

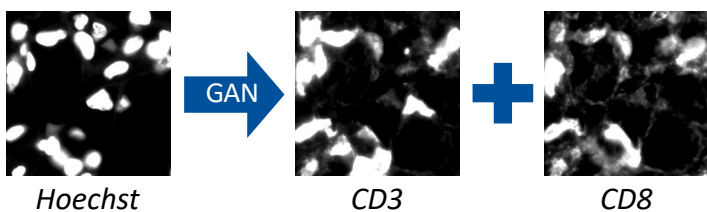
## Virtual Lymphocyte Staining Using Generative Adversarial Networks

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### Task

**Virtual staining** aims to transform a whole slide image from one stain to another.



Given a kidney cancer tissue slide stained with *Hoechst*, we **synthesise** *CD3* and *CD8* stains.

### Contributions

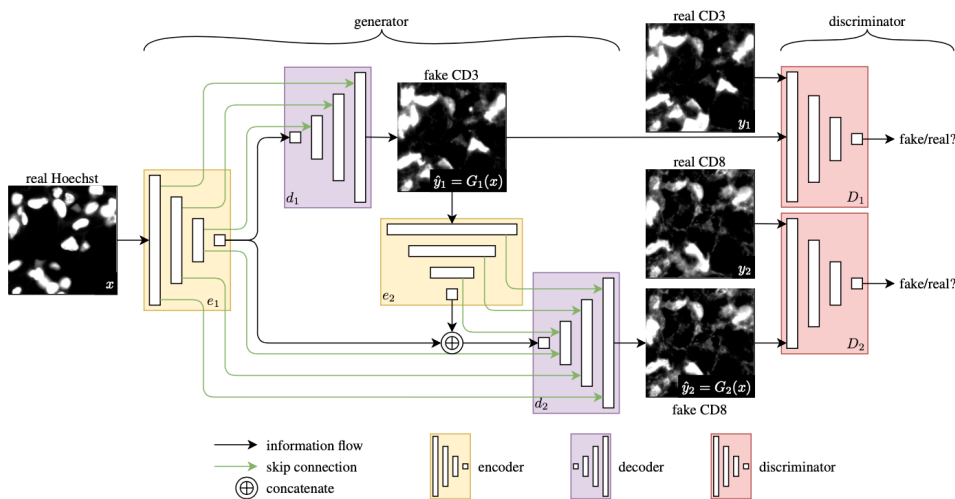


### Method

We propose a novel GAN architecture based on **pix2pix** [Isola *et al.*, 2017] that has **two separate output branches**.

The generator's encoder learns **mutually beneficial representations** because it is **shared** across both branches.

Information from the CD3 branch flows back to **assist** CD8 generation as well.



$$\text{GAN loss: } \mathcal{L}_{\text{cGAN}}(G_1, G_2, D) = \mathbb{E}_x [\log(1 - D(x, G_1(x), G_2(x)))] + \mathbb{E}_{x, y_1, y_2} [\log D(x, y_1, y_2)]$$

$$\text{objective: } G^* = \arg \min_{G_1, G_2} \max_D \mathcal{L}_{\text{cGAN}}(G_1, G_2, D) + \lambda \mathcal{L}_{L_1}(G_1) + \lambda \mathcal{L}_{L_1}(G_2)$$

### Results

We introduce a novel evaluation metric, the **relative masked intensity ratio** ( $MIR_{rel}$ ), for assessing virtual staining **quality**.

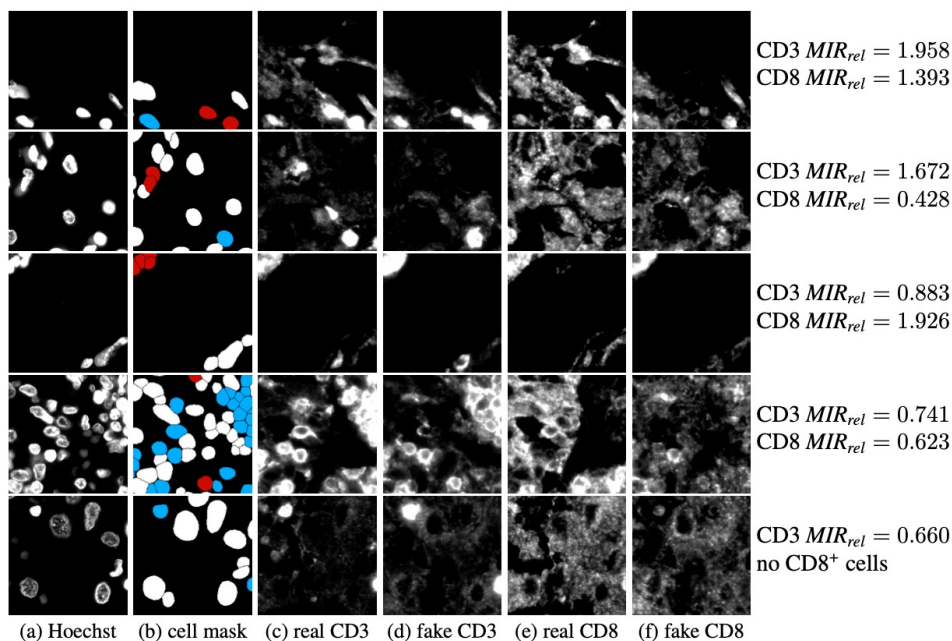
$$MIR = \frac{\text{mean pixel intensity within mask}}{\text{mean pixel intensity outwith mask}}$$

$$MIR_{rel} = \frac{MIR_{fake}}{MIR_{real}}$$

Our model **outperforms** U-Net regression and pix2pix baselines.

In an **ablation** study, we find that CD8 stain generation is able to learn from CD3.

We achieve best results with a **joint discriminator**.



Method	Parameters ↓	Train CD3 $MIR_{rel}$ ↑	Test CD3 $MIR_{rel}$ ↑	Train CD8 $MIR_{rel}$ ↑	Test CD8 $MIR_{rel}$ ↑
HoechstGAN-MCD	$2.16 \times 10^8$	<b><math>1.23 \pm 1.21</math></b>	<b><math>1.48 \pm 1.27</math></b>	$1.07 \pm 0.97$	$1.43 \pm 1.04$
HoechstGAN-MC	$2.19 \times 10^8$	$0.87 \pm 1.07$	$1.12 \pm 1.23$	<b><math>1.22 \pm 0.88</math></b>	$1.39 \pm 1.02$
HoechstGAN-MD	$2.16 \times 10^8$	$0.88 \pm 1.07$	$1.15 \pm 1.21$	$1.01 \pm 0.93$	<b><math>1.45 \pm 1.03</math></b>
HoechstGAN-M	$2.19 \times 10^8$	$0.89 \pm 1.26$	$1.24 \pm 1.30$	$0.84 \pm 0.97$	$1.26 \pm 1.06$
HoechstGAN-D	<b><math>9.20 \times 10^7</math></b>	$0.90 \pm 1.01$	$1.14 \pm 1.17$	$0.87 \pm 0.94$	$1.33 \pm 1.05$
pix2pix	$1.14 \times 10^8$	$0.89 \pm 1.04$	$1.15 \pm 1.23$	$0.83 \pm 0.87$	$1.16 \pm 1.00$
Regression-MC	$2.13 \times 10^8$	$0.62 \pm 0.74$	$0.92 \pm 0.75$	$0.57 \pm 0.58$	$1.09 \pm 0.73$

### Discussion

- This paper introduces a **virtual staining model** for ccRCC.
- HoechstGAN architecture is **applicable** to any image-to-images problem.
- We suggest our **evaluation metric**  $MIR_{rel}$  as a criterion to benchmark future virtual staining models.
- Future research: investigate **symmetric architectures** where CD3/CD8 information flows both ways.

### More information

project page with data, code and more



[georg.woelflein.eu/hoechstgan](http://georg.woelflein.eu/hoechstgan)