

The purpose of this study is to investigate the effect of the administration of Liriope tuber extract, which contains much oligosaccharide and is used as a body fluid supplement in traditional medicine, on exercise ability in swim-trained rats by evaluating maximum exercise time, blood fatigue elements (lactate, ammonia, inorganic phosphate, pH). The exercise regimen was designed as swimming loaded with 10g weight to the base of the rat's tail. Experimental groups were trained swimming on schedule for 4 weeks and divided into 6 groups: (water (control), 5% water fraction (A), 10% water fraction (B), 5% crude extract (C), 10% crude extract (D), commercial beverage (E)). In 5-8 week study, we investigated the effect of only one administration (10ml/body weight (kg) before swimming) and in 9-10 week study, we investigated the effect of administration for two weeks.

Obtained results were as follows:

1. In only one administration study, A group and C group significantly improved exercise performance and reduced blood fatigue elements, but B and D showed no significant differences.
2. In two weeks administration study, A, B, C, and D groups all significantly improved exercise performance and reduced blood fatigue elements.

[PA1-59] [10/18/2001 (Thu) 14:00 - 17:00 / Hall D]

Pharmacological Action of *Cordyceps scarabaeicola*

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Dongchunghacho, one of folk medicines, is traditionally believed to be effective against various diseases. It includes many different genera such as *Cordyceps*, *Paecilomyces*, *Torrubiella* and *Podonectria*. *Cordyceps scarabaeicola* is one of well-known species. The 70% ethanolic extract was prepared from two different sources of *C. scarabaeicola*, fruiting bodies devoid of host materials (CS) and liquid medium-cultured cells (SC). Anti-angiogenic activity was determined by the chick embryo chorioallantoic membrane assay. Both CS and SC were found to contain strong anti-angiogenic activities. The extracts at the dose of 10 µg showed anti-angiogenic activity comparable to that of retinoic acid (dose, 1 µg), used as a control agent. Anti-angiogenic activities of CS and SC appeared to be dose-dependent. No significant differences were found between the effects of CS and SC. *Cordycepin*, an inhibitor of RNA synthesis identified in some *Dongchunghacho* species, showed anti-angiogenic activity. These results might suggest the plausible anti-tumor activity of *C. scarabaeicola*. Other pharmacological actions of *C. scarabaeicola* were examined.

[PA1-60] [10/18/2001 (Thu) 14:00 - 17:00 / Hall D]

The pharmacological profile of JOINS (SKI 306X) II : the potentiality as a curative therapeutics of rheumatoid arthritis

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Rheumatoid arthritis is a chronic multisystemic disease of unknown etiology and its characteristic feature is persistent inflammatory synovitis. Since the etiology and pathogenesis are not clear, the therapeutic approaches of these days are not curative, but just relieving the signs and symptoms of the disease. JOINS is a purified extract from a mixture of three oriental herbs, *Clematis mandshurica*, *Trichosanthes kirilowii*, and *Prunella vulgaris*, which have been widely used for the treatment of inflammatory diseases such as lymphadenitis and arthritis in Far East Asia. JOINS showed excellent analgesic and anti-

inflammatory activities in several animal models and in a clinical study on patients with OA, JOINS was revealed to have a good analgesic efficacy and safety profile. In this study, we tried to evaluate the possibility of JOINS as a curative therapeutics of rheumatoid arthritis using several in vitro and in vivo models. JOINS inhibited adjuvant-induced arthritis in rats and reduced inflammatory pouch volume and capillary permeability. JOINS also attenuated the PMA-stimulated chemiluminescence in neutrophils and degradation of articular cartilage by oxygen radicals. JOINS decreased conA-stimulated T cell proliferation and LPS-stimulated B cell proliferation. In conclusion, JOINS has a strong potentiality to be developed as a safe drug for rheumatoid arthritis.

[PA1-61] [10/18/2001 (Thr) 14:00 - 17:00 / Hall D]

Mucogen Ameliorates the Fibrosis and Inflammation of Chronic Pancreatitis in Mice

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In the present study, we established experimental chronic pancreatitis model in mice through repetitive induction of acute pancreatitis with intraperitoneal injections of cerulein (40 mcg/kg, 6 times every hours twice per week for 5 or 10 weeks), which led to chronic pancreatitis with fibrosis. Severe pancreatic acinar atrophy, trans-differentiation of acinar to duct like tubular complexes, islets hyperplasia, and dilatation of intraacinar lumina developed. Masson-Trichrome staining demonstrated progressive accumulation of extracellular matrix in interlobar and interacinar spaces. The extents of pancreatic fibrosis were statistically significantly decreased in accordance with lessened pancreatic inflammations after treatment of Mucogen (DA-9601), phytopharmaceutical showing antioxidative and cytoprotective actions. Using nuclear extracts from pancreas and radiolabeled NF-kappaB probe, EMSA was done, which showed the increased NF-kappaB binding in chronic pancreatitis and significantly attenuated NF-kappaB binding activities after mucogen treatment. The levels of myeloperoxidase and iNOS activities were also significantly decreased in mucogen treated group compared to pancreatitis control group. Cytoprotective proteins such as heat shock protein-70 and metallothioneine were significantly increased in mucogen-treated group. Mucogen decreased the expressions of alpha-SMA and type I collagen in cultured pancreatic stellate cells. Conclusively, we could establish the mouse model of chronic pancreatitis and mucogen might be considered as therapeutics in the prevention and treatment of chronic pancreatitis with fibrosis.

[PA1-62] [10/18/2001 (Thr) 14:00 - 17:00 / Hall D]

G protein-coupled phosphoinositide 3-kinase γ is required for autotaxin-mediated tumor cell motility

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Cell motility is a fundamental process required during normal embryonic development, inflammatory responses, wound healing, and tumor metastasis. Autotaxin (ATX) is a 125-kDa glycoprotein secreted by the human melanoma cell line A2058. This autocrine motility factor has been shown to stimulate random and directed motility of human tumor cells at high picomolar to low nanomolar concentrations (ED50 = ~300-500 pM). In the present study, we have shown that G protein-coupled phosphoinositide 3-kinase (PI-3 kinase) γ is involved in the signal transduction of Autotaxin (ATX), a novel tumor cell motility-stimulating factor. Pretreatment of the cells with PI-3 kinase inhibitors, wortmannin or LY294002 inhibits ATX-induced motility. Reverse transcriptase PCR and Western blot analysis showed that human melanoma cells have PI-3 kinase, p110 γ . ATX increased the PI3-kinase activity in p110 γ , but not p85, immunoprecipitates, which can be abolished by pretreatment of PI-3 kinase inhibitors (wortmannin, LY294002) or pertussis toxin. Collectively these results strongly suggest that PI-3 kinase p110 γ is