

## Department of Internal Medicine

### Division of Neurology

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#### General Summary

Our researches in 2017 consist of the following areas: 1) cerebrovascular disease 2) neurodegenerative disease. We did not only clinical researches but started to do basic researches regarding these areas.

#### Research Activities

##### *Cerebrovascular disease*

Firstly, we continued to assume a leading role of multi-center study about the clinical characteristics in juvenile stroke and to take part in other multi-center studies (study about wake-up stroke, study about embolic stroke undetermined sources and study about stroke under anticoagulation therapy). Secondly, we have tried to establish intra-hospital system to treat hyper acute stroke patients, consisting of stroke coordinate nurses who support the hyper acute treatment including thrombolysis and new smartphone application (JOIN<sup>®</sup>) that can share text, neuroimaging, photos, and videos among stroke team. By using smartphone App (JOIN<sup>®</sup>), we are able to evaluate the stroke volume and the site of occlusion with the same quality as PC monitor. Thirdly, we did several prospective and retrospective studies from our stroke care unit (SCU) registry. The main theme of our clinical studies are follows: 1) right-to-left shunt (RLS) evaluation using a novel probe (pasteable soft ultrasound probe; PSUP), 2) insertable cardiac monitor for the patients with embolic stroke of undetermined source (ESUS), 3) nonstenotic carotid plaque of ipsilateral ESUS, 4) clinical characteristics of acute stroke patients with venous thromboembolism, 5) pre-operative double antiplatelet therapy for the patients with unruptured brain aneurysm who underwent coil embolization. On the other hand, we continued the basic research to establish animal model (especially primate model) of cerebral infarction with middle cerebral artery occlusion.

##### *Neurodegenerative disease*

###### 1. Parkinson's disease (PD) and the related disorders

We continue some clinical studies about the cardiovascular autonomic dysfunction in patients with PD and the related disorders. At first, we reported that the patients with reduced nocturnal blood pressure fall had a low cardiac uptake in <sup>123</sup>I-MIBG scintigraphy.

We also evaluated the influence of dopamine agonist for nocturnal blood pressure fall obtained from 24-hour ambulatory blood pressure monitoring test. Nocturnal blood pressure fall may improve in patients which some dopamine agonist had administered. Secondary, we clarify that PD patients with high norepinephrinergetic orthostatic hypotension have the characteristics of cognitive decline and impaired vasopressin release. On the other hand, We conducted a pilot study to evaluate a new olfactory threshold measurement device (FDL-1; Shimadzu, Kyoto, Japan) for differential diagnosis of parkinsonian disorders.

## 2. Dementia and the related disorders

We did some studies to clarify the associations between neuroimaging and neuropathological feature. We conduct a Dat SPECT examination on dementia with grain (DG) to clarify the difference between groups with parkinsonism. We also demonstrate a finding called cingulate island sign (CIS) on SPECT is seen in a group of patients neuropathologically diagnosed as pure dementia with Lewy bodies (DLB).

## 3. Amyotrophic lateral sclerosis

We clarified that percutaneous endoscopic gastrostomy (PEG) with noninvasive positive pressure ventilation seemed to be a valid method for dysphagic ALS patients with respiratory failure. We also started basic research about ALS pathogenesis.

## Publications

**Omoto S, Utsumi T, Matsuno H, Terasawa Y, Iguchi Y.** Thrombotic Microangiopathy Presenting with Intestinal Involvement Following Long-term Interferon-beta1b Treatment for Multiple Sclerosis. *Intern Med.* 2018; **57**: 741-4.

**Sakuta K, Iguchi Y, Sato T, Sakai K, Terasawa Y, Mitsumura H.** Chronic kidney disease is independently associated with acute recurrent cerebral infarct in patients with atrial fibrillation. *J Clin Neurosci.* 2017; **40**: 97-101.

**Sato T, Terasawa Y, Mitsumura H, Komatsu T, Sakuta K, Sakai K, Matsushima S, Iguchi Y.** Venous Stasis and Cerebrovascular Complications in Cerebral Venous Sinus Thrombosis. *European Neurology.* 2017; **78**: 154-60.

**Sato T, Sakuta K, Komatsu T, Sakai K, Terasawa Y, Mitsumura H, Iguchi Y.** Yield of combined MRI sequences in isolated cortical vein thrombosis diagnosis. *J Neurol Sci.* 2017; **381**: 328-30.

**Wang Z, Sawaguchi S<sup>1</sup>, Hirose H<sup>2</sup>, Ohara K<sup>2</sup>, Sakamoto S<sup>2</sup>, Mitsumura H, Ogawa T, Iguchi Y, Yokoyama M (Nihon Pharmaceutical Univ, Kaneka Corporation, Tokyo, Japan).** An In Vitro Assay for Sonothrombolysis Based on The Spectrophotometric Measurement of Clot Thickness. *J Ultrasound Med.* 2017; **36**: 681-98.

**Wang Z, Komatsu T, Mitsumura H, Nakata N, Ogawa T, Iguchi Y, Yokoyama M.** An Uncovered Risk Factor of Sonothrombolysis: Substantial Fluctuation of Ultrasound Transmittance through the Human Skull. *Ultrasonics.* 2017; **77**: 168-75.

**Aoki J<sup>1,2</sup>, Kimura K<sup>1,2</sup>, Morita N<sup>2</sup>, Harada M<sup>4</sup>,**

**Metoki N<sup>2</sup>, Tateishi Y<sup>2</sup>, Todo K<sup>7</sup>, Yamagami H<sup>7</sup>, Hayashi K<sup>6</sup>, Terasawa Y<sup>1</sup>, Fujita K<sup>4</sup>, Yamamoto N<sup>1</sup>, Deguchi I<sup>2</sup>, Tanahashi N<sup>2</sup>, Inoue T<sup>2</sup>, Iwanaga T<sup>10</sup>, Kaneko N<sup>11</sup>, Mitsumura H, Iguchi Y, Ueno Y<sup>12</sup>, Kuramoto Y<sup>12</sup>, Ogata T<sup>13</sup>, Fujimoto S<sup>14</sup>, Yokoyama M<sup>15</sup>, Nagahiro S<sup>4</sup>; on behalf of YAMATO Study Investigators (Nippon Med Sch, <sup>2</sup>Kawasaki Med Sch, <sup>3</sup>Natl Cerebral Cardiovasc Ctr, Suita, Japan, <sup>4</sup>Tokushima Univ, <sup>5</sup>Hirosaki Stroke and Rehabilitation Center, <sup>6</sup>Nagasaki Univ, <sup>7</sup>Kobe City Med Ctr General Hosp, <sup>8</sup>Fukui-ken Saiseikai Hosp, <sup>9</sup>Saitama Med Univ, <sup>10</sup>Jpn Red Cross Okayama Hosp, <sup>11</sup>Okinawa Kyodo Hosp, <sup>12</sup>Shinko Hosp, <sup>13</sup>Fukuoka Univ, <sup>14</sup>Steel Memorial Yawata Hosp, <sup>15</sup>Yokohama Sakae Kyosai Hosp).** YAMATO Study (Tissue-Type Plasminogen Activator and Edaravone Combination Therapy). *Stroke.* 2017; **48**: 712-9.

**Yamashita T<sup>1,2</sup>, Miki A<sup>1,2</sup>, Goto K<sup>2</sup>, Araki S<sup>2</sup>, Takizawa G<sup>2</sup>, Ieki Y<sup>2</sup>, Kiryu J<sup>2</sup>, Tabuchi A<sup>1</sup>, Iguchi Y, Kimura K<sup>3</sup>, Yagita Y<sup>2</sup> (Kawasaki Univ Med Welfare, <sup>2</sup>Kawasaki Med Sch, Okayama, Japan, <sup>3</sup>Nippon Med Sch).** Preferential atrophy of the central retinal ganglion cells in homonymous hemianopia due to acquired retrogeniculate lesions demonstrated using swept-source optical coherence tomography. *Acta Ophthalmol.* 2017; **96**: e538-9.

**Mishina M<sup>1,2</sup>, Ishii K<sup>2</sup>, Kimura Y<sup>2,3</sup>, Suzuki M, Kitamura S<sup>1</sup>, Ishibashi K<sup>2</sup>, Sakata M<sup>2</sup>, Oda K<sup>2,4</sup>, Kobayashi S<sup>1</sup>, Kimura K<sup>1</sup>, Ishiwata K<sup>2,5,6</sup> (Nippon Med Sch, <sup>2</sup>Tokyo Metropolitan Institute of**

*Gerontology*, <sup>3</sup>Kinki Univ, <sup>4</sup>Hokkaido Univ, <sup>5</sup>Southern TOHOKU Research Institute for Neuroscience, <sup>6</sup>Fukushima Med Univ). Adenosine A1 receptors measured with 11 C-MPDX PET in early Parkinson's disease. *Synapse*. 2017; **71**. doi: 10.1002/syn.21979. Epub 2017 Apr 25.

**Takei J, Irie K, Tanaka H, Okuno K, Hasegawa I, Shimoyama T, Yaguchi H, Hatano K, Maruyama F, Yamamoto Y, Tochigi S, Hasegawa Y, Murayama Y.** Evaluation of the CT high-density area after endovascular treatment for acute ischemic stroke. *J Neuroendovascular Therapy*. 2017; **11**: 227-34.

**Tanabe H<sup>1</sup>, Higuchi Y<sup>1</sup>, Yuan JH<sup>1</sup>, Hashiguchi A<sup>1</sup>, Yoshimura A<sup>1</sup>, Ishihara S<sup>1,2</sup>, Nozuma S<sup>1</sup>, Okamoto Y<sup>1</sup>, Matsuura E<sup>1</sup>, Ishiura H<sup>3</sup>, Mitsui J<sup>3</sup>, Takashima R<sup>4</sup>, Kokubun N<sup>4</sup>, Maeda K<sup>5</sup>, Asano Y<sup>6</sup>, Sunami Y<sup>6</sup>, Kono Y, Ishigaki Y<sup>7</sup>, Yanamoto S<sup>8</sup>, Fukae J<sup>8</sup>, Kida H<sup>9</sup>, Morita M<sup>10</sup>, Tsuji S<sup>1</sup>, Takashima H<sup>1</sup>** (<sup>1</sup>Kagoshima Univ, <sup>2</sup>Univ the Ryukyus, Okinawa, Japan, <sup>3</sup>University Tokyo, <sup>4</sup>Dokkyo Medical Univ, <sup>5</sup>Natl Hosp Organization Higashi-ohmi General Med Ctr, Shiga, Japan, <sup>6</sup>Tokyo Metropolitan Neurologi-

<sup>cal Hosp, <sup>7</sup>Coral Clinic, Tokyo, Japan, <sup>9</sup>Fukuoka Univ Sch Med, <sup>10</sup>Kurume Univ, <sup>11</sup>Jichi Med Univ</sup>). Clinical and genetic features of Charcot-Marie-Tooth disease 2F and hereditary motor neuropathy 2B in Japan. *J Peripher Nerv Syst*. 2018; **23**: 40-8.

## Reviews and Books

**Suzuki M, Mitoma H<sup>1</sup>, Yoneyama M<sup>2</sup>** (<sup>1</sup>Tokyo Med Univ, <sup>2</sup>MCHC R&D Synergy Center, Inc., Yokohama, Japan). Quantitative analysis of motor status in Parkinson's disease using wearable devices: From methodological considerations to problems in clinical applications. *Parkinsons Dis*. 2017; 6139716. doi: 10.1155/2017/6139716. Epub 2017 May 18.

**Yogo M, Suzuki M.** Findings of 123I-MIBG cardiac scintigraphy: Parkinson's disease and related disorders and others (RBD, Cardiac Diseases, DM, etc.). In: Iwase S, Hayano J, Orimo S. Clinical assessment of the autonomic nervous system. Springer, Tokyo. 2017; **15**: 231-51. DOI [https://doi.org/10.1007/978-4-431-56012-8\\_15](https://doi.org/10.1007/978-4-431-56012-8_15).