## Machine Learning & Understanding

Working with manually annotated whole slide images (WSIs)

- annotations of cancer vs. non-cancer regions, exclude regions
- using non-cancer regions from healthy patients
- rate limiting steps are manual annotations

A prostate sample WSI



### Machine Learning & Understanding

- Developing models that can process gigapixel images (e.g., 20 Gpix) efficiently
  - tile-based input into deep convolutional networks
  - experiments with different architectures



# Pipeline

Developed complete modular pipelines:

- from reading/tiling big WSIs dynamically and reproducibly based on annotations to generate positive/negative tile sets
- processing tiles augmentations
- training
- evaluation
- Storing compressed history of each run
- Issues with reproducibility



https://github.com/baidu-research/NCRF

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### **Training Data**

- 698 slides; 156 patients
- The training set: 6,513,435 negative patches and 1,365,240 positive patches.
- augmentation techniques:
  - random vertical and horizontal flips with 50
  - random brightness perturbations in range [-64,64],
  - random hue perturbations in range [-10,10],
  - random saturation perturbations in range [-64,64] and
  - random contrast perturbations in range [0.5,2.0].
- All patches were scaled to [-1,1] range.
- Training data three step sampling:
  - 1. a label is selected uniformly at random
  - 2. single slide is picked uniformly at random from all the slides containing at least one patch with label

3. a patch with label is selected uniformly at random from the slide A performance issue: A new slide (i.e. something large) is opened every time a new patch is sampled.

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# Model

Convolutional Neural Networks (CNNs)

- VGG16
- combined with ImageNet initial weights
- experimenting with different ways how to include bigger context
  - aggregating 3  $\times$  3 vs. big tiles with evaluating centers only



• 14. prosince 2020ps://mc.ai/extract-features-visualize-filters-and-feature-maps-in-vgg16-and-vgg19-5cm/8 models,

# **Training & Testing**

- RMSprop optimizer with the following parameters: momentum = 0.9,  $\rho$  = 0.9, initial learning rate of  $\eta$  = 5 \* 10<sup>-5</sup>
- Learning rate was halved after every 5 consecutive epochs without improvement on a validation data.
- The training would stop if no improvement on a validation data is made for 10 consecutive epochs.

Testing:

Test set: 87 slides; 10 patients; 193,235 tiles

# Machine Learning & Understanding

Understanding how the deep neural networks work – explainability

- to gain trust in the diagnostic process
- to assess limitations of what the network can assess

Here gradient-based saliency maps





#### **Interactive Visualizations**

- Interaction with pathologists
  - visualizing results vs. original annoations
  - visualizing explainability information
  - annotating all these results
- Developed a visualization pipeline: from automated harvesting of trained models

