CURRICULUM VITAE

1 NAME:

Masaaki Watanabe

2 BIRTH DATA:

15th October, 1974

3 ADDRESS:

Residence: postal code:001-0011, N11W1 1-7-1101, Sapporo, Hokkaido, Japan **Workplace:** postal code: 060-8638, N15W7, Sapporo, Hokkaido, Japan Department of Transplant Surgery Faculty of Medicine and Graduate School of Medicine, Hokkaido University

4 PHONE AND EMAIL:

Phone: +81-80-9615-1015 (mobile) Email: masaaki@w8.dion.ne.jp

5 COURSES AND DEGREES

Clinical training:

Hokkaido University School of Medicine: 1995-2001 The degree of medicine, Hokkaido University, School of medicine, 23rd March, 2001 Full Medical License (Japan) #420359: 17th May, 2001

Residency:

2001-2007: Resident at general surgery, Hokkaido University Hospital, Sapporo, Japan **Fellowship and specialist training:**

2007-2012: Clinical and research fellowship at department of transplantation, Hokkaido University Hospital, Sapporo, Japan

2012-2016 Clinical Fellowship and post-doctoral fellowship at Department of transplant surgery, Karolinska University, Sweden

The degree of Doctor of philosophy:

25th September, 2013 Doctor of philosophy, Ph.D., Hokkaido University School of Medicine Title of the thesis: ASKP1240, a Fully Human Anti-CD40 Monoclonal Antibody, Prolongs Pancreatic Islet Allograft Survival in Nonhuman Primates (*American Journal of Transplantation. 2013 Aug;13(8):1976-88.*)

Specialist training:

Board certificated transplant physician: 1st August, 2012: Japan Society of Transplantation

Board certificated surgeon: 1st January, 2019: Japan Surgical Society

Board certificated gastrointestinal surgeon: 1st January, 2020: Japan Surgical Society

Board certificated specialist physician of regenerative medicine: 1st January, 2021: Japanese Society for Regenerative Medicine

Academic Research:

2007-2012: Research Fellow of transplantation/islet transplantation at 1st Department of Surgery, Hokkaido University School of medicine, Japan

2012-2016: Research Fellow and post-doctoral fellow at Department of transplant surgery, Karolinska University, Sweden

2016-present: Research of transplantation/islet and cell transplantation at Department of Transplant Surgery, Faculty of Medicine and Graduate School of Medicine, Hokkaido University, Japan

6 DOCTORAL DEGREE

The degree of Doctor of philosophy

Year: 25th September, 2013

Discipline or subject area: Transplantation

Thesis title: ASKP1240, a Fully Human Anti-CD40 Monoclonal Antibody, Prolongs Pancreatic Islet Allograft Survival in Nonhuman Primates.

Name of the academic institution: Hokkaido University School of Medicine Names of the main supervisor: Professor Satoru Todo Names of co-supervisor: Professor Kenichiro Yamashita

7 POSTDOC APPOINTMENTS

Time period: 2012-2016

Name of the academic institution: Department of transplant surgery, Karolinska University, Sweden

Name of the supervisor: Professor Bo-Göran Ericzon

8 DOCENT-LEVEL COMPETENCE

Time period: 2016 -

Subject: tolerance, pancreatic islet transplantation, hepatocyte transplantation

Name of the academic institution: Department of Transplant Surgery, Faculty of Medicine and Graduate School of Medicine, Hokkaido University, Japan

9 CURRENT POSITION

Current employment, title: Lecturer at Department of Transplant Surgery, Faculty of Medicine and Graduate School of Medicine, Hokkaido University

Place of work: postal code: 060-8638, N15W7, Sapporo, Hokkaido, Japan

Period of the appointment: 1st April, 2018 - 31st March, 2025

Proportions of research, teaching and clinical activity involved in the position:

Research: Researching for tolerance induction after transplantation, and new treatment options for cell transplantation.

Teaching: As a P.I., teaching for Ph.D. program students, and teaching for medical students and training surgeons as a sub-chief of transplant group at Hokkaido University hospital.

Clinical activity: As a sub-chief of transplant group at Hokkaido University hospital, involving clinical liver, pancreas, kidney, and pancreatic islet transplantation.

10 PRIOR POSITIONS

1995-2001: Hokkaido University School of Medicine

2001-2007: Residency and training program at general surgery, Hokkaido University Hospital, Sapporo, Japan

2007-2012: Clinical and research fellowship at department of transplant surgery, Hokkaido University Hospital, Sapporo, Japan

2012-2016: Research Fellow and post-doctoral fellowship at Department of transplant surgery, Karolinska University, Sweden

2016-2018: Research of transplantation, islet, and cell transplantation at Department of Transplant Surgery, Faculty of Medicine and Graduate School of Medicine, Hokkaido University, Japan

11 TIME DEDUCTED FROM ACTIVE RESEARCH TIME

2012-2016: Research Fellow and post-doctoral fellowship at Department of transplant surgery, Karolinska University, Sweden.

September 15, 2013. Three weeks of parental leave on my third son's birthday.

12 SELECTED ACADEMIC DISTINCTIONS AND OTHER MERITS

Organ transplantation, cell transplantation, tolerance induction

SCIENTIFIC PORTFOLIO

1 CURRENT SCIENTIFIC ACTIVITY

Area of research and the title: transplantation, tolerance, cell transplantation,

- Tolerance induction following liver transplantation
- Tolerance induction following pancreatic islet/cell transplantation
- New treatment strategy for preventing the early cell loss following cell transplantation
- Immunological evaluation for tolerance patients following liver transplantation

Position associated with your current research team affiliation or the equivalent: Lecturer and P.I. at Department of Transplant Surgery, Faculty of Medicine and Graduate School of Medicine, Hokkaido University, Japan (<u>https://surg1.med.hokudai.ac.jp/en.html</u>)

2 SCIENTIFIC PUBLICATIONS

2.1 The ten most-cited publications

- 1. Todo S, Yamashita K, Goto R, Zaitsu M, Nagatsu A, Oura T, <u>Watanabe M</u>, et al. A pilot study of operational tolerance with a regulatory T-cell-based cell therapy in living donor liver transplantation. *Hepatology. 2016 Aug;64(2):632-43. doi: 10.1002/hep.28459. Epub 2016 Mar 10. PMID: 26773713* number of citations: 141
- Oura T, Yamashita K, Suzuki T, Fukumori D, <u>Watanabe M</u>, et al. Long-term hepatic allograft acceptance based on CD40 blockade by ASKP1240 in nonhuman primates. *Am J Transplant.* 2012 Jul;12(7):1740-54. doi: 10.1111/j.1600-6143.2012.04014.x. Epub 2012 Mar 15.PMID: 22420525 number of citations: 46
- **3.** *<u>M. Watanabe</u>, K. Yamashita, T. Suzuki, H, et al. ASKP1240, a Fully Human Anti-CD40 Monoclonal Antibody, Prolongs Pancreatic Islet Allograft Survival in Nonhuman Primates. *Am J Transplant.* 2013 Aug;13(8):1976-88. doi: 10.1111/ajt.12330. Epub 2013 Jul 10. *PMID:* 23841873 number of citations: 41
- 4. Oura T, Taniguchi M, Shimamura T, Suzuki T, Yamashita K, Uno M, Goto R, <u>Watanabe M</u>, et al. Does the permanent portacaval shunt for a small-for-size graft in a living donor liver transplantation do more harm than good? *Am J Transplant. 2008 Jan;8(1):250-2. doi: 10.1111/j.1600-6143.2007.02045.x. Epub 2007 Dec 18. PMID: 18093277* number of citations: 36
- Jorns C, Nowak G, Nemeth A, Zemack H, Mörk LM, Johansson H, Gramignoli R, <u>Watanabe</u> <u>M</u>, et al. De Novo Donor-Specific HLA Antibody Formation in Two Patients With Crigler-Najjar Syndrome Type I Following Human Hepatocyte Transplantation With Partial Hepatectomy Preconditioning. Am J Transplant. 2016 Mar;16(3):1021-30. doi: 10.1111/ajt.13487. Epub 2015 Nov 2. PMID: 26523372 number of citations: 31
- 6. Taniguchi M, Shimamura T, Suzuki T, Yamashita K, Oura T, <u>Watanabe M</u>, et al. Transient portacaval shunt for a small-for-size graft in living donor liver transplantation. *Liver Transpl.* 2007 Jun;13(6):932-4. doi: 10.1002/lt.21080. PMID: 17538989 number of citations: 26
- 7. Takahashi T, Matsumoto S, Matsushita M, Kamachi H, Tsuruga Y, Kasai H, <u>Watanabe M</u>, et al. Donor pretreatment with DHMEQ improves islet transplantation. J Surg Res. 2010 Sep;163(1):e23-34. doi: 10.1016/j.jss.2010.04.044. Epub 2010 May 21. PMID: 20638688 number of citations: 17
- 8. Daisuke Kuraya, <u>Masaaki Watanabe</u>, et al. The efficacy of DHMEQ, a NF-κB inhibitor, in islet transplantation: I. HMGB1 suppression by DHMEQ prevents early islet graft damage. *Transplantation.* 2013 Sep 15;96(5):445-53. doi: 10.1097/TP.0b013e31829b0744. PMID: 23900151 number of citations: 16

- Masaaki Watanabe, Torbjörn Lundgren, Yu Saito, et al. A non-hematopoietic erythropoietin analogue, ARA 290, inhibits macrophage activation and prevents damage to transplanted islets. *Transplantation*. 2016 Mar;100(3):554-62. doi: 10.1097/TP.000000000001026. PMID: 26683514 number of citations: 11
- Masaaki Watanabe, Kenichiro Yamashita, Hirofumi Kamachi, et al. The efficacy of DHMEQ, a NF-κB inhibitor, in islet transplantation: II. Induction DHMEQ treatment ameliorates subsequent allo-immune responses, and permits a long-term islet allograft acceptance. *Transplantation.* 2013 Sep 15;96(5):454-62. doi: 10.1097/TP.0b013e31829b077f. PMID: 23860082 number of citations: 9

2.2 The ten most important publications

Pancreatic islet transplantation; new strategies for long-term acceptance after transplantation

- Daisuke Kuraya, <u>Masaaki Watanabe</u>, Yasuyuki Koshizuka, et al. The efficacy of DHMEQ, a NF-κB inhibitor, in islet transplantation: I. HMGB1 suppression by DHMEQ prevents early islet graft damage. *Transplantation*. 2013 Sep 15;96(5):445-53. doi: 10.1097/TP.0b013e31829b0744. PMID: 23900151 number of citations: 16 Transplantation. Impact factor = 4.546
- Masaaki Watanabe, Torbjörn Lundgren, Yu Saito, et al. A non-hematopoietic erythropoietin analogue, ARA 290, inhibits macrophage activation and prevents damage to transplanted islets. *Transplantation.* 2016 Mar;100(3):554-62. doi: 10.1097/TP.000000000001026. PMID: 26683514 number of citations: 11 Transplantation. Impact factor = 4.546
- 3. T. Yoshida, K. Yamashita, <u>M. Watanabe</u>, et al. The Impact of c-Fos/Activator Protein-1 Inhibition on Allogeneic Pancreatic Islet Transplantation. Am J Transplant. 2015 Oct;15(10):2565-75. doi: 10.1111/ajt.13338. Epub 2015 May 26. PMID: 26012352 number of citations: 4 Am J Transplant. Impact factor = 7.338
- Ming Yao, <u>Masaaki Watanabe</u>, Sune Sun, et al. Improvement of Islet Allograft Function Using Cibinetide, an Innate Repair Receptor Ligand. *Transplantation*. 2020 Oct;104(10):2048-2058. doi: 10.1097/TP.00000000003284. PMID: 32345869 number of citations: 1 Transplantation. Impact factor = 4.546
- Masaaki Watanabe, Kenichiro Yamashita, Hirofumi Kamachi, et al. The efficacy of DHMEQ, a NF-κB inhibitor, in islet transplantation: II. Induction DHMEQ treatment ameliorates subsequent allo-immune responses, and permits a long-term islet allograft acceptance. *Transplantation. 2013 Sep* 15;96(5):454-62. *doi:* 10.1097/TP.0b013e31829b077f. PMID: 23860082 number of citations: 9 Transplantation. Impact factor = 4.546
- *<u>M. Watanabe</u>, K. Yamashita, T. Suzuki, et al. ASKP1240, a Fully Human Anti-CD40 Monoclonal Antibody, Prolongs Pancreatic Islet Allograft Survival in Nonhuman Primates. *Am J Transplant.* 2013 Aug;13(8):1976-88. doi: 10.1111/ajt.12330. Epub 2013 Jul 10. PMID: 23841873 number of citations: 41 Am J Transplant. Impact factor= 7.338

Tolerance induction after liver transplantation; for wide application of the cell therapy

- Todo S, Yamashita K, Goto R, Zaitsu M, Nagatsu A, Oura T, <u>Watanabe M</u>, et al. A pilot study of operational tolerance with a regulatory T-cell-based cell therapy in living donor liver transplantation. *Hepatology. 2016 Aug;64(2):632-43. doi: 10.1002/hep.28459. Epub 2016 Mar 10. PMID: 26773713* number of citations: 141 *Hepatology.* Impact factor = 14.679
- Masaaki Watanabe, Makiko Kumagai-Braesch, Ming Han Yao, et al. *Ex-vivo* generation of donor antigen-specific immunomodulatory cells: a comparison study of anti-CD80/86 mAbs and CTLA4-lg costimulatory blockade. *Cell Transplant.* 2018 Nov;27(11):1692-1704. doi: 10.1177/0963689718794642. *Epub* 2018 Sep 27. PMID: 30261751 number of citations: 0 Cell Transplant. Impact factor = 3.341

Hepatocyte transplantation; as a new generation cell transplantation

- 9. Jorns C, Nowak G, Nemeth A, Zemack H, Mörk LM, Johansson H, Gramignoli R, <u>Watanabe</u> <u>M</u>, et al. De Novo Donor-Specific HLA Antibody Formation in Two Patients With Crigler-Najjar Syndrome Type I Following Human Hepatocyte Transplantation With Partial Hepatectomy Preconditioning. *Am J Transplant.* 2016 Mar;16(3):1021-30. doi: 10.1111/ajt.13487. Epub 2015 Nov 2. PMID: 26523372 number of citations: 31 Am J Transplant. Impact factor = 7.338
- Masaaki Watanabe, Louise Hagbard, Helene Johansson, et al. Maintenance of hepatic functions in primary human hepatocytes cultured on xeno-free and chemical defined human recombinant laminins. PLoS One. 2016 Sep 6;11(9):e0161383. doi: 10.1371/journal.pone.0161383. PMID: 27598296 number of citations: 6 PLoS One. Impact factor = 2.74

3 INTERNATIONAL SCIENTIFIC CONGRESSES

3.1 Invited speaker or chair

Invited speaker:

A-PHPBA September 4th – 7th , 2019, Seoul, Korea

"Operational tolerance with a donor antigen specific immunomodulatory cell therapy in living donor liver transplantation"

Masaaki Watanabe, Ryoichi Goto, Kenichiro Yamashita, Satoru Todo, Tsuyoshi Shimamura, and Akinobu Taketomi

Chair person:

A-PHPBA September 4th -7th , 2019, Seoul, Korea Chair person, session pp20, September 6th

3 SCIENTIFIC DISTINCTIONS

- **1.** Young investigator award, American Transplant Congress, 30th May, 2009 "Combined treatment with a nuclear factor (NF)-κB inhibitor, DHMEQ, and tacrolimus permits a long-term allogeneic islet engraftment in mice"
- 2. Young investigator award, American Transplant Congress, 2011 "Long-term acceptance of islet allografts by ASKP1240 (4D11), a fully human anti-CD40 monoclonal antibody in cynomolgus monkeys"
- 3. Mentor and mentee award, Transplantation Science Symposium, 11th November, 2015 "Ex vivo generation of alloantigen-specific T regulatory cells using selective T-cell costimulation blockade"