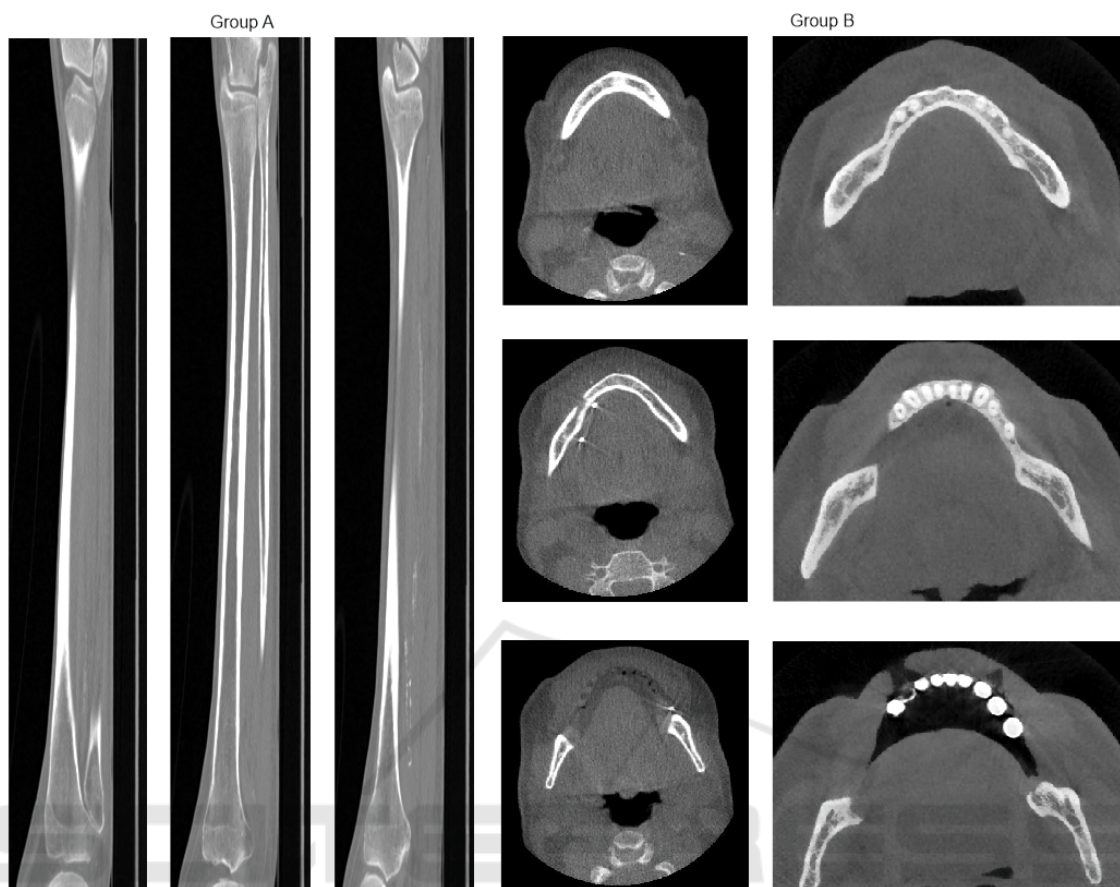


Figure 3: Example of CT scans on which the experiments were done. 3 lateral slices through a tibia in group A (left) and 3 axial slices of each mandible in group B (right).

to segment tissue of arbitrary shape. The third group (C) consisted of 9 standard kidney scans with 0.7 mm voxel spacing and 1.6 mm slice thickness. This dataset was also fully manually segmented to provide ground truth. All scans were cropped around the objects of interest as an image pre-processing step. Examples of the scans in each group are shown in Figure 3.

The method was implemented in Python and Matlab programming languages. Scikit-learn library (Pedregosa et al., 2011) was used for implementation of the random forests and Visualization Toolkit (Schroeder et al.,) for data visualization. Matlab implementation of the Maxflow algorithm (Boykov and Kolmogorov, 2004) was used for the graph-cut optimization. Tests were run on a laptop with 2.50GHz i5-3210M processor and 8GB RAM. The required computing time ranged from 4 minutes on smaller data volumes such as single teeth scans to 15 minutes on larger volumes such as shin bones.

For the scans in group A with full manual segmentations available, Dice coefficient (Dice, 1945) was used to quantitatively determine accuracy of the pro-

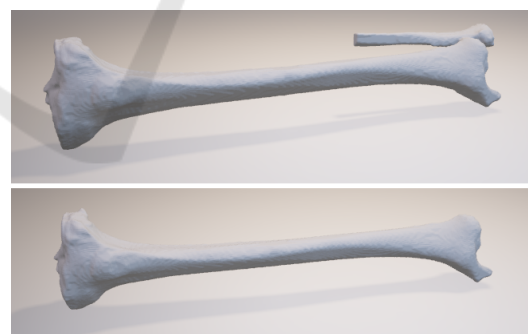


Figure 4: Example of false negative segmentation of part of splint-bone during shin-bone segmentation with Dice coefficient of 0.944 (upper) and correct shin-bone segmentation after adding an additional input slice yielding Dice coefficient of 0.987 (lower).

posed method and to study dependency of its success rate on the number of user-annotated slices used as an input. The experiments showed a very high precision of the proposed method with Dice coefficient of 0.976 ± 0.016 when using 5 user-annotated slices as input, specifically 3 orthogonal slices through middle

Table 1: Dice coefficients (DC) of final result and computation times of the classification step of the method for individual objects of interest. Training times for cases where 5 input slices were used are presented.

	DC (2 slices)	DC (3 slices)	DC (5 slices)	Total slices	Training (s)	Classification (s)
Shin bone 1	0.983	0.983	0.984	770	95	423
Shin bone 2	0.959	0.968	0.991	530	182	764
Shin bone 3	0.944	0.944	0.987	540	134	386
Humerus	0.967	0.968	0.968	587	57	218
Radial bone	0.907	0.935	0.952	497	60	256

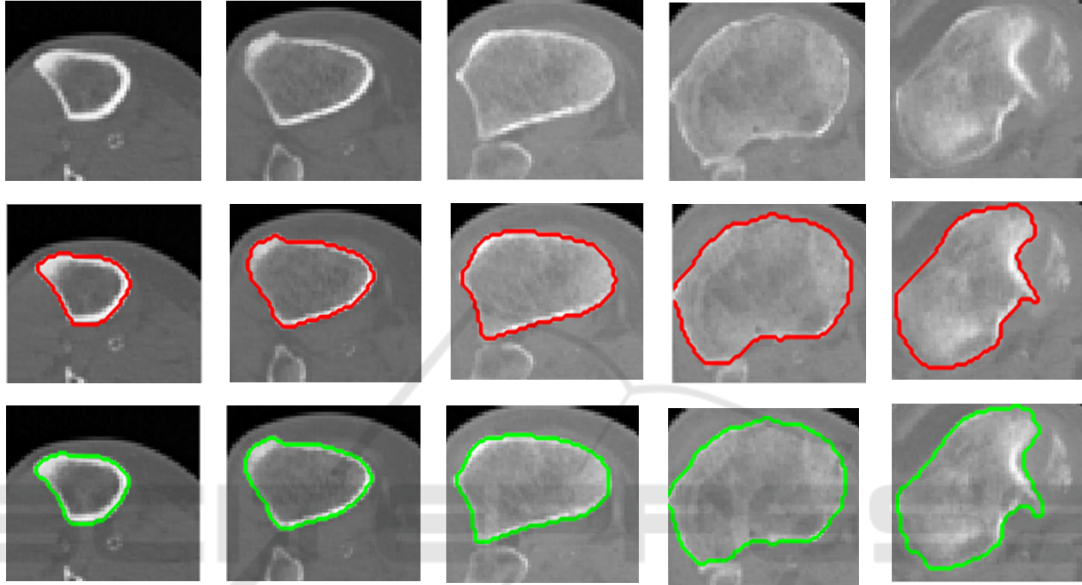


Figure 5: Example of result in 5 axial slices for the case where shin-bone was correctly segmented using only single frontal and lateral slice as input. Red contour marks expert-annotated segmentation in these slices, green contour marks result of our method.

of the bone and 2 additional axial slices through bone proximities. When using more than 5 input slices, the precision usually didn't improve anymore. The specific results and computing times for each bone are shown in Table 1. When using fewer user-annotated slices, i.e. only one sagittal and one frontal slice, the method still achieves high accuracy but can fail in separating adjoining structures such as part of splint-bone in case of shin-bone segmentation as shown in Figure 4. This happens mostly when the adjoining structure isn't present in the annotated slices and can be avoided by choosing the slices appropriately or by allowing the user to interact with the graph-cut step of the method. By marking only several voxels as being false-positive detections, the graph-cut can be reevaluated in several seconds, incorporating the newly added hard-constraints into the minimum cut search.

For the scans in group B, only several expert-annotated slices were available and used as an input. Although quantitative accuracy assessment is not pos-

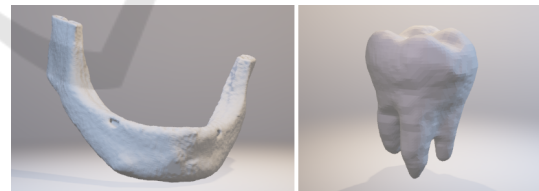


Figure 6: Examples of the segmentation result for mandible (left) and molar tooth (right). 11 axial slices were used as an input for the mandible segmentation and 3 orthogonal slices for the molar.

sible in these cases, the method yielded visually plausible results in all cases when using the available user-annotated slices as shown in Figure 6.

In experiments with the kidney scans in group C, 3 orthogonal expert-annotated slices were used as an input for each scan. The total number of analyzed slices per scan ranged from 59 to 83. The mean Dice coefficient of the segmentation results was 0.978 ± 0.008 achieving competitive results with ot-

Table 2: Dice coefficients (DC) of various kidney segmentation methods.

Method	Input	DC
(Glisson et al., 2011)	Seed points	0.93
(Sharma et al., 2015)	1 annotated slice	<0.9*
(Sharma et al., 2017)	None	0.86*
(Khalifa et al., 2017)	None	0.97
Our method	3 annotated slices	0.98

* Results on CT scans of kidneys with autosomal dominant polycystic kidney disease

her automatic and semi-automatic kidney segmentation methods as shown in Table 2. The example result of segmentation by our method is shown in Figure 7.

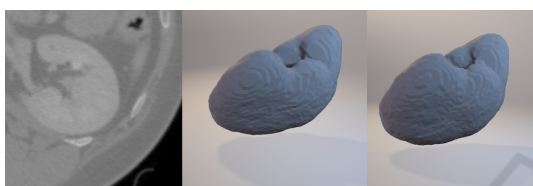


Figure 7: Example of kidney segmentation. Original CT data (left), manual segmentation (middle) and the output of our method (right).

5 CONCLUSIONS

In this paper we proposed a method able to perform segmentation of given tissue by first interpolating the user-annotated slices using random forest classifiers and then refining their output with graph-cut. We also introduce a new methodology of incorporating an output of voxel-wise classifier into boundary-term energy of the graph structure generated from the image in addition to the region-term energy.

The experiments show very good fit of the results with manually segmented tissues. After using hard tissue dataset to tune and analyze the algorithm performance, we tested it on the kidney dataset to show that the output of our universal method is comparable with state-of-the-art methods designed for the specific task.

Although it takes more user input, i.e. several manually segmented 2D slices, when compared to other methods, it brings significant advantages. First, it allows the user to segment an arbitrary tissue despite its shape or intensity as the random forest classifier infers the appropriate object and edge features on its own as long as the variability is captured in the annotated slices. Second, it is possible for the user to correct any faulty segmented regions by including them in the annotated slices and then either reevaluating the graph-cut optimization step in several seconds, or running

the full process again in case of larger errors. While this takes several minutes for every iteration, it still saves a lot of time and manual work when correcting the segmentation result in every slice individually. Also, the computational complexity could be further reduced by parallelizing the random forest classification step which takes majority of the computing time.

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