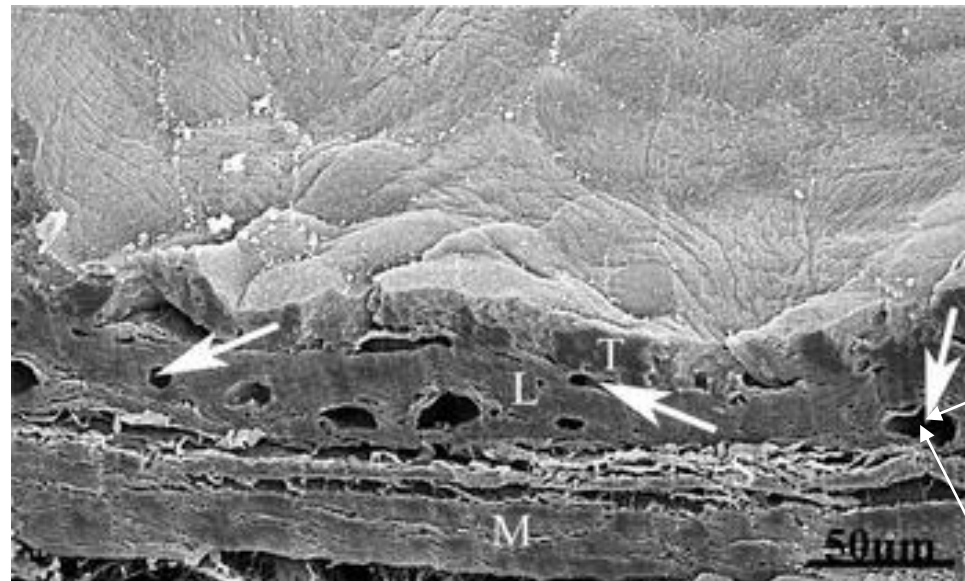
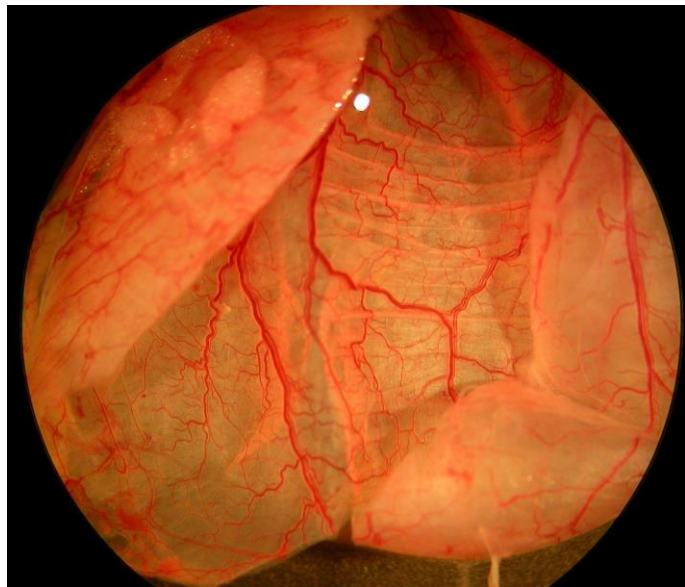


Crossing the Disciplinary Divide - To Chase The Phantom of Bladder Permeability

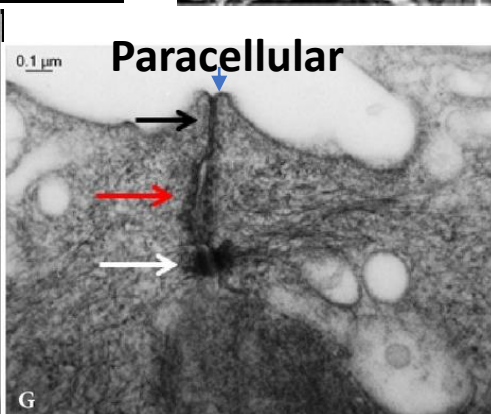
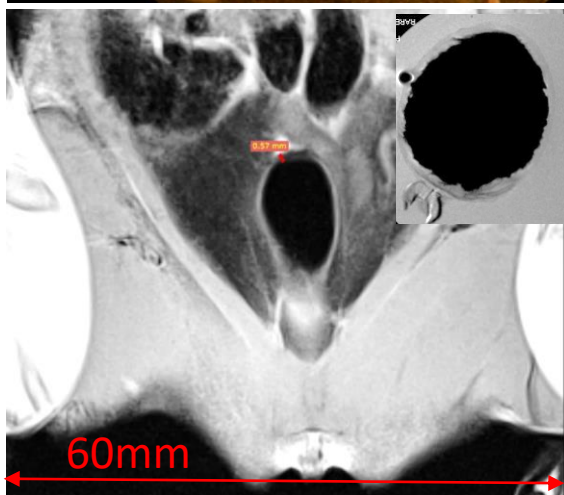
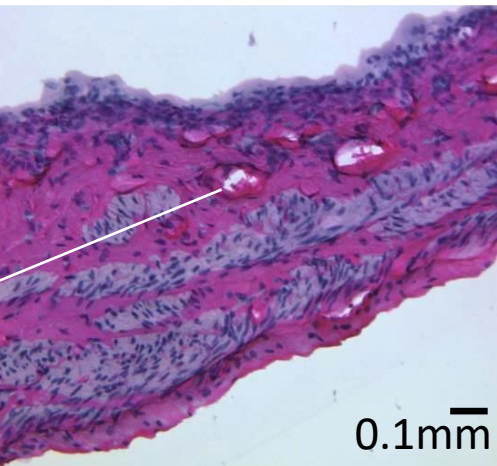
Pradeep Tyagi PhD
University of Pittsburgh



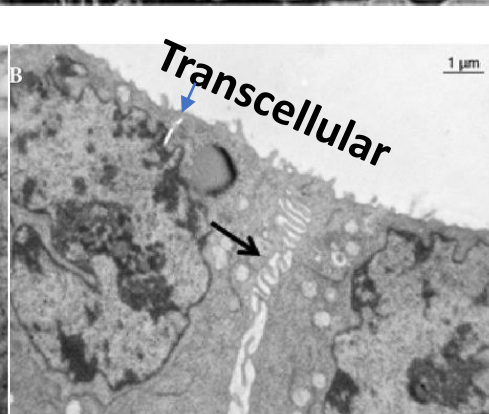
Bladder Wall Architecture & Permeability



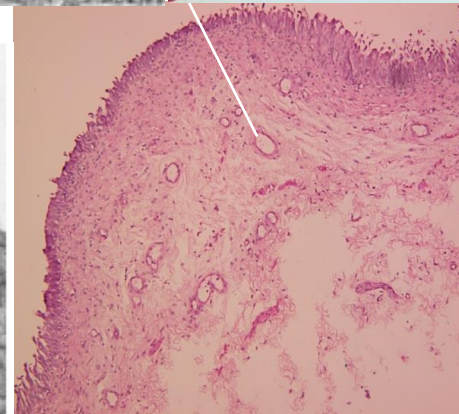
- Conc. In **Blood** ?
- Conc. In Bladder



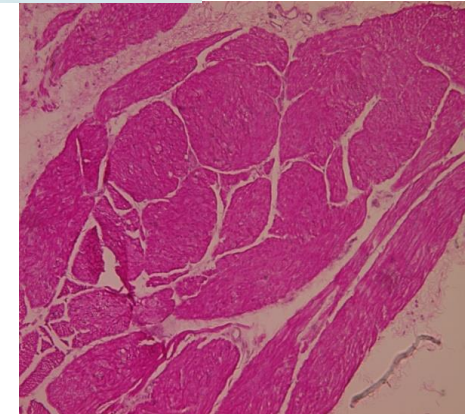
Normal



Diseased



1.2mm



10mm

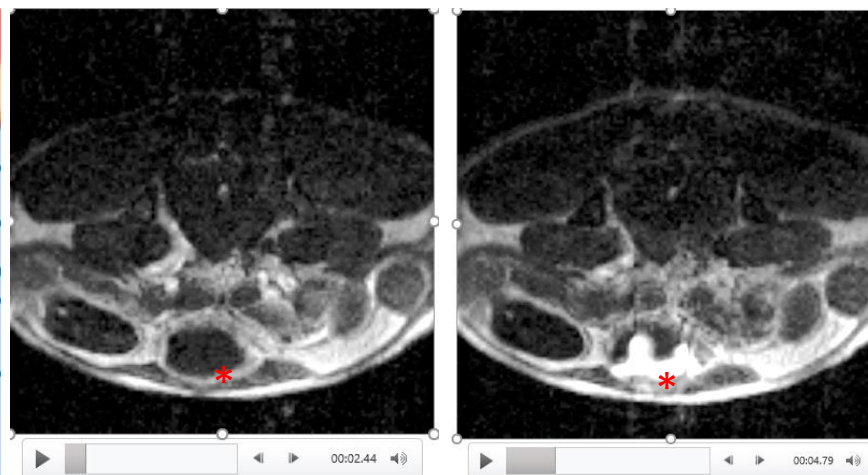
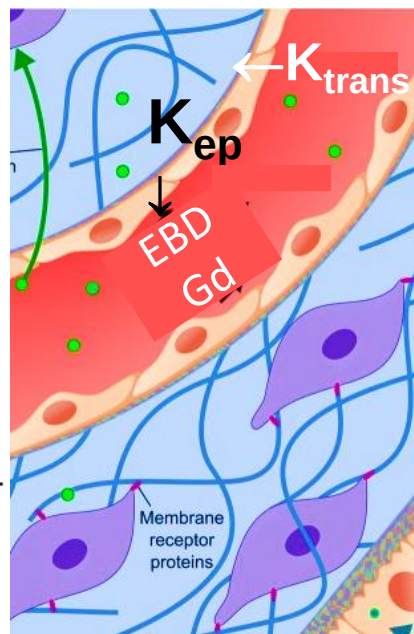
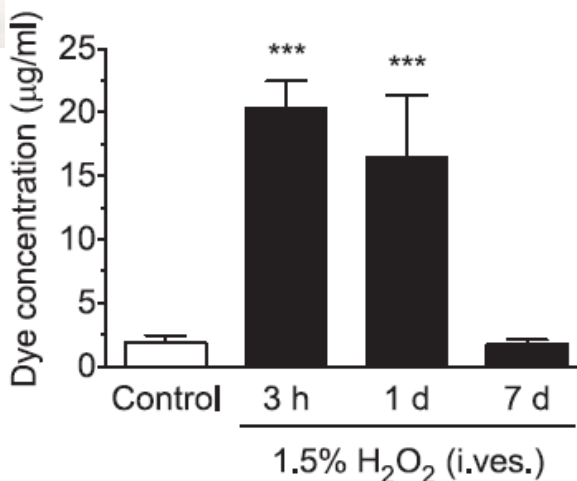
Why Cross the Disciplinary Divide ?

Evans blue dye EBD



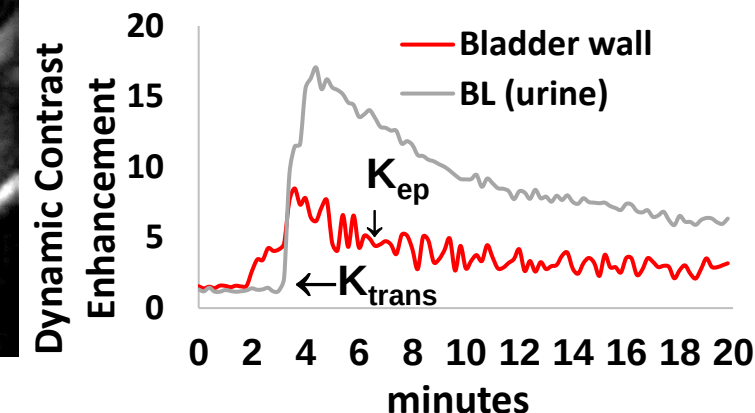
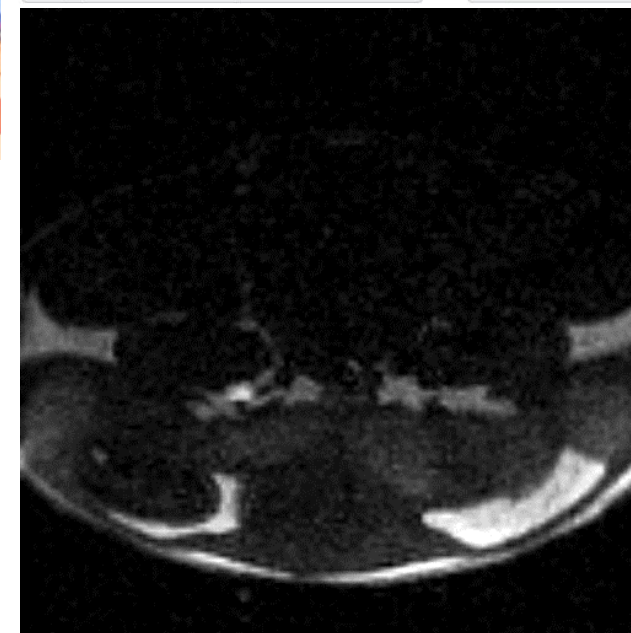
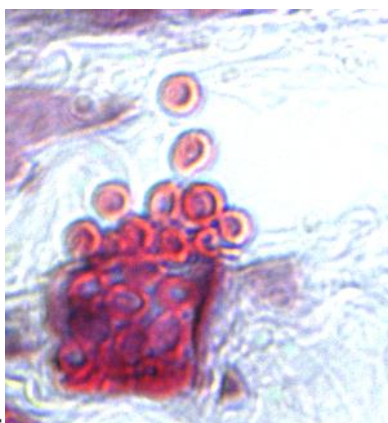
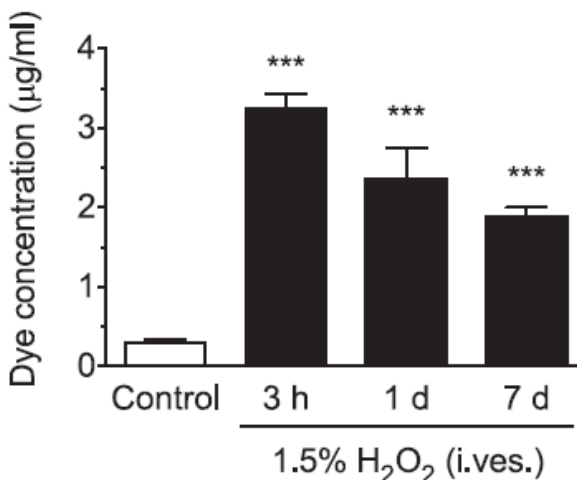
Paracellular 961 Daltons

A) Urothelial permeability



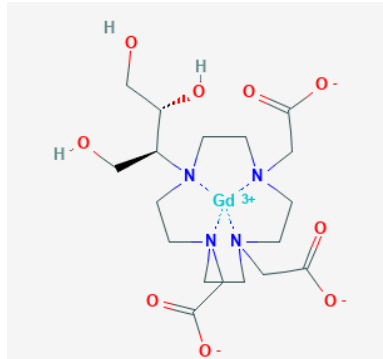
Gadobutrol
607 Daltons
Paracellular

B) Vascular permeability

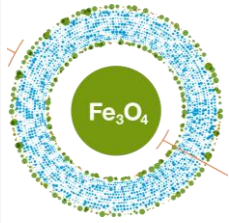


Homan et al J Pharmacol Sci, **121**: 327, 2013;

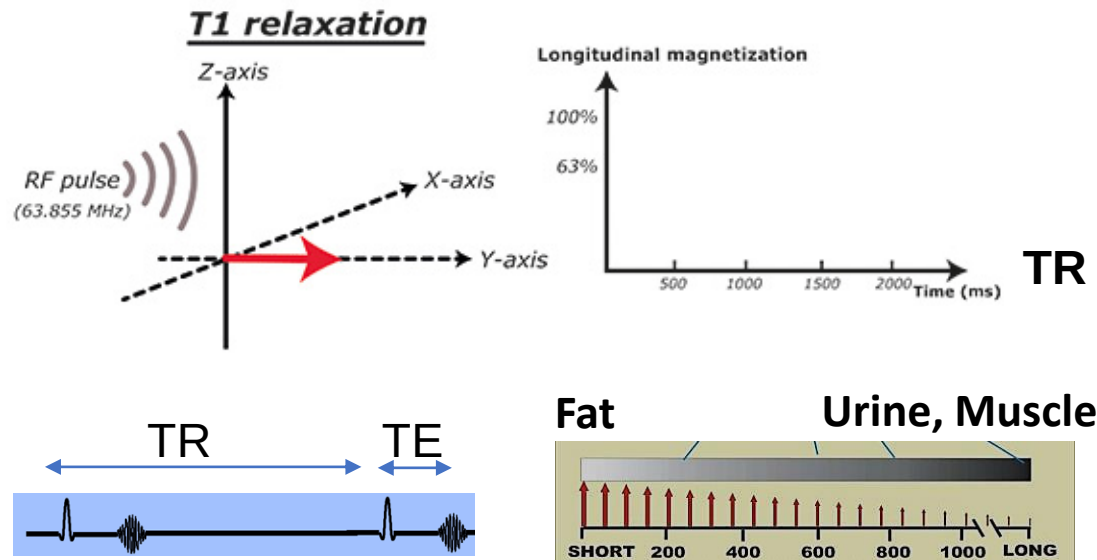
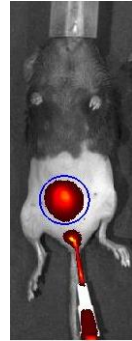
Instillation of Gadobutrol Mixture



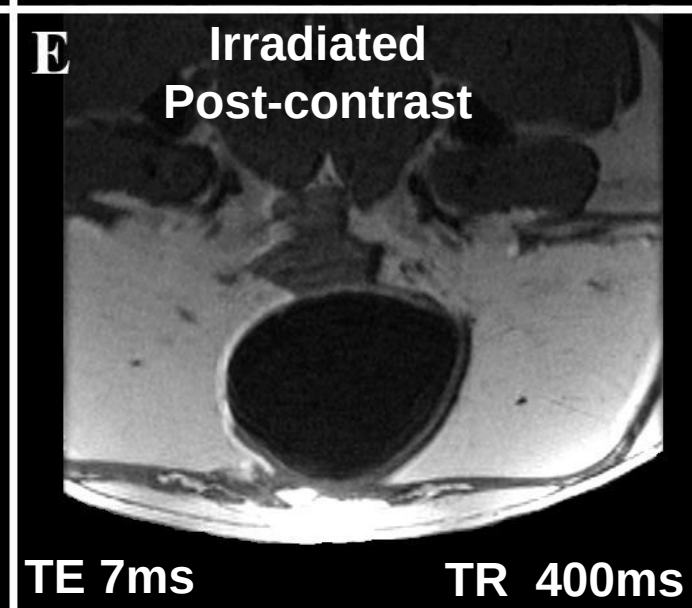
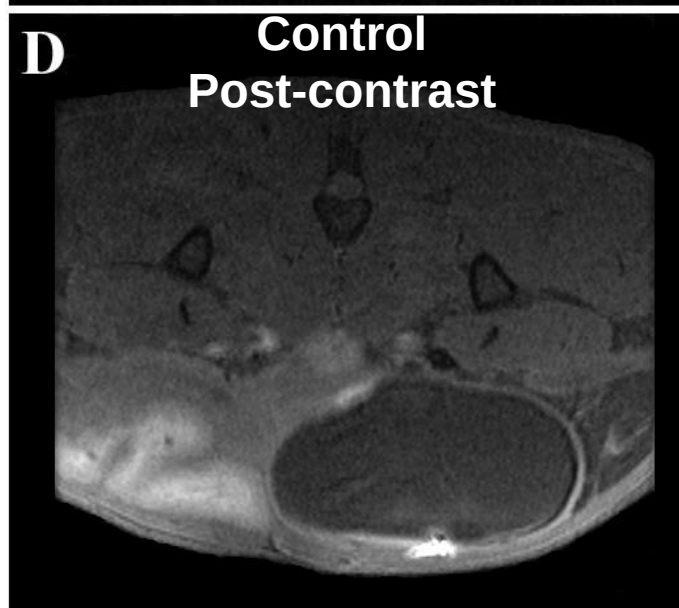
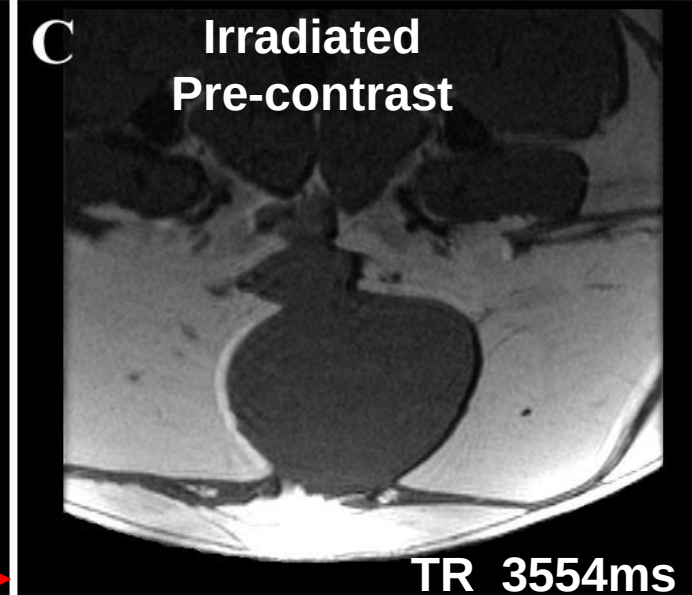
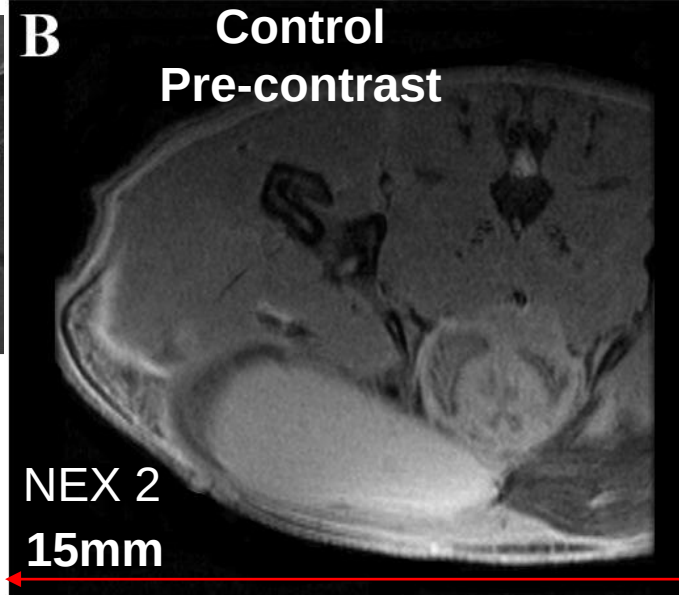
Gadobutrol 4mM
607 Daltons



+ Ferumoxytol 5mM
731 KiloDaltons

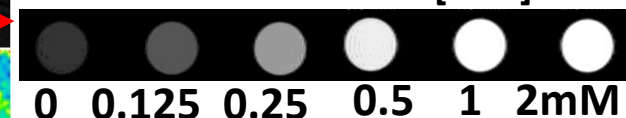
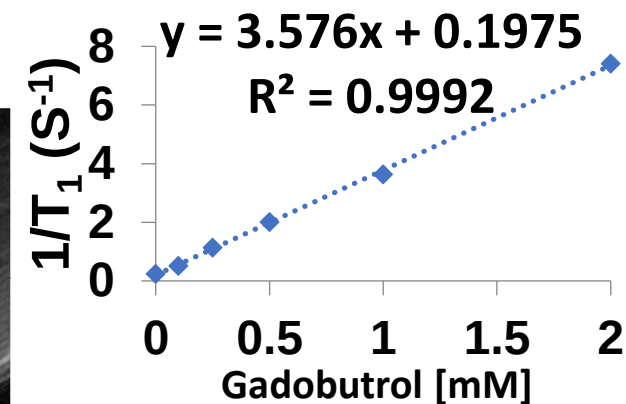
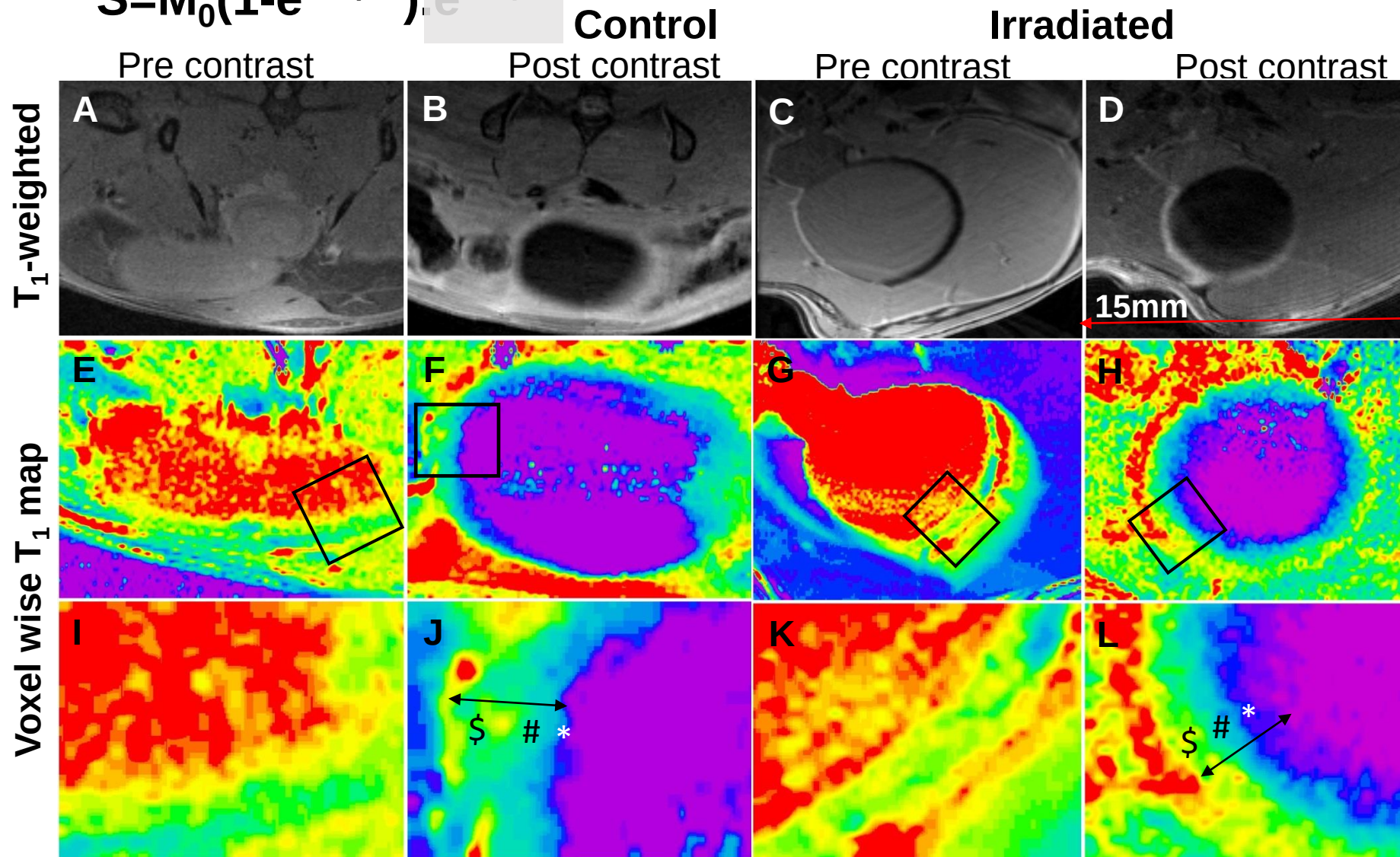


$$S = M_0(1 - e^{-TR/T1}) \cdot e^{-TE/T2}$$



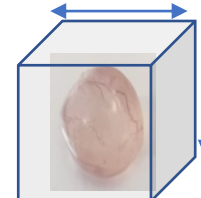
Voxel-wise T_1 mapping at 9.4T

$$S = M_0(1 - e^{-TR/T_1}) \cdot e^{-TE/T_2}$$



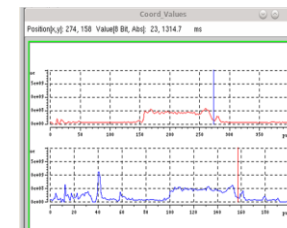
5000ms

Matrix 200

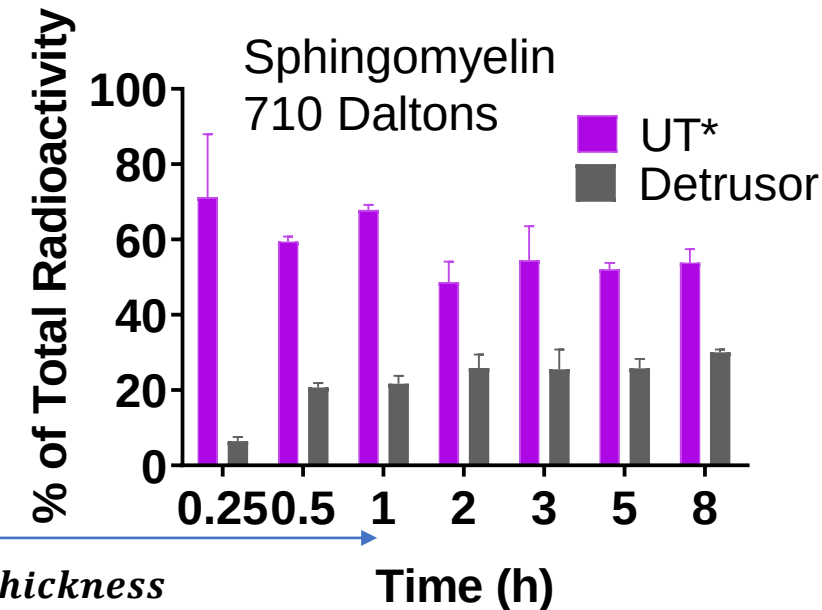
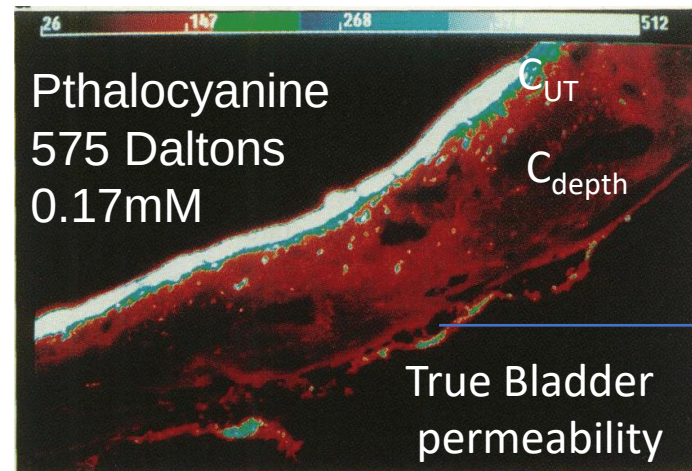
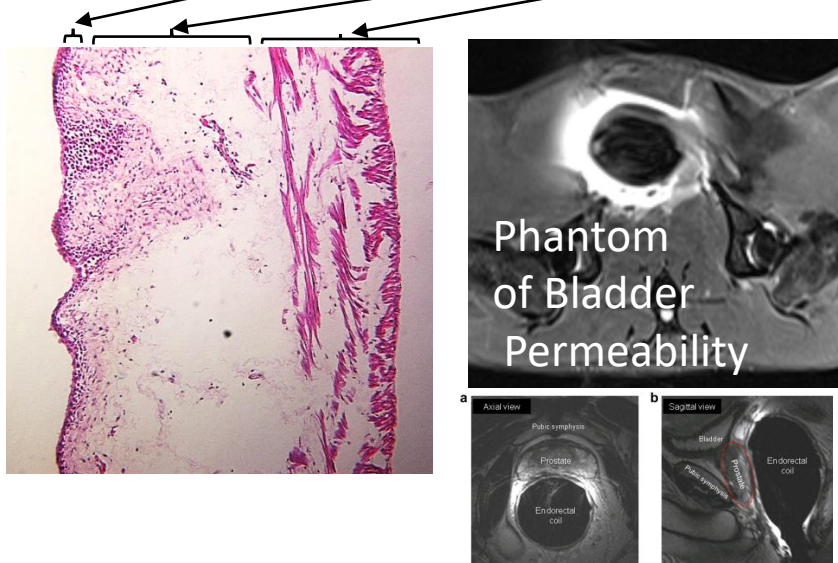
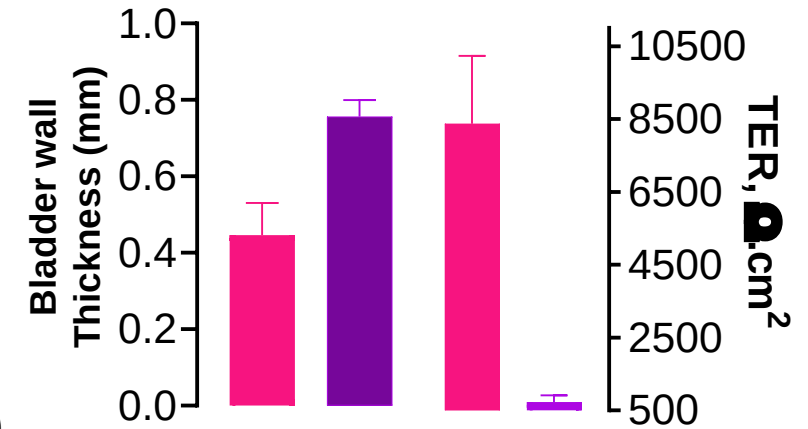
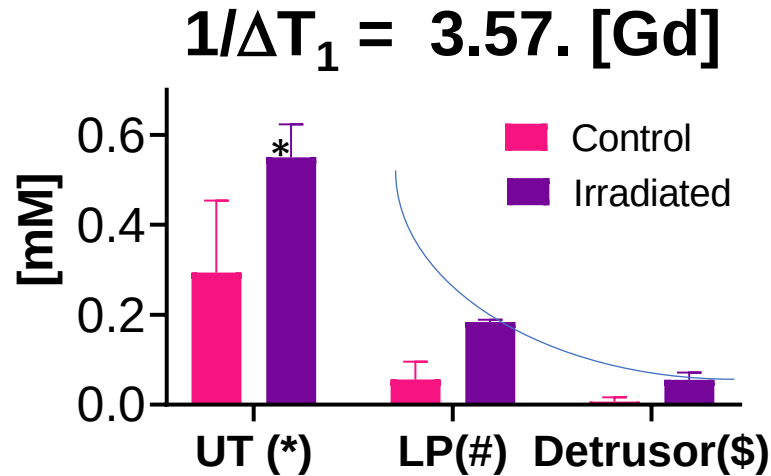
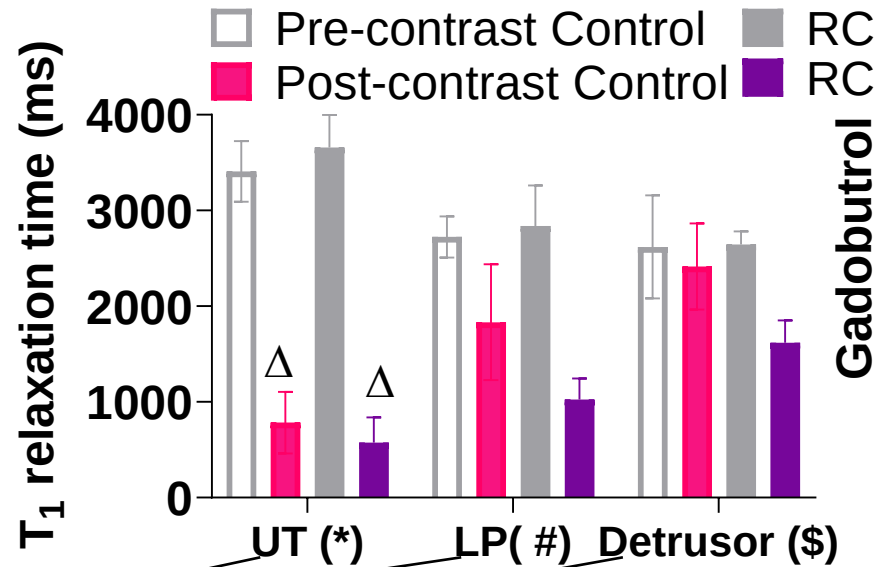


Slice
Thickness
0.7mm

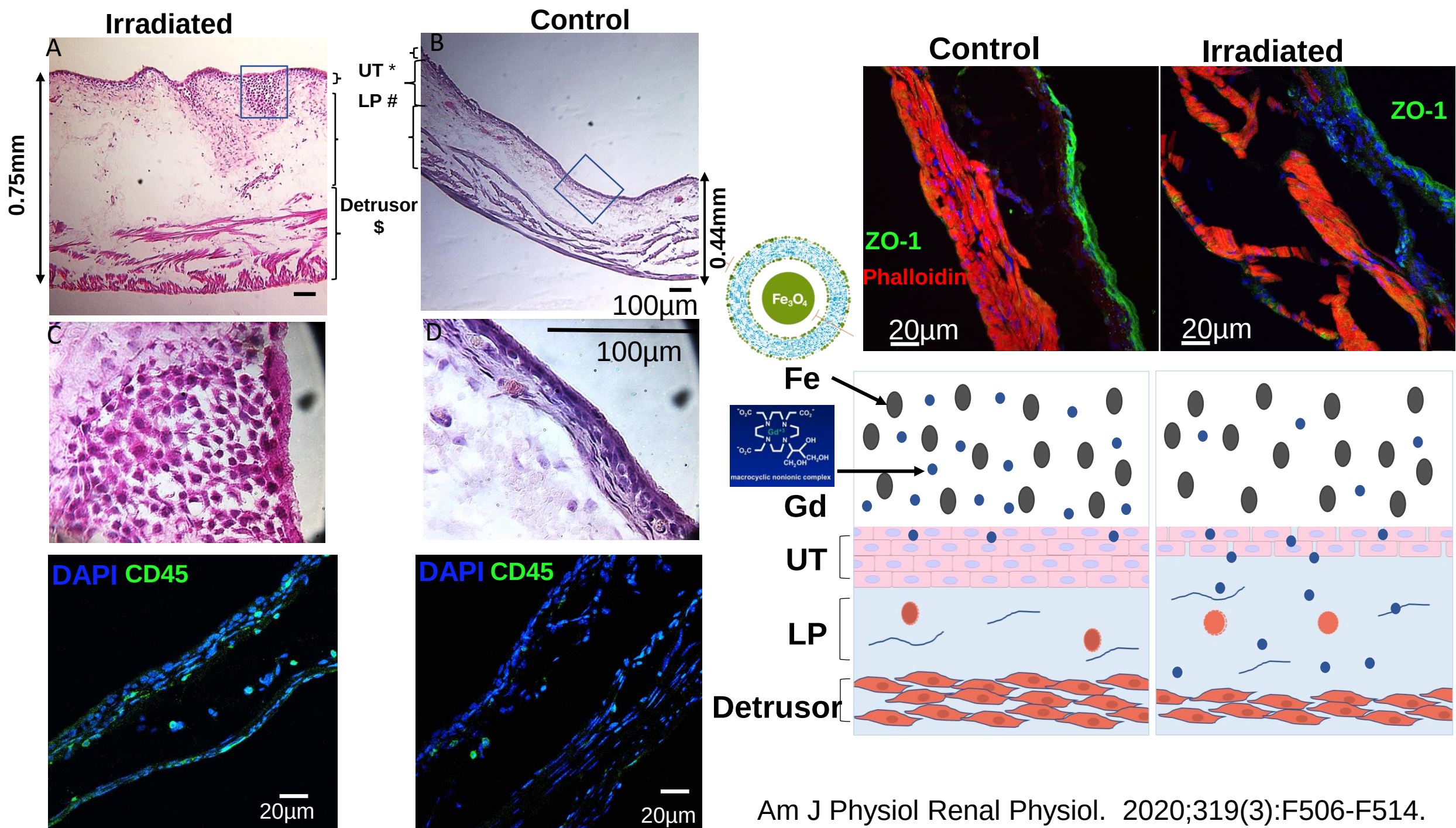
0 ms
 T_1
relaxation time



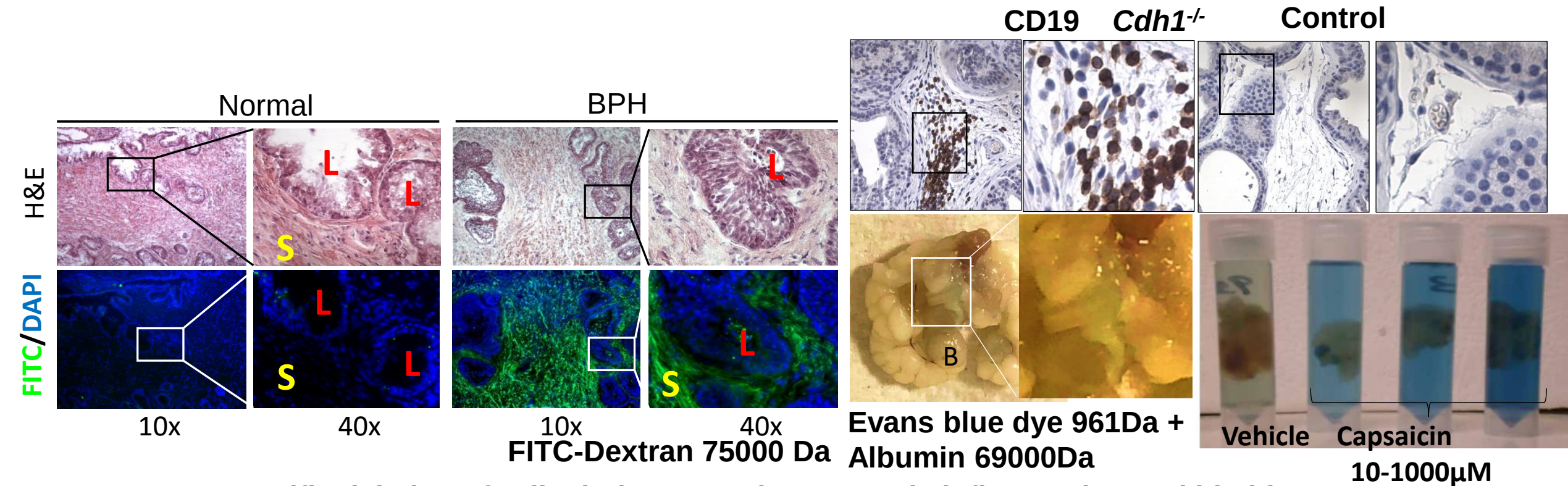
Data Mining for Permeability Assay



$$C_{Depth} = (C_{UT} - C_b) e^{-\frac{0.693}{w1} UT \text{ thickness}} + C_b$$



Bladder To Prostate (BPH)



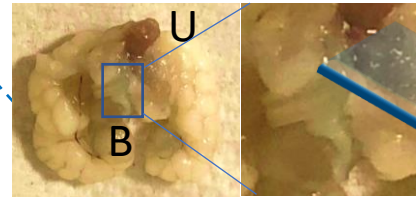
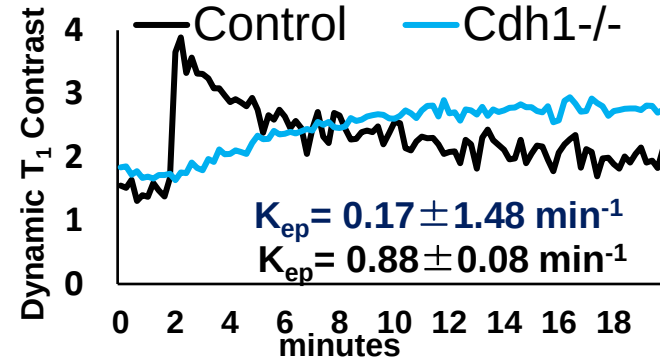
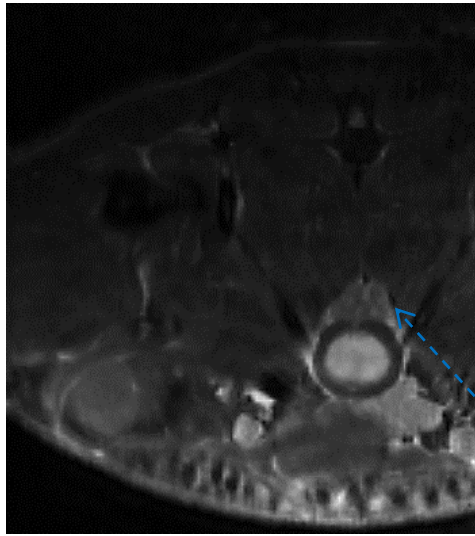
Prostate-specific deletion of *Cdh1* induces murine prostatic inflammation and bladder overactivity- Endocrinology

- What comes first- prostatic inflammation or prostatic permeability ?
- What is the effect of molecular targeted drugs on PI and permeability ?
- Clinical feasibility of MRI based permeability assay using gadolinium analog?

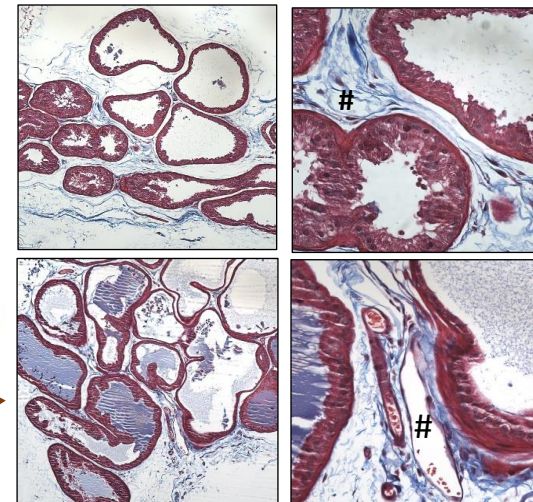
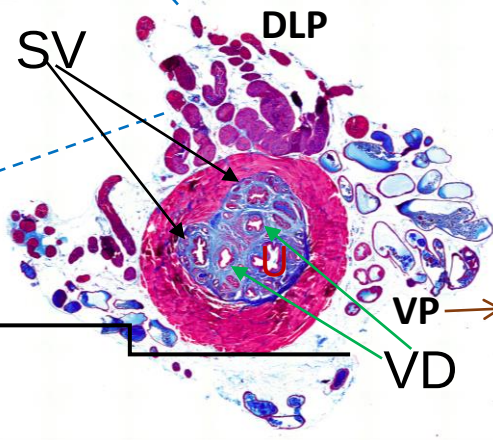
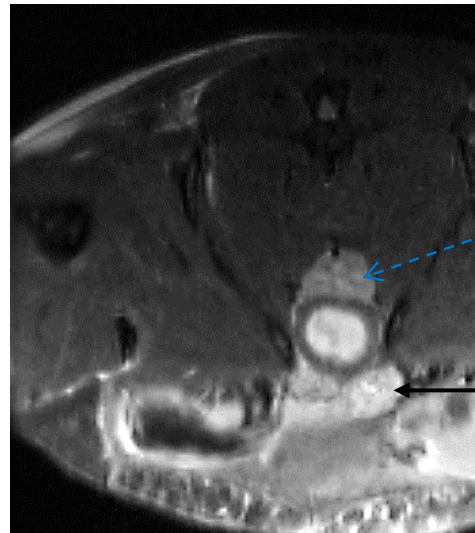
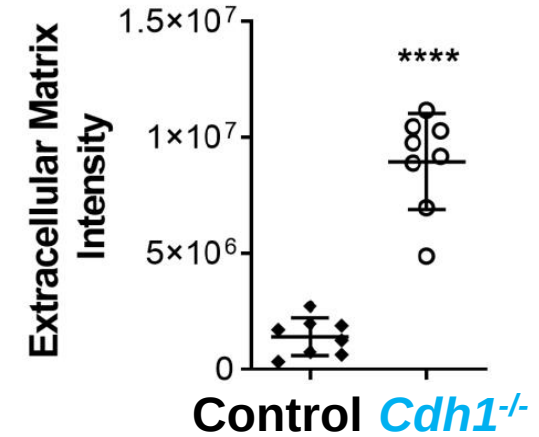
Endocrinology 2020 Nov 19:bqaa212. doi: 10.1210/endocr/bqaa212.
Eur Urol. 2007;51(4):1119-27

Replace EBD with Gadobutrol Injection at 7T

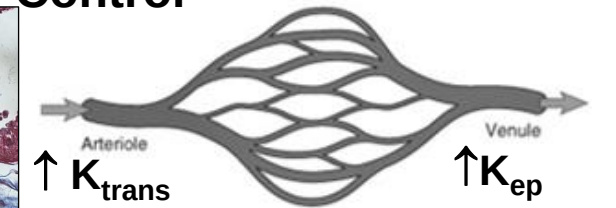
Axial



Axial



Control



Cdh1^{-/-}



Conclusions

- Crossing the disciplinary divide can virtually predict histology and function of diseased bladder and prostate
- Voxel-wise T_1 mapping can be a potential pain-free alternative to PST and biopsy of BPS/IC patients
- mpMRI for prostate cancer- when tweaked with basic research insights can advance our understanding of BPH/LUTS

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- Lesley Foley

NIDDK- DK 108397, U54 DK112079,
NCI- CA252590, P30CA047904 , Hillman Cancer center
NIA- AG062971