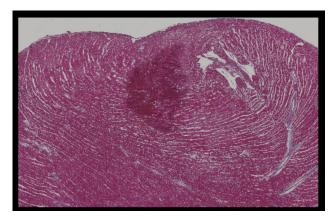
Automated Image Alignment for Multimodal Biological Imaging

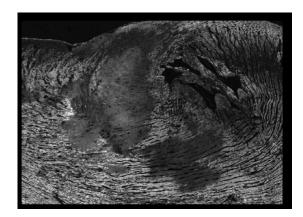
Cole Diianni, Cate Eberman, Helen Wilson

Background

- Histology images provide information about the cells, collagen content, and other tissue properties
 - Polarized collagen images provide context for the structural properties of the tissue
 - Location of the cells on the brightfield images



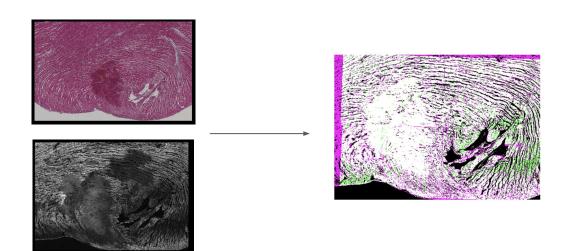
Brightfield H&E



Polscope H&E

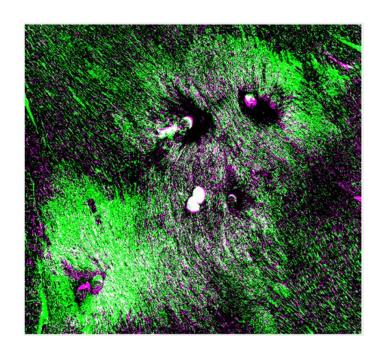
Project Goals

- Compare histological samples with images taken using different imaging modalities and on different scales
- Develop workflow to import and edit the images before alignment, then print out the final corresponding images



Challenges

- Need to register images from different modalities even though they contain different information
- There is global, translation, rotation and scaling
- Regional variation in translation, rotation, shear → Global homography won't match up across whole sample
- Very large images
- Images are taken at different scales



Global Homography Leads to Poor Local Matching

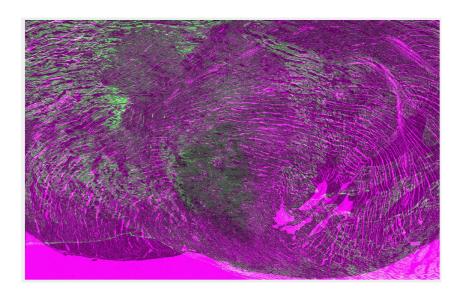
Current Methods

MATLAB

- Automatic Registration
- Control Point Registration
- Is extremely slow due to large image size

ImageJ

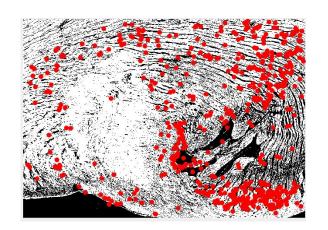
- Was only able to handle images from the same sample and modalities at different scales
- These methods struggled with the high variation between samples and the sample size

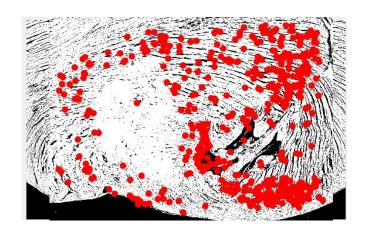


MATLAB Image Automatic Image Registration

Our Approach

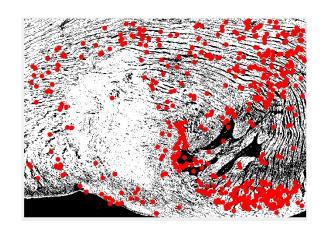
- Import, resize, and threshold the images so interesting landmarks are clear
- Used automated SIFT algorithm to match landmarks in the images
- Crop images to align area of interest
- Compute homography matrices to map sections of one image to the other
- Display the result

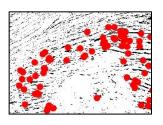


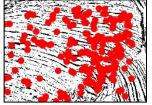


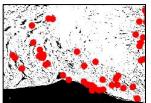
Our Approach

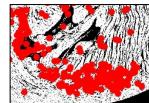
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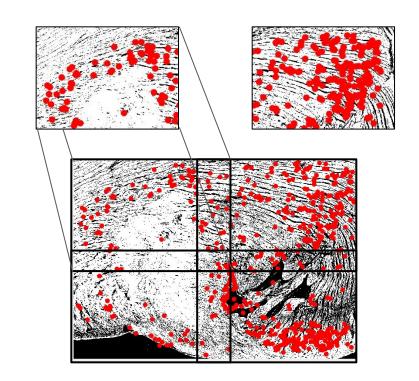


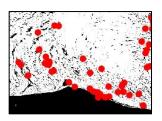


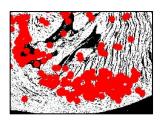


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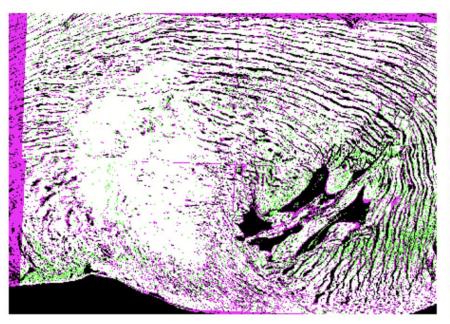


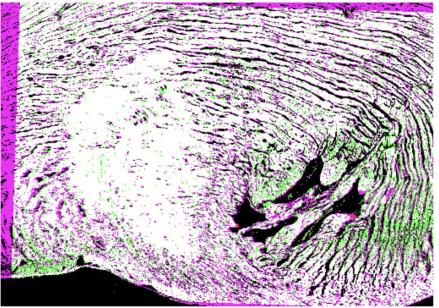




Results

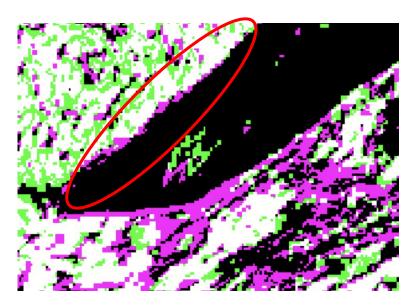
 Comparison of registration with patches (left) and with no patches (right)



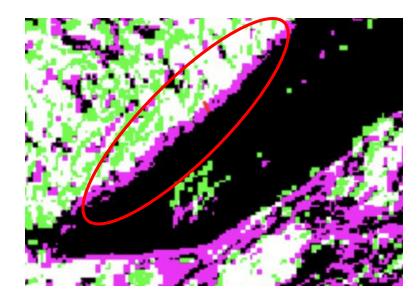


Results

Patches

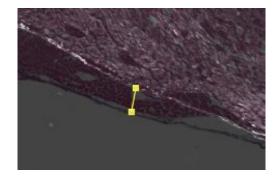


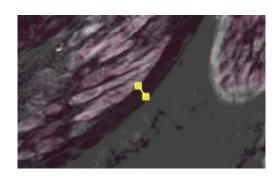
No Patches

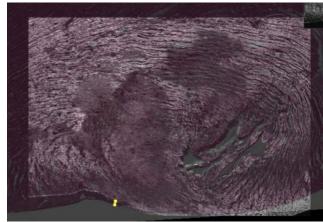


Comparison with Manual Registration

- Performed manual affine transform based on selected points in the image
- Accuracy is heavily dependent on the number of points
- Generally, affine accuracy was within 10s - 100s of pixels, while our algorithm was within 10s of pixels more globally
- Affine transform has more distortion away from selected points







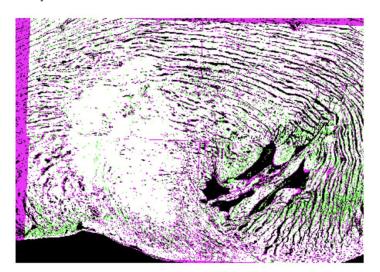
Limitations

- Works well when the tissue has well defined features that appear in both images, but not as well when the information contrasts (light in one, dark in other)
- Also works better when there are features in all four corners of the image



Takeaways

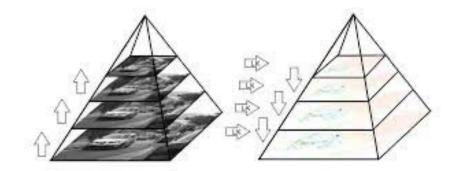
- Enables matching images with different information, scales, and warpings
- Runtime is dependent on number of patches and number of iterations, generally we ran in under 2-3 minutes, which is significant improvement over manual or MATLAB implementation



Future Work

- Homographies can be rescaled to apply to the original image enabling faster processing
- Compute homographies at iteratively finer scales to produce better local alignment

$$\begin{bmatrix} h_{11} & h_{12} & h_{13} \\ h_{21} & h_{22} & h_{23} \\ h_{31} & h_{32} & h_{33} \end{bmatrix} . * \begin{bmatrix} \frac{s_2}{s_1} & \frac{s_2}{s_1} & s_2 \\ \frac{s_2}{s_1} & \frac{s_2}{s_1} & s_2 \\ \frac{1}{s_1} & \frac{1}{s_1} & 1 \end{bmatrix}$$



Rescaling Homography Matrix

References

- Dan Mueller, Dirk Vossen, Bas Hulsken. (2011). Real-time deformable registration of multi-modal whole slides for digital pathology.
 Computerized Medical Imaging and Graphics. ISSN 0895-6111, Volume 35, Issues 7–8, Pages 542-556,
 https://doi.org/10.1016/j.compmedimag.2011.06.006.
- Jeremy L Muhlich, Yu-An Chen, Clarence Yapp, Douglas Russell, Sandro Santagata, Peter K Sorger, Stitching and registering highly multiplexed whole-slide images of tissues and tumors using ASHLAR, Bioinformatics, Volume 38, Issue 19, 1 October 2022, Pages 4613–4621, https://doi.org/10.1093/bioinformatics/btac544
- Krishna, V. et al. (2021). GloFlow: Whole Slide Image Stitching from Video Using Optical Flow and Global Image Alignment. In: , et al. Medical Image Computing and Computer Assisted Intervention MICCAI 2021. MICCAI 2021. Lecture Notes in Computer Science(), vol 12908. Springer, Cham. https://doi.org/10.1007/978-3-030-87237-3_50
- MATLAB, imregister, https://www.mathworks.com/help/images/ref/imregister.html
- Lowe, D. G. (2004). Distinctive Image Features from Scale-Invariant Keypoints. International Journal of Computer Vision, 60(2), 91–110. doi:10.1023/b:visi.0000029664.99615.94
- Bankhead, P. et al. QuPath: Open source software for digital pathology image analysis. *Scientific Reports* (2017).
- https://doi.org/10.1038/s41598-017-17204-5
- Bankhead, P. "QuPath Extension Align," https://github.com/qupath/qupath-extension-align

Thank you! Any questions?