# Deep Active Learning for Interactive 3D Segmentation Of Medical Images

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# **Table of contents**

- 1. 3D Image Segmentation
- 2. Convolutional Neural Networks
- 3. Aims and Design
- 4. Results
- 5. Further Work
- 6. Conclusion

# 3D Image Segmentation

# **Medical Image Segmentation**

# 3D Medical Image Segmentation

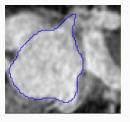


Figure 1: Segmentation of the left atrium

Main Problem: Laborious, particularly in 3D

# **Medical Image Segmentation**

# 3D Medical Image Segmentation

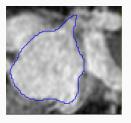


Figure 1: Segmentation of the left atrium

Main Problem: Laborious, particularly in 3D

How can we automate this?

**Aim:** Separate the set of *background* pixels (B) from the set of *object* pixels (O)

Define a cost function to be minimized over image X

$$E(\mathbf{X}) = \sum_{i} R_{i} + \lambda \sum_{i} \sum_{j} B_{ij}$$
 (1)

- R<sub>i</sub> Regional Penalty Term penalty for assigning pixel i to either the background (B) or object (O)
- B<sub>ij</sub> Boundary Penalty Term penalty depending on properties of pixels i and j
- $\lambda > 0$  a relative weighting

# R<sub>i</sub> - Regional Penalty Term

Map (0,1) probabilities to  $(\infty, 0)$  penalties with negative log

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Map (0,1) probabilities to  $(\infty, 0)$  penalties with negative log

$$R_{i} = \begin{cases} -\log P(i \in O) & i \in \mathbf{O} \\ -\log P(i \in B) & i \in \mathbf{B} \end{cases}$$
 (2)

B<sub>ij</sub> - Boundary Penalty Term

Want to encourage boundaries at discontinuities in intensity

Penalize boundaries that are placed between similar pixels

# B<sub>ij</sub> - Boundary Penalty Term

Want to encourage boundaries at discontinuities in intensity

# Penalize boundaries that are placed between similar pixels

$$B_{ij} = \begin{cases} 0 & y_i \neq y_j \\ \frac{1}{r_{ij}} e^{\frac{-(X_i - X_j)^2}{2\sigma^2}} & y_i \neq y_j \end{cases}$$
 (3)

- X<sub>i</sub> grayscale value at pixel i
- $y_i$  binary label for pixel i (0 means  $i \in \mathbf{B}$ )
- $r_{ij}$  euclidean distance between pixels i and j

#### How do we solve it?

Represent the image as a graph with

- One node per pixel
- One source node (object)
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Represent the image as a graph with

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and

- Edges between pixels have a cost  $\lambda \frac{1}{r_{ij}} e^{-(X_i X_j)^2}$
- ullet Edges between pixels and the source terminal cost  $-\log P(i\in B)$
- Edges between pixels and the sink terminal cost  $-\log P(i \in O)$

A minimum cost cut which severs the source from the sink will now minimize the cost function

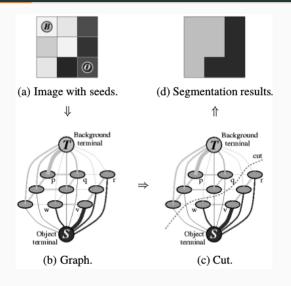


Figure 2: Minimum cost cut

Maxflow: Finding the maximal flow from source to sink

The edges that are saturated in maxflow are those in the minimal cost cut.

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The edges that are saturated in maxflow are those in the minimal cost cut.

Solve with Breadth First Search, by gradually saturating edges

How do we get  $P(i \in B)$  and  $P(i \in O)$ ?

#### User interaction

- User gives seed points
- Probability distribution produced from seed points
- Probabilities of other pixels inferred from distribution

Works well, but...

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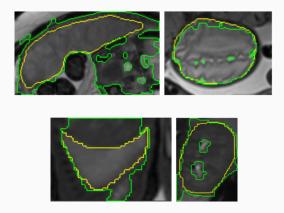


Figure 3: Grabcut algorithm in difficult cases

Requires object and background pixels to have distinct intensity values

This is the case for most traditional segmentation methods

# **Advantages**

- Spatial operations
- Learning
- Performance

# Disadvantages

- Computationally expensive
- Poor generalisation to unseen types
- Need large datasets expensive for medical imagery

Classification CNNs - gradually extract larger features from smaller ones

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**U-Net** - gradually re-combine those features to produce a segmentation map

# **Segmentation CNNs**

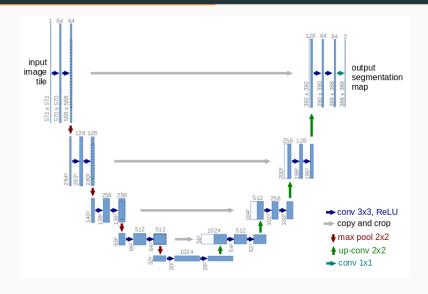


Figure 4: The U-Net architecture

# **Segmentation CNNs**

# Improvements since original U-Net:

- Residual blocks
- Spatial Dropout
- Segmentation Layers

# **Segmentation CNNs**

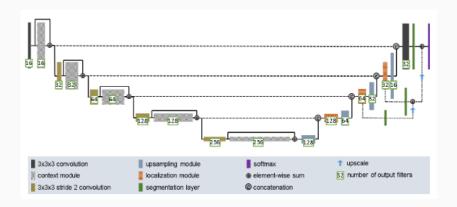


Figure 5: The Isensee et al. architecture

Segmentation Layers encourage earlier layers to produce segmentations

Aims and Design

# **Aims**

#### Aims:

Minimize the disadvantages of CNNs so that they can be used for 3D image segmentation

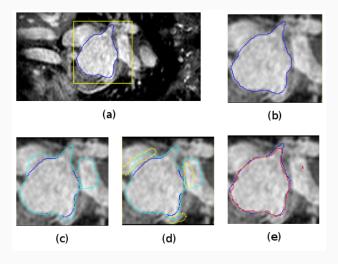
# Disadvantages

- 1. Computationally expensive
- 2. Poor generalisation to unseen types
- 3. Need large datasets expensive for medical imagery

# **Bounding-box Image-specific Fine-Tuning Segmentations** (BIFSeg) Combines CNN and CRF

Leverages user corrections

- 1. Draw bounding box around area of interest
- 2. CNN and CRF give initial guess
- 3. User provides correction scribbles
- 4. CNN fine-tunes using corrections and gives corrected result



**Figure 6:** The steps in the BIFSeg framework. In *blue* is the ground truth, in *cyan* the initial guess, in *yellow* pixels relabelled as background and in *red* is the final result

# Two things going on:

Combining CNN and CRF
 CNN softmax output used as probability values for calculating R<sub>i</sub>

### Two things going on:

- Combining CNN and CRF
  CNN softmax output used as probability values for calculating R<sub>i</sub>
- Fine-Tuning CNN using CRF
  - 1. CNN, CRF and scribbles produce segmentation
  - 2. CNN trained to match segmentation
  - 3. Repeat steps 1 and 2 n times

### Fine-Tuning CNN using CRF

CNN loss function is **weighted** on a per-pixel basis during fine-tuning, such that the weight for pixel i is

$$w_i = \begin{cases} 0 & i \in \mathbf{U} \\ w & i \in \mathbf{S} \\ 1 & \text{otherwise} \end{cases} \tag{4}$$

- U set of pixels for which the current segmentation is uncertain
- S set of user scribbles

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Pixels in  $\mathbf{U}$  are either **geodesically** near a scribble of opposite label or have  $P_i$  near 0.5

# Design

**BIFSeg works on unseen object types** - though it needs a CNN that has learned **generalisable features** 

Performs better and with less user interaction on seen object types

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# Transfer learning

- 1. Start by using a CNN trained in generalised segmentation
- 2. After *n* images have been segmented, add them to a dataset
- 3. Load a new CNN with the generalised CNN weights and train it on the dataset
- 4. Repeat steps 2 and 3 until all images have been segmented

# Results

Evaluate performance on seen objects

Train and test on left atrium images

### **Dice Scores**

- Initial Prediction  $0.872 \pm 0.025$
- ullet After User Interaction 0.9112  $\pm$  0.018

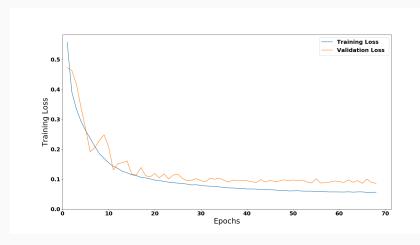


Figure 7: Loss plot for weighted dice coefficient loss on left atrium data

Evaluate performance on seen objects

First train a generalised CNN

| Organ Type  | Dice Score        |
|-------------|-------------------|
| Left Atrium | $0.616 \pm 0.025$ |
| Prostate    | $0.873 \pm 0.054$ |
| Hippocampus | $0.487 \pm 0.125$ |
| All         | $0.617 \pm 0.054$ |

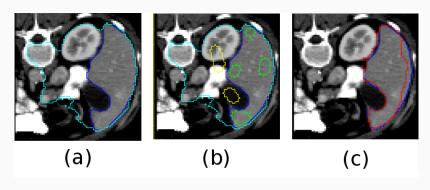
**Table 1:** Validation scores for the generalized segmentation CNN on the organs it was trained on

| Organ Type | Dice Score (CNN)  | Dice Score (CNN+CRF) |
|------------|-------------------|----------------------|
| Liver      | $0.516 \pm 0.019$ | $0.532 \pm 0.022$    |
| Spleen     | $0.417 \pm 0.033$ | $0.484 \pm 0.037$    |

**Table 2:** Dice scores by the generalized segmentation CNN on unseen object types with and without the CRF

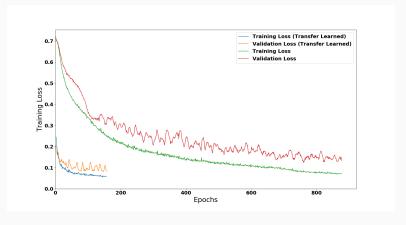
| Organ Type | Dice Score (10 scribbles) | Dice Score (15 scribbles) |
|------------|---------------------------|---------------------------|
| Liver      | $0.873 \pm 0.029$         | $0.891 \pm 0.021$         |
| Spleen     | $0.892 \pm 0.018$         | $0.918 \pm 0.016$         |

**Table 3:** Dice scores by the generalized segmentation CNN on unseen object types, after fine-tuning with user interaction



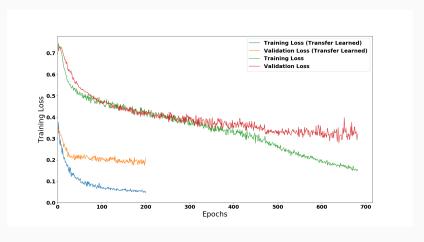
**Figure 8:** Segmentation of unseen object type (liver) using BIFSeg. In *blue* is the ground truth, in *yellow* are background scribbles, in *green* foreground scribbles and in *red* the final segmentation. Final Dice score over the 3D image was 0.884

# **Transfer Learning - Liver**



**Figure 9:** Comparison of transfer learning and learning from scratch learning on the liver with 4 training images and 6 validation images

# **Transfer Learning - Spleen**



**Figure 10:** Comparison of transfer learning and learning from scratch learning on the spleen with 4 training images and 6 validation images

**Further Work** 

# Conclusion