

Keita Tsujimura, Ph.D

Chief Scientific Director

Rett syndrome Organization Japan (RSOJ)

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WORK EXPERIENCE:

2023- Present Pharmaceutical company Japan
Research Scientist - Department of Pharmacology, Drug Discovery & Disease Research Laboratory

2020-2023 Nagoya University Aichi, Japan
Group Director (Designated Lecturer/ Laboratory Head) - Group of Brain Function and development, Institute of Neuroscience, Graduate school of Science

2020-2023 Nagoya University Aichi, Japan
Unit Leader (Frontier special unit) - Research unit for developmental disorders, Institute for Advanced Research

2021-2022 Harvard Medical School / Massachusetts General Hospital
Boston, United States
Visiting Lecturer - Department of Radiology, Athinoula A. Martinos Imaging Center

2017-2020 Nagoya University Aichi, Japan
Unit Leader - Research unit for developmental disorders, Institute for Advanced Research

2015-2020 Nagoya University Nagoya, Japan
Designated assistant professor - Graduate school of Medicine
1. Department of Psychiatry (4/2017-9/2020)
2. Department of Pharmacology (10/2015-3/2017)

2013-2015 Kyushu University Fukuoka, Japan
Designated assistant professor - Graduate school of Medicine

2010-2013. Nara Institute for Science and Technology (NAIST) Nara, Japan
Researcher - Graduate school of Biological Science

EDUCATION:

- Nara Institute for Science and Technology, Ph.D. in Biological Science (Apr 2007–Mar 2010)
- Nara Institute for Science and Technology, M.S. in Biological Science (Apr 2005–Mar 2007)
- Tokyo University of Science, B.A. in Chemistry (Apr 2001–Mar 2005)

QUALIFICATIONS:

- Business English level, PROGOS English Speaking Skill: CEFR(CEFR-J) Level”B1 High” (2023).
- Ph.D. degree (Biological Science; Mar 2010)

Research achievements

Publications

➤ Research article

● First author, Corresponding author, or Last author

- 1) **Tsujimura K.** (First author), Abematsu M., Kohyama J., et al., Neuronal differentiation of neural precursor cells is promoted by the methyl-CpG-binding protein MeCP2. *Exp Neurol* 219, 104-111 (2009)
- 2) Guo W., **Tsujimura K.**(Co-First author), Otsuka MI., et al. VPA Alleviates Neurological Deficits and Restores Gene Expression in a Mouse Model of Rett Syndrome. *PLOS ONE*, 9(6), e100215 (2014)
- 3) **Tsujimura K.**, Irie K., Nakashima H., et al., miR-199a links MeCP2 with mTOR signaling and its dysregulation leads to Rett syndrome phenotypes. *Cell Rep*, 12(11), 1887-1901 (2015)
- 4) Irie K., **Tsujimura K.**(Corresponding author), Nakashima H., et al. MicroRNA-214 promotes dendritic development by targeting the schizophrenia-associated gene Quaking (Qki). *J Biol Chem*, 291, 13891-13904 (2016)
- 5) Nakashima, H., **Tsujimura, K.** (Corresponding author), Irie, K., et al., Canonical TGF- β Signaling Negatively Regulates Neuronal Morphogenesis through TGIF/Smad Complex-Mediated CRMP2 Suppression. *J Neurosci* 38, 4791-4810 (2018)
- 6) Nakashima, H., **Tsujimura, K.** (Corresponding author), Irie, K., et al., MeCP2 controls neural stem cell fate specification through miR-199a-mediated inhibition of BMP-Smad signaling. *Cell Rep* 35(7), 109124 (2021)
- 7) Akaba, Y., **Tsujimura, K.**(Corresponding/Last author), et al., Comprehensive Volumetric Analysis of Mecp2-null Mouse Model for Rett syndrome by T2 Weighted 3D Magnetic Resonance Imaging. *Front*

Neurosci 16, 885335 (2022)

8) Akaba Y., **Tsujimura, K.** (Corresponding/Last author), et al., miR-514a promotes neuronal development in human iPSC-derived neurons. *Front Cell Dev Biol* 7:11:1096463 (2023)

9) **Tsujimura, K.** (First author), et al., Abnormal Structural Alteration of SVZ tractography fibers in Autism Spectrum Disorder Brain. *In preparation*

10) Irie K., **Tsujimura, K.** (Corresponding author), et al., MeCP2 controls dendritic morphogenesis via miR-199a-mediated Qki downregulation. *In preparation*

11) Akaba Y., **Tsujimura, K.** (Corresponding/Last author), et al., Dysregulation of MeCP2/miR-199a pathway contributes MECP2-duplication syndrome phenotypes. *Submitted*

12) Narita H., **Tsujimura, K.** (Last author), et al., C Diffuse but non-homogeneous brain atrophy in Rett syndrome: MRI volumetric study *Submitted*

● **Co-author**

13) Kohyama J, **Tsujimura K.**, et al., BMP-induced REST regulates establishment and maintenance of astrocytic identity. *J Cell Biol* 189, 159-170 (2010)

14) Abematsu M., **Tsujimura K.**, Yamano M., et al., Neurons derived from transplanted neural stem cells restore disrupted neuronal circuitry in a mouse model of spinal cord injury. *J Clin Invest*, 120 (9), 3255-66 (2010)

15) Juriandi B., **Tsujimura K.**, et al., Induction of superficial cortical layer neurons from mouse embryonic stem cells by valproic acid. *Neurosci Res* 72, 23-31 (2012)

16) Fujimoto Y., **Tsujimura K.**, et al. Treatment of a mouse model of spinal cord injury by transplantation of human induced pluripotent stem cell-derived long-term self-renewing neuroepithelial-like stem cells. *Stem Cells* 30, 1163-1173 (2012)

17) Juriandi B, **Tsujimura K.**, et al. Reduced adult hippocampal neurogenesis and cognitive impairments following prenatal administration of the antiepileptic drug, valproic acid. *Stem Cell Reports*, 5, 1-14 (2015)

18) Sekiguchi, M., **Tsujimura, K.**, et al. ARHGAP10, which encodes Rho GTPase-activating protein 10, is a novel gene for schizophrenia risk. *Transl Psychiatry* 10, 247 (2020)

19) Kato, H., **Tsujimura, K.**, et al. Rare genetic variants in the gene encoding histone lysine demethylase 4C (KDM4C) and their contribution to susceptibility to schizophrenia and autism spectrum disorder. *Transl Psychiatry* 10, 421 (2020)

20) Shiohama, H., **Tsujimura, K.**, et al. Small Nucleus Accumbens and Large Cerebral Ventricles in Infants and Toddlers Prior to Receiving Diagnoses of Autism Spectrum Disorder. *Cereb Cortex* 283, (2021)

21) Takeguchi R, **Tsujimura K.**, et al. Structural and functional changes in the brains of patients with Rett syndrome: A multimodal MRI study. *J Neurol Sci* 441, 120381 (2022)

22) Suzuki T, **Tsujimura K.**, et al. Pathological gait in Rett syndrome: Quantitative evaluation using three-dimensional gait analysis. *Eur J Paediatr Neurol* 42, 15-21 (2023)

23) Yonemoto K, **Tsujimura K.**, et al. Heterogeneity and mitochondrial vulnerability configurate the divergent immunoreactivity of human induced microglia-like cells. *Clin Immunol* 255, 109756 (2023)

24) Taira R, **Tsujimura K.**, et al. Gnao1 is a molecular switch that regulates the Rho signaling pathway in differentiating neurons. *Sci Rep* 14, 17097 (2024)

➤ **Review**

- 1) **Tsujimura K.** (Corresponding author), Nakashima H., Irie K., et al. Emerging roles for miRNA-based post-transcriptional regulation in neuronal morphogenesis and neurodevelopmental disorders. *RNA&DISEASE*, **3**, e1456 (2016)
- 2) Shiohama, H., **Tsujimura, K.**, Quantitative Structural Brain Magnetic Resonance Imaging Analyses: Methodological Overview and Application to Rett Syndrome. *Front Neurosci* (2022)
- 3) **Tsujimura, K.**, et al. microRNA biology on Brain Development and Neuroimaging Approach. *Brain Sci* (2022)

➤ **Book**

- 1) **Tsujimura, K.**, Nakashima, K. Rett Syndrome and Stem cell Research. “Stem Cell Genetics for Biomedical Research”, *Springer*, 27-41 (2018)
- 2) **Tsujimura, K.** MicroRNA and human diseases; MicroRNAs in Neurological Diseases, “MicroRNA: From Bench to Beside”, *Elsevier*, 317-329 (2022)

Patent

1. Patent 6869587 • **Keita Tsujimura**, Norio Ozaki, Hiroshi Abe, Yasuaki Kimura “Method for detection of miRNA processing and its application” • Nagoya University • 14/03/2019
2. Patent pending PCT/JP2020/2568 • **Keita Tsujimura**, Norio Ozaki, Hiroshi Abe, Yasuaki Kimura “Method for detection of miRNA processing and its application” • Nagoya University • 14/03/2019
3. Patent pending 2021-088243 • **Keita Tsujimura**, “Method for detection of Protein-Protein Interaction and its application” • Nagoya University • 26/05/2021
4. Patent pending 2022-070553 • **Keita Tsujimura**, Jun Natsume “Method for evaluation of efficacy of drugs and diagnose for Rett syndrome” • Nagoya University • 22/04/2022

Research Funds (Total more than 200,000,000 yen [JPY])

Japan Agency for Medical Research and Development (AMED) The iD3 booster DNW-21014 (Validation stage) : 2021-2022 (Principal Investigator)

Project Name : Drug discovery for developmental disorders

Amount : 8,800,000 yen (JPY)

Japan Agency for Medical Research and Development (AMED) Practical Research Project for Rare/Intractable Diseases, Research and Development (R&D) on rare and/or intractable diseases; Drug Discovery (Discovery phase: Step 0) : 2021-2023 (Principal Investigator)

Project Name : Development of miRNA pathology-based effective therapeutic drug for Rett syndrome

Amount : 93,210,000 yen (JPY)

Japan Agency for Medical Research and Development (AMED) Practical Research Project for Rare/Intractable Diseases, Research and Development (R&D) on rare and/or intractable diseases; Regenerative/Gene/Cell therapy (Discovery phase: Step 0) : 2021-2023 (Co-Principal Investigator; PI: Keiichiro Suzuki [Osaka University])

Project Name : Development of innovative gene editing technology for developmental disorders
Amount : 19,500,000 yen (JPY)

Japan Science and Technology Agency (JST) Program for Creating Start-ups from Advanced Research and Technology (START; SCORE University promotion type) : 2021 (Principal Investigator)
Project Name : Innovative drug discovery platform for rare diseases
Amount : 10,400,000 yen (JPY)

Japan Society for the Promotion of Science (JSPS) Grants-in-Aid for Scientific Research (KAKENHI) Grant-in-Aid for Scientific Research (C) : 2021-2023 (Principal Investigator)
Project Name : Understanding mechanisms of brain function and development by non-coding RNA
Amount : 4,030,000 yen (JPY)

Riken-Nagoya university joint research program grant: 2021-2022 (Principal Investigator)
Project Name : Elucidation of pathogenesis of developmental disorders by precise control of MECP2 gene expression
Amount: 500,000 yen (JPY)

Tokai Network for Global Leading Innovators, GAP Fund for Startup preparation: 2021-2022 (Principal Investigator)
Project Name : Development of drug discovery platform by non-coding RNA detection technique
Amount: 1,500,000 yen (JPY)

Tokai National Higher Education and Research system, Startup preparation fund: 2021 (Principal Investigator)
Project Name : Commercialization of drug discovery platform by microRNA processing detection technique
Amount: 1,000,000 yen (JPY)

Rett syndrome support organization, 10th Memorial Research grant : 2021 (Principal Investigator)
Project Name : Development of innovative gene therapy method for Rett syndrome
Amount : 1,000,000 yen (JPY)

Japan Society for the Promotion of Science (JSPS) Grants-in-Aid for Scientific Research (KAKENHI) Grant-in-Aid for Scientific Research (A) : 2021-2024 (Co-Principal Investigator; PI: Keiichiro Suzuki [Osaka University])
Project Name : Development of innovative genome editing therapeutic avenue at embryonic stage
Amount : 8,580,000 yen (JPY)

Japan Agency for Medical Research and Development (AMED) Practical Research Project for Rare/Intractable Diseases, Research and Development (R&D) on rare and/or intractable diseases; Genome medicine project (G) : 2020-2022 (Co-Principal Investigator; PI: Kenjiro Kosaki [Keio University])
Project Name : Achieving a diagnosis for all through deciphering structural variants and making sense of

non-coding mutations

Amount : 30,000,000 yen (JPY)

Nagoya University • Program for Promoting the Enhancement of Research Universities • Setting up young researcher units • ”frontiers” for the advancement of new and undeveloped fields : 2020-2021 (Principal Investigator)

Project Name : Research Unit for Developmental Disorders

Amount : 3,000,000 yen (JPY)

Japan Agency for Medical Research and Development (AMED) Practical Research Project for Rare/Intractable Diseases : 2019-2021 (Principal Investigator)

Project Name : Elucidation of the molecular pathogenesis of developmental disorders caused by MECP2 variations and development of novel diagnostic and therapeutic avenues

Amount : 24,000,000 yen (JPY)

Japan Agency for Medical Research and Development (AMED) The iD3 booster DNW-17001 (Screening stage) : 2020 (Principal Investigator)

Project Name : Exploration of innovative drug for developmental disorders

Amount : 10,000,000 yen (JPY)

Japan Agency for Medical Research and Development (AMED) The iD3 booster DNW-17001 (Screening stage) : 2019 (Principal Investigator)

Project Name : Exploration of innovative drug for developmental disorders

Amount : 20,000,000 yen (JPY)

Japan Agency for Medical Research and Development (AMED) The iD3 booster DNW-17001 (Validation stage) : 2018 (Principal Investigator)

Project Name : Exploration of innovative drug for developmental disorders

Amount : 16,000,000 yen (JPY)

Japan Agency for Medical Research and Development (AMED) The iD3 booster DNW-17001 (Validation stage) : 2017 (Principal Investigator)

Project Name : Exploration of innovative drug for developmental disorders

Amount : 10,000,000 yen (JPY)

Nagoya University • Program for Promoting the Enhancement of Research Universities • Setting up young researcher units for the advancement of new and undeveloped fields : 2017-2019 (Principal Investigator)

Project Name : Innovative drug discovery research unit for mental disorders

Amount : 3,000,000 yen (JPY)

Japan Society for the Promotion of Science (JSPS) Grants-in-Aid for Scientific Research (KAKENHI) Grant-in Scientific Research on Innovation Areas “Constructive understanding of multi-scale dynamism of neuropsychiatric disorders” : 2019-2020 (Principal Investigator)

Project Name : Constructive understanding of multi-scale dynamism of Rett syndrome

Amount : 7,800,000 yen (JPY)

Rett syndrome support organization, Research grant : 2019 (Principal Investigator)
Project Name : Elucidation of pathogenic neural circuits of Rett syndrome and development of therapeutic avenue
Amount : 1,000,000 yen (JPY)

Japan Society for the Promotion of Science (JSPS) Grants-in-Aid for Scientific Research (KAKENHI) Grant-in-Aid for Scientific Research (C) : 2018-2020 (Principal Investigator)
Project Name : Elucidation of common molecular mechanism of MECP2 abnormal disorders
Amount : 3,400,000 yen (JPY)

Kawano Masanori Memorial Public Interest Incorporated Foundation for Promotion of Pediatrics : 2018 (Principal Investigator)
Project Name : Elucidation of common molecular mechanism of Rett syndrome and MECP2 duplication syndrome
Amount : 500,000 yen (JPY)

Japan Society for the Promotion of Science (JSPS) Grants-in-Aid for Scientific Research (KAKENHI) Grant-in-Aid for Young Scientist (B) : 2016-2017 (Principal Investigator)
Project Name : Regulation of axon formation by developmental disorder causative gene MeCP2
Amount : 3,380,000 yen (JPY)

Award

5th Japanese Society of RNAi/Extracellular Vesicles, Best presentation award, Hiroshima, 29-31/08/2013

Outreach activities/Social activities

1. **Opinion exchange meeting between AMED project group (PI: Keita Tsujimura) and MECP2 duplication syndrome support organization**, Online, 05/09/2021
2. **Opinion exchange meeting between AMED project group (PI: Keita Tsujimura) and Rett syndrome support organization**, Online, 29/08/2021
3. **Opinion exchange meeting between AMED project group (PI: Keita Tsujimura) and MECP2 duplication syndrome support organization**, Online, 21/02/2021
4. **Keita Tsujimura**, “Unraveling the Mechanisms of Brain Development from Genes”, Aichi Science Festival 2020, Sakae, Nagoya, 02/11/2020
5. **Opinion exchange meeting between AMED project group (PI: Keita Tsujimura) and Investigation Team of Rett syndrome support organization**, Nagoya University, 15/07/2020
6. **Keita Tsujimura**, “Elucidation of mechanisms of developmental disorders -Challenging for development of therapeutic methods”, Aichi Science Festival 2019, Nagoya University Café “ Science and Me”, 20/11/2019
7. **Keita Tsujimura**, “Brain and diseases”, Nagoya University Open Lecture, 21/03/2019
8. **Keita Tsujimura**, “Common molecular mechanisms of developmental psychiatric disorders” Aichi Developmental Disability Center • Division of Genetics, Lecture for medical doctor 27/12/ 2018

Educational activities

1. Basic Biochemistry, November-December /2020 Undergraduate school of Science, Nagoya University
2. Special Lecture of Biological Science 2/5 (English), November 11-12/2020 Undergraduate school of Science and Graduate school of School (G30 International program Graduate student), Nagoya University
3. Basic Biology course June-July/2020, Undergraduate school of Science, Nagoya University
4. Neuroscience Course (English) Basic medical course, October 11/2019, Graduate school of medicine, Nagoya University
5. Integrative Graduate Education and Research Program in Green Natural Sciences, October /2019 Graduate school of science, Nagoya University

Academic activities

1. 2021-2023 Ethical review board (IRB) member, Nagoya University
2. Organizer, Workshop for the advancement of new and undeveloped fields, “Understanding brain development and disease pathogenesis”, Nagoya University, February 3/2021

Peer reviews

Psychiatry and Clinical Neuroscience, August/2021
Psychiatry and Clinical Neuroscience, July/2021
Cellular Signaling, April/2021
Cellular Signaling, March/2021
Molecular Psychiatry, March/2021
Cellular Signaling-1, February2021
Cellular Signaling-2, February2021
Journal of Neurochemistry, January/2021
Cellular Signaling-2, January/2021
Stem Cell, December/2020
Journal of Neurochemistry, December/2020
Stem Cell, November/2020
Molecular Psychiatry, September/2020
Frontiers in Cell and Developmental Biology, May/2020
Cellular Signaling, September/2019
Psychiatry and Clinical Neuroscience, December/2018

Achievements of Industry-academia Collaboration

Joint research contract with Olympus corporation (2017-2021)
Joint research contract with Oji Holdings Corporation (2021-2022)