

Microscopy Cell Segmentation via Convolutional LSTM Networks

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1 Methods

We address individual cells' segmentation from microscopy sequences using C-LSTM. The main challenge in this type of problems is not only foreground-background classification but also the separation of adjacent cells. As in [1] we suggest to enhance individual cells' delineation by a partitioning of the image domain $\Omega \in \mathbb{R}^2$ into three classes: foreground, background, and edges. We set $C = \{0, 1, 2\}$ to denote these classes, respectively. Let $\{I_t\}_{t=1}^T$ be the input image sequence, where $I_t : \Omega \rightarrow \mathbb{R}$ is a grayscale image. We define a network f_Θ with parameters Θ as follows:

$$(o_t, h_t) = f_\Theta(I_t, o_{t-1}, h_{t-1}), \quad c \in C \quad (1)$$

where, the output $o_t : \Omega \rightarrow \mathbb{R}^3$ is a three-dimensional feature vector corresponding to each input pixel $\mathbf{x} \in \Omega$ and h_t are the hidden variables of the network. We define the segmentation as the pixel label probabilities using the softmax equation:

$$p(c|o_t(\mathbf{x})) = \frac{\exp\{[o_t(\mathbf{x})]_c\}}{\sum_{c' \in C} \exp\{[o_t(\mathbf{x})]_{c'}\}} \quad (2)$$

The final segmentation is defined as all pixels where

$$\Gamma_t = \arg_{c \in C} \max p(c|o_t(\mathbf{x})) \quad (3)$$

Each connected component of the foreground class is given a unique label and is regarded as a separate cell.

Network Architecture The proposed network f_Θ incorporates C-LSTM blocks into the U-Net architecture. While the U-Net [5] and C-LSTM [6] are widely used, their composition is suggested here for the first time and is shown to be powerful. The U-Net architecture, built as an encoder-decoder with skip connections, is able to extract meaningful descriptors at multiple image scales. However, this alone does not account for the cell specific history that can significantly support the segmentation. The introduction of C-LSTM blocks into the network's decoder allows considering past cell appearances by holding their compact representations in the memory units. The network is fully convolutional

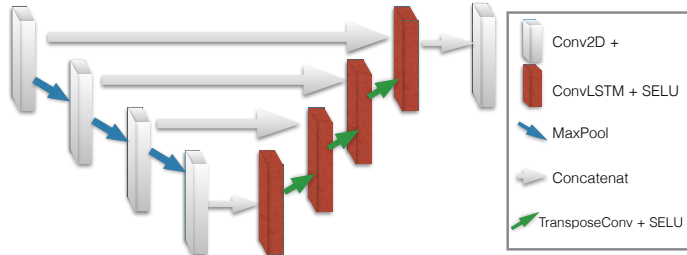


Fig. 1: The U-LSTM network architecture. The down-sampling path (left) consists of convolutional layers, SeLU activations, the output is then down-sampled and passed to the next layer. The up-sampling path (right) consists of a concatenation of the input from the lower layer with the parallel layer from the down-sampling path followed by a C-LSTM layer and a transpose-convolution up-sampling layer followed by SeLU activations.

and, therefore, can be used with any image size¹ during both training and testing. Figure 1 illustrates the full network architecture detailed in Section 2.

Training During the training phase the network is presented with a full sequence and manual annotations $\{I_t, \Gamma_t\}_{t=1}^T$, where $\Gamma_t : \Omega \rightarrow [0, 1, 2]$ are the ground truth (GT) labels for the pixels in image I_t . The network is trained using Truncated Back Propagation Through Time (TBPTT). At each back propagation step the network is unrolled to τ time-steps. The loss is defined using the cross-entropy loss:

$$L = \sum_{t'=t}^{t+\tau} \sum_{\mathbf{x} \in \Omega} p(\Gamma_{t'}(\mathbf{x}) | o_{t'}(\mathbf{x})) \quad (4)$$

and the weights are updated using gradient descent. Note that this loss differs from the U-Net loss, where boundary pixels are labelled as background weighted by their proximity to the two nearest cells [5]. Here, since a separate class for boundary pixels is defined, weighting is not required. Figure 2 show a visual example of the annotations and the network output. Comparisons of the two losses using the proposed network architecture is reported in the supplementary material.

2 Implementation Details

Architecture The network comprises four down-sampling blocks and four up-sampling blocks. Each block in the down-sampling branch is composed of a convolutional layer [3], and leaky SeLU activation function [4]. The up-sampling

¹ In order to avoid artefacts it is preferable to use image sizes which are multiples of eight due to the three max-pooling layers.

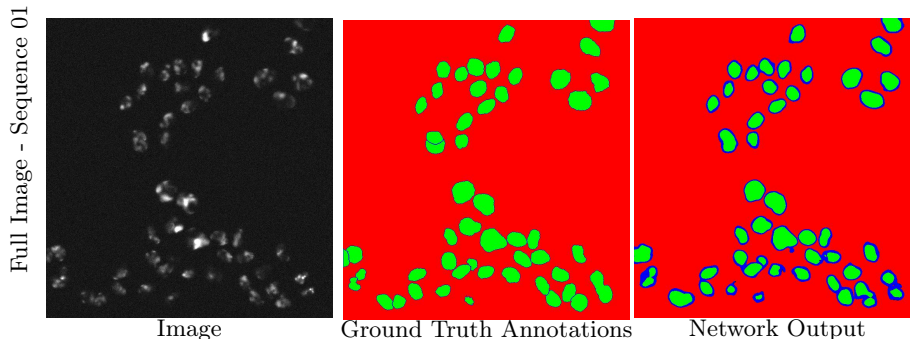


Fig. 2: Annotation Example: The visualization of the annotations as a three-class segmentations. On the left an image from the Fluo-N2DH-SIM+ data set, the center image is the GT annotation and the right image is the network output. The colors red, green and blue represent the three classes, background, foreground and cell contour, respectively. It is evident that the network learned to classify the contour of the cells as class edge allowing the separation of individual cells.

blocks consist of a transpose-convolution, a concatenation with the parallel down-sample block and a C-LSTM. All convolutional layers use kernel size 3×3 with layer depths (128, 256, 512, 1024). All maxpool layers use kernel size 2×2 without overlap. All C-LSTM and transpose convolutions kernels are of size 3×3 and 5×5 respectively with layer depths (1024, 512, 256, 128). The last convolutional layer uses kernel size 1×1 with depth 3 followed by a softmax layer to produce the final probabilities (see Figure 1).

Training Regime We trained the networks for approximately 100K iterations with an RMS-Prop optimizer [2] as implemented in Tensorflow (version 1.4.0) with learning rate of 0.0001. The unroll length parameter τ was set to five (Section 1) and the batch size was set to three sequences.

References

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