

# DEEP NEURAL NETWORK FOR NUCLEI SEGMENTATION IN HISTOPATHOLOGY IMAGES

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## ABSTRACT

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**Index Terms**— Deep Networks, Nuclei Segmentation, Histopathology. Maybe more?

## 1. INTRODUCTION

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## 2. RELATED WORK

Talk about related work. Recent work in deep learning for segmentation, histopathology analysis helped by images. Maybe not too much as the paper is small..

## 3. DATASET

One of the main contributions of this paper is the now public available nuclei detection dataset within HE stained histopathology images which can be found at [www.cbio.enscm.fr/pnaylor/download/dataset.zip](http://www.cbio.enscm.fr/pnaylor/download/dataset.zip) (not a true address). This annotated dataset provides images clustered by patients. Each patient has at least 3 annotated  $512 \times 512$  HE stained histopathology images with their associated ground truth. Each ground truth image is a  $512 \times 512$  where each pixel value above 0 is considered as a pixel belonging to a nuclei. The differences in values of these pixels denote different nucleus, such an annotation can be used in several processing step which does or doesn't take into account clustered nuclei. See figure 1 for an example of three annotated images. This annotation was conducted via the

help of the software ITK-snap and were annotated by the authors of the paper.

These patients were randomly picked from an unpublished study on tripple negative breast cancer. For each of these patients we had access to their biopsy sample as a whole slide image (WSI). WSI enables a medical practitioner to digitize the huge amount of information that can be found in glass slides. WSI can be up to 60 GB big uncompressed and can't be stored in RAM on a standard computer. Given the WSI of a patient, we randomly cropped  $512 \times 512$  samples from the WSI. 3 to 7 images were choosen from the randomly sampled images to try and give the most diversified dataset among these patients. Once the samples were choosen, we fully annotated each nuclei via the software ITK-snap and touching nuclei are differentiate via a different annotation value.

We can describe the dataset provided.

## 4. METHODOLOGY

Let  $A$  be the space of RGB images,  $A$  can typically be  $\mathbb{R}^{n \times p \times 3}$  and let  $B$  the space of annotation images, in our case  $\{0, 1\}^{n \times p}$ . We have a set of  $(A_l, B_l)_{l \in [1, N]}$  for a supervised learning approach. Our goal is a prediction task named as semantic segmentation, we wish to maximize the prediction of an unseen element belonging to  $B$  given an new element in  $A$ . We maximize thus prediction by modelizing our prediction function as the softmax output of a deep neural network. We find the model parameters by minimizing a log loss function defined as:  $\frac{1}{\sum_{i,j} \sum_k w_{i,j} t_{i,j,k}} \sum_{i,j} \sum_k w_{i,j} t_{i,j,k} \log(\widehat{p_{i,j,k}})$ , where  $k$  designates a certain label,  $w_{i,j}$  is certain weight given to pixel  $i, j$ ,  $t_{i,j,k}$  is equal to 1 if pixel  $i, j$  is of class  $k$  and  $\widehat{p_{i,j,k}}$  designates the estimated probability of pixel  $i, j$  of being  $k$  via the softmax output of the neural network. We minimize the loss function via stochastic gradient descent.

We have our training set  $(A_l, B_l)_{l \in [1, N]}$  where  $N$  is equal to 34, how ever each element  $A_l$  belongs to a certain patients and several elements  $A_l$  can belong to the same patient. As we are dealing with histopathology images, it is know that samples can widely vary from one patient to the other. We thus validate our model by a leave one patient out scheme. Our validation scheme is as followed, for a given set of type

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parameters, we train our model on every patient except one that is used for validation. Our final score is averaged over all patients. Several metrics assess the quality of the model, the accuracy (Acc), the F1 score and a performance (Perf) score which is the mean between the true positive rates and the true negative rates.

To train our models, as the number of available annotated is scarce, we used a great number of transformation for the data augmentation. From a original size of 33 annotated images, adding flips, rotations, blurriness and random elastic deformations enabled us to have more than 400000 training images per patient. We also try out several hyperparameter configurations: the learning rate and momentum for the stochastic gradient descent, the weight decay value. In practice, we found that hyperparameters tuning didn't influence the scores much, the exception being the learning rate. If the learning rate was not of the right magnitude the given network did not seem to learn. We also experienced with different initialization value and if possible, we also considered pretrained layers. Using pretrained layers made learning more efficient and made score values more robust.

## 5. DIFFERENT ARCHITECTURES/RESULTS

We experience with 4 know architectures in semantic segmentation, one will be named BaochuanNet, Fully Convolutional Net (FCN), DeconvNet, UNet. The most basic architecture, BaochuanNet, consists of 4 convolutional layer where each convolutional layer has 8 feature map. This net, being not deep, has the advantage of being less computationally intensive. FCN is a first attempt of applying "deep feature" representations to the task of semantic segmentation. This architecture has the advantage of re-using a classical deep learning architecture with added on upsampling layer and skip layers. The upsampling layers enable the network to learn a pixel level classification and the skip layers enable the network to fuse different levels of abstraction to the final prediction. This model can be fine tuned with a set of pretrained-weights extracted from the classical deep learning architecture. DeconvNet is also based on a classical architecture, however, in this network there are no skip layer as one intends to learn the proper upsampling through repeated deconvolution and convolution layers. This model can also be fine tuned.

## 6. RESULT

table of values

## 7. DISCUSSION

What's next?

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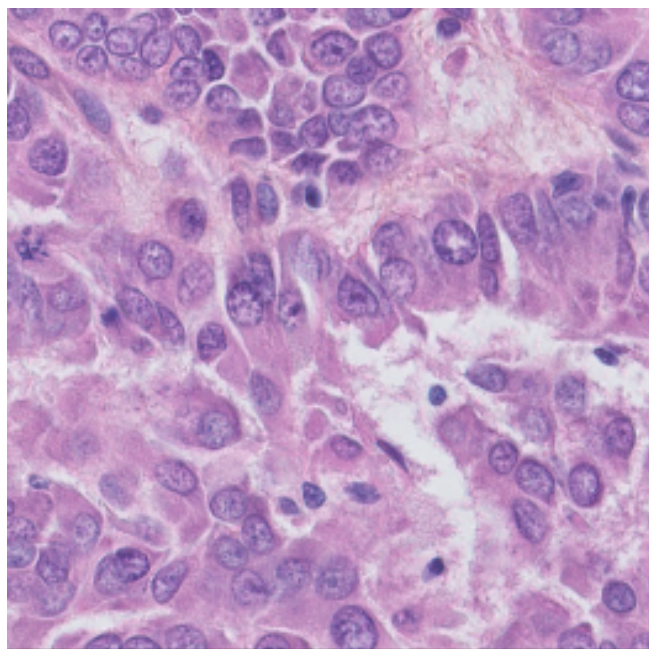
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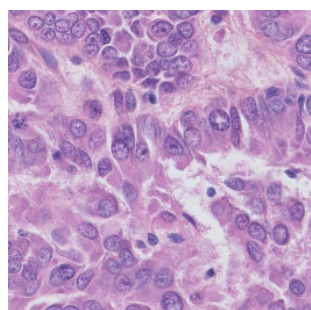
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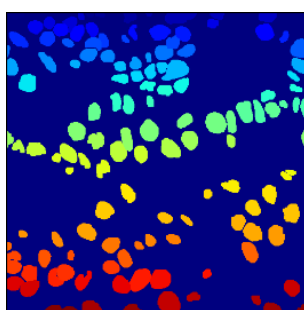
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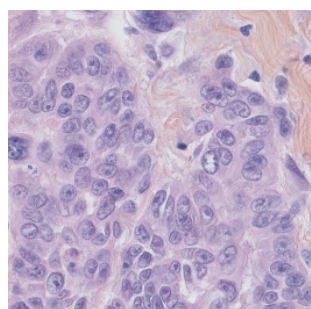
Patient 498959



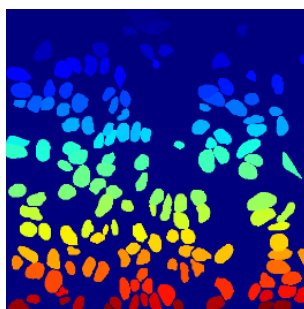
Sample from patient 498959



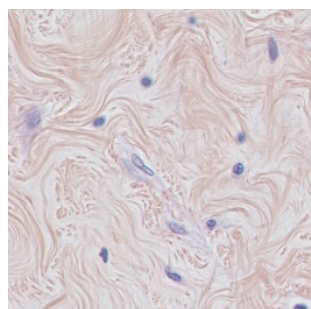
Associated ground truth



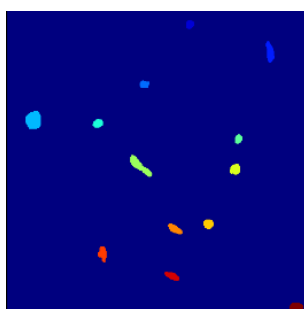
Sample from patient 581910



Associated ground truth



Sample from patient 581910



Associated ground truth

**Fig. 1.** Random annotated samples from the dataset