第1 4 9 回

東北大学加齢医学研究所



プログラム

149th IDAC Biannual Meeting Program



日時:平成 30 年 1 月 26 日 (金曜日) 13:00~

場所:加齡医学研究所

スマート・エイジング研究棟1階 国際会議室

January 26, 2018,13:00~ Center for Smart Aging Research 1F, IDAC

共催:東北大学加齢医学研究所

Institute of Development, Aging and Cancer, Tohoku University 東北大学加齢医学研究所研究会同窓会

Society of Institute of Development, Aging and Cancer, Tohoku University

13:00— Opening remarks Dr. Ryuta Kawashima

第25回加齢医学研究所研究奨励賞授与式・受賞記念講演 25th IDAC Young Investigator Award Ceremony and Lecture

13:00-13:15 Ceremony Dr. Ryuta Kawashima

13:15-13:35 Lecture Chair: Dr. Kozo Tanaka

Development of PET tracers for imaging neuropathological hallmarks — From development to validation —

Department of Geriatrics and Gerontology, Institute of Development, Aging and Cancer, Tohoku University Department of Pharmacology, Tohoku University Graduate School of Medicine

Ryuichi Harada

Alzheimer's disease (AD), which is the most common cause of dementia, is an irreversible and progressive neurodegenerative diseases clinically characterized by memory loss and cognitive decline. Neuropathological hallmarks in AD are not only abundant neuronal loss and gliosis but also protein accumulation of amyloid- β and tau protein in the brain. Clinicopathological studies have established evidences that tau deposition is correlated with neuronal loss and the severity of cognitive impairment, but not amyloid deposition. Therefore, non-invasive imaging of tau pathology would provide new insights into the pathogenesis of AD. Through screening and the compound optimization, we developed several ¹⁸F-labeled quinoline derivatives as tau selective PET tracers. In clinical PET study, the latest derivative, ¹⁸F-THK5351 demonstrated elevated tracer retention in site susceptible to tau deposition in patients with AD. However, there are unexplained tracer retention in the basal ganglia and thalamus (i.e. off-target binding). Recent blocking studies identified monoamine oxidase-B (MAO-B) as an off-target substrate of ¹⁸F-THK5351. An imaging-autopsy validation of a patient with AD who underwent ¹⁸F-THK5351 PET prior to death demonstrated that *in vivo* ¹⁸F-THK5351 retention significantly correlated with MAO-B levels in the whole brain and tau aggregates in the neocortex. MAO-B was dominantly expressed in astrocytes and its level increased in patients with AD, which is associated with neuroinflammatory changes characterized by reactive astrocytes. Astrogliosis is linked to neurodegeneration, although it is still debate regarding relationship between gliosis and AD pathophysiology and its time course. Now we are trying to develop selective tau and astrogliosis (MAO-B) through lead optimization to elucidate its relationship by longitudinal PET studies.

13:35–13:40 **break**

13:40-14:40 Sessions $1\sim 4$

Chair: Masanori Ikeda

1 . Metformin directly binds the alarmin HMGB1 and inhibits its proinflammatory activity

Natsumi Sakata¹, Takahiro Horiuchi¹, Yoshihiro Narumi¹, Tomohiro Kimura¹, Takashi Hayashi², Keisuke Nagano³, Keyue Liu⁴, Masahiro Nishibori⁴, Sohei Tsukita⁵, Tetsuya Yamada⁵, Hideki Katagiri⁵, Ryutaro Shirakawa¹, and Hisanori Horiuchi¹ ¹Department of Molecular and Cellular Biology, Institute of Development,

Aging and Cancer, Tohoku University

²Biomedical, Technology Research Center and

³First Institute of New Drug Discovery, Tokushima Research Institute,

Otsuka Pharmaceutical Co., Ltd.

⁴Department of Pharmacology, Okayama University Graduate School of

Medicine, Dentistry and Pharmaceutical Sciences

⁵Department of Metabolism and Diabetes, Tohoku University Graduate School of Medicine

2. A mathematical model describing difference in chromosome dynamics between normal and cancer cells

Manuel A Campos, Kenji Iemura, Kozo Tanaka

Department of Molecular Oncology, Institute of Development, Aging and Cancer, Tohoku University

3 New methodology to detect and quantitate extracellular vesicles in blood plasma

Masashi Takao¹, Tetsuhiko Ohba², and Yutaka Nagai³ ¹Department of Project Program, IDAC, Tohoku University ²Department of Physics, Graduate School of Science and Faculty of Science, Tohoku Univerisy ³IVD Operations, Nihon Kohden Corporation

4 、 A novel and feasible antibody-based medicine for cancers targeting vasohibin-2

EunSeo LEE¹, Yasuhiro Suzuki¹, Hironori Nakagami², Hideki Tomioka³, Yasufumi Sato¹

¹Department of Vascular Biology, IDAC, Tohoku University

²Department of Health Development & Medicine, Osaka University. Graduate School of Medicine

³Department of Research & Development, FunPep Co.,Ltd

14:40-14:50 *Coffee break*

14:50-15:35 Sessions 5~7

Chair: Akihiro Yamada

5 Nrf2 plays a role in ischemia-reperfusion lung injury after lung transplantation

Takeo Togo¹, Yasushi Hoshikawa², Keiko Taguchi³, Hiroshi Yabuki ¹, Hideki Mitomo¹, Tatsuaki Watanabe¹, Masafumi Noda¹, Junichi Funahashi¹, Masayuki Yamamoto³, Yoshinori Okada ¹

¹Department of Thoracic Surgery, Institute of Development, Aging and Cancer, Tohoku University

²Department of Thoracic Surgery, Fujita Health University School of Medicine ³Department of Medical Biochemistry, Tohoku University Graduate School of Medicine

6. Identification of molecular biological factors on effect of anti-EGFR treatment in colorectal cancer for a biomarker development

Akira Okita, Shin Takahashi, Kota Ouchi, Chikashi Ishioka Department of Clinical Oncology, Institute of Development, Aging and Cancer, Tohoku University

7 、Gene regulatory mechanisms by mechanical forces in cardiovascular system

Atsushi Kubo, Makoto Kanayama, Kakeru Watanabe, Takahiro Niida, Yusuke Watanabe, Ken Matsumoto, Toshihiko Ogura Department of Developmental Neurobiology, Institute of Development, Aging and Cancer, Tohoku University

15:35–15:40 **break**

15:40-16:25 Sessions $8\sim 10$ Chair: Hiromitsu Ota

8 、Involvement of IL-32 in the regulation of malignant mesothelioma cell growth and VEGF and IL-8 secretion

Muneo Numasaki, Jyuri Ueda, Aiko Ishiki, Naoki Tomita, Shoji Okinaga and Hiroyuki Arai

Department of Geriatrics and Gerontology, Institute of Development, Aging and Cancer, Tohoku University

9 、Activation of NRF2 Alleviates Lethal Autoimmune Inflammation in Scurfy Mice

Takuma Suzuki^{1,2}, Shohei Murakami¹, Shyam S. Biswal³, Shimon Sakaguchi⁴,

Hideo Harigae², Masayuki Yamamoto⁵ and Hozumi Motohashi¹

¹Department of Gene Expression Regulation, Institute of Development, Aging and Cancer, Tohoku University

²Department of Hematology and Rheumatology, Tohoku University Graduate School of Medicine

³Department of Environmental Health Sciences, Bloomberg School of Public Health, Johns Hopkins University

⁴Experimental Immunology, World Premier International Research Center,

Immunology Frontier Research Center, Graduate School of Medicine, Osaka University

⁵Department of Medical Biochemistry, Tohoku University Graduate School of Medicine

1 0 、 Single cell RNA-Seq analysis defines distinct gene-expression profiles of tissue-specific plasma cells

Ari Itoh-Nakadai, Atsuko Kayaba, Toshiyuki Takai Department of Experimental Immunology, Institute of Development, Aging and Cancer, Tohoku University

一般口演について

発表時間12分,討論3分とします。時間厳守にてお願いします。座長は 研究員会委員の集談会コンテスト係が行ないます。

16:25-16:30 Closing remarks Dr. Hozumi Motohashi

終了後

加齢研実験研究棟7階セミナー室(1)におきまして18時から研究員会 主催新年会を開催いたします。皆様、多数ご参加くださいますようご案内 いたします。