Application Note: Traumatic Brain Injury





Introduction

Traumatic Brain Injury (TBI) is a significant health issue with neuroscience underlying its management and treatment. In the United States alone, there are approximately 2.5 million new cases each year, and about 5.3 million people living with long term disabilities as a result of TBI. In 2014, President Obama hosted the first summit on Healthy Kids and Safe Sports to raise awareness for sports-related concussions, while U.S. organizations committed over \$70 million to researching and raising awareness for TBI.

The public is growing to understand the long-term risks for youth who play contact sports and other groups who often suffer from concussions, such as soldiers. In clinical management of these cases, non-invasive imaging techniques such as CT and MRI are used to monitor patients and the extent of injury. These techniques also allow researchers to study the effects of TBI on cognitive tasks. However, non-invasive techniques are limited by the resolution they're able to provide. On the other hand, histology images tissue at a much higher resolution, and can validate results from other modalities, but only post-mortem. Therefore, methods like MRI and CT have to be combined with post-mortem histology to further our understanding of the brain. Of the technologies available for histology, whole-mount slide imaging such as Huron Digital Pathology's TissueScope™ solution provides researchers with the opportunity to effectively map the entire brain, and link detailed cytoarchitectural maps with functional imaging data.

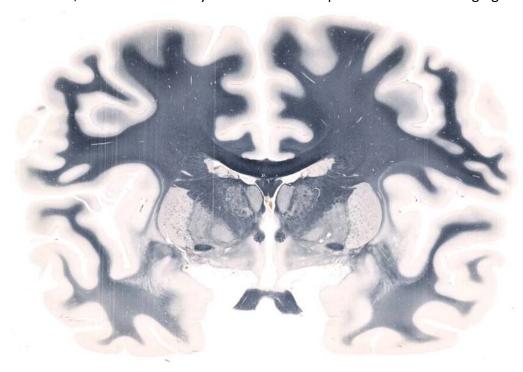


Figure 1. Whole-mount Brain slice scanned on a 5" x 7" slide at 20X resolution





Figure 2. The image in figure 1 magnified to highlight regions of interest in the left hemisphere.

Whole-mount slide imaging enables innovative processes and outcomes for brain research. Because the TissueScope can fit slides of up to 6" x 8", an entire slice of the human brain can be scanned at once, making a precise 3D reconstruction of the whole brain possible. Unlike traditional microscopy, sections do not have to be excluded, so a global image is more easily acquired. Reconstructing the brain from histology slides involves thinly slicing the brain along one axis. A tiling algorithm can be used so that for each slice of the brain thousands of uniformly spaced scans are taken at a high resolution.

With the TissueScope, this resolution can reach up to 0.2µm at 40x magnification, or 0.4µm at 20x. This provides researchers with a detailed, cellular mapping of the brain, with the potential to clarify uncertainties present in lower resolution in-vivo imaging modalities. Through software, brain slice images can be compiled into a 3D volume. Huron offers 3D histology software to provide volumetric reconstruction of whole-mount slides, so that hundreds of sections can be processed and viewed in minutes. Additionally, volumetric images from confocal and optical z-stacks can be read, visualized, and explored. From here, researchers can count cells, measure volume of specific regions, and more.





Figure 3. The Huron TissueScope LE, capable of scanning any size slide up to 6" X 8".

Additionally, the TissueSnap™ workflow accessory improve throughput by offloading preprocessing and preview scanning, so that the main scanner operates at maximum speed, running without interruption.

Because the whole-mount method allows for high resolution, 3D reconstruction, it enables new insights when combined with other modalities. In historically significant, long term cases, studies performed with non-invasive techniques like MRI or CT can be put into greater context in comparison to wholemount histology imaging. In patients with TBI, whole-mount imaging can reveal diffuse pathology in the brain's deep white matter, and allow the boundaries of lesions or injury to be more precisely determined.

Acquiring this high-resolution reference, further studies can lead to insight on neural circuits used in perceptual tasks and how those are affected by TBI. By combining whole-slide imaging with molecular methods, the effect that TBI has on global brain connectivity can be better understood.

Overall, a microscopic map of the brain can be linked to a variety of other structural and functional data. Histological study can validate the results of non-invasive MR-based imaging, improving the methods used in treating traumatic brain injury, ultimately to improve patient outcomes. Huron Digital Pathology has already supported a variety of human brain research through the acquisition of high resolution, whole brain images. Want to see more? Visit Huron's image gallery.



About Huron Digital Pathology

We believe that innovation is the key to making digital pathology a ubiquitous reality. But for us, innovation is about more than just delivering great image quality and fast scanning speeds. It's also about designing products that are simple to use, easy to integrate with your workflow, and have attractive price-performance characteristics.

Based in Waterloo, Ontario, Canada, Huron Digital Pathology has a 20 year history designing sophisticated imaging instrumentation. Our end-to-end digital whole slide scanning solutions for digital pathology incorporate our award-winning TissueScope™ digital slide scanners and our TissueSnap™ preview scanning station. Visit us at www.hurondigitalpathology.com.

