

ANNUAL PUBLICATIONS

2nd Department of Oral Surgery

1993.1.-2000.3.

I. Staffs and Students (April, 1999)

Professor, Chairman	Shoji Enomoto
Associate Professor	Yuzo Takahashi
Associate Professor (Lecturer)	Masatoshi Ishii
Associate Professor (Lecturer)	Izumi Asahina
Research Associate	Kiyoshi Harada
	Tetsuya Yoda
	Masaji Mori
	Eiki Sakai
	Maimi Tsukahara
	Yasuhiro Okada
	Kazunori Ohkura
	Takashi Miki
Clinical Staff	Mitsuo Oda
	Akimune Kikegawa
	Yoshimasa Ishii
	Yusuke Nakajima
	Masato Abe
	Makiko Higashinakagawa
	Seiki Mogi
	Kentaro Hoshi
	Hiroshi Sato
	Akemi Gunji
	Wataru Ishiguro
Postdoctoral Fellow	Yutaka Maruoka
	Makiko Fujii
	Eiki Nishito
	Kozue Masaka
Graduate Student	Takashi Moriyama
	Jinkyō Sakurai
	Hideki Imai
	Ichiro Seto
	Satoru Hoshino
	Minoru Ikuta
	Yuji Kabasawa
	Hiroaki Shimamoto
	Keiichi Morita
	Hisanobu Yonezawa
	Hideyo Ihara
	Eriko Marukawa
	Alam Md Imranul
	Etsuko Takahashi
	Hitoshi Kurokawa
	Seiko Morishima
	Fumiko Imaizumi
	Fumihiko Tsushima
	Masato Hosoya
Reserch Student	Noriko Taguchi

Ken Hamazaki
Yasushi Sekine
Yuki Kawanobe
Masaru Harashima
Kohtaro Hayashi
Yuka Watanabe
Osamu Watanabe
Yoko Ichihara
Emi Sumida
Yoshitaka Shirakura
Naoto Chikamori
Akira Misaki
Yu Shinjo
Daisuke Tutui
Masaru Sato
Kaori Yoshida
Miki Suzuki
Michio Saito
Yuji Ichikawa
Masashi Sakagami
Masatoshi Watanabe
Furi So
Hiroyasu Tsukahara
Kozue Meguro
Wakana Daigo
Kiyoko Hattori
Kazumi Ide
Tadahiro Suzuki
Ichiro Sakamoto
Yoshinobu Honma
Shigemi Tasaki

II. Educational Outline of Graduate Course

In the field of oral and maxillofacial surgery, there are various kinds of diseases such as malformation, malignant tumor, benign tumor, trauma, deformity of face, dental infection and temporomandibular disorders. Accordingly, it is necessary for the graduate course students to have an outlook about oral and maxillofacial surgery at first. They have to be educated and trained for the future of oral and maxillofacial surgery in order to promote the basic and clinical standard of this field. First year, they will take part in the clinical rotation for the outpatient and the hospitalized patient. And they can understand the problem from which the patient is suffered. During this rotation, they can find the research problem which is necessary to be studied for the patients in this field. The aim of the research in the clinical department should be related to the patients who are suffering for the disease in this field.

From these points, we have prepared the following research projects. 1) Functional reconstruction of mandible using growth factors, 2) Application of Distraction Osteogenesis for Treating Dento-Facial Deformity Patients, 3) Basic and clinical study of cleft lip and palate, 4) Studies on pathogenesis and treatment of internal derangement or osteoarthritis in temporomandibular joint, 5) Molecular Analysis of Oral Cancer for Biological Diagnostic Markers, 6) Tissue engineering for oral mucosa, 7) Immunological studies for oral mucosal disease.

III. Research Subjects

1) Functional reconstruction of mandible using growth factors :

We are trying to develop the new method for functional reconstruction of mandible using growth factors and tissue engineering. Each project is as followings: 1) Development of the new carrier for BMP. 2) Evaluation of the combination graft of BMP, FGF and/or TGF- β for bone formation. 3) Reconstruction of mandible bone defect in large experimental animals. 4) Development of the

artificial bone substitute containing osteoblasts.

2) Application of Distraction Osteogenesis for Treating Dento-Facial Deformity Patients :

Orthognathic surgery has been a method with high surgical stress for patients. Distraction osteogenesis will be able to decrease this surgical stress. In our department, an experimental study for mandibular distraction osteogenesis on rabbit mandible has been performed to apply the distraction technique to the treatment for dentofacial deformity patients. However, the treatment period of the distraction method is relatively long (about 2 months). Therefore, we are applying bone-morphogenetic-protein (BMP) to the distracted region to shorten the treatment period.

3) Basic and clinical sutudy of cleft lip and palate :

Basic and clinical studies on cleft lip and palate are performed at our clinic as follows. 1. Experimental studies using monkeys with simulated alveolar clefts are performed. Autogenous cancellous bone and bone morphogenetic protein are packed into the simulated alveolar clefts in order to investigate the bone formation of the alveolar bone grafting. 2. Studies on artificial mucosa and cultured mucosa which will be applied for the defect after oral surgery especially palatoplasty. 3. Basic research on scar tissue developed after palatoplasty. 4. Clinical studies on secondary bone grafting into the alveolar clefts. 5. Clinical studies on patients with cleft lip and palate.

4) Studies on pathogenesis and treatment of internal derangement or osteoarthritis in temporomandibular joint (TMJ):

Studies are as follows: 1. Development of a new treatment i.e. disk repositioning therapeutic exercise for internal derangement in TMJ. 2. Clinical and histological studies on morphological bone changes of the condyles in temporomandibular disorders. 3. Histological investigation on healing process of injured TMJ disk and synovial tissue by means of organ culture. 4. Elucidation on pathogenesis of internal derangement in TMJ using our rabbit models brought about anterior disk displacement.

5) Molecular Analysis of Oral Cancer for Biological Diagnostic Markers :

As the human cancers are genetical disease with several abnormalities of tumor related genes, identification of the tumor responsible genes is necessary to understand the nature of tumor cells, for clinical diagnosis, application to treatment. In 1992, we first reported that two-thirds of oral squamous cell carcinoma have p53 gene mutation. As none of other genes was reported to have mutation so frequently as p53 gene, this genetical alteration characterize the oral squamous cell carcinoma. Now, we are studying the method of genetical diagnosis based on the mutation of p53 tumor suppressor gene, the malignancy grade based on the character of telomerase gene products. Furthermore, we are identifying the metastasis related genes by comparing the primary tumors with metastasized tumors in same patients. In oral surgery, for the prevention and treatment of congenital and acquired disease arising in oral and maxillofacial lesion, we are performing education to be understood the cause, pathology, diagnosis, treatment and prognosis of such diseases.

IV. Publications (January, 1993-March, 2000)

A. Original Articles

- 1) Asahina I, Sampath T. K, Nishinura I. and Hauschka P. V. : Human Osteogenic Protein-1 Induces both Chondroblastic and Osteoblastic Differentiation of Osteogenic Cells Derived from Newborn Rat Calvaria. *Journal of Cell Biology* 123 : 921 - 933, 1993.
- 2) Asahina I, Sampath TK, Hauschka PV: Human osteogenic protein-1 induces chondroblastic , and/or adipocytic differentiation of clonal murine target cells. *Experimental Cell Research* 222:38-47, 1996.
- 3) Asahina I., Watanabe M., Sakurai N., Mori M., and Enomoto S.: Repair of Bone Defect in Primatemandible using a Bone Morphogenic protein (BMP)-Hydroxyapatite-Collagen Composite.. *J Med Dent Sci* 44: 63, 1997.
- 4) Asahina, I., Sakakibara, T., Miyashita, M., Tachikawa, N., Enomoto, S.: Congenital Midline Simus of the upper lip: Case report and review of literature. *Cleft Palate-Craniofacial Journal* 34:83-85,1997
- 5) Asahina, I., Abe, M. : Sialoadenoma papilliferum of the hard palate: A case report and review of literature. *J. Oral Maxillofac. Surg.* 55:1000-1003, 1997
- 6) Chang H.Y., Nishitoh H., Yang X., Ichijo H., Baltimore D.: Activation of apoptosis signal-regulating kinase 1 (ASK1) by the adapter protein Daxx. *Science* 281(5384):1860-1863, 1998.
- 7) Enomoto S. : Investigation of dental implants up to the present. *Journal of Japanese Association For Dental Science.* 13 : 53-75, 1994. (in Japanese)
- 8) Fujibayashi T., Gotoh S., Takahashi Y., Mori Y., Minato S. and Enomoto S. : Cell-Mediated Immunity in Recurrent Malignant

- Tumors of the Oral Cavity. *Biotherapy* 7 (2) : 150 - 157, 1993. (in Japanese)
- 9) Fujibayashi T., Gotoh S., Takahashi Y., Mori Y., Minato S. and Enomoto S. : Effectiveness of OK-432 for Cell-Mediated Immunity in Malignant Tumors of the Oral Cavity. *Biotherapy* 7 (2) : 150 - 157, 1993. (in Japanese)
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 - 13) Harada, K., Ono, J., Okada, Y., Nagura, H., Enomoto, S.: Postoperative stability after sagittal split ramus osteotomy with condylar-positioning appliance and screw fixation: Asymmetric versus symmetric cases. *Oral Surg. Oral Med. Oral Pathol.* 83:532-536, 1997
 - 14) Harada, K., Watanabe, M., Okada, Y., Enomoto, S. : Mandibular stability after sagittal split ramus osteotomy without post-operative maxillomandibular fixation in the treatment of prognathic patients with symmetric mandibles. *Clinical Orthodontics and Research* 1:44-51, 1998
 - 15) Harada, K., Tajima, K., Enomoto, S.: Stability after sagittal split ramus osteotomy without postoperative maxillomandibular fixation in the treatment of prognathic patients with asymmetric mandibles. *Clinical Orthodontics and Research* 2: 27-33, 1999
 - 16) Harada K., Torikai K. and Funaki J. : Augmentation Genioplasty with Hydroxyapatite Blocks. *Journal of Oral Maxillofac Surg.* 22 : 265-266, 1993.
 - 17) Harada K. and Enomoto S. : A New Method of Tongue Reduction for Macroglossia. *Journal of Oral and Maxillofacial Surgery* 53 : 91-92, 1995.
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 - 21) Harada K., Okada Y., Okura K., Nagisa M., Ono J., Watanabe M. and Enomoto S. : Changes in preoperative and postoperative temporomandibular joint symptoms in patients treated by sagittal split ramus osteotomy of the mandible using an appliance for repositioning the proximal segment with titanium screw fixation. *Jpn. J. Oral Maxillofac. Surg.* 43 (9) : 667-673, 1997. (in Japanese)
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