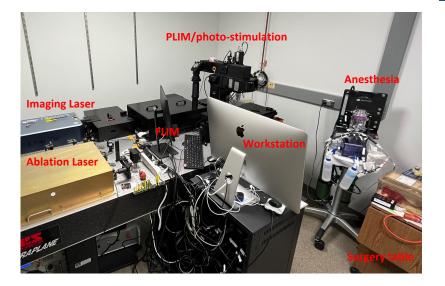
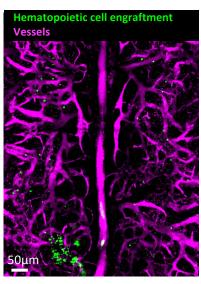
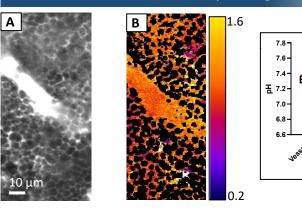
Intravital Multiphoton Imaging and Manipulation



Spatiotemporal tracking of rare cells

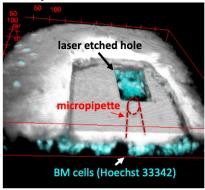


Quantification of interstitial calcium/pH at single cell resolution [1]



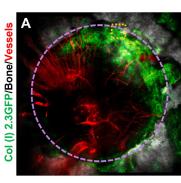
Plasma-mediated laser ablation and micromanipulaton [2]

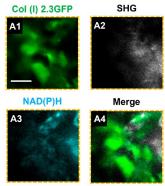
bone etching for micro-orthotopic injection or cell extraction



- (A) Intravenous delivery of cell impermeable, fluorescent pH or calcium probes that fill in the extracellular space of the bone marrow.
- (B) Ratiometric map and conversion to absolute pH based on calibration curves.

Fluorescence and phosphorescence lifetime measurements in vivo^[3,4]





p< 0.0001

B

2.5
2.0

Next 1.5

Defect Nearby Native Bone Osteoblast Location

Defect Nearby Native Bone Measurement Location

D Autofluorescence lifetime imaging of a label-free bone marrow

Bone

50 μm

- Zoomed image
- (A) 1 mm cranial defect with bone, osteoblast, and vasculature visualized intravital two-photon imaging.
- (B) In vivo fluorescence lifetime of NAD(P)H from osteoblasts
- (C) In vivo phosphorescence lifetime measurements for direct quantification of pO₂.
- Autofluorescence lifetime of bone lining populations and hematopoietic cells from a label-free bone marrow

References

- [1] Yeh SCA, Hou J et al, Nat Comm, 13(1):393, 2022
- [2] Haase C. et al, Nat Methods, 19(12):1622-1633, 2022
- [3] Schilling K, Zhai Y et al, *Elife*, 11:e83146, 2022
- [4] Schilling K, Brown E, and Zhang X, Bone, 116257, 2021