Refinement-Cut: User-Guided Segmentation Algorithm for Translational Science

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In this contribution, a semi-automatic segmentation algorithm for (medical) image analysis is presented. More precise, the approach belongs to the category of interactive contouring algorithms, which provide real-time feedback of the segmentation result. However, even with interactive real-time contouring approaches there are always cases where the user cannot find a satisfying segmentation, e.g. due to homogeneous appearances between the object and the background, or noise inside the object. For these difficult cases the algorithm still needs additional user support. However, this additional user support should be intuitive and rapid integrated into the segmentation process, without breaking the interactive real-time segmentation feedback. I propose a solution where the user can support the algorithm by an easy and fast placement of one or more seed points to guide the algorithm to a satisfying segmentation result also in difficult cases. These additional seed(s) restrict(s) the calculation of the segmentation for the algorithm, but at the same time, still enable to continue with the interactive real-time feedback segmentation. For a practical and genuine application in translational science, the approach has been tested on medical data from the clinical routine in 2D and 3D.

Nowadays, the most clinics – at least in the western world – have in general several medical scanners, like computed tomography (CT) or magnetic resonance imaging (MRI), which produce every day a massive amount of medical patient data. In addition, new scanner generations get more and more precise, and thus produce more and more data. However, there is by far not the time and manpower for a precise manual analysis of this important and critical data. Therefore, approximations are often used, like the estimated calculation of a tumor volume via its maximal diameter in a 2D view, which may not very accurate and can lead to inaccurate treatment decisions. A solution could be to support and automate medical image analysis with segmentation algorithms, like Active Contours in 2D or 3D, Active Appearance Models, graph-based approaches, fuzzy-based approaches, or neural networks. But after observing dozen of interventions in several clinics and different departments, I never met a physician who used any segmentation algorithm. The main reason was, that the segmentation approaches are not stable enough and fail far too often, especially for fully automatic algorithms. This may also be the reason that major manufacturers of medical imaging equipment don’t really offer sophisticated segmentation options within their workstations and software packages. Additionally, the existing approaches are often not user friendly and intuitive implemented, e.g. they need a precise definition of “mystic” parameters for an accurate segmentation result. A temporal solution to speed-up a segmentation task, until (fully) automatic algorithms provide reliable results, are semi-automatic methods, like interactive segmentation approaches. Thereby, the user supports and guides the algorithm by interactive input. This can be carried out by marking parts of the pathology and the surrounding background with a simple brush. Thus providing the segmentation algorithm with information about the pathology’s location in the image and information about the texture of the pathology and background. An overview about several interactive medical image segmentation approaches has recently been published by Zhao and Xie, where they also classify the approaches by their type of interactions:

- Pictorial input on an image grid, like Seeds for region growing,
- Parameter tuning using slider, dial, or similar interface, like the maximum size of segmented regions,
- Menu option selection by mouse clicking, like Accept/reject the segmentation results.
The paper is organized as follows: The Materials and Methods section presents the details of the proposed algorithm and online resources where medical data can be found; the Results section displays the outcomes of my experiments; and the Discussion section concludes the paper and outlines areas for future research.

Results

Figure 1 presents the interactive refinement segmentation of a vertebral body contour in 2D from a MRI acquisition. The leftmost image shows the native scan and the second image from the left displays the initial user-defined seed point (white) that has been placed inside the vertebral body for the interactive segmentation. However, due to the bright region inside the vertebral body, the average gray value – which is automatically calculated from the region around the user-defined seed – is not calculated “correctly”, and thus the resulting contour (red) leaks in the upper area and misses an edge in the lower left (note: for the interactive segmentation of the vertebral body, a rectangle was used as template to construct the graph. Thereby, the center of the rectangle is the user-defined seed point and the yellow crosses in the two rightmost images display the four corners of the rectangle). Finally, the rightmost image presents the result of the refined segmentation. Therefore, the user simply placed three additional seeds (white dots on the contour of the vertebral body), and thus forced the algorithm to perform the min-cut at these positions – which also influences the cuts along the neighboring rays. Furthermore, additional gray value information can be extracted around these extra seeds that the user placed on the contour of the vertebral body.
Figure 2 | Interactive refinement segmentation of the rectum from an intraoperative gynecological 3-Tesla magnetic resonance imaging dataset. The leftmost image shows the native scan and the second image from the left presents the initial seed point (white) for the interactive segmentation placed by the user inside the rectum. The red dots present the segmentation outcome with regard to the current seed point position (note: for the interactive segmentation of the rectum, a triangle was used as template to construct the graph). Thereby, the center of the rectangle is the user-defined seed point and the yellow crosses display the three corners of the triangle. In the third image from the left, an additional seed point (white) has been placed in the upper left contour of the rectum. This additional seed forces the algorithm to perform the min-cut at this position. In the fourth image from the left, the user has interactively repositioned the initial seed point inside the rectum to find a better segmentation outcome. However, the additional seed at the contour stays fixed during the interactive repositioning of the initial seed and still forces the algorithm to perform the min-cut at its position in the upper left contour of the rectum. In the rightmost image, the user further refined the segmentation outcome with two additional seed points.

Figure 3 | Interactive segmentation of a stented lumen and the thrombus from a postoperative computed tomography angiography (CTA) scan from a patient with an abdominal aortic aneurysm (AAA). The leftmost image shows the original scan and the second image from the left presents the segmentation of the stented lumen (red) with the initial user-defined seed point (green) placed inside the lumen (note: for the interactive segmentation a circle was used as template to construct the graph). The following three images show how the user places a second seed point and interactively drags it to the contour of the thrombus. However, the graph is still constructed from the initial seed point that has been placed at first inside the lumen. The second seed point forces the algorithm to perform the min-cut at its position and therefore also influences the positions of the min-cut in the neighboring rays. During the interactive dragging of the second seed inside the thrombus (images three and four from the left), the algorithm tries to adapt to other structures appearing in the thrombus. In this example, contrast enhanced blood from an endoleak is visible (elongated bright area inside the thrombus), and the resulting contour partly fits to this endoleak in the third and the fourth image in the lower right area: once to the left contour of the endoleak (third image) and once to the right contour of the endoleak (fourth image). In the rightmost image, the segmentation outcome has furthermore been refined with an additional seed that has been placed by the user in the lower right within the sagittal view.

Figure 4 presents the interactive segmentation of the prostate central gland (PCG) in 3D with a spherical template. The leftmost images show the original scan in axial (top), coronal (middle) and sagittal (bottom) views. The second image from the left presents the segmentation outcome (red) for a user-defined seed point (blue) placed inside the prostate (note: the seed point has been placed in the axial view, even if it is also displayed in the coronal and sagittal views). For comparison, the green masks display the outcome of a manual slice-by-slice segmentation from an expert. However, as the initial seed point is placed close to the right border of the prostate, the algorithm missed the contours on the left side of the PCG (axial and coronal views). Though, the interactive real-time behavior of the approach makes a repositioning easy, and thus it is also easy to find a good segmentation outcome for the axial, coronal and sagittal views (third image from the left). In the rightmost image, the segmentation result has been further refined with an additional seed that has been placed by the user in the lower right within the sagittal view.

Figure 5 presents different views – axial (top), coronal (middle) and sagittal (bottom) – of the 3D segmentation outcome from Figure 4. The left images show the last nodes (red) that still belong to the foreground (PCG) after the min-cut, and therefore defining the prostate central gland. In the images displayed in the middle column, the segmentation result has been superimposed with the manual mask (green) from the slice-by-slice expert segmentation. Finally, the rightmost images present a closed surface form the graph’s nodes, which can be used to generate a solid mask of the segmentation outcome for further processing.

In addition, performance tests have been carried out with a square template for vertebral body segmentation on a laptop with Intel Core i5-750 CPU, 4 x 2.66 GHz, 8 GB RAM running Windows 7 Professional x64 Version. Thereby, the computation time included the graph construction (sending out the rays from the user-defined seed point, sampling the nodes along these rays and constructing the
**Figure 4** Interactive segmentation of the prostate central gland (PCG) in 3D with a spherical template. The leftmost images show the original scan in axial (top), coronal (middle) and sagittal (bottom) views. The second image from the left presents the segmentation outcome (red) for a user-defined seed point (blue) that has been placed inside the prostate (note: the seed point has been placed in the axial view, but it is also displayed in the coronal and sagittal views). For comparison, the green masks display the outcome of a manual slice-by-slice segmentation from an expert. However, as the initial seed point is placed close to the right border of the prostate, the algorithm missed the contours of the PCG on the left (axial and coronal views). Though, the interactive real-time behavior of the approach makes a repositioning easy, and thus it is also easy finding a good segmentation outcome for the axial, coronal and sagittal views (third image from the left). In the rightmost image, the segmentation result has been further refined with an additional seed that has been placed in the lower right within the sagittal view.

**Figure 5** Different views – axial (top), coronal (middle) and sagittal (bottom) – of the 3D segmentation outcome from Figure 4. The left images present the last nodes (red) that still belong to the foreground after the min-cut, and therefore they define the segmented prostate central gland contour. In the images of the middle column, the segmentation result has been superimposed with the manual mask (green) from the slice-by-slice expert segmentation. Finally, the rightmost images present a closed surface form the graph’s nodes, which can be used to generate a solid mask of the segmentation outcome for further processing.
The evaluation has been performed on medical data from the clinical routine. There are several areas for future work: in particular, supporting manual strokes from the user which have been drawn along the border of the object to segment – instead of “only” single seed points. Albeit, this may “break” the real-time feedback you get from single seed points even if these are dragged on the image. As shown in the result section, a single seed point can still be moved around to find a better segmentation result, this is not so easy and intuitive anymore if the user has once drawn a stroke. Though, a solution may be an iterative adaption to the manually sketched parts of the user22.

Furthermore, a detailed study for the end user (which are primarily physicians in case of medical data) is necessary. Even if several physicians from different fields already tested the approach and responded positively, it’s of course not certain that they will use it for own research (e.g. for a time-consuming analysis of medical data for own research purpose) or even in the clinical routine. However, after carrying out several studies with a typical stroke-based approach22,23, it is clear that they would only accept such a course of action (the initialization) if the segmentation outcome is afterwards always satisfying (note: for the automatic segmentation the participating physicians had only to mark parts of the fore- and background with a simple brush; no other settings or parameters had to be defined). However, a long-term end user study regarding the presented approach already has been started within two European funded projects ClinicIMPPACT (www.clinicimppact.eu) and GoSmart (www.gosmart-project.eu), where post-interventional radiofrequency ablation (RFA) zones are segmented22.

Figure 6 | Semi-automatic segmentation of the prostate where several seed points (white) have been placed to get a segmentation result (red) that matches almost perfect with the manual segmentation (green).
Methods

Data. For a practical and genuine application in translational science, the elaborated approach has been tested with three-dimensional medical data from the clinical routine. Intraoperative gynecological 3-Tesla magnetic resonance imaging datasets that have been used for this study can be found here\textsuperscript{22,23,34}. MRI datasets of the spine, which are public available for research purposes, can be downloaded here\textsuperscript{35,36}. Pre- and intra-procedural MR-guided prostate biopsy datasets with manual segmentations are freely available here\textsuperscript{37,38}.

Software. The presented approach has been implemented as own MeVisLab C++ module and is a combination and extension of the Template-Cut\textsuperscript{22,23} and the Interactive-Cut\textsuperscript{33} approaches, and the refinement method introduced in \textsuperscript{26}. The new algorithm (Refinement-Cut), as well as the predecessor methods it builds up, belong to the graph-based approaches. Here, an image is interpreted as graph \( G(V,E) \) which consists of nodes \( v \in V \) sampled in the image and edges \( e \in E \) establishing connections between nodes. After graph construction a minimal \( s-t \)-cut\textsuperscript{13} is calculated on the graph, that is considering the two disjoint sets, whereby one set the segmented objects and one set the background represents – note: for the calculation of the minimal \( s-t \)-cut, two additional virtual nodes \( V_{\text{source}} \) and \( V_{\text{sink}} \) are used. The minimal \( s-t \)-cut returns the global optimum on a constructed graph, in contrast to iterative approaches, like the Active Contours, which in general find a solution stepwise, and thus can get stuck during this process in a local minimum. However, the immediate calculation of a global optimum, like the minimal \( s-t \)-cut, makes graph-based approaches in particular eligible for an interactive real-time application. First of all, for the graph construction, the nodes \( v \in V \) are sampled along rays which are sent out from one single seed point and with regards to a certain template. This template represents the basic shape of the segmented object, like described in the Template-Cut approach. Examples are

- A rectangle shape for vertebra segmentation in 2D\textsuperscript{26};
- A circle template for prostate central gland segmentation in 2D\textsuperscript{26};
- A cubic shape for vertebreal body segmentation in 3D\textsuperscript{40,41};
- A spherical shape for prostate central gland or brain tumor segmentation in 3D\textsuperscript{41,42};
- Or even a user-defined shape for objects that vary too much to be predefined by a simple shape\textsuperscript{37}.

After the nodes and the underlying texture values within the image have been sampled, the graph’s edges \( E \) are generated, that establish the connections between the (virtual) nodes, and an edge \( (v_i,v_j) \in E \) defines the connection between the two nodes \( v_i, v_j \). Taking over the notation of Li et al.\textsuperscript{39,40}, there are two types of \( s-t \)-weighted edges: Intra-edges which connect nodes along the same ray to ensure that the minimal \( s-t \)-cut runs through only one edge within this ray;

- Inter-edges which connect nodes from different rays under a smoothness value \( \Delta_s \), which influences the number of possible \( s-t \)-cuts and therefore the flexibility of the resulting segmentation.

Furthermore, there are edges between the sampled nodes and the virtual nodes (\( s \) and \( t \)) for the graph construction established, and the weights of these edges depend on the sampled texture values within the image and a cost function. For more detail about the graph construction the reader is referred to the previous Template-Cut\textsuperscript{22,23} publication. However, the specific graph construction, which basically starts from one single seed point inside the segmentation object, is particularly suitable for an interactive real-time segmentation, because the user has only to drag this one single seed point over the image – in contrast, to approaches where more input like information about foreground and background or strokes are needed. Moreover, the user can easily add more seed points on the object’s contour, which modify the graph and force the minimal \( s-t \)-cut to go through this additional seeds. Thereto, the user can easily add more seed points on the object’s contour, which modify the input like information about foreground and background or strokes are needed. Moreover, the intra-edge between \( s \) and \( t \) is forced to be at this position. In order to ensure that, the graph’s node \( c \) and all its predecessors within the same ray are connected via \( \cong \)-weighted edges to the source \( s \), and all successor of \( c \) within the same ray are connected via \( \cong \)-weighted edges to the sink \( t \). Furthermore, the intra-edge between \( s \) and \( t \) spiral to the minimal \( s-t \)-cut.

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**Author contributions**

Conceived and designed the experiments: J.E. Performed the experiments: J.E. Analyzed the data: J.E. Contributed reagents/materials/analysis tools: J.E. Wrote the paper: J.E.

**Additional information**

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