## WATARU OTSU, DVM, Ph.D.

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### **Education**

2013	Doctor of Philosophy, Graduate School of Veterinary Medicine, Hokkaido University
2009	Doctor of Veterinary Medicine, Graduate School of Veterinary Medicine, Hokkaido University

## **Positions and Employment**

2019-present	Research Assistant Professor, Department of Biomedical Research, Gifu Pharmaceutical University
2019	Research Associate, Department of Ophthalmology, Weill Cornell Medical College
2014-2018	Postdoctoral Associate, Department of Ophthalmology, Weill Cornell Medical College
2013	Visiting Researcher, Graduate School of Veterinary Medicine, Hokkaido University

#### **Research interests**

- Membrane trafficking and intracellular organelle dynamics
- Protein quality control and degradation pathway
- Development of the central nervous system
- Physiology of cone photoreceptors and retinal diseases

## **Publications**

- 1. Ando S., <u>Otsu W.</u>, Osanai D., Kamiya S., Ishida K., Nakamura S., Shimazawa M., and Hara H. Survival motor neuron protein modulates lysosomal function through the expression of transcription factor EB in motoneurons. *BPB. Reports*, 3(4): 130-137, **2020**.
- 2. Saito Y., Okuyoshi H., Nakamura S., <u>Otsu W.</u>, Yamaguchi A., Hitchcock P.F., Nagashima M, Shimazawa M., and Hara H. Tauroursodeoxycholic acid promotes neuronal survival and proliferation of tissue resident stem and progenitor cells in retina of adult zebrafish. *BPB. Reports*, 3(3): 92-96, **2020.**
- Inagaki S., Shimazawa M., Hamaguchi K., <u>Otsu W.</u>, Araki T., Sasaki Y., Numata Y., Tsusaki H., and Hara H. Anti-VGEF antibody inhibits vascular leakage and decreases subretinal fibrosis to some degree in a cynomolgus monkey CNV model. *Curr. Neurovasc. Res.*, May 23 2020.
   PubMed PMID: 32445455
- 4. Kiatpakdee B., Sato K., Otsuka Y., Arashiki N., Chen Y., Tsumita T., <u>Otsu W.</u>, Yamamoto A., Kawata R., Yamazaki J., Sugimoto Y., Takada K., Mohandas N., and Inaba M. Cholesterol-binding protein TSPO2 coordinates maturation and proliferation of terminally differentiating erythroblasts. *J. Biol. Chem.*, 295(23):8048-8063, **2020**.
  - PubMed PMID: 32358067.
- Otsu W., Ishida K., Nakamura S., Shimazawa M., Tsusaki H., and Hara H. Blue light-emitting diode irradiation promotes transcription factor EB-mediated lysosome biogenesis and lysosomal cell death in murine photoreceptor-derived cells. *Biochem. Biophys. Res. Commun.*, 526(2):479-484, 2020.
   PubMed PMID: 32234235.

- 6. Saito Y., Yako T., Otsu W., Nakamura S., Inoue Y., Muramatsu A., Nakagami Y., Shimazawa M., and Hara H. A triterpenoid NRF2 activator, RS9, promotes LC3-associated phagocytosis of photoreceptor outer segments in a p62-independent manner. *Free Radic. Biol. Med.*, 152:235-247, 2020. PubMed PMID: 32217192.
- 7. Hsu KS\*, Otsu W\*, Li Y., Wang HC., Chen S., Tsang S., Chuang, JZ, and Sung, CH. CLIC4 regulates late endosomal trafficking and matrix degradation activity of MMP14 at focal adhesions in RPE cells. Sci. Rep., in press, \*contributed equally.
- 8. <u>Otsu W.</u>Hsu YC, Chuang, JZ, and Sung, CH. The Late Endosomal Pathway Regulates the Ciliary Targeting of Tetraspanin Protein Peripherin 2. *J. Neurosci.* 2019 May 1;39(18). 3376-3393. PubMed PMID: 30819798.
- 9. <u>Otsu W,</u> Chuang, JZ, and Sung, CH. Establishment of a novel cone transfection technique for cone-rod dystrophy mouse models. (under preparation)
- 10. Saito M, Sakaji, K, <u>Otsu W</u>, and Sung CH. Ciliary Assembly/Disassembly Assay in Non-transformed Cell Lines. *Bio-protocol* 2018 8(6): e2773. DOI: <u>10.21769/BioProtoc.2773</u>.
- 11. Saito M, Otsu W, Hsu KS, Chuang JZ, Yanagisawa T, Shieh V, Kaitsuka T, Wei FY, Tomizawa K, Sung CH. Tctex-1 controls ciliary resorption by regulating branched actin polymerization and endocytosis.

  EMBO Rep. 2017 Aug;18(8):1460-1472. PubMed PMID: 28607034; PubMed Central PMCID: PMC5538770.
- 12. Chou SY\*, Hsu KS\*, Otsu W\*, Hsu YC, Luo YC, Yeh C, Shehab SS, Chen J, Shieh V, He GA, Marean MB, Felsen D, Ding A, Poppas DP, Chuang JZ, Sung CH. CLIC4 regulates apical exocytosis and renal tube luminogenesis through retromer- and actin-mediated endocytic trafficking. *Nat Commun.* 2016 Jan 20;7:10412. PubMed PMID: 26786190; PubMed Central PMCID: PMC4736046. \*contributed equally.
- 13. Otsu W, Kurooka T, Otsuka Y, Sato K, Inaba M. A new class of endoplasmic reticulum export signal PhiXPhiXPhi for transmembrane proteins and its selective interaction with Sec24C. *J Biol Chem.* 2013 Jun 21;288(25):18521-32. PubMed PMID: 23658022; PubMed Central PMCID: PMC3689993.
- 14. Sato K, <u>Otsu W</u>, Otsuka Y, Inaba M. Modulatory roles of NHERF1 and NHERF2 in cell surface expression of the glutamate transporter GLAST. *Biochem Biophys Res Commun.* 2013 Jan 11;430(2):839-45. PubMed PMID: <u>23200831</u>.
- 15. Wang CC, Sato K, Otsuka Y, <u>Otsu W</u>, and Inaba M. Clathrin-mediated endocytosis of mammalian erythroid AE1 anion exchanger 1 facilitated by a YXXPhi or a noncanonical YXXXPhi motif in the Nterminal stretch. *J. Vet. Med. Sci.* 2012 74: 17-25. PubMed PMID: <u>21873807</u>.
- 16. Adachi H, Kurooka T, <u>Otsu W</u>, and Inaba M. The forced aggresome formation of a bovine anion exchanger 1 (AE1) mutant through association with deltaF508-cystic fibrosis transmembrane conductance regulator upon proteasome inhibition in HEK293 cells. *Jpn. J. Vet. Res.* 2010 58:101-110. PubMed PMID: 20715420.

### **Awards**

1. Student Poster Presentation award in the 31st annual meeting, the Membrane Society of Japan (2009)

## **Completed Research Support**

1. Research Fellowship for Young Scientist (DC1), the Japan Society for the Promotion Sciences (approx. \$30,000 USD per year, 2010-2013)

## **Professional Skills**

## In vitro (molecular cloning, biochemistry, imaging in cell culture)

- 1. Gene Engineering: fundamental techniques for PCR and subcloning (>100 plasmid vectors), genomic DNA or RNA isolation, cDNA synthesis, real time PCR (StepOne, invitrogen)
- 2. Cell Biology: general skills for cell culture (293, HeLa, MEF, LLC-PK1, MDCK, RPE-1, ARPE19, 661W), plasmid transfection (Lipofectamine, electroporation), gene knockdown (siRNA, miRNA), lentivirus isolation and infection, pulse-chase analysis using radio-isotope, ciliary assembly/disassembly assay
- 3. Biochemistry: cell/tissue lysis, immunoblotting, immunoprecipitation, cell surface biotinylation assay, sucrose gradient fractionation, protein purification with affinity chromatography
- 4. Proteomics: liquid chromatography tandem-mass spectrometry (Dionex UltiMate 3000 and LTQ Orbitrap Fourier transform-MS/MS system, Thermo Fisher)
- 5. Imaging: confocal microscopy (Zeiss LSM 5 and LSM880; Leica TCS SP2), live imaging with tracer (MitoTracker, fluorescence tagged EGF, dextran, transferrin), photobleaching after recovery assay

## In vivo (handling and tissue sampling)

- 6. General skills for handling laboratory animals (mouse, rat, dog, and cattle), maintaining colonies of transgenic mice (conditional knock-out in rod, cone, or RPE), and harvesting tissues (blood, kidney, retina)
- 7. Cardiac perfusion fixation in mouse and rat
- 8. Mouse retina vibratome sectioning and analysis by immunofluorescence or immunoelectron microscopy
- 9. Mouse rod photoreceptor perikarya sorting (FACS Aria II, Becton Dickinson)

# In vivo (surgery, functional analysis, and imaging in live animals)

- 10. Laparotomy and in utero electrophoresis (cone transfection) in mouse (>100 mice)
- 11. Electroretinography (ColorDome, Diagnosys LLC, >100 mice)
- 12. Fundus imaging and fluorescein angiography (Micron III, Phoenix Research Labs, >50 mice)
- 13. Optical coherence tomography and indocyanine green angiography (Spectralis HRA+OCT, Heidelberg Engineering, >30 mice)

## **Others**

14. Softwares for data analysis, and organizing figures: Microsoft Office, Pages, Numbers, Keynote, EndNote, Adobe Photoshop, Illustrator, Prism 7, Image J, G\*Power 2, Proteome Discoverer 1.0, SnapGene

## **Teaching experience**

- 1. Teaching assistant in the Veterinary Clinical Examination Class (for undergraduate students in Japanese) and in the Practice of Analytical Machines Class (for graduate students in English), in Hokkaido University (2009-2012).
- 2. Lab mentor of technicians and rotation/summer students (2014-2018).

### **Conference Poster/Talk Presentations**

#### International

- 1. Otsu W, Hsu KS, Dunaief JL, Li Y, Tsang SH, Chuang JZ, and Sung CH. Mice with RPE-specific CLIC4 deficiency exhibit AMD-like changes in the retina-RPE-choroid complex. ARVO Annual Meeting 2019, Vancouver, Canada, Apr 28-May 2, 2019.
- 2. <u>Otsu W</u>, Chuang JZ, and Sung CH. Establishment of a novel cone transfection technique for cone-rod dystrophy mouse models. ARVO Annual Meeting 2018, Honolulu, USA, Apr 29-May 3, 2018.

3. Otsu W, Otsuka Y, Miyazono K, Sato K, and Inaba M. Different roles of Derlin-1 and Derlin-2 in the ubiquitiylation-independent ER-associated degradation of a bovine AE1 anion exchanger mutant. The 35th Naito Conference on The Ubiquitin-Proteasome System: From Basic Mechanisms to Pathophysiological Roles, Sapporo, Japan, July 9-12, 2013.

## **Domestic**

4. Made oral or poster presentations in almost all the annual meetings of The Membrane Society of Japan, The Japanese Biochemical Society, and The Japanese Society of Veterinary Science, from 2008 to 2013.

## **Professional Memberships**

2018-present	The Association for Research in Vision and Ophthalmology
2019-present	The Molecular Biology Society of Japan
2008-2013	The Japanese Biochemical Society
2008-2013	The Membrane Society of Japan
2008-2013	The Japanese Society of Veterinary Science

## **Outreach activity**

- 1. Japanese Association of Scholars in Science (JASS, <a href="https://jass-newyork.webnode.com/">https://jass-newyork.webnode.com/</a>), committee (2018-present, maintenance of website)
- 2. Japanese Medical Society of America (JMSA, <a href="https://www.jmsa.org/ja/">https://www.jmsa.org/ja/</a>), committee for JMSA New York Life Science Forum 2019

## References

Available Upon Request