

Project 15:

Mouse segmentation and optical properties for bioluminescence tomography (BLT)

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Topic and Goals

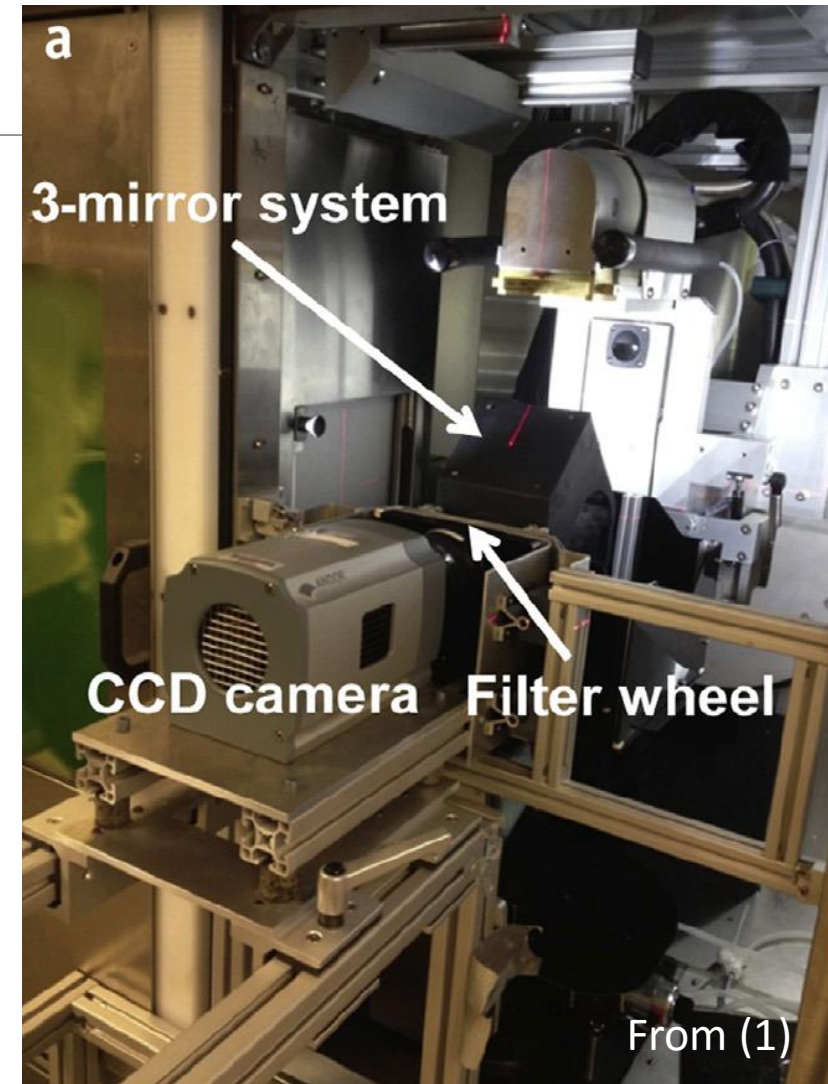
- Gather literature values of mouse organ optical properties and evaluate their distribution
- Expedite the segmentation of cone beam computed tomography (CBCT) images of mice.
- Modify existing BLT reconstruction to address optical property heterogeneity.
 - Experiment with optical property sets for best reconstruction.



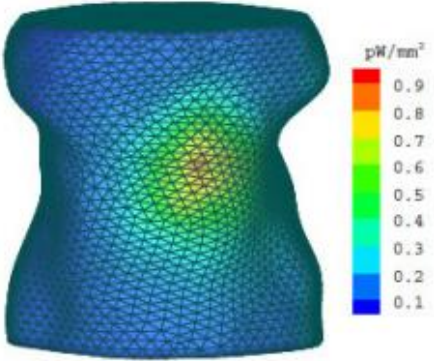
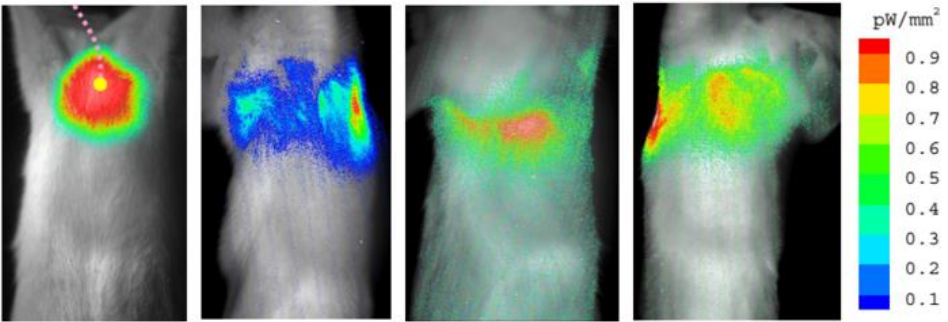
Statement of Relevance

Small Animal Radiation Research Platform (SARRP)

- Tool for preclinical radiation research
- CBCT for radiation delivery guidance
- CBCT has less utility for localizing small/low-contrast targets in soft tissue
- Hence, incorporated bioluminescence tomography (BLT)



Basic Idea of BLT



(b)



(c)

Images from (16)

Basic Idea of BLT

Diffusion Approximation

Equations from (1):

$$\begin{cases} -\nabla \cdot D(r) \nabla \Phi(r) + \mu_a(r) \Phi(r) = S(r), & r \in \Omega \\ \Phi(\xi) + 2A\hat{n} \cdot D(\xi) \nabla \Phi(\xi) = 0, & \xi \in \partial\Omega \end{cases}$$



Green's Functions

$$\begin{bmatrix} \varphi_1 \\ \vdots \\ \varphi_M \end{bmatrix} = \begin{bmatrix} G_{1,1} & \cdots & G_{1,N} \\ \vdots & \ddots & \vdots \\ G_{M,1} & \cdots & G_{M,N} \end{bmatrix} \begin{bmatrix} s_1 \\ \vdots \\ s_N \end{bmatrix}$$



Multispectral Approach (590, 610, 630, 650 nm)

$$\begin{bmatrix} \varphi(\lambda_1) \\ \vdots \\ \varphi(\lambda_k) \end{bmatrix} = \begin{bmatrix} \eta(\lambda_1) G(\lambda_1) \\ \vdots \\ \eta(\lambda_k) G(\lambda_k) \end{bmatrix} [s]$$

Project 15:

To address optical property heterogeneity in reconstruction:

Summary of Technical Approach

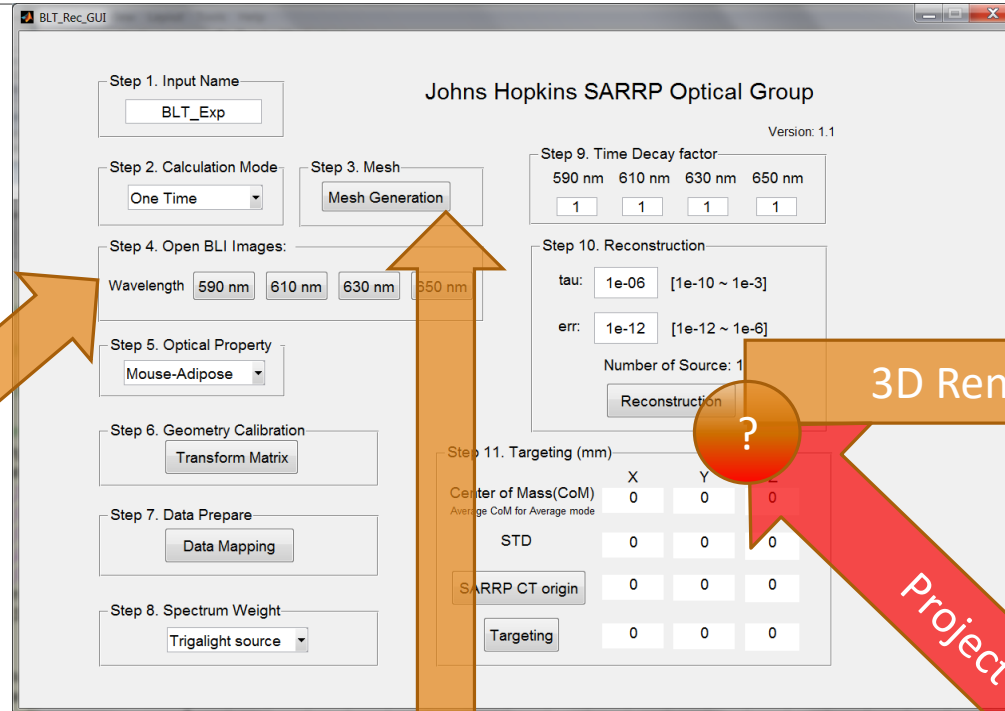


Image Prepared Mouse

CBCT

Combine DICOM slices to single .img volume

→

Crop and segment mouse body from background

→

Meshes Defining Organ Spatial Distributions

→

Map from Organ Type to Optical Properties

→

3D Rendering of Reconstructed Source

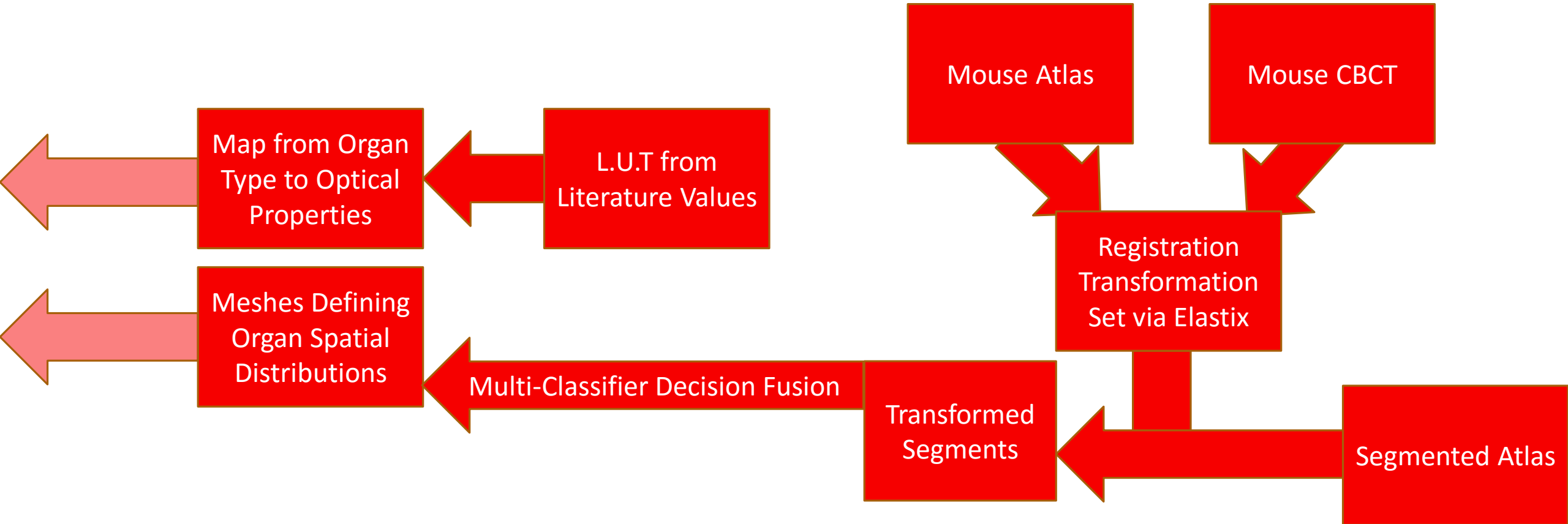
BLI

3D Rendering of Reconstructed Source

Project 15



Project 15 Approach



Deliverables

Minimum Deliverables

- Tabulate literature values for optical properties
- Manually segment mouse images for atlas and simulated source
- Modify Matlab code to incorporate organ specific optical properties
- Test code under simulation conditions

Expected Deliverables

- Workflow for registering new images to atlas set using elastix
- Matlab code for multi-classifier decision fusion strategy

Maximum Deliverables

- Perform BLT experiment on implanted light source in specific organ
- Determine optimal optical property value sets for reconstruction

Project Timeline

Key Milestones Highlighted	Week of:										
	February		March				April				May
	21	28	06	13	20	27	03	10	17	24	01
Read <u>Elastix</u> Manual (2-3)											
Read Core Literature (5-13)		1									
Project Plan + Presentation											
Read BLT documentation											
Run BLT on Sample Images			2								
Seminar Presentation											
Manual Segment Atlas Set											
Checkpoint Presentation											
Second Literature Round						3					
Modify BLT code											
Test BLT in Simulation								4			
Try <u>Elastix</u> Parameters											
Multi-class decision fusion											
Experiments with new sets										5	
Final Exam + Poster Session											

1. **3/5**: Finished tabulating core literature results and main reading phase.
 - Ready for seminar presentation for week of 3/06
2. **3/12**: Able to execute existing BLT workflow and begin modification
3. **3/27**: Modified BLT code to incorporate optical properties information
 - Manual segmentations for atlas completed
 - Finished optical property data gathering
 - Ready for checkpoint presentation for week of 3/27
4. **4/16**: Tested modified BLT with light source simulation
 - Decided on Elastix registration parameters
5. **4/30**: Finished experimenting with reconstruction on new data from implanted sources
 - Ready to produce final report and presentation

Dependencies

Resource	Status	Comment
Mouse image set for initial BLT practice	Received	
Mouse image sets for atlas + experiments	Unknown	To be discussed w/ mentor 2/29 Digimouse alternative/complement (14,15)
Matlab source code	Received	
SAARP/BLT workflow documentation	Received	
Elastix registration software	Installed	
Nirfast light transport modeling software	Received	

Management Plan

- Weekly meetings with mentor
- Regular e-mail correspondence
- Progress updates on project wiki

Skills

- CIS Prerequisites

Preliminary Reading List

- 1) Zhang, B., Wang, K.K., Yu, J., Eslami, S., Iordachita, I., Reyes, J., ... Wong, J.W. Bioluminescence Tomography–Guided radiation therapy for preclinical research. *International Journal of Radiation Oncology *Biology *Physics*, doi:<http://dx.doi.org/10.1016/j.ijrobp.2015.11.039>
- 2) Klein, S., Staring, M., Murphy, K., Viergever, M.A., Pluim, J.P.W. "elastix: a toolbox for intensity based medical image registration," *IEEE Transactions on Medical Imaging*, vol. 29, no. 1, pp. 196 - 205, January 2010. [download doi](#)
- 3) Shamonin, D.P., Bron, E.E., Lelieveldt, B.P.F., Smits, M., Klein, S., Staring, M. "Fast Parallel Image Registration on CPU and GPU for Diagnostic Classification of Alzheimer's Disease", *Frontiers in Neuroinformatics*, vol. 7, no. 50, pp. 1-15, January 2014. [download doi](#)
- 4) Rohlfing, T., Brandt, R., Menzel, R., & Maurer Jr., C. R. (2004). Evaluation of atlas selection strategies for atlas-based image segmentation with application to confocal microscopy images of bee brains. *Neuroimage*, 21(4), 1428-1442. doi:<http://dx.doi.org/10.1016/j.neuroimage.2003.11.010>
- 5) Honda N, Ishii K, Terada T, Nanjo T, Awazu K; Determination of the tumor tissue optical properties during and after photodynamic therapy using inverse monte carlo method and double integrating sphere between 350 and 1000 nm. *J. Biomed. Opt.* 0001;16(5):058003-058003-7. doi:10.1117/1.3581111.
- 6) Kienle, A., Lilge, L., Patterson, M.S., Hibst, R., Steiner, R., and Wilson, B.C. "Spatially resolved absolute diffuse reflectance measurements for noninvasive determination of the optical scattering and absorption coefficients of biological tissue," *Appl. Opt.* 35, 2304-2314 (1996)
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- 8) Bashkatov, A.N., Genina, E.A., Tuchin, V.V. (2011). Optical properties of skin, subcutaneous, and muscle tissues: A review. *Journal of Innovative Optical Health Sciences*, 04(01), 9-38.
- 9) Cheong, W., Prah, S.A., & Welch, A. J. (1990). A review of the optical properties of biological tissues. *Quantum Electronics, IEEE Journal of*, 26(12), 2166-2185.
- 10) Firbank, M., Hiraoka, M., Essenpreis, M., and Delpy, D.T. (1993). Measurement of the optical properties of the skull in the wavelength range 650-950 nm. *Physics in Medicine and Biology*, 38(4), 503.
- 11) Sandell, J.L., & Zhu, T.C. (2011). A review of in-vivo optical properties of human tissues and its impact on PDT. *Journal of Biophotonics*, 4(11-12), 773-787.
- 12) Jacques, S.L. (2013). Optical properties of biological tissues: A review. *Physics in Medicine and Biology*, 58(11), R37
- 13) Welch, A.J, Gemert, M.J.C. Optical-thermal response of laser-irradiated tissue. Dordrecht: Springer; 2011
- 14) B. Dogdas, D. Stout, A. Chatziioannou, RM Leahy, Digimouse: A 3D Whole Body Mouse Atlas from CT and Cryosection Data, *Phys. Med. Bio*, 52: 577-587, 2007. <http://dx.doi.org/10.1088%2F0031-9155%2F52%2F3%2F003>
- 15) D. Stout, P. Chow, R. Silverman, R. M. Leahy, X. Lewis, S. Gambhir, A. Chatziioannou, Creating a whole body digital mouse atlas with PET, CT and cryosection images, *Molecular Imaging and Biology*.2002; 4(4): S27
- 16) Ge Wang, Wenxiang Cong, Kumar Durairaj, Xin Qian, Haiou Shen, Patrick Sinn, Eric Hoffman, Geoffrey McLennan, and Michael Henry, "In vivo mouse studies with bioluminescence tomography," *Opt. Express* 14, 7801-7809 (2006)