Department of Orthopaedic Surgery

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General Summary

Basic Research

The research performed in our Department of Orthopaedic Surgery, from studies on connective tissue cells to clinical application of basic research data, continues to be held in high regard both in Japan and abroad. This is reflected by the large numbers of papers published in English-language journals and of competitive scientific grants awarded to department members. The number of scientific awards received by our researchers has further increased in the last year, reaching 11 prizes accepted during the last 5 years. The highest priority of our research team is to carry out basic research that is readily applicable to clinical practice, not just that which deals only with basic science itself.

Clinical Research

An approach in which basic studies are carried out with clinical practice in mind and in which clinical research is strongly based on basic science has been proven valid by the results of many collaborative multicenter projects. We continue to cultivate and to present to the world our department's vision that an orthopaedic surgeon dealing with bones, cartilages, blood vessels, ligaments, tendons, and skin shall find answers to the questions of everyday clinical practice through basic laboratory research. This new concept of orthopedic surgical practice has started to be incorporated into evidencebased research: 2 of our researchers have been appointed authors of bone metabolismrelated clinical guidelines.

Research Activities

Arthroscopic Bankart repair for recurrent shoulder dislocation in patients older than 40 years

Clinical features and results of arthroscopic surgery were evaluated in patients older than 40 years who had sustained recurrent dislocations of the shoulder without a full-thickness tear of the rotator cuff. Three-fourths of the patients had sustained the first dislocation in their 20 s. The mean Japan Shoulder Society Shoulder Instability Score was 51 points before surgery and reached 92 points at the final follow-up. The postoperative limitation of range of motion was decreased with the procedure.

Isolated avulsion fracture of the lesser tuberosity of the humerus in an adolescent tennis player

An avulsion fracture of the lesser tuberosity of the humerus was diagnosed in a 17-year-old tennis player. He began to feel pain on stroking 4 years before admission. Pain was induced by elevation and internal rotation of the shoulder. Arthroscopy showed impingement of a bony fragment between the glenoid rim and the lesser tuberosity of the humerus. The bony fragment was surgically removed, and the subscapularis muscle tendon was repaired.

Spondylolisthesis of the lumbar spine in patients with severe osteoarthritis

We investigated the incidence of spondylolisthesis of the lumbar spine in patients with severe osteoarthritis. Spondylolisthesis was identified in 57% of patients with osteoarthritis of the knee and in 20% of patients with osteoarthritis of the hip. Lumbar disorders should be considered when patients complain of pain around the knee and when severe osteoarthritis is diagnosed with radiographic findings.

Prevention of dislocation after primary total hip arthroplasty with a modular femoral stem

Rotational deformity of the proximal femur remains a common problem after primary total hip arthroplasty in Japanese patients. The deformity can result in postoperative dislocation. The rotation-free modular system used for total hip arthroplasty in Japanese patients produces promising results in regard to preventing postoperative dislocation.

Patient-specific templating method in total knee arthroplasty: A prospective study of the accuracy of different patient-specific bone cutting guides

Preoperative and intraoperative patient-specific templating has gained attention as the next technological development after computer-assisted surgery navigation systems in knee surgery. In our department, we have been evaluating the accuracy of implant positioning during total knee arthroplasty with patient-specific bone cutting guides and are performing a comparative study against the computer-assisted navigation system. The evaluation also includes a comparative trial against conventional surgery, analysis of 3-dimensional reconstructions, and the development of more precise preoperative planning software. The comparative analysis on accuracy of different patient-specific cutting guides is being performed in a prospective manner.

Examination of hallux rotation in hallux valgus

We examined rotations of the hallux in patients with hallux valgus by comparing coronal computed tomographic images of these patients with those of a control group of patients and examined the degrees of rotation of the first and second metatarsals and the proximal phalanx of the hallux. Our results indicated that hallux rotation is caused by strain of the soft tissue around the metatarsophalangeal joint.

Collagen cross-linking and abnormalities of bone metabolism in osteoporosis and diabetes

Collagen cross-linking, a major posttranslational modification of collagen, plays many important roles in the biological and biomechanical features of bone. Our recent basic and clinical investigations of collagen cross-links seem to open a new chapter for general medicine as well as for orthopaedic practice. For example, serum or urine levels of pentosidine are now used to estimate future fracture risk in osteoporosis and diabetes. In our research, we demonstrated age-related changes in collagen cross-links in bone and abnormalities of cross-links in osteoporosis and diabetes.

Effects of alendronate on bone formation induced by recombinant human bone morphogenetic protein 2

Local alendronate administration could inhibit β -tricalcium phosphate (TCP) resorption or bone formation or both induced by recombinant human bone morphogenetic protein 2.

Bone formation and resorption was examined in patients after implantation of β -TCP blocks with 60% and 75% porosity in opening wedge high tibial osteotomy. The β -TCP was completely resorbed within 3.5 years.

Publications

Chazono M, Masui F, Kawaguchi Y, Hazama H, Ueda J, Saito S, Ito Y, Kasama K, Liu K, Marumo K. Dumbbell-shaped osteochondroma of 5th rib causing spinal cord compression:a case report. J Orthop Sci 2009; **14:** 336-8.

Mori Y, Kuriyama G, Tanaka T, Tajima N. Usefulness of aggressive lipid-lowering therapy with rovastatin in hypercholesterolemic patients with concomitant type 2 diabetes. *Endocrinology* 2009; **36:** 412–8.

Takahashi Y, Ikeda R, Kato F. Synaptic potentiation in the central amygdale involves different mechanisms depending on pain model. *Pain Res* 2009; **24:** 137-46.

Narita N, Kobayashi Y, Nakamura H, Maeda K, Ishihara A, Mizoguchi T, Usui Y, Aoki K, Simizu M, Kato H, Ozawa H, Udagawa M, Endo M, Takahashi N, Saito N. Multiwalled carbon nanotubes specifically inhibit osteoclast differentiation and function. Nano Lett 2009; 9: 1406-13.

Koide M, Kinugawa S, Ninomiya T, Mizoguchi T, Yamashita T, Maeda K, Maeda K, Yasuda H, Kobayashi Y, Nakamura H, Takahashi N, Udagawa M. Diphenylhydantoin inhibits osteoclast differentiation and function through suppression of NFATc1 signaling. J Bone Miner Res 2009; 24: 1469-80.

Murakami K, Inagaki J, Saito M, Ikeda Y, Tsuda C, Noda Y, Kawakami S, sawa sawa T, Shimizu T. Skin atrophy in cytoplasmic SOD-deficient mice and its complete recovery using a vitamin C derivative. Biochem Biophys Res Commun 2009; **382:** 457-61.

Saito M, Marumo K, Soshi S, Kida Y, Ushiku C, Shinohara A. Raloxifene ameliorates detrimental collagen cross-link formation and bone strength in rabbits with hyperhomocysteinemia. *Osteoporos Int* 2010; **21:** 655–66. Epub 2009 May 30.

Saito M, Shiraishi A, Ito M, Sakai S, Hayakawa N, Mihara M, Marumo K. Comparison of effects of alfacalcidol and alendronate on mechanical properties and bone collagen cross-links of callus in the fracture repair rat model. *Bone* 2010; **46:** 1170-9. Epub 2009 Dec 22.

Kii I, Nishiyama T, Li M, Matsumoto K, Saito M, Amizuka N, Kudo A. Incorporation of tenascin-C into the extracellular matrix by periostin underlies an extracellular meshwork architecture. J Biol Chem 2010; **285:** 2028-39.

Eda H, Aoki K, Kato S, Okawa Y, Takada K, Tanaka T, Marumo K, Ohkawa K. The proteasome inhibitor bortezomib inhibits FGF-2induced reduction of TAZ levels in osteoblast-like cells. Eur J Haematol 2010; 85: 68-75.

Maruhashi T, Kii I, Saito M, Kudo A. Interaction between periostin and BMP-1 promotes proteolytic activation of lysyl oxidase. *J Biol Chem* 2010; **285:** 13294-303. Epub 2010 Feb 24.

Funasaki H, Yoshida M, Kan I, Kato S, Kasama K, Ishizuka R, Marumo K. The effectiveness of conservative treatment in non-operative patients with a full-thickness tear of rotator cuff (in Japanese). *Katakansetsu* 2009; **33:** 697-700.

Chazono M, Soshi S, Inoue T, Nakamura Y, Shinohara A, Marumo K. Anatomical considerations for the accurate free-handed placement of cervical pedicle screws (in Japanese). *Rinsho Seikeigeka* 2009; **44**: 257-62.

Ueno Y, Otani T, Fujii H, Kawaguchi Y, Kato T, Tamegai H, Marumo K. Utilities of computer simulation and individual bone model in the hip surgeries (in Japanese). *Seikeigeka* 2009; **60**: 1395-400.

Otani T, Fujii H, Ueno Y, Kawaguchi Y, Kato T, Marumo K. Short-term clinical results of the detorsion femoral osteotomy for hip osteoarthritis (in Japanese). Seikeigeka 2010; 61: 11-6.

Chazono M, Soshi S, Inoue T, Kida Y, Shinohara A, Marumo K. Surgical treatment for posterior recapping laminectomy with tumor resection using two different threadwire saw (in Japanese). *Nippon Sekitsui Installmentation Gakkaishi* 2009; **8:** 52–6.

Chazono M, Soshi S, Inoue T, Kida Y, Nakamura Y, Shinohara A, Marumo K. Use of a digital skeletal age (DSA) staging system to assess skeletal maturity in Japanese patients with idiopathic scoliosis (in Japanese). *Sekitsui Henkei* 2009; **24:** 34-9.

Kawaguchi Y, Otani T, Fujii H, Ueno Y, Kato T, Tamegai H, Marumo K. Follow-up study of surgical treatment for slipped capital femoral epiphysis (in Japanese). *Hip Joint* 2009; **35**: 605-9.

Tanaka T, Kumagae Y, Chazono M, Kikuchi T, Ishikawa Y, Mitsuhashi M. Bone formation and resorption after implantation of Beta-TCP with 60% and 75% porosity in knee and hip joint. *Orthopaedic Ceramic Implants* 2009; **28:** 19-22.

Chazono M, Saito S, Ryu K, Hattori H, Soshi S, Marumo K. Clinical outcomes and surgical invasiveness for lumbar disc herniation using expandable tubular retractor (in Japanese). *Seikei Saigai Geka* 2009; **52:** 1211–5.

Otani T, Fujii H, Ueno Y, Kawaguchi Y, Kato T, Tamegai H, Marumo K. Prevention of dislocation after primary THA with posterior approach using rotationally adjustable modular femoral stem (in Japanese). *Nippon Jinko Kansetsu Gakkaishi* 2009; **39:** 16–7.

Ueno Y, Otani T, Fujii H, Kawaguchi Y, Kato T, Tamegai H, Marumo K. Computer simulation and individual bone model in total hip arthroplasty for Crowe IV osteoarthritis (in Japanese). *Nippon Jinko Kansetsu Gakkaishi* 2009; **39**: 412–3.

Marumo K, Kurosaka D. Patient-specific templating technique in total knee arthroplasty (in Japanese). *Seikei Saigai Geka* 2010; **53:** 49– 52.

Kan I, Funasaki H, Yoshida M, Kato S, Kasama K, Marumo K. Morphological and immunohistological study of the acromion in patients with rotator cuff tear (in Japanese). Kanto Seikei Saigai Geka Gakkai Zasshi 2010; **41:** 11-7.

Kawaguchi Y, Otani T, Fujii H, Kato T, Tamegai H, Marumo K. Manipulative reduction and internal fixation for unstable-type slipped capital femoral epiphysis (in Japanese). *Seikeigeka (Bessatsu)* 2010; **57:** 28–31.

Sai T, Iwayama K, Otsu M, Inoue N, Kasai H. Benzyloxycarbonyl-Leucyl-Leucinal, Z-LLL-H, induces the differentiation of mouse embryonic stem cell-derived neural stem cells neurons (in Japanese). Nippon Hoken Kagaku Gakkaishi 2010; **12:** 230-6.

Reviews and Books

Saito M. The latest term explanation: Pentosidine (in Japanese). Kotsu Soshosho chiryo 2009; 8: 61-4.

Saito M. Clinical practice for rehabilitation: Effect of mechanical stress on bone quality as a determinant of bone strength (in Japanese). *J Clin Rehabil* 2009; **18**: 509–15.

Saito M. Role of nutrition status on bone quality in osteoporosis (in Japanese). *Rinsho Eiyo* 2009; **114:** 490-5.

Saito M. Collagen cross-links in bone and bone strength (in Japanese). *Jin to Kotsutaisha* 2009; **22:** 195–205.

Saito M. A medical examination and treatment guide: deficiency of vitamin B and fracture risk (in Japanese). *chiryo* 2009; **91:** 1919-26.

Saito M. Progress in clinical practice in osteoporosis; what is bone quality? (in Japanese). *InternMed* 2009; **104:** 516-20.

Saito M. Biochemical markers of bone turnover. New aspect. Bone collagen metabolism: new biological markers for estimation of bone quality (in Japanese). *Clin Calcium* 2009; **19:** 1110–7.

Saito M. Nutrition and bone health. Roles of vitamin C and vitamin B as regulations of bone mass and quality (in Japanese). *Clin Calcium* 2009; **19:** 1192–9.

Saito M. Poor bone quality in diabetes and arterioscelerosis (in Japanese). *Clin Calcium* 2009; **19:** 1257-68.

Saito M. Bone quality is a determinant of bone strength and the role of nutrition status on bone quality (in Japanese). *Shokuhin to Kaihatsu* 2009; **44:** 9–15.

Saito M. Collagen cross-links (in Japanese). SERM 2009; **7:** 2–5.

Saito M. Raloxifene improves bone quality (in Japanese). SERM 2009; **7:** 12-8.

Saito M. Bone quality markers: pentosidine, homocysteine, and MTHFR polymorphism (in Japanese). *Rinsho Byori* 2009; **57:** 876-83.

Saito M. A plan for improvement of postmenopausal osteoporotic women: the agent for treatment of osteoporosis (in Japanese). *Sanfujinka Chiryo* 2009; **99:** 461–8.

Saito M. Vitamin K and osteoporosis: vitamin K2 improves bone quality (in Japanese). *Kotsu Soshosho Chiryo* 2009; **8:** 205-11.

Saito M. Effect of vitamin K on bone material

properties (in Japanese). *Clin Calcium* 2009; **19:** 1797–804.

Saito M. The latest novel knowledge of basic science in orthopaedics: collagen cross-links (in Japanese). *Rinsho Seikeigeka* 2009; **44:** 1218-24.

Kubota M, Taguchi T. Achilles tendon injury: Cause and diagnosis (in Japanese). *Orthopaedics* 2009; **22:** 1–5.

Omori T, Marumo K. Diagnosis of anterior cruciate ligament injuries (in Japanese). Orthopaedics 2009; **22:** 1-8.

Suzuki H, Marumo K. Anterior cruciate ligament reconstruction using semitendinosus and gracilis tendon autografts with poly-L-lactic acid ligament augmentation device (in Japanese). Orthopaedics 2009; 22: 22-35.

Shinohara A, Saito M, Soshi S, Marumo K. Dialysis related osteoarthropathy and rehabilitation in hemodialysis patients (in Japanese). Jin to Toseki 2009; 66: 825–30.

Maeda K. Vocabulary Wnt5a (in Japanese). Seikeigeka 2009; **60:** 1100.

Otani T. Primary total hip arthroplasty with a

modular femoral stem: A way to maximize its advantage of modularity (in Japanese). *Seikei Saigai Geka* 2010; **53:** 54-5.

Saito M, Marumo K. Collagen cross-links as a determinant of bone quality: a possible explanation for bone fragility in aging, osteoporosis, and diabetes mellitus. *Osteop Int* 2010; **21:** 195-214.

Saito M. Bone quality in life-style related diseases (in Japanese). Jin to kotsutaisha 2010; 23: 15-26.

Saito M. SERMs and bone: Foreword; new era in the treatment of osteoporosis (in Japanese). *Clin Calcium* 2010; **20:** 7.

Saito M. Effects of SERMs on bone health. Effects of raloxifene and bisphosphonate on bone quality in osteoporosis: collagen cross-links, mineralization, and bone strength (in Japanese). *Clin Calcium* 2010; **20:** 345–54.

Saito M. Musculoskeletal rehabilitation and bone. Mechanical stress and bone quality: do mechanical stimuli alter collagen cross-link formation in bone? "Yes" (in Japanese). *Clin Calcium* 2010; **20:** 345-354.