

## 抗リン脂質抗体症候群の治療方針試案

---

### 〈静脈血栓症〉

- ・ワーファリンが第一選択（INR 約 2.0）
- ・少量アスピリン（81-100 mg/日）の併用

### 〈動脈血栓症〉

- ・少量アスピリン必須
- ・血小板凝集抑制剤の併用：シロスタゾール 200mg/日、塩酸チクロピジン 100-200 mg/日、または他の血小板凝集抑制剤でもよい
- ・症例により<sup>1)</sup> ワーファリンの併用（INR 約 2.0）

### 〈妊娠合併症〉

- 1) 妊娠合併症の既往のある場合
    - (1) 少量アスピリン
    - (2) (1) が無効のとき、ヘパリン（または低分子ヘパリン）の併用
  - 2) 血栓症の既往のある場合  
少量アスピリンとヘパリン（または低分子ヘパリン）併用
- 

1) 動脈血栓症でワーファリンが必要な場合：弁膜合併症の存在する時、明らかなトロンビン生成の亢進を認める場合、血小板凝集抑制剤を使用しても血栓症が再発するとき、など

## IgG aPS/PT (U)

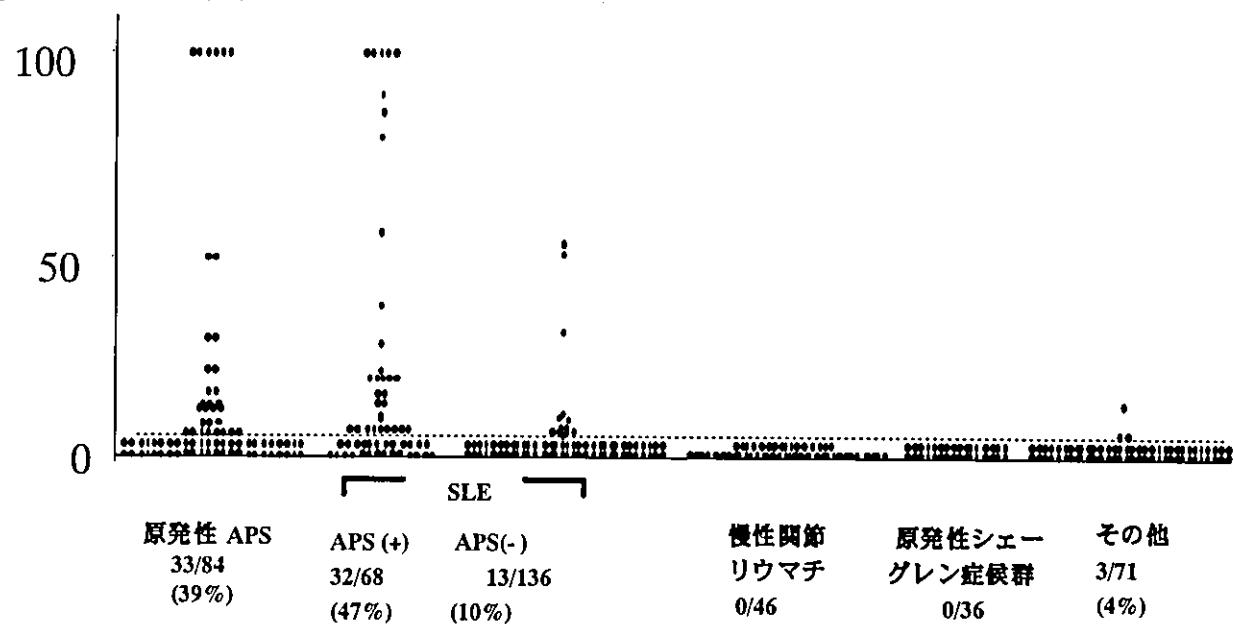


図1. ホスファチジルセリン依存性抗プロトロンビン抗体の分布

## F. 研究発表

### 1. 論文発表

1. Yasuda S, Atsumi T, Matsuura E, Kaihara K, Yamamoto D, Ichikawa K, Koike T. Significance of valine/leucine<sup>247</sup> polymorphism of  $\beta_2$ -glycoprotein I in antiphospholipid syndrome: increased reactivity of anti- $\beta_2$ -glycoprotein I autoantibodies to the valine<sup>247</sup>  $\beta_2$ -glycoprotein I variant. *Arthritis Rheum* (in press)
2. Atsumi T, Matsuura E, Koike T. Immunology of anti-phospholipid antibodies and cofactors. RB Lahita ed in "Systemic Lupus Erythematosus" 1081-1105, 2004. Elsevier Academic Press
3. Li,N., Nakamura,K., Jiang,Y., Tsurui,H., Matsuoka,S., Abe,M., Ohtsuji,M., Nishimura, H., Kato,K., Kawai,T., Atsumi,T., Koike,T., Shirai,T., Ueno,H., Hirose,S. Gain-of-function polymorphism in mouse and human Ltk: implications for the pathogenesis of lupus erythematosus. *Hum Mol Genet* 13:2 171-179.2004
4. Shimizu,C., Koike,T., Sawamura,Y. Double pituitary adenomas with distinct histological features and immunophenotypes. *J Neurol Neurosurg Psychiatry* 75:140.2004
5. Endo,T., Nakao,S., Koizumi,K., Nishio,M., Fujimoto,K., Sakai,T., Kuwano,K., Obara,M., Koike,T. Successful treatment with rituximab for autoimmune hemolytic anemia concomitant with proliferation of Epstein-Barr virus and monoclonal gammopathy in a post -non myeloblastic stem cell transplant patient. *Ann Hematol.* 83. 114-116.2004
6. Koizumi,K., Fujimoto,K., Haseyama,Y., Endo,T., Nishio,M., Yokota,K., Itoh,K., Sawada,K., Koike,T.: Effective high-dose chemotherapy combined with CD34+-selected peripheral blood stem cell transplantation in a patient with cutaneous involvement of nasal NK/T-cell lymphoma. *Eur J Haematol.* 72:140-144. 2004
7. Nagai,S., Shimizu,C., Umetsu,M., Taniguchi,S., Endo,M., Miyoshi,H., Yoshioka,N., Kubo,M., Koike,T.: Identification of a functional peroxisome proliferator-activated receptor responsive element within the murine perilipin gene. *Endocrinology*. 145. 2346-2356.2004
8. Yasuda,S., Atsumi,T., Ieko,M., Matsuura,E., Kobayashi,K., Inagaki,J., Kato,H., Tanaka,H., Yamakado,M., Akino,M., Saitou,H., Amasaki,Y., Jodo,S., Amengual,O., Koike,T.: Nicked  $\beta_2$ -glycoprotein I: a marker of cerebral infarct and a novel role in the negative feedback pathway of extrinsic fibrinolysis. *Blood*. 103:10. 3766-3772.2004
9. Kubo,M., Shimizu,C., Kijima,H., Nagai,S., Koike,T.: Alternate promoter and 5'-Untranslated exon usage of the mouse adrenocorticotropin receptor gene in adipose tissue. *Endocrinol J.* 51.25-30. 2004
10. Endo,T., Mogi,Y., Koizumi,K., Nishio,M., Fujimoto,K., Sakai,T., Kumano,K., Obara,M., Ikeda,H., Koike,T.: Peripheral blood stem cell mobilization following plus rituximab therapy combined with G-CSF in patients with B-cell non-Hodgkin's lymphoma. *Bone Marrow Transpl.* 33.703-707. 2004
11. Yamamoto,S., Tsuji,T., Matsuzaki,J., Zhang,Y., Chamoto,K., Kosaka,A., Togashi,Y., Sekikawa,K., Sawada,K., Takeshima,T., Koike,T., Nishimura,T.: Unexpected role of TNF- $\alpha$  in graft versus host reaction (GVHR): donor-derived TNF- $\alpha$  suppresses GVHR via inhibition of IFN- $\gamma$ -dependent donor type-1 immunity. *Int Immunol.* 16: 811-817. 2004
12. Endo,T., Mogi,Y., Koizumi,K., Nishio,M., Fujimoto,K., Sakai,T., Kumano,K., Obara,M., Ikeda,H., Koike,T.: Peripheral blood stem cell mobilization following plus rituximab therapy combined with G-CSF in patients with B-cell non-Hodgkin's lymphoma. *Bone Marrow Transpl.* 33.703-707. 2004
13. Hashimoto,S., Kawata,T., Schnermann,J., Koike,T.: Chloride Channel Blockade Attenuates the Effect of Angiotensin II on Tubuloglomerular Feedback in WKY but not Spontaneously Hypertensive Rats. *Kidney Blood Press R.* 27:35-42.2004
14. Yasuda,S., Atsumi,T., Ieko,M., Matsuura,E., Kobayashi,K., Inagaki,J., Kato,H., Tanaka,H., Yamakado,M., Akino,M., Saitou,H., Amasaki,Y., Jodo,S., Amengual,O., Koike,T.: Nicked  $\beta_2$ -glycoprotein I: a marker of cerebral infarct and a novel role in the negative feedback pathway of extrinsic fibrinolysis. *Blood*. 103:10. 3766-3772.2004
15. Yasuda,S., Ogura,N., Horita,T., Yasuda,I., Hioka,T., Kondo,N., Fujisaku,A. Abacterial prostatitis and primary biliary cirrhosis with Sjogren's syndrome. *Mod Rheumatol* 14:70-72.2004
16. Das,H., Atsumi,T., Fukushima,Y., Shibuya,H., I-

- to,K., Yamada,Y., Amasaki,Y., Ichikawa,K., Amengual,O., Koike,T.: Diagnostic value of antiagalactosyl IgG antibodies in rheumatoid arthritis. *Clin Rheumatol* 23:218-222.2004
17. Ieko,M., Tarumi,T., Takeda,M., Nito,S., Nakabayashi,T., Koike,T.: Synthetic selective inhibitors of coagulation factor Xa strongly inhibit thrombin generation without affecting initial thrombin forming time necessary for platelet activation in hemostasis. *J Thrombosis and Haemostasis* 2: 612-622.2004
18. Amengual,O., Atsumi,T., Koike,T.: Antiprothrombin antibodies and the diagnosis of antiphospholipid syndrome. *Clin Immunol* 112: 144-149.2004
19. Suzuki,F., Shimizu,C., Umetsu,M., Nagai,S., Takeuchi,J., Endo,M., Miyoshi,H., Yoshioka,N., Mitsumasa,K., Koike,T.: Adult-onset idiopathic hypogonadotropic hypogonadism due to isolated pituitary gonadotropin deficiency. *Internal med* 43:7. 571-574.2004
20. Endo,T., Koizumi,K., Nishio,M., Fujimoto,K., Sakai,T., Kumano,K., Obara,M., Minauchi,K., Koike,T.: Localized relapse in bone marrow of extremities after allogeneic stem cell transplantation for acute lymphoblastic leukemia. *Am J Hematol* 76: 279-282.2004
21. Yang,L., Hakoda,M., Iwabuchi,K., Takeda,T., Koike,T., Kamatani,N., Takada,K.: Rheumatoid Factors Induce Signaling from B cells, leading to Epstein-barr virus and b-cell activation. *J Virol* 78:18. 9918- 9923.2004
22. Kataoka,H., Koike,T.: Lupus mortality in Japan. *Autoimmunity Reviews* 3: 421-422.2004
23. Endo,T., Sato,N., Koizumi,K., Nishio,M., Fujimoto,K., Yamamoto,S., Sakai,T., Bohgaki,T., Sawada,K., Koike,T.: A preliminary analysis of the balance between Th1 and Th2 cells after CD34+ cell-selected autologous PBSC transplantation. *Cytotherapy* 6:4.337-343.2004
24. Xiao,S., Deshmukh,S.U., Jodo,S., Koike,T., Sharma,R., Furusaki,A., Sung,J.S., Ju,Shyr-Tu.: Novel negative regulator of expression in Fas Ligand(CD178)Cytoplasmic tail: Evidence for Translational Regulation and against Fas Ligand Retention in secretory lysosomes. *Journal of Immunology* 173: 5095-5102.2004
25. Yasuda,S., Atsumi,T., Ieko,M., Koike,T.: $\beta$ 2-glycoprotein I, anti- $\beta$ 2-glycoprotein I, and fibrinolysis. *Throm Res* 114: 461- 465.2004
26. Astumi,T., Amengual,O., Yasuda,S., Koike,T.: Antiprothrombin antibodies—are they worth assaying? *Throm Res* 114: 533-538.2004
27. Hashimoto,S., Ogawa,Y., Ishida,T., Mochizuki, T., Koike,T., Sato,H., Ueda,T.: Steroid-sensitive nephrotic syndrome associated with positive C1q immunofluorescence. *Clin Exp Nephrol* 8: 266- 269.2004
28. Bohgaki,M., Atsumi,T., Yamashita,Y., Yasuda, S., Sakai,Y., Furusaki,A., Bohgaki,T., Amengual,O., Amasaki,Y., Koike,T.: The p38 mitogen-activated protein kinase(MAPK)pathway mediates induction of the tissue factor gene in monocytes stimulated with human monoclonal anti- $\beta$ 2Glycoprotein I antibodies. *Int Immunol* 16:11. 1633- 1641.2004
29. Sugiura-ogasawara.M., Atsumi,T., Ozaki,Y., Koike,T., Suzumori,K.: Phosphatidylserine-dependent antiprothrombin antibodies are not useful markers for high-risk woman with recurrent miscarriages. *Fertil Steril* 82:51440-1442.2004
30. Higuchi,M., Ishizu,A., Ikeda,H., Hayase,H., Fugo,K., Tsuji,M., Abe,A., Sugaya,T., Suzuki,A., Takahashi,T., Koike,T., Yoshiki,T.: Functional alteration of peripheral CD25<sup>+</sup>CD4<sup>+</sup> immunoregulatory T cells in a transgenic rat model of autoimmune diseases. *J Autoimmunity* 20. 43-49. 2003.
31. Wu,R., Shoenfeld,Y., Shere,Y., Patnaik,M., Matsuura,E., Gilburd,B., Koike,T., Peter,B.J.: Anti-idiotypes to oxidized LDL antibodies in intravenous immunoglobulin preparations—possible immunomodulation of atherosclerosis. *Autoimmunity*. 36; 91-97. 2003.
32. Sakai,Y., Atsumi,T., Itoh,T., Koike,T.: Uveitis, pancarditis, haemophagocytosis, and abdominal masses. *Lancet* 361.834. 2003.
33. Hashimoto,S., Shiroshita,K., Sakurai,T., Takeda,A., Kawata,T., Koike,T., Sato,H., Ueda,T.: Unilateral renal cortical necrosis with contralateral hydronephrosis after surgery for uterus carcinoma. *Clin Exp Nephrol*. 7. 72-76. 2003.
34. Koizumi,k., Nakao,S., Haseyama,Y., Kato,H., Ohi,M., Motohara,T., Endo,T., Sawada,K., Koike,T.: Severe aplastic anemia associated with thymic carcinoma and partial recovery

- of hematopoiesis after thymectomy. *Ann Hematol*. 82:367-370. 2003.
35. Kawakubo,K., Endo,T., Koizumi,K., Nishio,M., Fujimoto,K., Sakai,T., Sawada,K., Koike,T.: Granular lymphocyte proliferative disorder after autologous peripheral blood stem cell transplantation for multiple myeloma. *Eur J Haematol*. 71: 1-4. 2003.
36. Shoenfeld,E. ,Krause,I., Kvapil,F., Sulkes,J., Lev,S., Von Landenberg,P., Font,J., Zaech,J., Cervera,R. ,Piette,J.C., Boffa,M.C., Khamashta,M.A., Bertolaccini,M.L., Hughes,G.R.V., Y ouinou,P., Meroni,P.L., Pengo,V., Alves,J.D., Tincani,A., S zegedi,G. , Lakos,G. ,Sturfelt,G., Jonsen,A., Koike,T., Sanmarco,M. ,Ruffatti,A. ,Ulcova-Gallova,Z., Praprotnic,S., Rozman,B., Lorber,M., Vriezman,V.B., Blank,M.: Prevalence and clinical correlantions of antibodies against six  $\beta$ 2-glycoprotein-I-related peptides in antiphospholipid syndrome. *J Clin Immunol*. 23: 375-381. 2003.
37. Oku,K., Atsumi,T., Furukawa,S., Horita,T., Sakai,Y., Jodo,S., Amasaki,Y., Ichikawa,K., Amengual O., Koike,T.: Cerebral imaging by magnetic resonance imaging and single photon emission computed tomography in systemic lupus erythematosus with central nervous system involvement. *Rheumatol*. 42; 773-777. 2003.
38. Nagasaka,K. ,Harigai,M., Tateishi,M., Hara,M., Y Oshizawa,Y., Koike,T., Miyasaka,N: Efficacy of combination treatment with cyclosporin A and corticosteroids for acute interstitial pneumonitis associated with dermatomyositis. *Mod Rheumatol*. 13:231-238. 2003.
39. Horita,T., Tsutsumi,A., Takeda,T., Yasuda,S., Takeuti,R., Amasaki,Y., Ichikawa,K., Atsumi,T., Koike,T.: Significance of magnetic resonace imafing in diagnosis of nodular regenerative hyperplasia of the liver complicated with systemic lupus erythematosus:a case report and review of the literature. *Lupus*. 11:193-196.2002
40. Okamoto,T., Tanaka,S., Stan.C.A., Koike,T., Kas,M., Makita,Z., Sawa,H., Nagashima,K.:Advanced glycation end products induce angiogenesis in vivo. *Mol Res* 3:186-195. 2002.
41. Ieko,M., Nakabayashi,T., Takeda,T., Naitoh,S., Atsumi,T., Koike,T.: The inhibition of protein C amticoagulamt activity by anti- $\beta$ 2-glycoprotein I( $\beta$ 2GPI)antibodies isolated from patients with antiphospholipid syndrome by chromatography methods. *Mod Rheumatol*. 12:44-49.2002.
42. Yasuda,S., Tsustumi,A., Atsumi,T., Bertolaccini,M.L., Ichikawa,K., Khamashta,M.A., HugesR.V., Koike,T.: Gene polymorphisms of tissue plasminogen activator and plasminogenactivator inhibitor-1 in patients with antiphospholipid antibodies. *J .Rheumatol*. 29:6.1192-1197.2002.
43. Amborozic,A., Avicin,T., Ichikawa,K., Kveder,T., Mastuur,E., Hojnik,M., Astumi,T., Rozman,B., Koike,T.: Anti- $\beta$ 2-glycoprotein I antibodies in children with atopic dermatitis. *Int.Immunol*.14:7.823-830.2002.
44. Koizumi,K. ,Haseyama,Y., Machino,R. ,Sato,Y., Sawada,K., Koike,T.: The hemophagocytic syndrome in prostate cancer renealed by disseminated carcinoinatosis of the bone marrow. *J Urology*. 168, 1101-1102. 2002.
45. Yamaguchi,M. ,Hirayama,F., Murahashi,H., Azuma,H., Sato,N. ,Miyazaki,H., Fukazawa,K., Sawada,K., Koike,T., Ikeda,H., Ikebuchi,K.: Ex vivo expansion of human UC blood primitive hematopoietic progenitors and transplantable stem cells using human primary BM stromal cells and human AB serum. *Cyotherapy*. 4:2.109-118 .2002.
46. Miyoshi,H., Taguchi,T., Sugiura,M., Takeuchi,M., Yanagisawa,K., Watanabe,Y., Miwa,I., Makita,Z., Koike,T.: Aminoguanidine pyridoxal adduct is superior to aminoguanidine for preventing diabetic nephropathy in mice. *Horm Metab Res*. 34. 371-377. 2002.
47. Atsumi,T., Koike,T. :Clinical relevance of antiprothrombin antibodies. *Autoimmunity Reviews*.1:49-53.2002.
48. Matsuura,E., Kobayashi,K., Kasahara,J., Yasuda,T., Makino,H., Koike,T., Shoenfeld,Y.: anti- $\beta$ 2-glycoprotein I antibodies and atherosclerosis. *Int. Rev. Immunol*.21.51-66.2002.
49. Tyndall A. , Koike,T.: High -Dose immunoablative therapy with hematopoietic stem cell support in the treatment of severe autoimmune disease: current status and future direction. *Int Med*. 41: 608-612. 2002.

50. Amengual O, Atsumi T, Koike T: Specificities, properties, and clinical significance of antiprothrombin antibodies. *Arthritis Rheum* 48:886-895, 2003
51. Pang M, Setoyama Y, Tsuzaka K, Yoshimoto K, Amano K, Abe T, and Takeuchi T. Defective expression and tyrosine phosphorylation of the T cell receptor zeta chain in peripheral blood T cells from systemic lupus erythematosus patients. *Clin Exp Immunol* 129: 160-169, 2002.
52. Tsuzaka K, Fukuhara I, Setoyama Y, Yoshimoto K, Suzuki K, Abe T, and Takeuchi T. Forced expression of TCR $\zeta$  mRNA with alternatively spliced 3' untranslated region found in SLE patients lead to decreased production and cell surface expression of TCR $\zeta$  and TCR-CD3 complex. *J Immunol* 171:2496-2503, 2003.
53. Takeuchi T, Tsuzaka K, and Abe T. Altered expression of the T cell receptor-CD3 complex in systemic lupus erythematosus. *Int Rev Immunol* 23:273-291, 2004.
54. Takeuchi T, Tsuzaka K, Abe T, Yoshimoto K, Shiraishi K, and Amano K. T cell abnormalities in systemic lupus erythematosus. *Autoimmunity* in press.
55. Higuchi, T., Aiba, Y., Nomura, T., Matsuda, J., Mochida, K., Suzuki, M., Kikutani, H., Honjo, T., Nishioka, K. and Tsubata T. 2002: Cutting Edge: Ectopic expression of CD40 ligand on B cells induces lupus-like autoimmune disease. *J. Immunol.* 168: 9-12.
56. Wakabayashi, C., Adachi, T., Wienands, J., and Tsubata, T. 2002: A distinct Signaling Pathway Used by the IgG-Containing B Cell antigen receptor. *Science*. 298: 2392-2395
57. Suzuki, A., Kaisho, T., Ohishi, M., Tsukio-Yamaguchi, M., Tsubata, T., Koni, P.A., Sasaki, T., Mak, T.W., and Nakano, T. 2003: Critical roles of Pten in B cell homeostasis and immunoglobulin class switch recombination. *J. Exp. Med.* 197: 657-667
58. Hokazono, Y., Adachi, T., Wabl, M., Tada, N., Amagasa, T., and Tsubata T. : Inhibitory co-receptors activated by antigens but not by anti-immunoglobulin heavy chain antibodies install requirement of co-stimulation through CD40 for survival and proliferation of B cells. *J. Immunol.* 171:1835-1843, 2003
59. Kawamura, T., Kanai, T., Dohi, T., Uraushihara, K., Totsuka, T., Iiyama, R., Taneda, C., Yamazaki, M., Nakamura, T., Higuchi, T., Aiba, Y., Tsubata, T. and Watanabe, M.: Ectopic CD40 ligand expression on B cells trigger intestinal inflammation. *J. Immunol.* 172: 6388-6397, 2004
60. Yokota S, Miyamae T, Imagawa T, Iwata N, Katakura S, Mori M, Woo P, Nishimoto N, Yoshizaki K, Kishimoto T. Therapeutic Efficacy of Humanized Recombinant Anti-IL 6-Receptor Antibody for Children with Systemic-Onset Juvenile Idiopathic Arthritis. *Arthritis Rheum* (in press)
61. Nishimoto N, Yoshizaki K, Miyasaka N, Yamamoto K, Kawai S, Takeuchi T, Hashimoto J, Azuma J, Kishimoto T. Treatment of Rheumatoid Arthritis with Humanized Anti-interleukin 6 Receptor Antibody. *Arthritis Rheum* 50: 1761-1769, 2004.
62. Becker C, Fantini MC, Schramm C, Lehr HA, Wirtz S, Nikolaev A, Burg J, Strand S, Kiesslich R, Huber S, Ito H, Nishimoto N, Yoshizaki K, Kishimoto T, Galle PR, Blessing M, Rose-John S, Neurath MF. TGF-beta suppresses tumor progression in colon cancer by inhibition of IL-6 trans-signaling. *Immunity* 21:491-501, 2004.
63. Nakahara H, Song J, Sugimoto M, Hagiwara K, Kishimoto T, Yoshizaki K, Nishimoto N. Anti-interleukin-6 receptor antibody therapy reduces vascular endothelial growth factor (VEGF) production in rheumatoid arthritis. *Arthritis Rheum* 48:1521-1529, 2003.
64. Nishimoto N, Yoshizaki K, Maeda K, K.Kuritani T, Deguchi H, Sato B, Imai N, Kakehi T, Takagi N, Suemura M, Kishimoto T. Toxicity, pharmacokinetics, and dose-finding study of repetitive treatment with the humanized anti-interleukin 6 receptor antibody MRA in rheumatoid arthritis. Phase I/II clinical study. *J. Rheumatol* 30:1426-1435, 2003.
65. Li N, Nakamura K, Jiang Y, Tsurui H, Matsuoka S, Abe M, Ohtsuji M, Nishimura H, Kato K, Kawai T, Atsumi T, Koike T, Shirai T, Ueno H, Hirose S: Gain-of-function polymorphism in mouse and human *Ltk*: implications for the pathogenesis of systemic lupus erythematosus. *Hum Mol Genet*, 13:171-179, 2004.
66. Zhang D, Fujio K, Jiang Y, Zhao J, Tada N, Sudo K, Tsurui H, Nakamura K, Yamamoto

- K, Nishimura H, Shirai T, Hirose S: Dissection of the role of MHC class II A and E genes in autoimmune susceptibility in murine lupus models with intragenic recombination. *Proc Natl Acad Sci USA*, 101:13838-13843, 2004.Nakamura K, Xiu Y, Ohtsuji M, Sugita G, Abe M, Ohtsuji N, Hamano Y, Jiang Y, Takahashi N, Shirai T, Nishimura H, Hirose S. Genetic dissection of anxiety in autoimmune disease. *Hum Mol Genet*, 12:1079-1086, 2003.
67. Miura-Shimura Y, Nakamura K, Ohtsuji M, Tomita H, Jiang Y, Abe M, Zhang D, Hamano Y, Tsuda H, Hashimoto H, Nishimura H, Taki S, Shirai T, Hirose S: *C1q* regulatory region polymorphism down-regulating murine *C1q* protein levels with linkage to lupus nephritis. *J Immunol* 169:1334-1339, 2002.
68. Xie Y, Nakamura K, Abe M, Li N, Wen X-S, Jiang Y, Zhang D, Tsurui H, Matsuoka S, Hamano Y, Fujii H, Ono M, Takai T, Shimokawa T, Ra C, Shirai T, Hirose S: Transcriptional regulation of *Fcgr2b* gene by polymorphic promoter region and its contribution to humoral immune responses. *J Immunol*, 169:4340-4346, 2002.
69. Kudo, H., Matsuoka, T., Mitsuya, H., Nishimura, Y. and Matsushita, S. Cross-linking HLA-DR molecules on Th1 cells induces anergy in association with increased level of cyclin-dependent kinase inhibitor p27Kip1. *Immunol. Lett.* 81: 149-155, 2002
70. Ohyama, H., Nishimura, F., Meguro, M., Takashiba, S., Murayama, Y. and Matsushita, S. Counter-antigen presentation: fibroblasts produce cytokines by signaling through HLA class II molecules without inducing T-cell proliferation. *Cytokine* 17: 175-181, 2002
71. Meguro, M., Nishimura, F., Ohyama, H., Takashiba, S., Murayama, Y. and Matsushita, S. Ligation of HLA-DR molecules on fibroblasts induces rantes expression via c-Jun N-terminal kinase (JNK) pathway. *Cytokine* 22: 107-115, 2003.
72. Matsushita, S., Ohyama, H., Kudo, H., Tabata, H. and Matsuoka, T. HLA-mediated signaling via HLA-peptide-TCR complex determines immune responses of antigen-presenting cells. *Current Topics in Peptide & Protein Research*.6: 1-20, 2004.
73. Liu, T., Kohsaka, H., Suzuki, M., Takagi, R., Hashimoto, K., Uemura, Y., Ohyama, H., and Matsushita, S. Positional effect of amino acid replacement on peptide antigens for the increased IFN $\gamma$  production from CD4T cells. *J. Allergy Clin. Immunol.* 113: 216-217, 2004
74. Fugo K, Ishizu A, Ikeda H, Hayase H, Sugaya T, Higuchi M, Tsuji M, Abe A, Suzuki A, Shibata M, Takahashi T, Yoshiki T. The role of the thymus in development of necrotizing arteritis in transgenic rats carrying the env-pX gene of human T cell leukemia virus type I. *Am J Pathol* 161: 755-761, 2002.
75. Higuchi M, Ishizu A, Ikeda H, Hayase H, Fugo K, Tsuji M, Abe A, Sugaya T, Suzuki A, Takahashi T, Koike T, Yoshiki T. Functional alteration of peripheral CD25 $^{+}$ CD4 $^{+}$  immunoregulatory T cells in a transgenic rat model of autoimmune diseases. *J Autoimmun* 20: 43-49, 2003.
76. Ishizu A, Tsuji T, Abe A, Saito S, Takahashi T, Ikeda H, Meruelo D, Yoshiki T. Transduction of dominant negative ATF-1 suppresses the pX gene expression in joint fibroblastic cells derived from HTLV-I transgenic rats. *Exp Mol Pathol* 74: 309-313, 2003.
77. Tsuchikawa T, Ikeda H, Kikuchi K, Tsuji T, Baba T, Ishizu A, Tanaka Y, Kato H, Yoshiki T. Hematopoietic progenitor cells as possible origins of epithelial thymoma in a human T lymphocyte virus type I pX gene transgenic rat model. *Lab Invest* 84: 245-252, 2004.
78. Abe A, Ishizu A, Ikeda H, Hayase H, Tsuji T, Miyatake Y, Tsuji M, Fugo K, Sugaya T, Higuchi M, Matsuno T, Yoshiki T. Bone marrow cells carrying the *env-pX* transgene play a role in the severity but not prolongation of arthritis in human T-cell leukemia virus type-I transgenic rats: a possible role of articular tissues carrying the transgene in the prolongation of arthritis. *Int J Exp Pathol* 85: 191-200, 2004.
79. Kobayashi T, Ito S, et al. Risk of Periodontitis in Systemic Lupus Erythematosus is Associated with Fcg Receptor Polymorphisms. *J Periodontology* 74:378-384, 2003
80. Ito S, et al. Patient with diffuse mesangial and endocapillary proliferative glomerulonephritis with hypocomplementemia and elevated anti-streptolysin O treated with prednisolone, angiotensin-converting enzyme inhibitor, and

- angiotensin II receptor antagonist. *Clin Exp Nephrol* 7:290-295, 2003
81. Takahashi R, Tsutsumi A, Ohtani K, Goto D, Matsumoto I, Ito S, et al. Anti-mannose binding lectin antibodies in sera of Japanese patients with systemic lupus erythematosus. *Clin Exp Immunol* 136:585-590, 2004
82. Ida H, Kawakami A, Eguchi K. ReGranzyme B leakage-induced cell death: a new type of activation-induced natural killer cell death. *Eur J Immunol*. 33 (12): 3284-3292, 2003.
83. Shigeno M, Nakao K, Ichikawa T, Suzuki K, Kawakami A, Abiru S, Miyazoe S, Nakagawa Y, Ishikawa H, Hamasaki K, Nakata K, Ishii N, Eguchi K. Interferon- $\alpha$  sensitizes human hepatoma cells to TRAIL-induced apoptosis through DR5 upregulation and NF- $\kappa$ B inactivation. *Oncogene* 22 (11): 1653-1662, 2003.
84. Nakashima K, Kawakami A, Hida A, Yamasaki S, Nakamura H, Kamachi M, Miyashita T, Tanaka F, Izumi Y, Tamai M, Ida H, Furuyama M, Koji T, Nakamura T, Migita K, Origuchi T, Eguchi K. Protection of mitochondrial perturbation by human T-lymphotropic virus type 1 tax through induction of Bcl-xL expression. *J Lab Clin Med* 142 (5): 341-347, 2003.
85. Origuchi T, Kawasaki E, Ide A, Kamachi M, Tanaka F, Ida H, Kawakami A, Migita K, Eguchi K. Correlation between interleukin 10 gene promoter region polymorphisms and clinical manifestations in Japanese patients with Sjögren's syndrome. *Ann Rheum Dis* 62 (11): 1117-1118, 2003.
86. Kamachi M, Kawakami A, Yamasaki S, Hida A, Nakashima T, Nakamura H, Ida H, Furuyama M, Nakashima K, Shibatomi K, Miyashita T, Migita K, Eguchi K. Regulation of apoptotic cell death by cytokines in a human salivary gland cell line: distinct and synergistic mechanisms in apoptosis induced by tumor necrosis factor alpha and interferon  $\gamma$ . *J Lab Clin Med* 139 (1): 13-19, 2002
87. Okazaki, T. et al., Autoantibodies against cardiac troponin I are responsible for dilated cardiomyopathy in PD-1-deficient mice. *Nat. Med.* 9:12, 1477-1483.2003
88. Iwai, Y. et al., PD-1 inhibits antiviral immunity at the effector phase in the liver. *J. Exp. Med.* 198:1, 39-50.2003
89. Iwai, Y. et al., Involvement of PD-L1 on tumor cells in the escape from host immune system and tumor immunotherapy by PD-L1 blockade. *Proc Natl Acad Sci U.S.A.* 99:19, 12293-12297.2002
90. Okazaki, T. et al., New regulatory co-receptors: inducible co-stimulator and PD-1. *Curr Opin Immunol.* 14:6, 779-782.2002
91. Ishida, M. et al., Differential expression of PD-L1 and PD-L2, ligands for an inhibitory receptor PD-1, in the cells of lymphohematopoietic tissues. *Immunol Lett.* 84:1, 57-62.2002
92. Okamoto N, Yotsuyanagi H, Ooka S, Matsui T, Suzuki-Kurokawa M, Suzuki M, Jino S, Nishioka K, Kato T. Autoantibodies to CD69 in patients with chronic hepatitis type C: A candidate marker for predicting the response to interferon therapy. *Intervirology*;46(1);56-65;2003.
93. Yuan GH, Masuko-Hongo K, Kato T, Nishioka K. Immunological intervention in the Pathogenesis of Osteoarthritis. *Arthritis Rheum*;48(3);602-611;2003.
94. Ooka S, Matsui T, Nishioka K, Kato T. Autoantibodies to low-density-lipoprotein-receptor-related protein 2 (LRP2) in systemic autoimmune diseases. *Arthritis Res Ther*; 5(3);R174-R180;2003.
95. Xiang Y, Sekine T, Nakamura H, Imajoh-Ohmi S, Fukuda H, Nishioka K, Kato T. Proteomic Surveillance of Autoimmunity in Osteoarthritis - Identification of Triosephosphate Isomerase as an Autoantigen in Patients With Osteoarthritis. *Arthritis Rheum*;50(5);1511-1521;2004.
96. Shan ZZ, Masuko-Hongo K, Dai SM, Nakamura H, Kato T, Nishioka K. A potential role of 15-deoxy- $\Delta$ 12,14-prostaglandin J2 for induction of human articular chondrocyte apoptosis in arthritis. *J Biol Chem*; 279(36);37939-37950;2004,
97. Yoshida K, Arai T, Kaburaki J, Ikeda Y, Kawakami Y, Kuwana M. Restricted T cell receptor  $\beta$ -chain usage by T cells autoreactive to  $\beta_2$ -glycoprotein I in patients with antiphospholipid syndrome. *Blood* 2002; 99(7): 2499-2504.
98. Kuwana M. Autoreactive CD4 $^{+}$  T cells to  $\beta_2$ -glycoprotein I in patients with antiphospholipid syndrome. *Autoimmun Rev*. 2003; 2(4): 192-198.

99. Kuwana M, Nomura S, Fujimura K, Nagasawa T, Muto Y, Kurata Y, Tanaka S, Ikeda Y. The effect of a single injection of humanized anti-CD154 monoclonal antibody on the platelet-specific autoimmune response in patients with immune thrombocytopenic purpura. *Blood*. 2004; 103(4): 1229-1236.
100. Kuwana M.  $\beta_2$ -glycoprotein I: antiphospholipid syndrome and T-cell reactivity. *Thromb. Res.* 2004; 114(5-6): 347-355.
101. Kuwana M, Matsuurra E, Kobayashi K, Okazaki Y, Kaburaki K, Ikeda Y, Kawakami Y. Binding of  $\beta_2$ -glycoprotein I to anionic phospholipids facilitates processing and presentation of a cryptic epitope that activates pathogenic autoreactive T cells. *Blood*. In press.
102. N.Takasawa, Y.Munakata, K.K.Ishii, Y.Takahashi, M.Takahashi, Y.Fi, T.Ishii, H.Fujii, T.Saito, H.Takano, T.Noda, M.Suzuki, M.Nose, S.Z.Patzner, and T.Sasaki. :Human parvovirus B19 transgenic mice become susceptible to polyarthritis. *J Immunol.* 173: 4675-4683, 2004
103. T.Saito, Y.Munakata, Y.Fi, K.Ishii, T.Sasaki.:Evaluation of anti-parvovirus B19 neutralizing activity by quantitative polymerase chain-reaction – based assay. *J. Virol. Meth.* 107: 81-87, 2003
104. Y.Minami, T.Sasaki, S.Hisamichi.: Dietary factors in relation to clinical manifestations of systemic lupus erythematosus. *J.Rheumatol.* 30: 747-754, 2003
105. H.Harigae, O.Nakajima, N.Suwabe, H.Yokoyama, K.Furuyama, T.Sasaki.: Aberrant iron accumulation and oxidized status of erythroid-specific  $\delta$ -aminolevulinate synthase (ALAS2)-deficient definitive erythroblasts. *Blood*. 101: 1188-1193, 2003
106. Y.Fu, K.K.Ishii, Y.Munakata, T.Saitoh, T.Sasaki.: Regulation of TNF- $\alpha$  promoter by human parvovirus B19 NS1 through activation of AP-1 and AP-2. *J.Virol.* 76: 5395-5403, 2002
107. Ogawa N, Sugai S. Reply to the letter by Sfriso et al. *Arthritis Rheum.* 48(8):2391-2392, 2003
108. Ogawa, N., Kawanami, T., Shimoyama, K., Ping, L., Sugai, S. Expression of I-TAC (interferon-inducible T cell  $\alpha$  chemoattractant, CXCL11) in the salivary glands of patients with Sjogren's syndrome. *Clin. Immunol.* 112(3):235-238, 2004.
109. Ping, L., Ogawa, N., Sugai, S. Novel role of CD40 in Fas-dependent apoptosis in cultured salivary epithelial cells from patients with Sjogren's syndrome. *Arthritis Rheum.* in press, 2005.
110. Tsujimura S, Saito K, Tokunaga M, Nakatsuka K, Nakayamada S, Nakano K, Tanaka Y. Overcoming treatment unresponsiveness mediated by P-glycoprotein overexpression on lymphocytes in refractory active systemic lupus erythematosus. *Mod Rheumatol* (in press)
111. Tsujimura S, Saito K, Nakayamada S, Nakano K, Tsukada J, Kohno K, Tanaka Y. Transcriptional regulation of multidrug resistance-1 gene by interleukin-2 in lymphocytes. *Genes Cells* 9, 1265-1273
112. Nakano K, Okada Y, Saito K, Tanaka Y. Fibroblast growth factor-2 induces receptor activator of nuclear factor kappa B ligand expression and osteoclast maturation by binding to heparan sulfate proteoglycan on rheumatoid synovial fibroblasts. *Arthritis Rheum.* 50, 2450-2458.2004
113. Nakayamada S, Okada Y, Saito K, Tamura M, Tanaka Y.  $\beta$ 1 integrin/focal adhesion kinase-mediated signaling induces intercellular adhesion molecule 1 and receptor activator of nuclear factor  $\kappa$ B ligand on osteoblast and osteoclast maturation. *J Biol Chem* 278: 45368-45374. 2003
114. Tanaka Y, Nakayamada S, Fujimoto H, Okada Y, Umehara H, Kataoka T, Minami Y: H-Ras/mitogen-activated protein kinase pathway inhibits integrin-mediated adhesion and induces apoptosis in osteoblasts. *J Biol Chem* 277, 21446-21452.2002
115. Kyogoku C, Tsuchiya N, Wu H, Tsao BP, Tokunaga K : Association of Fc $\gamma$  receptor IIA, but not of IIB and IIIA, polymorphisms with systemic lupus erythematosus: A family-based association study in Caucasians. *Arthritis Rheum* 50:671-673.2004
- 116.5. Hitomi Y, Tsuchiya N, Kawasaki A, Kyogoku C, Ohashi J, Suzuki T, Fukazawa T, Bejrchandra S, Siriboonrit U, Chandanayong D, Suthipinittharm P, Tsao BP, Hashimoto H, Honda Z, Tokunaga K: CD72 polymorphisms associated with alternative splicing modify susceptibility to human systemic lupus erythematosus through epistatic interaction with FCGR2B. *Hum Mol Genet* 13: 2907-2917.2004Chu ZT,

- Tsuchiya N, Kyogoku C, Ohashi J, Qian YP, Xu SB, Mao CZ, Chu JY, Tokunaga K :Association of Fc $\gamma$  receptor IIb polymorphism with susceptibility to systemic lupus erythematosus in Chinese: a common susceptibility gene in the Asian populations. *Tissue Antigens* 63:21-27.2003
- 117.Kyogoku C, Dijstelbloem HM, Tsuchiya N, Hatta Y, Kato H, Yamaguchi A, Fukazawa T, Jansen MD, Hashimoto H, van de Winkel JGJ, Kallenberg CGM, Tokunaga K : Association of Fc $\gamma$  receptor gene polymorphisms in Japanese patients with systemic lupus erythematosus: Contribution of FCGR2B to the genetic susceptibility to SLE. *Arthritis Rheum* 46: 1242-1254.2002
- 118.Tsuchiya N, Ohashi J, Tokunaga K:Variations in immune response genes and their associations with multifactorial immune disorders. *Immunol Rev* 190: 169-181.2002
- 119.Suzuki, Y., Wakita, D., Chamoto, K., Narita, Y., Tsuji, T., Takeshima, T., Gyobu, H., Kawarada, Y., Kondo, S., Akira, S., Katoh, H., Ikeda, H.,Nishimura, T.: Liposome-encapsulated CpG oligodeoxynucleotides as a potent adjuvant for inducing type 1 innate immunity. *Cancer Res*, 64: 8754-8760, 2004.
- 120.Yamamoto, S., Tsuji, T., Matsuzaki, J., Zhange, Y., Chamoto, K., Kosaka, A., Togashi, Y., Sekikawa, K., Sawada, K. I., Takeshima, T., Koike, T.,Nishimura, T.: Unexpected role of TNF- $\alpha$  in graft versus host reaction (GVHR): donor-derived TNF- $\alpha$  suppresses GVHR via inhibition of IFN- $\gamma$ -dependent donor type-1 immunity. *Int Immunol*, 16: 811-817, 2004.
- 121.Sato, T., Saito, R., Jinushi, T., Tsuji, T., Matsuzaki, J., Koda, T., Nishimura, S., Takeshima, H.,Nishimura, T.: IFN- $\gamma$ -induced SOCS-1 regulates STAT6-dependent eotaxin production triggered by IL-4 and TNF- $\alpha$ . *Biochem Biophys Res Commun*, 314: 468-475, 2004.
- 122.Hoshino, A., Tsuji, T., Matsuzaki, J., Jinushi, T., Ashino, S., Teramura, T., Chamoto, K., Tanaka, Y., Asakura, Y., Sakurai, T., Mita, Y., Takaoka, A., Nakaike, S., Takeshima, T., Ikeda, H.,Nishimura, T.: STAT6-mediated signaling in Th2-dependent allergic asthma: critical role for the development of eosinophilia, airway hyper-responsiveness and mucus hypersecretion, distinct from its role in Th2 differentiation. *Int Immunol*, 16: 1497-1505, 2004.
- 123.Gyobu, H., Tsuji, T., Suzuki, Y., Ohkuri, T., Chamoto, K., Kuroki, M., Miyoshi, H., Kawarada, Y., Katoh, H., Takeshima, T.,Nishimura, T.: Generation and targeting of human tumor-specific Tc1 and Th1 cells transduced with a lentivirus containing a chimeric immunoglobulin T-cell receptor. *Cancer Res*, 64: 1490-1495, 2004.
- 124.Murashima A, Fukazawa T, Hirashima M, Takasaki Y, Oonishi M, Niijima S, Yamashiro Y, Yamataka A, Miyano T, Hashimoto H. Long term prognosis of children born to lupus patients. *Ann rheum dis* 63(1): 50-53, 2004
- 125.Hirashima M, Fukazawa T,Abe K, Morita Y, Kusaoi M, Hashimoto H. Expression and activity analyses of CTLA4 in peripheralblood lymphocytes in systemic lupus erythematosus. *Lupus* 13, 24-31, 2004
- 126.Yamada M, Yagita H, Inoue H, Takanashi T, Matsuda H, Munechika E, Kanamaru Y, Shirato I, Tomino Y, Matushima K, Okumura K, Hashimoto H. Selective accumulation of CCR4+ T lymphocytes into renal tissue of patients with lupus nephritis. *Arthritis Rheum* 46(3), 735-740, 2003
- 127.Tokano Y, Ogasawara H, Ando S, Fujii T, Kaneko H, Tamura N, Yano T, Hirokawa K, Fukazawa T, Murashima A, Kobayashi S, Sekigawa I, Takasaki Y, Iida N, Hashimoto H. Cyclosporin A therapy for interstitial pneumonitis associated with rheumaticdisease. *ModRheumatol* 12, 305-310, 2003
- 128.Sugimoto K, Morimoto S, Kaneko H, Nozawa K, Tokano Y, Takasaki Y, Hashimoto H. Decreased IL-4 producing CD4 $^{+}$  T cells in patients with active systemic lupus erythematosus-relation to IL-12R expression. *Autoimmunity* 35(6): 381-387, 2002
- 129.Sugiur T et al: Increased IL-15 production of muscle cells in polymyositis and dermatomyositis. *Int Immunol* 14 (8), 917-924, 2002
- 130.Hirakata M, Suwa A, Kuwana M, Sato S, Mimori T, Hardin JA. Anti-Ku autoantibodies are associated with the DPB1 gene. *Arthritis Rheum.* (in press)
- 131.Miyachi K, Hirano Y, Horigome T, Mimori T, Miyakawa H, Onozuka Y, Shibata M, Hirakata M, Suwa A, Hosaka H, Matsushima

- S, Komatsu T, Matsushima H, Hankins RW. Autoantibodies from primary biliary cirrhosis patients with anti-p95c antibodies bind to recombinant p97/vcp and inhibit in vitro nuclear envelope assembly. *Clin Exp Immunol* 136:568-573, 2004
132. Fujii T, Mimori T, Kimura N, Hirakata M. Pseudoscleroderma associated with transforming growth factor beta1-producing advanced gastric carcinoma: comment on the article by Varga. *Arthritis Rheum.* 48(6):1766-1767, 2003
133. Yasuoka H, Ihn H, T.A. Medsger TA, Hirakata M, Kawakami Y, Ikeda Y, Kuwana M. A novel testicular protein highly expressed in testis is overexpressed in systemic sclerosis fibroblasts and targeted by autoantibodies. *J. Immunol.* 171:6883-6890, 2003
134. Kuwana M, Kimura K, Hirakata M, Kawakami Y, Ikeda Y. Differences in autoantibody response to Th/To between systemic sclerosis and other autoimmune diseases. *Ann. Rheum. Dis.* 61(9):842-846, 2002
135. Kuwana M, Matsuura E, Kobayashi K, Okazaki Y, Kaburaki J, Ikeda Y, Kawakami Y. Binding of  $\beta$ 2-glycoprotein I to anionic phospholipids facilitates processing and presentation of a cryptic epitope that activates pathogenic autoreactive T cells. *Blood* (in press)
136. Yasuda S, Atsumi T, Ieko M, Natsuura E, Inagaki J, Kato H, Tanaka H, Yamakado M, Akino M, Saito H, Amasaki Y, Jodo S, Amengual O, Koike T. Nicked  $\beta$ 2-glycoprotein I: A marker of cerebral infarct and role in the negative feedback pathway of extrinsic fibrinolysis. *Blood* 103: 3766-3772, 2004.
137. Kobayashi K, Kishi M, Atsumi T, Bertolaccini ML, Makino H, Sakairi N, Yamamoto I, Yasuda T, Khamashta MA, Hughes GRV, Koike T, Voelker DR, Matsuura E. Circulating oxidized low density lipoprotein forms complexes with  $\beta$ 2-glycoprotein I: implication as an atherogenic autoantigen. *J Lipid Res* 44: 716-726, 2003.
138. Born WK, Vollmer M, Reardon C, Matsuura E, Voelker DR, Giclas PC, O'Brien R.L. Hybridomas expressing  $\gamma\delta$  T-cell receptors respond to cardiolipin and  $\beta$ 2-glycoprotein I (apolipoprotein H). *Scand J Immunol* 58: 374-381, 2003
139. Liu Q, Kobayashi K, Furukawa J, Inagaki J, Sakairi N, Iwado A, Yasuda T, Koike T, Voelker DR, Matsuura E.  $\omega$ -Carboxyl variants of 7-ketocholesteryl esters are ligands for  $\beta$ 2-glycoprotein I and mediate antibody-dependent uptake of oxidized LDL by macrophages. *J Lipid Res* 43: 1486-1495, 2002.
140. Kageyama A, Torikoe K, Iwamoto M, Masuyama J, Shibuya Y, Okazaki H, Yazawa K, Minota S. Nocardia arthritidis sp.nov., a New Pathogen Isolated from a Patient with Rheumatoid Arthritis in Japan. *J Clin Microbiol* 42: 2366-2371, 2004.
141. Horie S, Nakada K, Minota S, Kano S. High proliferative potential colony-forming cells (HPP-CFCs) in the peripheral blood of rheumatoid arthritis patients with interstitial lung disease. *Scand J Rheumatol* 32: 273-276, 2003.
142. Yokota S, Kubota H, Matsuoka Y, Naitoh M, Hirata D, Minota S, Takahashi H, Fujii N, Nagata K. Prevalence of HSP47 antigen and autoantibodies to HSP47 in the sera of patients with mixed connective tissue disease. *Biochem Biophys Res Commun.* 303:413-418, 2003.
143. Okazaki H, Hirata D, Kamimura T, Sato H, Iwamoto M, Yoshio T, Masuyama J, Fujimura A, Kobayashi E, Kano S, Minota S. Effects of FTY720 in MRL-lpr/lpr mice for a therapeutic potential on systemic lupus erythematosus. *J Rheumatol.* 29:707-716, 2002.
144. Okazaki H, Kakurai M, Hirata D, Sato H, Kamimura T, Onai N, Matsushima K, Nakagawa H, Kano S, Minota S. Characterization of chemokine receptor expression and cytokine production in circulating CD4+ T cells from patients with atopic dermatitis: Up-regulation of C-C Chemokine Receptor 4 in atopic dermatitis. *Clin Exp Allergy.* 32:1236-1242, 2002.
145. Sato K, Sato U, Tateishi S, Kubo K, Horikawa R, Mimura T, Yamamoto K, Kanda H. Aire downregulates multiple molecules that have contradicting immune-enhancing and immune-suppressive functions. *Biochem Biophys Res Commun.* 2004, 318(4): 935-40
146. Hasegawa M, Yamazaki M et al. Autoantibodies against phosphatidylserine-prothrombin complex ion patients with systemic sclerosis. *Ann Rheum Dis* 2004, 63:

- 1514-1517, 2004
147. 小池 隆夫、渥美 達也:「抗リン脂質抗体症候群－最近の話題」、『Molecular Medicine: 臨時増刊号』、40: 328-335、2004
148. 坊垣 幸、小池 隆夫:「全身性エリテマトーデス」、『月刊 臨牀と研究』、81:2, 234-240、2004
149. 片岡 浩、市川 健司、小池 隆夫:「抗リン脂質抗体症候群」、『NEW MOOK 整形外科(金原出版株式会社)』、14: 71-176、2004
150. 渥美 達也、家子 正裕、保田 晋助、高橋 郁恵、小池 隆夫:「抗リン脂質抗体症候群の診断におけるホスファチジルセリン依存症プロトロンビン抗体の意義」、『医学と薬学』、52:4, 629-636、2004
151. 渡部 一郎、向井 正也、小池 隆夫:「関節リウマチ専門医からみた薬物療法のあり方」、『Pharma Media(メディカルレビュー社)』、22:12, 88-97、2004
152. 古川 真、渥美 達也、小池 隆夫:「抗リン脂質抗体関連血小板減少症」、『日本臨床免疫学会会誌』、26:5, 267-273、2003
153. 近 祐次郎、渥美達也、小池隆夫:「抗リン脂質抗体症候群の予防法と治療法」『Medical Practice』40, 625-8, 2003
154. 古川 真、小池隆夫:「抗リン脂質抗体症候群」『ESSENCE 膜原病・リウマチ -診断へのアプローチ-』診断と治療社、64-65、2003
155. 古川 真、小池隆夫:「抗リン脂質抗体症候群」『別冊・医学のあゆみ 免疫疾患-state of arts ver.2』医歯薬出版株式会社、460-465
156. 坊垣暁之、天崎吉晴、澤田賢一、小池隆夫:「幹細胞移植と自己免疫疾患」『メディカルサイエンスダイジェスト』28、No.10、4-7、2003
157. 坊垣暁之、天崎吉晴、小池隆夫:「自己免疫疾患の治療としての造血幹細胞移植」『臨床免疫』38、No.4、440-445、2003
158. 坊垣暁之、渥美達也、小池隆夫:「リウマチ性疾患に対する末梢血幹細胞移植－日本における現状と今後の展望－」『リウマチ科』28、No.4、382-387、2003
159. 中林 透、小池 隆夫:「膜原病の症状からみた診断へのアプローチ」、『日本内科学会雑誌』、92:10, 1904-1910、2003
160. 小池 隆夫:「抗リン脂質抗体症候群」:内科学 第八版(朝倉書店)』、1255-1257、2003
161. 萩原宏美、渥美達也、小池隆夫:「H アレルギー・膜原病、全身性エリテマトーデス」、『高久史磨総監修:外来診療のすべて 改訂第3版』、メディカルビュー社、東京、
- 888-9、2003
162. 坊垣暁之、天崎吉晴、澤田賢一、小池隆夫:「自己免疫疾患に対する造血幹細胞移植療法」『小澤 敬也編著:造血幹細胞移植基礎から遺伝子治療・再生医療へ』197-203、2003
163. 永井 聰、吉岡 成人、小池 隆夫:「女性における、砂糖の摂取と2型糖尿病のリスクに関する前向き研究」、『プラクティス』、20:6, 691-697、2003
164. 古川 真、渥美 達也、小池 隆夫:「抗リン脂質抗体関連血小板減少症」、『日本臨床免疫学会会誌』、26:5, 267-273、2003
165. 近 祐次郎、渥美達也、小池隆夫:「抗リン脂質抗体症候群の予防法と治療法」『Medical Practice』40, 625-8、2003 小池 隆夫:「抗リン脂質抗体症候群」、日本内科学会雑誌 創立 100 周年記念号、91:9、90-93、2002
166. 小池隆夫、渥美達也:「12. 抗リン脂質抗体症候群」、『日本リウマチ財団教育研修委員会:リウマチ基本テキスト』、東京、329-36、2002
167. 藤咲 淳、小池 隆夫:「全身性エリテマトーデス」、日常外来診療ハンドブック(メディカルトリューン社)、656-663、2002
168. 斎藤 幸、小池 隆夫:「膠原病と類縁疾患により腎障害(多発動脈炎、Wegener 肉芽腫症、Churg-Strauss 症候群)」、『下条 文武、内山 聖、富野康日己 編:専門医のための腎臓病学(医学書院)』、288-300、2002
169. 家子 正裕、小池 隆夫:「抗リン脂質抗体症候群」、『櫻川 信男、青崎正彦、上塙芳郎、和田英夫編著:抗凝固薬の適正な使い方(医歯薬出版株式会社)』、197-203、2002
170. 小池 隆夫:「抗リン脂質抗体症候群」、『日本内科学会雑誌 創立 100 周年記念号』、91:9、90-93、2002
171. 小池 隆夫:「1型糖尿病の発症と免疫異常」、『総合臨床』、51:7, 2195-2200、2002
172. 小池 隆夫:「モノクローナル抗体療法の有効性と問題点」、『炎症と免疫』、10:5, 39-41、2002
173. 遠藤 知之、小池 隆夫:「悪性リンパ腫と抗CD20 抗体」、『炎症と免疫』、10:5, 62-68、2002
174. 深江 淳、渥美 達也、小池 隆夫:「抗リン脂質抗体症候群の解析の進歩」、『アレルギー・免疫』、9:9, 62-28、2002
175. 古崎 章、渥美達也、小池隆夫:「大動脈炎症候群(高安動脈炎)の免疫抑制剤の使い方」『Heart View - 特集:大動脈炎症候群をどう診るか』6, 482-5、2002

- 176.酒井良江、渥美達也、小池隆夫:「抗リン脂質抗体症候群に伴う弁膜症」『Cardiac Practice - 特集:弁膜症-最近の動向』13、81-86, 2002
- 177.渥美達也:「抗リン脂質抗体症候群における血栓症の分子機構とその対策」『内科 - 特集:変貌する膠原病治療-最新治療の有用性と問題点』89、210-5, 2002
- 178.古川 真、渥美達也、小池隆夫:「抗リン脂質抗体症候群の合併症」『総合臨床 - 特集:膠原病と合併症』51、2195-200, 2002
- 179.酒井良江、渥美達也、小池隆夫:「抗リン脂質抗体症候群」『臨床医増刊号 - 検査値から読む病態と診断計画』28、1559-61, 2002
- 180.古川 真、渥美達也、小池隆夫:「抗リン脂質抗体の合併症」『総合臨床 - 膠原病の難治性合併症』51、2195-200, 2002
- 181.堀田 哲也、小池 隆夫:「アナフィラキシーショック」、臨床と研究, 79:2229, 229-232, 2002
- 182.天崎 吉晴、小池 隆夫:「「末梢血幹細胞移植と難治性自己免疫疾患」治療成績の実際と今後の展望」、炎症と免疫, 10:2, 135 - 144, 2002
- 183.向井正也・小池隆夫:「抗リン脂質抗体症候群」、60:1, 404- 409, 2002
- 184.小池 隆夫:「抗リン脂質抗体症候群」、日本内科学会雑誌 創立 100 周年記念号、9 1:9, 90-93, 2002
- 185.中山 玲慧、横山利幸、平塚義宗、中村淳夫、広瀬幸子、白井俊一:全身性エリテマトーデスにおける網脈絡膜症の発症機構の検討-(NZB x NZW)F1マウスを用いた解析-, 順天堂医学, 47:519-527, 2002
- 186.富田英明、三浦右子、中村和裕、津田裕士、橋本博史、広瀬幸子:C1q 遺伝子多型が C1q 産生ならびにループス腎炎におよぼす影響. 順天堂医学, 48:355-363, 2002
- 187.三浦浩二、修 岩、鶴井博理、高橋和子、張 丹青、広瀬幸子:SLE マウスにおける脾臓リンパ濾胞胚中心の免疫組織化学的解析. 順天堂医学, 48:384-395, 2002
- 188.赤倉玲子、文 香淑、阿部雅明、姜 奕、広瀬幸子:Fas 遺伝子多型による Fas 発現制御と自己免疫疾患. 順天堂医学, 48:364-374, 2002
- 189.趙京元、張丹青、黒澤尚、広瀬幸子:コラーゲン誘導関節炎の発症に及ぼす抗体遺伝子の影響 順天堂医学, 2004 印刷中
- 190.松下 祥:HLA による免疫応答の制御。ゲノム医学 4, 453- 458, 2004
- 191.松下 祥:Th2 応答とアジュバント。感染・炎症・免疫 34:192- 198, 2004
- 192.松下 祥 MHC クラス II 分子を介したシグナル伝達機構。臨床免疫 42:455- 463, 2004
- 193.伊藤 聰:難治性病態の治療戦略. ループス腎炎 内科 93:283-287, 2004
- 194.川上純、玉井慎美、中村英樹、江口勝美他. NF--B、MAPK 制御による抗炎症療法. 別冊・医学のあゆみ サイトカイン p386-390, 2004.
- 195.折口智樹、川上純、江口勝美、シェーグレン症候群とウイルス. シェーグレン症候群への Strategy 5: 22-29, 2004.
- 196.江口勝美. HTLV-I 感染トリウマチ性疾患. リウマチ科 27: 238-246, 2002.
- 197.江口勝美. アポトーシス抑制因子. リウマチ科 28: 184-195, 2002.
- 198.川上純、中島コト、江口勝美他. アポトーシス誘導療法. 内科 89: 309-312, 2002.
- 199.岡崎拓 PD-1 と自己免疫疾患。免疫 2005 41:327-333. 2004
- 200.岡崎拓、本庶佑 PD-1 と自己免疫疾患。最新医学 59(7): 1668-1672, 2004
- 201.岡崎拓、本庶佑 PD-1 欠損マウスと免疫異常。Annual Review 免疫 298-305. 2002
- 202.桑名正隆: 血小板減少症と抗リン脂質抗体、抗血小板抗体. リウマチ科 2002; 28(4):340-347.
- 203.鎌木淳一、桑名正隆、他: 膜原病における血栓症-抗リン脂質抗体症候群における血清中自己抗体の多様性. 臨床血液 2002; 43(6):454-458.
- 204.桑名正隆: 抗リン脂質抗体症候群. Medical Science Digest 2003; 29(1):4-5.
- 205.鎌木淳一、桑名正隆、他: 抗リン脂質抗体. 臨床病理 2003; 51(7):639-643.
- 206.桑名正隆: 抗リン脂質抗体の測定法. 検査と技術 2004; 32(1):11-16.
- 207.佐々木毅:自己免疫疾患治療の最近の進歩 日本医事新報 4100, 2002
- 208.佐々木毅:LE 細胞現象 日本内科学会雑誌 91: 41-45, 2002
- 209.佐々木毅:全身性エリテマトーデス 南江堂 93: 242-247, 2004
- 210.佐々木毅:抗 U1-RNP 抗体 Modern Physician 24: 905, 2004
- 211.奥田光崇、野村順、堀内高広、宮村耕一、佐々木毅:治療関連急性骨髓性白血病に対する HLA 一致同胞からの骨髓非破壊的同種造血幹細胞移植 臨床血液 45: 308-311, 2004
- 212.小川法良、菅井進 シェーグレン症候群の診療 リウマチ科 27(supple.1) :691-697, 2002.
- 213.小川法良 シェーグレン症候群の治療 内科 89(2) :288-293, 2002.

214. 小川法良、菅井進 膜原病と合併症、シェーグレン症候群。総合臨床 51(7):2149-2154, 2002.
215. 小川法良 シェーグレン症候群とケモカイン 臨床免疫 38:141-146, 2002.
216. 小川法良 病態とアボトーシス、シェーグレン症候群。臨床免疫 38 増刊、アボトーシスのすべて: 380-385, 2002.
217. 辻村静代、齊藤和義、河野公俊、田中良哉. SLE における多剤抵抗性遺伝子発現とその制御。臨床免疫(2004)42, 442-447
218. 田中良哉、辻村静代. P-糖蛋白質に対するシクロスボリンの作用:薬剤抵抗性の克服。医薬ジャーナル(2004)40, 178-182
219. 辻村静代、田中良哉. 多剤耐性遺伝子を標的とする新規治療。リウマチ科(2004)31, 55-61
220. 田中良哉. シクロスボリン療法の理論と実際。日本臨床(2002) 60, 2345-2350
221. 田中良哉、辻村静代、齊藤和義、河野公俊. 膜原病・リウマチ性疾患に於けるシクロスボリン療法の理論と実際。日本臨床免疫学会雑誌(2002) 25, 110-114
222. 土屋尚之 :全身性エリテマトーデス、関節リウマチ疾患感受性遺伝子研究の現状。医学のあゆみ 202: 815-820. 2002
223. 土屋尚之:SLE の疾患感受性遺伝子解析。臨床リウマチ 15: 87-101. 2003
224. 土屋尚之:ゲノム解析によるリウマチ性疾患の病因・病態解明へのアプローチ。日本臨床免疫学会誌 26: 103-114. 2003
225. 土屋尚之、京極千恵子:SLE 疾患感受性とFc・受容体 IIb 多型の関連。臨床免疫 42:435-441. 2004
226. 土屋尚之:免疫疾患の疾患感受性遺伝子。シミュレーション内科「リウマチ・アレルギー疾患を探る」(山本一彦編) 永井書店、大阪、(印刷中)
227. 許 志泉、戸叶嘉明、松平 蘭、金井美紀、竹内 健、高崎芳成、橋本博史: シェーグレン症候群における中国の診断基準の検討-感度と特異度について他の診断基準との比較。順天堂医学 48(4), 495-504, 2003
228. 許 志泉、竹内 健、松平 蘭、金井美紀、戸叶嘉明、高崎芳成、橋本博史: シェーグレン症候群における抗 SS-A/B 抗体の二重免疫拡散法、ウェスタンプロット法の比較検討。日臨免会誌 26(2), 74-79, 2003
229. 原まさ子. 膜原病におけるエンドキサンバルス療法の効用 日本臨床免疫学会会誌 25(1)115-120,2002.
230. 平形道人: 抗KS(アスパラギニルtRNA合成酵素)抗体。リウマチ科 27(6):591-597, 2002
231. 佐藤慎二、平形道人 :Amyopathic dermatomyositis に特異的に出現する自己抗体。臨床免疫 41(3):318-322, 2004
232. 平形道人: 抗アミノアシル tRNA 合成酵素抗体と多発性筋炎。炎症と免疫 12(3):308-316, 2004
233. 佐藤慎二、平形道人: Amyopathic dermatomyositis に特異的に出現する自己抗体-抗 US 抗体の臨床的意義を中心に-。リウマチ科 31(6):596-599, 2004
234. 平形道人: 筋炎患者血清に見出される特異自己抗体および関連自己抗体。Clinical Neuroscience 22 (10):1155-1157, 2004
235. 松浦栄次、小林和子、小池隆夫. 自己免疫と動脈硬化。日本アフェレシス学会学会誌、23: 167-175. 2004
236. 松浦栄次、小林和子. 酸化 LDL 研究の現状。アニムス、アニムス刊行会、東京. pp21-27, 2003.
237. 稲垣純子、青木耕治、松浦栄次. 子宮内膜症と抗ラミニン-1 抗体。臨床免疫 39: 575-581, 2003.
238. 松浦栄次. 抗リン脂質抗体症候群。岡山医学雑誌、114:19-26, 2002.
239. 小林和子、近藤朱音、松浦栄次. 脂質の酸化と生理活性。血小板と生理活性脂質、尾崎由基夫、池田康夫、島田和幸、高井義美、丸山征郎編、金芳堂、pp147-160, 2002
240. 河島洋平、山・雅英、ほか。「異なる疾患群における IgG 型抗プロトロンビン抗体測定キット2 社間の測定値の乖離」。日本血栓止血学会雑誌、印刷中。
241. 田中千洋、山・雅英、ほか。「広汎な分枝状皮斑と右正中神経栄養動脈閉塞を伴い、免疫吸着療法が奏効した原発性抗リン脂質抗体症候群の 1 例」。日本皮膚科学会雑誌, 112(12): 1611-1616, 2002

## 2. 学会発表

1. Koike,T. :" Antiphospholipid Syndrome" 7<sup>rd</sup> International Congress SLE and Related conditions , New York,U.S.A.,May 9-13, 2004
2. Koike,T. :" Antiphospholipid Syndrome,mechanism of thrombus and complication of reproductive system" 11<sup>rd</sup> European Congress on Reproductive Immunology, Pilzen,Czech ,June 30-July 3, 2004
3. Koike,T. :" Antiphospholipid Syndrome " 11<sup>rd</sup> Asia Pacific League Associations for Rheumatology Congress, Jeju, Korea ,September 11- September 15, 2004
4. Atsumi,T., Koike,T. :" Antiprothrombin-is it worth assaying? " 11<sup>rd</sup> International Congress on Antiphospholipid Antibodies, Sydney,

- Australia,12-19,November ,2004
5. Koike,T. :" Pathogenesis of antiprothrombin antibody " 4<sup>rd</sup> International Congress onAutoimmunity, Budapest, Hungary,3-7,November ,2004
  6. Tatsuya Atsumi, Masahiro Ieko, Satoshi Jodo, Yoshiharu Amasaki, Takao Koike: Sensitivity of semiquantitative lupus anticoagulant test for the diagnosis of antiphospholipid syndrome, 19th Congress of The International Society on Thrombosis and Haemostasis, Birmingham, UK, July 12-18, 2003
  7. Shinsuke Yasuda, Guang W. Wong, Steven A Krilis, Richard L. Stevens: Activation of Urokinase-type Plasminogen Activator (uPA) by Human Tryptase Epsilon/PRSS22, American association of immunology, Denver, CO, May 6-10, 2003
  8. Guang W. Wong, Yi Yanf, Shinsuke Yasuda, Richard L Stevens, Lixin Li. Mouse Mast cells (MCs) Express the Tryptic-like Serine Protease Neuropsin/KLK8/Prss19, American association of immunology, Denver, CO, May 6-10, 2003
  9. Toshiyuki Bohgaki, Yoshiharu Amasaki, Miyuki Bohgaki, Akira Furusaki, Yoshie Sakai, Shin Furukawa, Jun Fukae, Kazuko Matsumura, Tatsuya Atsumi, Takao Koike. Hematopoietic stem cell transplantation for three patients with systemic sclerosis. The 5th Korea-Japan combined meeting of rheumatology (KJCMR), Seoul, Korea, May 22, 2003
  10. Koike,T. :" Anticardiolipin and anti-  $\beta$ 2-glycoprotein I antibodies:application in the clinic" 3<sup>rd</sup> International Congress on Autoimmunity, Geneva, Switzerland, February, 20~24, 2002.
  11. Koike,T. :" Antiphospholipid antibodies; in arterial thrombosis." 1<sup>st</sup> Tuzing Antiphospholipid, Conference, Tuzing, Germany, April , 22~25, 2002.
  12. Koike,T. :" Antiphospholipid antibodies; recent progress and future prospect ". SLE Respons Criteria Meeting, Dusseldorf, Germany, May , 9~11, 2002.
  13. Koike,T. :" Anti-prothronbin antibodies; pathojenesis and specificity." 6<sup>th</sup> Dresden symposium on Autoaintibodies, Dresden, Germany, September , 4~7, 2002.Kameda H,
  14. Furukawa S, Atsumi T, Bohgaki T, Fukae J, Amasaki Y, Ichikawa K, Koike T. Clinical significance of serum KL-6 as a marker of intestinal pneumonitis complicated with connective tissue diseases. 26th International Congress of Internal Medicine. Kyoto, Japan, 26-30 May 2002
  15. Oku K, Atsumi T, Furukawa S, Sakai Y, Furusaki A, Horita T, Jodo S, Amasaki Y, Ichikawa K, Koike T. Significance of brain imaging in patients with systemic lupus erythematosus complicated with central nerve system manifestations. 26th International Congress of Internal Medicine. Kyoto, Japan, 26-30 May 2002
  16. Amengual O, Atsumi T, Komano Y, Jodo S, Amasaki Y, Ichikawa K, Koike T. Glycoprotein IIIA polymorphism HPA6Wa/b and aPL are independent risks for thrombocytopenia in patients with SLE. 10th International Congress on Antiphospholipid Antibodies. Taormina, Italy, 29 September - 3 October, 2002
  17. Sakai Y, Atsumi T, Furukawa S, Ieko M, Jodo S, Amasaki Y, Ichikawa K, Koike T. Associaticn of aPL and valvular heart diseases in Japanes patients with autoimmune diseases. 10th International Congress on Antiphospholipid Antibodies. Taormina, Italy, 29 September - 3 October, 2002
  18. Furukawa S, Atsumi T, Ieko M, Sakai Y, Jodo S, Amasaki Y, Ichikawa K, Koike T. Antiphospholipid associated thrombocytopenia: As a risk for thrombosis and pregnancy morbidity. 10th International Congress on Antiphospholipid Antibodies. Taormina, Italy, 29 September - 3 October, 2002
  19. Atsumi T, Ieko M, Ichikawa K, Horita T, Jodo S, Amasaki Y, Koike T. A simple semi-quantitative method for lupus anticoagulant activity using phosphatidylserine dependent monoclonal antiprothrombin antibody. 10th International Congress on Antiphospholipid Antibodies. Taormina, Italy, 29 September - 3 October, 2002
  20. Ichikawa K, Horita T, Atsumi T, Jodo S, Amasaki Y, Koike T. Monoclonal antibodies against the complex of phosphatidylserine and prothrombin from patients with APS. 10th International Congress on Antiphospholipid Antibodies. Taormina, Italy, 29 September - 3 October, 2002
  21. Atsumi T, Ieko M, Ichikawa K, Horita T, Jodo S, Amasaki Y, Koike T. High prevalence of lupus anticoagulant detected by a novel semi-quantitative assay using phosphatidylserine dependent monoclonal antiprothrombin antibody in patients with

- antiphospholipid syndrome. American College of Rheumatology 66th National Scientific Meeting, New Orleans, USA. 24-29 October 2002
22. Bertolaccini ML, Atsumi T, Khamashta MA, Koike T, Hughes GRV. Antiprothrombin antibodies: Detection, immunological characteristics and clinical significance in patients with systemic lupus erythematosus. American College of Rheumatology 66th National Scientific Meeting, New Orleans, USA. 24-29 October 2002
  23. Amengual O, Atsumi T, Komano Y, Jodo S, Amasaki Y, Ichikawa K, Koike T. The human platelet antigen 6Wb represents a risk factor for thrombosis in patients with systemic lupus erythematosus. American College of Rheumatology 66th National Scientific Meeting, New Orleans, USA. 24-29 October 2002
  24. Tincani, M. Taglietti, M. Spunghi, M. Cinquini, F. Allegri, G. Balestrieri, G. Reber, T. Koike, K. Ichikawa, P. Meroni, M.C. Boffa. (aCL) and a- $\square$ 2GPI elisas: the state of the art in 2002. 10<sup>th</sup> International Congress on Antiphospholipid Antibodies, Taormina, Italy, October 3, 2002
  25. Y. Yamashita, Y. Amasaki, T. Yamashita, K. Ichikawa, T. Atsumi and T. Koike. ILT2/LIR1 inhibitory receptor, a new marker for cytotoxic CD8+ T cells, regulates cytotoxic T cell function. Experimental Biology 2002 , April, 21, 2002
  26. Ishigami H, Abe T, Takeuchi T: Expression of adapter proteins in rheumatoid synovial fibroblast-like cells and their involvements in the signaling from growth factor receptors. 67th Annual Meeting, Orlando, U.S.A., October, 2003
  27. Tsuzaka K, Fukuhara I, Setoyama Y, Yoshimoto K, Suzuki K, Abe T, Takeuchi T. A splice variant of the TCR $\zeta$  mRNA lacking exon 7 can lead to the down-regulation of TCR $\zeta$  protein and IL-2 production in SLE T cells. American College of Rheumatology, 67th Annual Meeting, Orlando, U.S.A., October, 2003
  28. Tsubata, T.: CD40 signaling perturbs peripheral deletion of self-reactive B cells. Workshop on the Genetics and Mechanisms of Autoimmunity, Princeton, March 21-23, 2003.
  29. Tsubata, T.: Immunoglobulin isotype-specific regulation of B cell antigen receptor signaling. Joint Meeting of STS and FZI, Weimar, November 6-8, 2003.
  30. Tsubata, T.: Immunoglobulin isotype-specific regulation of B cell antigen receptor signaling, Keystone Symposia, Steamboat Springs, January 8-13, 2004.
  31. Adachi, T., Hokazono, Y., Matthias Wabl, Tsubata, T.: Inhibitory co-receptors activated by antigens but not by Anti-immunoglobulin heavy chain antibodies Install requirement of co-stimulation through CD40 for survival and proliferation of B cells. Keystone Symposia, Steamboat Springs, January 8-13, 2004.
  32. Nishimoto N, et al. : 68th National Annual Scientific Meeting of American College of Rheumatology: Repeated treatment with anti-IL-6 receptor antibody (MRA) lead to extended clinical response rheumatoid arthritis even after cessation of MRA. San Antonio, USA 2004.10.16-22
  33. Nishimoto N, et al. : 67th National Annual Scientific Meeting of American College of Rheumatology: Long-term safety and efficacy of anti-interleukin 6 receptor antibody (MRA; Atlizumab) in patients with rheumatoid arthritis. Orland, USA 2003.10.23-28
  34. Nishimoto N, et al. : American Society of Hematology 45th ASH Annual Meeting and Exposition: The long-term safety and efficacy of humanized anti-interleukin-6 receptor monoclonal antibody, MRA in multicentric castleman's disease. San Diego, USA 2003.12.5-9
  35. Nishimoto N, et al. : 66th National Annual Scientific Meeting of American College of Rheumatology: A Multi-Center, Randomized, Double-Blind, Placebo-Controlled Trial of Humanized Anti-interleukin-6 (IL-6) Receptor Monoclonal Antibody (MRA) in Rheumatoid Arthritis (RA). New Orleans, USA.2002.10.25-30
  36. Nishimoto N, et al. : American Society of Hematology 44th ASH Annual Meeting and Exposition: Improvement of Wasting by Treatment with a Humanized
  37. Liu, T-Y., Kohsaka, H., Uemura, Y., Ohyama, H. and Matsushita, S. Positional effect of amino acid replacement on peptide antigens for the enhancement of IFN $\gamma$  production. In 60th AAAAI annual meeting, San Francisco, USA. 2004
  38. Matsushita, S., Liu, T-Y., Ohyama, H. and Uemura, Y. Signal transduction through HLA-DQ molecules alter dendritic cell function to enhance Th2 differentiation. In 23rd EAACI. Amsterdam. 2004

39. Ohyama, H., Takeuchi, K., Ogata, K., Namisato, Y., Fukutomi, Y. and Matsushita, S. The polymorphism on the 5' flanking region of IL-12 receptor  $\beta$ 2 gene confers the susceptibility to mycobacteria infection. In 12th ICI. Montrial, USA.2004
40. Hayase H, Ishizu A, Higuchi M, Abe A, Tsuji M, Ikeda H, Yoshiki T. Comparative characterization of CD25+ CD4+ T cells between HTLV-I transgenic and wild type rats. 11th International Conference on Human Retrovirology: HTLV and Related Viruses, San Francisco. AIDS Res Hum Retroviruses 19: S38-39, 2003.
41. Ikeda H, Abe A, Ishizu A, Hayase H, Yoshiki T. A transgenic rat model of HTLV-I induced immunological diseases: effect of the env-pX transgene to effector cells and target tissues in collagen induced arthritis. 11th International Conference on Human Retrovirology: HTLV and Related Viruses, San Francisco. AIDS Res Hum Retroviruses 19: S68, 2003.
42. Yamamoto K, Kobayashi T, Ito S, et al. Fcy receptor polymorphisms in systemic lupus erythematosus and chronic peritonitis. International Association for Dental Research/American Association for Dental Research 80th General Session San Diego, USA 2002
43. Kawakami A, Nakashima K, Tamai M, Nakamura H, Eguchi K, et al. Toll-like receptor in Sjögren's syndrome. 11<sup>th</sup> Asia pacific league of association for rheumatology congress, 2004.
44. Kawakami A, Nakashima K, Eguchi K, et al. Immunohistochemical detection of apoptosis-related molecules of the salivary glands in patients with Sjögren's syndrome. 第8回国際シェーグレン症候群シンポジウム, 2002.
45. Okazaki, T. et al., Role of PD-1 in immune tolerance and tumor surveillance. Workshop on molecular and genetic basis of autoimmune diseases: SLE and RA. Lisbon, 2003 Apr
46. Xiang Y,Sekine T,Nakamura H,Imajoh-Ohmi S,Fukuda H,Nishioka K,Kato T.Fibulin-4 is a Component of Cartilage and a Target of Autoantibodies in Osteoarthritis and Rheumatoid Arthritis. American College of Rheumatology, 67th Annual Scientific Meeting.
47. Xiang Y,Sekine T,Nakamura H,Imajoh-Ohmi S,Fukuda H,Nishioka K,Kato T.Proteomic Surveillance of Autoimmunity in Osteoarthritis: Identification of Triosephosphate Isomerase as an Autoantigen in Patients with Osteoarthritis. American College of Rheumatology, 67th Annual Scientific Meeting.
48. Karasawa R,Ooka S,Sekine T,Nishimura H,Nukina N,Mitsui K,Ozaki S,Nishioka K,Kato T.Targets of anti-endothelial cell antibodies in patients with systemic vasculitis: identification by the proteomic approach. American College of Rheumatology, 68th Annual Scientific Meeting.
49. Kato T,Nakamura H,Masuko-Hongo K,Yudoh K,Nishioka K.Proteome analysis of peripheral B cells in rheumatoid arthritis. American College of Rheumatology, 68th Annual Scientific Meeting.
50. Matsuo K,Nakamura H,Masuko-Hongo K,Yudoh K,Noyori K,Saito T,Nishioka K,Kato T.The phosphoproteome profile of synoviocytes in rheumatoid arthritis is distinct from that in osteoarthritis. American College of Rheumatology, 68th Annual Scientific Meeting.
51. Kuwana M, et al: Restricted T cell receptor  $\beta$ -chain usage by T cells autoreactive to  $\beta_2$ -glycoprotein I in patients with APS. 10<sup>th</sup> International Congress on Antiphospholipid Antibodies. 2002. 9.
52. Yasuoka H, Kuwana M, et al: HLA-B51-restricted T cells responsive to a polymorphic region of MHC class I chain-related gene A (MICA) in Behçet's disease. The 66th Annual Scientific Meeting of American College of Rheumatology New Orleans. 2002. 10.
53. Kuwana M, et al: Binding of  $\beta_2$ -glycoprotein I to anionic phospholipids facilitates processing and presentation of a cryptic epitope that activates pathogenic autoreactive T cells. The 67th Annual Scientific Meeting of American College of Rheumatology. Orlando) 2003. 10.
54. Kuwana M, et al: Impaired function of circulating endothelial precursors in patients with systemic sclerosis: Implication for pathogenesis of microvascular damage. The 67th Annual Scientific Meeting of American College of Rheumatology .Orlando. 2003. 10.
55. Kuwana M:  $\beta_2$ -glycoprotein I : Antiphospholipid syndrome and T-cell reactivity. 11<sup>th</sup> International Congress on Antiphospholipid Antibodies. Sydney. 2004. 11.
56. Ogawa, N., Ping, L., Zhenjun, L., Takada, T., Sugai, S. Involvement of IFN-gamma-induced T cell-attracting chemokines, IP-10 (CXCL10) and Mig (CXCL9), in the salivary gland lesions

- of Sjogren's syndrome. 第8回国際シェーグレン症候群シンポジウム 金沢、2002年5月16-18日 第8回国際シェーグレン症候群シンポジウム抄録集 p10、2002.
57. Ping, L., Ogawa, N., Sugai, S. Study of proliferation and apoptosis in cultured salivary epithelial cells from patients with Sjogren's syndrome. 第8回国際シェーグレン症候群シンポジウム 金沢、2002年5月16-18日 第8回国際シェーグレン症候群シンポジウム抄録集 p35、2002.
58. Ogawa, N., Ping, L., Sugai, S. Involvement of IFN $\gamma$ -induced T cell-attracting chemokines, IP-10 (CXCL10) and Mig (CXCL9), in the salivary gland lesions of Sjogren's syndrome. 10th APLAR, Bangkok, Thailand, 2002年12月1-6日 10th APLAR抄録集 p85、2002.
59. Ogawa, N., Ping, L., Shimoyama, K., Kawanami, T., Sugai, S. Novel role of CD40 in Fas-dependent apoptosis of cultured salivary epithelial cells from patients with Sjogren's syndrome. 68th ACR meeting, San Antonio, Texas, 2004年10月17-21日 Arthritis Rheum. 50:S580, 2004.
60. Ohashi, H., Ohta, Y., Ogawa, N., Saito, M., Dei, Y., Kondo, S., Hayashi, H. The role of CD40 ligand in patients with Sjogren's syndrome. 68th ACR meeting, San Antonio, Texas, 2004年10月17-21日 Arthritis Rheum. 50:S580-S581, 2004
61. Tsujimura S, Saito K, Azuma T, Nakayamada S, Nakano K, Tsukada J, Tanaka Y. Clinical relevance of Characteristically High Expression of P-glycoprotein on Peripheral Lymphocytes to Steroid-Resistance in Refractory SLE Patients The 68<sup>th</sup> National Meeting of American college of Rheumatology, San Antonio. 平成16年10月
62. Tanaka Y, Tokunaga M, Fujii K, Nawata M, Tsujimura S, Nakayamada S, Saito K. A pilot study of rituximab (anti-CD20) for refractory systemic lupus erythematosus: relevance of quantity and quality reduction of B cells to clinical efficacy. The 68<sup>th</sup> National Meeting of American college of Rheumatology, San Antonio. 平成16年10月
63. Kyogoku C, Kono H, Tsuchiya N, Suzuki T, Yamamoto K, Tokunaga K, Honda Z (2003) SLE-associated polymorphism of Fc $\gamma$ RIIB Ile232Thr affects localization at lipid rafts and attenuation of BCR signaling. American College of Rheumatology 2003 Annual Scientific Meeting. (Arthritis Rheum 48 (Suppl); S647.)
64. Chu ZT, Tsuchiya N, Kyogoku C, Qian YP, Xu SB, Mao CZ, Chu JY, Tokunaga K (2003) Association of Fc gamma receptor IIb polymorphism with susceptibility to systemic lupus erythematosus in Chinese: a common susceptibility gene in the Asian populations? American College of Rheumatology 2003 Annual Scientific Meeting. (Arthritis Rheum 48 (Suppl); S380.)
65. Kuroki K, Tsuchiya N, Maenaka K, Rasubala L, Shiroishi M, Yamashita Y, Matsuta K, Fukazawa T, Kohda D, Koike T, Juji T, Hashimoto H, Tokunaga K (2003) Distinct associations of the leukocyte immunoglobulin-like receptor (LIR)1 and LIR6 polymorphisms with susceptibility to rheumatoid arthritis (RA) and systemic lupus erythematosus (SLE). American College of Rheumatology 2003 Annual Scientific Meeting. (Arthritis Rheum 48 (Suppl); S197.)
66. Kawasaki A, Tsuchiya N, Fukazawa T, Matsuta K, Murakami Y, Hashimoto H, Tokunaga K (2004) APRIL (TNFSF13) polymorphisms: independent confirmation of association with susceptibility and new association with clinical characteristics. American College of Rheumatology 2004 Annual Scientific Meeting. (Arthritis Rheum 50(Suppl.); S120, 2004. Arthritis Rheum 50(Suppl.); S203.)
67. Hitomi Y, Tsuchiya N, Kawasaki A, Kyogoku C, Ohashi J, Suzuki T, Fukazawa T, Bejrachanlra S, Chandanayingyong D, Suthipinittharm P, Tsao BP, Hashimoto H, Honda Z, Tokunaga K (2004) Epistatic interaction between CD72 and FCGR2B polymorphisms in conferring susceptibility to human systemic lupus erythematosus (SLE). American College of Rheumatology 2004 Annual Scientific Meeting. (Arthritis Rheum 50(Suppl.); S120.)
68. Matsuzaki, J., et. al. Immuno-steroid as a novel factor for regulation of Th1/Th2 balance: Th2-dependently produced progesterone kill Th1 cells but not Th2 cells. 12<sup>th</sup> ICI/4<sup>th</sup> FOCIS 2004
69. Chamoto, K., et. al. Potentiation of tumor eradication by adoptive immunotherapy with T cell receptor gene-transduced T helper type 1 cells. 12<sup>th</sup> ICI/4<sup>th</sup> FOCIS 2004
70. Nihimura, T., et. al. A novel tumor cells-therapy using T helper type 1 cells: The critical role of dendritic cell type 1/T helper type 1 cell-cell interaction for the initiation of

- antitumor immunity in vivo. 12<sup>th</sup> ICI/4<sup>th</sup> FOCIS 2004
71. Tsuji, T., et. al. Generation and targeting of human tumor-specific Tc1 and Th1 cells transduced with a lentivirus containing a chimeric immunoglobulin T cell receptor. 12<sup>th</sup> ICI/4<sup>th</sup> FOCIS 2004
72. Nishimura, T., et. al. A novel tumor cell therapy using Th1 cells: The critical role of APC/Th1 cell-cell interaction for the initiation of antitumor immunity in vivo. AACR Annual Meeting 2004
73. Kusaoi M, Fukazawa T, Hirashima M, Morita Y, Morita T, Tsuchiya N, Tokunaga K, Inoko H, Hashimoto H. Genomic screening with high density microsatelite markers for systemic lupus erythematosus on chromosome 1. American College of Rheumatology, San Antonio, USA, Oct. 2004
74. Kobayashi S, Tamura N, Akimoto T, Takaya M, Ikeda M, Hashimoto H. Clinical manifestations of Churg-Strauss syndrome (CSS) following treatment with cysteinyl leukotriene type I receptor antagonist. 11th International Vasculitis and ANCA Workshop, Prague Oct 2-5,2003
75. Abe K, Hashimoto H. Pregnancy in systemic lupus erythematosus.67th Annual Scientific Meeting of American College of Rheumatology, Orlando, FloridaOct, 2003
76. Miyashita R, Tsuchiya N, Hikami K, Kuroki K, Fukazawa T, Bijl M, Kallenberg C G M, Hashimoto H Molecular genetic analyses of human NKG2C gene deletion.67th Annual Scientific Meeting of American College of Rheumatology, Orlando, FloridaOct, 2003
77. Tsuchiya N, Kobayashi S, Kawasaki A, Kyogoku C, Arimura Y, Yoshida M, Tokunaga K, Hashimoto H. Genetic background of Japanese patients with ANCA-associated vasculitis: Association of HLA -DRB1 0901 with microscopic polyangiitis. American College of Rheumatology, Louisiana, USA, Oct. 2002
78. Katsumata Y, et al: Attenuation of experimental autoimmune myositis by blocking inducible costimulator (ICOS) - ICOS-ligand interaction. The 68th ACR Annual Scientific Meeting. 2004.
79. Katsumata Y, et al: Immunohistochemical analysis of recombinant-C-protein-induced experimental autoimmune myositis (EAM) in rats. 5th Korea-Japan Combined Meeting of Rheumatology, 2003
80. Hirakata M, Takada T, Suwa A, et al. Histopathological Features of Myopathies in Patients with Anti-SRP Autoantibodies. 68th Annual meeting of American College of Rheumatology, Oct., 2004
81. Hirakata M, Suwa A, Sato S, et al. New profiles of autoantibodies in the sera of Japanese patients with myositis. 67th Annual meeting of American College of Rheumatology, Oct., 2003
82. Hirakata M, Suwa A, Nojima T, et al. Clinical and immunological features in patients with anti-fer (Elongation factor 1 alpha) autoantibodies. 67th Annual meeting of American College of Rheumatology, Oct., 2003
83. Hirakata M, Suwa A, Sato S, et al. Clinical features of Japanese patients with anti-SRP autoantibodies: The immunogenetic backgrounds. 66th Annualmeeting of American College of Rheumatology, Oct., 2002
84. 2. Sato S, Suwa A, Hirakata M, et al. Clinical and immu.nological features of autoantibodies to the 140kDa polypeptide in patients with amyopathic dermatomyositis. 66th Annual meeting of American College of Rheumatology, Oct., 2002
85. Matsuura E, Kobayashi K, Kishi M, Atsumi T, Bertolaccini ML, Kasahara J, Makino H, Sakairi N, Yamamoto I, Yasuda T, Khamashta MA, Hughes GRV, Koike T, Voelker DR. Detection and clinical significance of oxidized LDL complexes with  $\beta$ 2-glycoprotein I and/or C-reactive protein. Atherosclerosis, Gordon Research Conference, Meriden, USA, 2003.
86. Kuwana M, Okazaki Y, Kaburaki J, Kawakami Y, Matsuura E. Binding of  $\beta$ 2-glycoprotein I to anionic phospholipids facilitates processing and presentation of a cryptic epitope that activates pathogenic autoreactive T cells. Annual Meeting of American College of Rheumatology, USA, 2003.
87. E. Matsuura E, Kobayashi K, Shoenfeld Y, Lopez LR. Role of natural antibodies and autoantibodies on development of atherosclerosis. Int Congress on Autoimmunity, Budapest, Hungary, 2003.
88. Inagaki J, Kondo A. Inagaki J, Lopez LR, Shoenfeld Y, Matsuura E. Anti-laminin-1 autoantibodies, pregnancy loss and endometriosis. Int Congress on Autoimmunity, Budapest, Hungary, 2003.
89. Ames PRJ, Delgado Alves J, Lopez LR, Iannaccone L, Margarita A. Scenna G, Brancaccio V, Matsuura E. Pro-atherogenic role

- for IgG anti-oxidised-low-density-lipoprotein- $\beta$ 2-glycoprotein-I complex in primary antiphospholipid syndrome. Int Congress on Autoimmunity, Budapest, Hungary, 2003.
90. Matsuura E, Kobayashi K, Kishi M, Atsumi T, Bertolaccini ML, Kasahara J, Makino H, Sakairi N, Yamamoto I, Yasuda T, Khamashta MA, Hughes GRV, Koike T, Voelker DR. Detection and clinical significance of oxidized LDL complexes with  $\beta$ 2-glycoprotein I and/or C-reactive protein. Atherosclerosis, Gordon Research Conference, Meriden, USA, 2003.
91. Kuwana M, Okazaki Y, Kaburaki J, Kawakami Y, Matsuura E. Binding of  $\beta$ 2-glycoprotein I to anionic phospholipids facilitates processing and presentation of a cryptic epitope that activates pathogenic autoreactive T cells. Annual Meeting of American College of Rheumatology, USA, 2003.
92. E. Matsuura E, Kobayashi K, Shoenfeld Y, Lopez LR. Role of natural antibodies and autoantibodies on development of atherosclerosis. Int Congress on Autoimmunity, Budapest, Hungary, 2003.
93. Inagaki J, Kondo A, Inagaki J, Lopez LR, Shoenfeld Y, Matsuura E. Anti-laminin-1 autoantibodies, pregnancy loss and endometriosis. Int Congress on Autoimmunity, Budapest, Hungary, 2003.
94. Ames PRJ, Delgado Alves J, Lopez LR, Iannaccone L, Margarita A, Scenna G, Brancaccio V, Matsuura E. Pro-atherogenic role for IgG anti-oxidised-low-density-lipoprotein- $\beta$ 2-glycoprotein-I complex in primary antiphospholipid syndrome. Int Congress on Autoimmunity, Budapest, Hungary, 2003
95. Nagashima, T., Okazaki, H., Minota, S.: Therapeutic effects of fluvastatin on the clinical course of MRL-lpr/lpr mice. American College of Rheumatology, 67<sup>th</sup> National Meeting, San Antonio, Texas, USA, Oct, 19, 2004.
96. Okazaki, H., Takahashi, M., Hirata, D., Kano, S., Minota, S.: Apoptosis of activated human T cells by statins through blocking protein farnesylation: A possible mechanism of their newly recognized immunomodulator activities. 26<sup>th</sup> International Congress of Internal Medicine, Kyoto, Japan, May, 28, 2002.
97. Nagashima, T., Okazaki, H., Hirata, D., Iwamoto, M., Matsuno, H., Kano, S., Minota, S.: Apoptosis of Rheumatoid Synovial Cells by Statins through Blocking Protein-geranylgeranylation: Potential Therapeutics for Rheumatoid Arthritis. American College of Rheumatology, 65<sup>th</sup> National Meeting, New Orleans, Louisiana, USA, Oct, 26, 2002
98. Sato K, Kanda H, Mimura T, Taniguchi T, Takayanagi H, Yamamoto K :Modulated Cytokine Production from APCs and a Novel Negative Feedback System Mediated by CD25+CD4+T Cells. 12th International Congress of Immunology and 4th Annual Conference of FOCIS.2004
99. Yamazaki M. Antiphospholipid syndrome, 3<sup>rd</sup> Asian-Pacific Congress on Thrombosis and Haemostasis (Symposium), Oct., 2004, Bangkok, Thai
100. Yamazaki M. Roles of adhesion molecules in the development of thrombosis and pregnant morbidity in antiphospholipid syndrome (APS): studies using APS-model mice deficient of adhesion molecules. 11<sup>th</sup> International Congress on Antiphospholipid Syndrome, Nov, 2004, Sydney, Australia.
101. Yamazaki M, et al. Roles of adhesion molecules in the development of thrombosis and pregnant morbidity in antiphospholipid syndrome (APS): studies using APS-model deficient of adhesion molecules. 45<sup>th</sup> America Soceity of Hematology, Dec., 2003, San Giego, USA
102. 小池隆夫:「抗リン脂質抗体症候群と血栓症」第 26 回日本医学会総会、福岡市、2004 年 3 月 3 日～5 日
103. 小池隆夫:「難治性の膠原病における医療の取り組み」第 58 回日本リウマチ学会総会・学術集会、岡山市、2004 年 4 月 15 日～17 日
104. 小池隆夫:「抗 Pathogenesis of Antiphospholipid Syndrome: p38 MAPK as a possible therapeutic target」第34回日本免疫学会総会・学術集会、札幌市、2004年12月1日～3日
105. 小池隆夫:「リウマチと免疫」第34回日本免疫学会総会・学術集会・市民公開講座、札幌市、2004年12月1日～3日
106. 小池隆夫:「自己抗体と血栓症・動脈硬化」第 26 回日本医学会総会、福岡市、2003 年 3 月 3 日～5 日
107. 小池隆夫:「抗リン脂質抗体症候群と血栓症」第 26 回日本静脈学会総会シンポジウム、東京都、2003 年 4 月 10 日～11 日
108. 小池隆夫:「抗リン脂質抗体症候群:オーバービュー」第26回日本血栓・止血学会総会、東京都、2003 年 11 月 27 日
109. 小池隆夫:「抗リン脂質抗体症候群研究の新展