1861 Comparative Efficacy of Lactulose and Bifidum-Bacteriae in the Treatment of Disturbances in the Microecological System of Large Intestine in Premature Infants

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The period of adaptation to the extrauterine life is commonly complicated by disturbances in the microecological system of the large intestine (MSLI) that diminish the physiological resistance of the premature infants.

A comparison of Lactulose syrup (LS; Inalco, Italy) and lypo-phillized Bifidum-bacteriae (LB; Russia. 1 dose is 10^8 B. bifidum I) effects on 75 premature infants at the gestational age 32.7 ± 0.2 weeks with the disturbances in MSLI was made. The Group A of 35 neonates received every day 1.0 ml of LS Ag. The Group B of 40 infants received 2.5 doses of LB3 times a day. The course of the treatment lasted for 14 days. All the babies were fed with native breast milk.

The study of the infants included a general clinical examination; bacteriological examination of the feces; the study of the coprogramm, immunoglobulin A, G, M in coprofiltrates, and fecal pH (p < 0.05) was considered to be significant.

After the treatment dyspeptic disorders disappeared in 90% and intestinal microflora normalized in 85% of neonates of Group 1. Normalization of above mentioned disorders after LB treatment was observed respectively only in 56% and 25% of cases in Group 2. LS treatment led also to coprogramm normalization as well as to the significant decrease in immunoglobulin A content in coprofiltrates and fecal pH values.

Thus utilization of LS in the treatment of the premature infants with disturbances in MSLI was more effective than that of LB.

1862 Helicobacter Pylori in HIV-1 Seropositive Patients

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Two hundred and ten dyspeptic patients, both HIV-1 +ve and HIV-1 -ve, were endoscoped in a closed Community for ex drug abusers were investigated for Helicobacter pylori (HP) infection by upper gastrointestinal endoscopy and serology.

Control group by serology were: A) 259 asymptomatic subjects, both HIV-1 +ve and HIV-1 -ve, living in the same Community, B) 219 age and sex matched dyspeptic patients, C) 322 asymptomatic blood donors.

Results: The table shows the endoscopic findings in the endoscoped patients according to HIV and HP status (N = normal, OCV = oesophageal candidosis and/or varices, AG = antral gastritis, PU = peptic ulcer, GC = gastric cancer, ED = erosive duodenitis).

<table>
<thead>
<tr>
<th>N</th>
<th>OCV</th>
<th>AG</th>
<th>PU</th>
<th>ED</th>
<th>GC</th>
</tr>
</thead>
<tbody>
<tr>
<td>HP+ (n = 44)</td>
<td>5</td>
<td>4</td>
<td>17</td>
<td>9</td>
<td>9</td>
</tr>
<tr>
<td>HP− (n = 67)</td>
<td>23</td>
<td>26</td>
<td>11</td>
<td>0</td>
<td>7</td>
</tr>
<tr>
<td>HIV− (n = 99)</td>
<td>3</td>
<td>0</td>
<td>32</td>
<td>22</td>
<td>8</td>
</tr>
<tr>
<td>HP+ (n = 653)</td>
<td>12</td>
<td>1</td>
<td>16</td>
<td>0</td>
<td>4</td>
</tr>
</tbody>
</table>

The overall prevalence of HP infection in the endoscoped patients was 52%. Infection was more common in HIV-1 -ve (65/99, 66%) than the HIV-1 +ve patients (44/111, 40%) (P < 0.001). HIV-1 +ve patients with AIDS had a lower prevalence of HP infection (24/73, 33%) compared to HIV-1 -ve patients without AIDS (20/38, 53%) (P = 0.05).

The seropositivity rates of HP infection were similar in the dyspeptic patients (52%) and asymptomatic subjects (56%) living in S. Patrignano, and in the dyspeptic control patients not living in the Community (55%). Asymptomatic blood donors showed a lower seroprevalence of infection (33%, P < 0.001).

Conclusion: (1) HIV-1 positivity does not play a role in the development of HP infection, whereas the pival role is probably due to life in community (2) Apart from candidiasis unusual gastroduodenal pathology is rare in HIV-1 +ve dyspeptic patients.

1863 Functional and Structural Changes of Small Intestine in HIV Infected Patients with Chronic Diarrhea

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HIV infection is often associated with unexplained diarrhea and weight loss. This study was designed to see if abnormalities of the intestinal mucosal architecture could explain the enteric dysfunction found in HIV infected patients with diarrhea.

20 far coming HIV infected patients (19 men, 1 woman, CD4 0.01-0.2) with apparently non-infectious chronic diarrhea were evaluated with a new intestinal function test (D-xylose breath test) and, for fifteen of the subjects, with an upper intestinal endoscopy with biopsy specimens taken from the duodenal mucosa.

The function test showed that the D-xylose uptake was largely decreased in the same manner as for patients with celiac disease. The severe malabsorption could be hardly be explained by the slight mucosal damage seen at light microscopy with partial villous atrophy and unspecific inflammation. However, electron microscopy showed enterocytes with signs of hypofunction and degeneration.

The ultrastructural changes of the mucosa appear to be the best morphological explanation to the intestinal malabsorption found in far coming HIV infected patients with chronic diarrhea.

1864 Serum Lipids in Patients with Acute Pancreatitis

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Hyperlipidemia may be the cause of acute pancreatitis (AP) or a sequela of peripancreatic fat necrosis.

The aim of this study was the evaluation of serum lipides, ie cholesterol (CH), triglycerides (TG) and HDL-cholesterol (HDL-CH) in patients with AP, in relation to the time course of the disease. 37 patients were studied. In 32 patients AP was due to gallstones (Group A) and hyperlipidemia was the causative factor in 5 Group B).

Methodology included biochemical tests (WSB serum and urine amylase, serum creatinine, bilirubin, SGOT, SGPT), ultrasonography, CT scan of the abdomen, ERC and determination of serum lipids on 1, 3 and 15, after the onset of AP Results were as follows (mean ± SD).

<table>
<thead>
<tr>
<th>Serum lipids in Group A (mg/dl)</th>
<th>CH</th>
<th>TG</th>
<th>HDL-CH</th>
</tr>
</thead>
<tbody>
<tr>
<td>Day 1</td>
<td>188 ± 45</td>
<td>112 ± 25</td>
<td>34 ± 5</td>
</tr>
<tr>
<td>Day 3</td>
<td>215 ± 47</td>
<td>118 ± 25</td>
<td>35 ± 6</td>
</tr>
<tr>
<td>Day 15</td>
<td>201 ± 51</td>
<td>126 ± 35</td>
<td>36 ± 6</td>
</tr>
</tbody>
</table>

In summary: (1) No significant changes in serum lipids were noted in patients with AP due to gallstones during the time course of the disease (day 1, 3, 15), (2) In patients with AP due to hyperlipidemia, a significant elevation of serum CH and TG was noted on day 1 but did not return to normal. TG levels declined significantly on day 3 and 15.

Conclusions: (1) Serum lipids fluctuate during the course of hyperlipemic pancreatitis, but remain elevated, (2) no significant changes of serum lipids were noted in biliary pancreatitis.

1865 Is Bilary Lithiasis Associated with Pancreatographic Changes?


We analyzed retrospectively the endoscopic retrograde cholangiopancreatography of 165 patients presenting biliary lithiasis and of 53 controls. Among the 165 patients, 35 had gallstones, 53 had gallstones and cholecystolithiasis, 10 had cholecystolithiasis without gallstone, 50 cholesterol-stones patients had cholecystolithiasis, 17 had undergone cholecystectomy without recurrence of biliary lithiasis. Analyzing the pancreatography, we measured the diameter of the pancreatic duct in the head, the body and the tail of the pancreas, the regularity of the main pancreatic duct and the presence of stenosis, the regularity or the dilatation of secondary ducts, the presence of cysts. In addition, we calculated a score based on the above parameters: pancreatographies were classified as normal or with mild, intermediate, moderate, severe abnormalities. In patients, no statistical difference for the pancreatographic features was found in case of cholecystolithiasis compared to patients without cholecystolithiasis. Patients were comparable to controls for sex, alcohol consumption but not for age (68 vs 55 years, p < 0.01). In patients and in controls, the presence of abnormalities of the pancreatography well correlated with age. The pancreatographic features of patients and controls were not found to be significantly different, even in younger controls.

Finally, multivariate analysis (stepwise multiple discriminant analysis) was performed. The age appeared as the only significant factor which was able to predict the presence of pancreatographic abnormalities.
Synthesis of Platelet – Activating Factor (PAF) in Trypsin-Induced Acute Pancreatitis in Rabbit

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Trypsin activation represents the final mechanism of cellular damage in most experimental models of acute pancreatitis. PAF, a potent phospholipid mediator with a wide spectrum of biological activities, can induce the development of acute pancreatitis in rabbits. Moreover, some of our observations demonstrate that trypsin stimulates PAF synthesis by human cultured endothelial cell. In this study we evaluated the synthesis of PAF in pancreatic tissue after a single trypsin injection into the superior pancreatico-duodenal artery (SPDA) in rabbits. Different doses of trypsin were injected into SPDA: 1, 10, 100, 1000 mcg (4 animals/group). Sterile saline was injected in ten animals as control. At 24 hours, the pancreatic damage (edema, damage of acinar cell necrosis) was evaluated by light microscopy: severity of pancreatic damage was dose dependent. Acinar cell necrosis was present in all animals treated with 100 and 1000 mcg. In control group, no animals showed histological lesions. PAF tissue levels were determined in pancreatic tissue after 30 min from injection of trypsin 100 mcg or saline alone. The pancreatic tissue supplied by SPDA (about 2.5 sq cm) was frozen in liquid N2 and then homogenized for lipid extraction. The organic phase was subjected to thin-layer chromatography and further characterized by high-performance liquid chromatography; PAF was quantitated by aggregation of washed rabbit platelets. PAF was detected in pancreatic tissue from 5 out of 6 rabbits treated with trypsin 100 mcg; the PAF level was 1566.10 ± 530.72 pgg wet weight of tissue, while PAF was not detectable in samples from controls.

In conclusion this study demonstrates that local injection of trypsin into the pancreatic arterial bloodstream (a) induces in rabbit an acute necrotizing pancreaticitis and (b) stimulates the tissue synthesis of PAF, which could be involved in the pathogenesis of acute pancreatic injury.

The Ratio Between Anionic and Cationic Trypsin in Pancreatic Extracts is Dependent on the Extent of CCK Stimulation

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The secretion of different pancreatic enzymes is supposed to be parallel in most situations i.e. in acute and chronic pancreatitis. During the rejection of pancreatic grafts after wholeorgan pancreatic transplantation however a marked dissociation is seen between immunoreactive anionic and cationic trypsin in serum in both pig and man. The same picture is seen after acute hyperamylaseemic pancreatitis with cerulein injection and we have seen that the ratio between anionic and cationic trypsin changes after a chronic infusion of CCK and after a chronic infusion of a CCK antagonist (Devazepide) in rat.

Methods: Male Sprague rats were given continuous infusion of CCK-85 (5 mcg/kg) or 0.5 devazepide (200 mcg/kg) n = 10 by a subcutaneous pump for 4 weeks. 11 control rats received saline. The rats were freely fed. After sacrifice the pancreases were harvested and weighed. Pancreatic extracts were performed in an acetate buffer at pH 4.3. Total trypsin activities were determined using BAPNA as substrate. Anionic and cationic trypsin activities were analysed separately after electrophoretic separation. Results: Bodyweights were similar in all three groups. Pancreatic weights were significantly higher in the CCK stimulated group 2.65 ± 0.28 g (mean ± SD) compared to the control group 1.18 ± 0.15 g. The pancreases in the devazepide group weighed significantly less than in the controls 0.92 ± 0.09 g. The total amount of trypsin activity in the control group was significantly higher in the CCK group 13.22 ± 0.06 compared to both the control group 4.31 ± 0.12 and the devazepide group 4.07 ± 0.32. The ratio between anionic and cationic trypsin in extracts from CCK stimulated rats were 1.08 ± 0.33 which was significantly higher than in the control group 0.71 ± 0.22. In the devazepide group this ratio was significantly lower 0.53 ± 0.17.

Conclusion: The secretion of anionic and cationic trypsin(ogen) seem to be dependent on “CCK tonus” in a anticoordinate way which may explain a nonparallel response in secretion in certain situations.

Diet of Patients with Alcoholic Chronic Pancreatitis in Poland

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The diet of patients with chronic pancreatitis (CP) in the period before the onset of the disease was many times studied but there are only a few studies concerning the diet in the time of developed CP. The purpose of the study was the comparison of the diet of patients with alcoholic CP with the diet of individuals selected from general population.

Material and Methods: The study group comprised 45 men with well confirmed alcoholic CP with the mean age of 42.9 years. The group of men chosen from general population (control group) consisted of 45 men with the mean age of 38.3 years. The dietary history was used as a tool in the research of the dietary data.

Results: Caloric value of the diet and main nutrients intake in both groups are presented in the table.

<table>
<thead>
<tr>
<th></th>
<th>CP group</th>
<th>control group</th>
</tr>
</thead>
<tbody>
<tr>
<td>caloric value, kcal</td>
<td>3197</td>
<td>3303</td>
</tr>
<tr>
<td>protein, g</td>
<td>120</td>
<td>114</td>
</tr>
<tr>
<td>fats, g</td>
<td>133</td>
<td>146</td>
</tr>
<tr>
<td>carbohydrates, g</td>
<td>373</td>
<td>384</td>
</tr>
</tbody>
</table>

These differences were not statistically significant. A detailed comparison revealed statistical differences only a small group of nutrients: pantethonic acid intake was higher in the study group and polyunsaturated fat, sodium and fiber intake was higher in the control group. The consumption of cottage cheese was twice higher in CP group but gryuze cheese intake was three times higher in control group. Even 72% of patients with steatorrhea consumed over 100 g of fat daily.

Conclusions: 1. The diet of the patients with alcoholic CP in the time of developed disease shows only a little differences with the diet of healthy men.

2. High mean fat intake is a factor which may be a factor of steatorrhea in many patients.

KSG-504, a Novel Cholecystokinin Receptor Antagonist Prevents the CCK-8-Induced Acute Pancreatitis in Rats

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The role of cholecystokinin (CCK) receptor blockade was studied in acute edematous pancreatitis evoked by high doses of CCK-8. The aim of our study was to evaluate the effects of KSG-504, a CCK receptor antagonist recently synthesized by the Kissele Pharmaceutical Co. (Masumoto, Japan) in this pancreatitis model.

Methods. Pancreatitis was induced with serial CCK-8 injections (75 ug/kg s.c. at 2-h intervals over 6 h). KSG-504 (5 or 50 mg/kg s.c.) was injected 0.5 h before and 3 h after the last CCK administration. Rats were examined at 6 h after pancreatitis induction. To assess the severity of inflammation, pancreatic edema measurement, histological grading, serum amylose determination and plasma CCK bioassay were performed.

Results: Administration of CCK-8 alone caused edematous pancreatitis with a significant elevation of pancreatic weight/body weight (pw/bw) ratio (5.65 ± 0.7) and the serum amylose level (38.7 ± 4.9 U/ml) vs. the controls (3.38 ± 0.26 and 2.9 ± 0.3 U/ml, respectively). Both 5 mg/kg and 50 mg/kg of KSG-504 significantly decreased the pancreatic enzyme (3.45 ± 0.25 and 2.4 ± 0.2, respectively) and serum amylose activity (14.2 ± 2.3 U/ml and 5.1 ± 0.8 U/ml, respectively). The histologic changes seen in the CCK-8-treated group (interstitial edema with neutrophilic cell infiltration, and pancreatic acinar cells with vacuoles of different sizes) were completely prevented by the high dose of KSG-504, but were only slightly influenced by the low dose. The plasma CCK level was significantly elevated in the CCK-8-treated group vs. the control (4.9 ± 1.4 pm and 0.5 ± 0.07 pm, respectively). No significant plasma CCK level changes were observed in the KSG-504-treated groups as compared to the control group.

Conclusion. These findings provide further evidence that the plasma CCK is involved in the pathogenesis of acute edematous pancreatitis. KSG-504 proved to be a potent CCK receptor antagonist in this acute pancreatitis model.

Malignant Pancreatic Neoplasms – Clinicomorphological Correlations

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The present research is based on 172 dead patient autopsies. In this study pancreatic neoplasms ranked at fourth place, after malignant neoplasms of the lung, the stomach and the colon. The men to women distribution was 80%-20%, respectively. They occurred most frequently in the 6th-7th decades – over 55%, while the 5th and the 8th decades were identically presented – 20%. The neoplastic process was mainly localized in the head of the pancreas – 71%, pure – 50%, body combined – 11%, body and tail – 10%. The predominant histological types were adenocarcinoma (75%) and papillary adenocarcinoma (20%). Mucus secretion was observed in 45%, anaplasia – in 30%, and desmoplastic reaction – in 40%. The first metastases were found in the regional lymph nodes – 75 %, the liver – 65% and the lung – 33%. Pancreatic and pleural carcinomatases were observed in 32% and 24%, respectively. As accompanying diseases were found: chronic pancreatitis – in 20%, peptic ulcer in 18%, diabetes – in 8% and atrophic chronic gastritis –
in 17%. The most frequently signs found were: icterus – 74%, ascites – 56% and thromboembolic disease – 54%. The death was connected to: lung embolism (35%), sheer cardiovascular failure (15%), liver failure (13%), liver kidney failure (12%), pentonitis and pneumonia (8%) and haemorrhage including the brain (6%). The proper ante-mortem diagnosis was made in 68% of the cases, the percentage of correct diagnoses being highest in patients with pure head of the pancreas localization and the lowest when the tumour was localized in the body tail or the tail of the pancreas.

1871 Endoscopic and Surgical Therapy of Chronic Pancreatitis
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Therapy of the particularly painful forms of chronic pancreatitis (ch.p.) is hard. Elimination of pain improves the quality of life of the patients, enables them to increase food intake and so adjust changes connected with nutritional disorder.

In 40 patients with clinically, morphologically and functionally proved ch.p. – the obstructive type ETP of the pancreatic duct was carried out and followed either by duct drainage or an attempt of stone removing from the duct. In 47 patients with chronic painful type of ch.p. and with dilatation of the pancreatic duct – resection of the duct was carried out. The subjective changes in pain, change in body weight, stool frequency and endocrine function were evaluated before and after the therapy.

Surgical and endoscopic treatment led in approximately 70% of patients to pain relief, to an increase of body weight of about more than 2 kg in 50% of patients. Drainage therapy influenced the endocrine function of the pancreas less frequently than the resection of the pancreas cauda.

Surgical therapy led to a longer duration of the favourable effect on the condition than the endoscopic one, but it was burdened with more severe complications.

Note: statistical evaluation by T-test.

1872 Assessment of the Effect of Undernutrition in Chronic Pancreatitis on Pancreatic Utilization of Amino Acids After Secretin and Cerulin Stimulation
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In undernutrition we observed disturbances of utilization of amino acids (AA) in the synthesis of pancreatic enzymes. Decreased AA level after stimulation of the pancreas was not different from that found by us in chronic pancreatitis (CP) with normal nutrition. For assessing the phenomenon of AA utilization by the pancreas after secretin and ceruline stimulation as a possible pancreatic function test it was tried to find out whether any significant difference existed in AA level decrease between patients with undernutrition in CP and those of other origin.

Material and methods. Three groups were studied: I – 13 CP patients with normal nutritional status, II – 8 CP patients with undernutrition, III – 11 undernourished patients without CP. Blood samples for AA determination (in automatic analyser) were obtained before and in the 45th minute of i. v. secretin (Sec) or ceruline (Cer) infusion (1 CU/kg) and ceruline (Cer) (75 ng/kg).

Results. The AA level decreased in 45 min. after i. v. Sec or Cer statistically significantly (p < 0.05) in group I (by 324 ± 40.7 nmol/ml i.e. 10.9%) and in group III (by 316 ± 50.53 nmol/ml i.e. 12.5%). In group II it decreased not significantly by 94 ± 16.7 nmol/ml, i.e. 3.8%. This decrease (in group II) was significantly lower (p < 0.001) than in group I and III.

Conclusion: After pancreatic stimulation in patients with chronic pancreatitis undernutrition the amino acids were found to be less extensively utilized for enzyme synthesis than in patients with this disease but with normal nutrition and in patients with undernutrition caused by other factors than pancreatic disease.

1873 Experience with Stenting of the Pancreatic Duct in Chronic Pancreatitis in a Central Hospital

The main goal of treatment is the relief of persistent pain or relapsing pain occurring as clinical attacks of pancreatitis. The classic treatment have been a surgical procedure. In the last 10 years a few reports have been published on the endoscopic management of chronic pancreatitis.

Material and method: The Central Hospital of Akershus serves 400,000 peoples and are a local hospital for about 200,000.

Since January 1986, patients (n = 16), m = 12, w = 4, age 33–73 (median 55) with chronic pancreatitis with ductal strictures and/or stone and upstream dilatation underwent stenting of the main pancreatic duct through the major papilla (n = 13) or the minor papilla (n = 3) in order to drain the duct and relief of pain. The etiology was alcohol abuse in 6, gallstones in 3 and unknown in 5 patients.

Four of the patients were earlier operated on pancreas. 3 due to the chronic pancreatitis and 1 patient had underwent Whipple’s operation (1973) due to a tumor of the papilla of Vater. We have been using 5–10 French Am-sterdam” or “Pigtail” endoprosthesis. The intension was to treat the patient with endoprosthesis for 1–2 year and replace it individually in case of relapse.

Results: One severe complication did occur to the method, a purulent in- fection of the pancreatic duct. Some patients did have more pain a few days after the stent-placement. All patients earlier operated on did get a good pain relief after stenting. One of the patients with pancreas division had a good pain relief with the endoprosthesis. The other two underwent surgery. One do still have pain and the other died in a pancreatic carcinoma 9 months after the operation.

The rest of the patients, n = 9, is still treated with endoprosthesis and do all have pain relief in shorter or longer periods after stent replacement.

Conclusion: Chronic pancreatitis with severe pain is a difficult problem. Endoprosthesys does not solve all problems but in selected patients it is the “treatment of choice”. In others it can be the way to select the right treatment.

1874 Prevalence and Incidence of Chronic Pancreatitis France. A Population Based Study
J. Faivre, P. Bernades, Observatoire de la Pancréat Chirurgique. Fauteuil de Médecine, Dijon et Hôpital Beaujon, Clichy, France

Very few data are available on the epidemiology of chronic pancreatitis (CP) Gastroenterologists, endocrinologists and radiologist from 7 french departe- ments (population 3,700,000) were asked to report from January 1990 to December 1992, all the patients seen with known or newly diagnosed CP Prevalence and incidence rates were standardized by the direct method us- ing the European standard population. Population used in calculating rates were based on data from the 1990 census. During these 3 years 1,293 cases were registered. Age standardized prevalence rates per 100,000 varied be- tween 77.8 and 37.8 in men according to the areas involved and between 14.1 and 4.1 in women. Incidence rates were a mean of three years regis- tration. Only cases diagnosed for the first time between 1990 and 1992 we involved in the analysis. Incidence rates varied between 10.6 and 6.1 in men and between 2.3 and 0.8 in women. Extrapolating these data to the french population it was estimated that the number of new cases each year was about 2,500 and that nearly 20,000 subjects were living with a CP. These data suggest that there are differences in prevalence and incidence rates of CP in France and that by its chronicity this illness represents an important health problem.

Supported by a grant of LTM-Laténa, France (Solvay-Pharma).

1875 Management of Multiple Abscesses Following Necrotizing Pancreatitis

Pancreatic abcess and sepsis are the leading causes of mortality in necrotiz- ing pancreatitis. Since 1986, 123 patients with infected pancreatic necrosis and several retroperitoneal absesses have been treated. In all cases, the in- fected necrosis or pancreatic abscess was combined with retroperitoneal ab- scesses. 80 were situated the right or left retrocolic area, 13 in the subphrenic region, and 30 in the retroperitoneal or subhepatic area. The surgical treatment was performed on average 18.5 days after the onset of acute pancreatitis. The operative management consisted of wide-ranging necrosectomy in the total affected area, combined with widespread washing and suction drainage. In 56 of the 123 cases (45.9%), some other surgical intervention (distal pancrea- tic resection, splenectomy, cholecystectomy, or colon resection) was also performed. Continuous washing and suction drainage was applied for an av- erage of 39.5 days, using ± 6 liters of saline solution per day. The bacte- riologic findings revealed mainly enteric bacteria, but Candida infection was detected frequently. The incidence of infected Candida infection was 21%. Dissemi- nated Candida infection was noted in 12 cases and 3 of these died. Recently capacity of cytokine production (TNF, IL-1, IL-6) has been investigated and it has shown correlation with the prognosis. The overall hospital mortality was 7.3% (9 patients died). In our experience, the procedure of choice for pan- creatic abscesses is aggressive surgical treatment, continuous, long-standing washing and suction drainage, together with supportive therapy combined with adequate antibiotic and antifungal medication. This surgical strategy is the only possibility for recovery in cases of pancreatic abscesses.
Operative Treatment of Chronic Pancreatitis in a Central Hospital

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The Central Hospital of Akerhus serves 400,000 people and is a local hospital for about 200,000.

Materials and Methods: From May 1983 to May 1993, 56 patients have been hospitalized due to chronic pancreatitis. Different operations have been performed in 17 patients, 9 men and 8 women, median age 48 years (33–68).

The indication was severe pain in all, but 2 had jaundice as well. The etiology was alcohol abuse in 9 patients, gallstone in 1 and unknown in 4. 3 patients had pancreatic carcinoma, in 2 the diagnosis were verified at autopsy, and in 1 at re-operation because of increasing pain.

The operations used where internal stonies because of pseudocysts in 4, resections of the tail and body of pancreas in 3, pancreasresections with pancreatico-jejunostomy a.m. Jordan in 4, pancreatico-jejunostomy a.m. Puestow in 3, pancreatico-gastrostomy in 1 and Whipple's operation in 2 patients. There were 2 severe post-operative complications; both external pancreatic fistula. One had to be re-operated, the other was treated with an endoprosthesis in the pancreatic duct.

Results: The outcome was good or excellent in 8, in 5 patients the pain relief was good for only a period of 6 to 4 years. 2 of these patients have later been treated with endoscopic endotheraphy in the pancreatic duct. 1 of our patients had no benefit from the operation.

Conclusion: Chronic pancreatitis with severe pain is a difficult problem. It may also be difficult to differentiate between pancreatic carcinoma and chronic pancreatitis. In 3 of our patients we missed this diagnose. 13 out of 17 patients had benefit from the operation.

Synthetic Proteinase Inhibitors Exert Oxidative Stress Ameliorating Effect in Acute Experimental Pancreatitis

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This study was undertaken to determine whether synthetic proteinase inhibitors - nafamostat mesilate (FUT-175) and gabexate mesilate (FOY) have any influence on multigram oxidant-antioxidant balance in acute experimental pancreatitis. Acute haemorrhagic pancreatitis was induced in male Wistar rats using a retrograde intraductal injection of 5% Na-taurocholate. Rats were treated with FUT-175* (25 mg/kg/h) or FOY** (2.5 mg/kg/h) given i.v. immediately after pancreatitis induction. Animals were sacrificed at 3 h; malondialdehyde (MDA) concentration, as an index of oxidative injury, and sulphhydril groups (SH-groups), known as a major nonenzytatic antioxidant, were determined in pancreatic, lung and tissue liver. Results are expressed as mean ± SD.

Dose- and Time-Dependent Effect of Alcohol and Acetaldehyde Exposition on the Amylase Secretion of Isolated Rat Pancreatic Acini Stimulated by Cholecystokinin Octapeptide (CCK 8)

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We have previously showed that intragastrically given alcohol has a non-competitive inhibitory effect on the amylase secretion of the isolated pancreatic acini stimulated by cholecystokinin octapeptide (CCK 8). In these experiments we studied the effect of alcohol and acetaldehyde preincubation on the amylase secretion of the pancreatic acini.

Methods: Isolated acini were prepared from male Wistar rats by a method as described by Hajnal et al., Gastroenterol. 1990; 98: 191. The acini were incubated with different doses of alcohol and acetaldehyde for 30-240 minutes. The acini were then stimulated by raising doses of CCK 8 and the amylase secretions were measured.

Results: Preincubation with 10, 20, 50, and 100 mM alcohol caused a significant increase in the threshold dose of 0.1 and a graded decrease in the amylase secretion at the higher secretagogue doses. Raising of preincubation time caused a proportional non-competitive inhibition of amylase release. After preincubation with 10 mM acetaldehyde also a non-competitive
inhibitory effect was observed in the amylase secretion stimulated by CCK.

Conclusion: The time- and dose-dependency of the alcohol and acetald- hyde exposition of the acini may refer to the progressive intoxication which is reflected by the declining enzyme production. The identical effects of al- cohol and its toxic metabolite may serve as indirect evidence for the alcohol metabolizing feature of the acinar cell.

**1881 Acute Pancreatitis in Buskerud County-Incidence and Etiology**

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The incidence of acute pancreatitis is not well established, but figures of 10-20/100,000 have usually been found in studies from Great Britain and USA, while the diagnosis has been made more frequently in urban Swedish populations.

We wanted to study the incidence of acute pancreatitis in a norwegian population, covering both rural and urban areas – the county of Buskerud with 224,000 inhabitants.

All three hospitals in the county participated. All patients presenting with symptoms consistent with acute pancreatitis and serum-amylase or serum- lipase values of more than twice the upper reference limit, i.e. 600 UI and 400 U/I, respectively, were prospectively included.

Hypermia was seen in 139 cases, giving an incidence of 61.9/100 000. Acute pancreatitis was diagnosed in 123 instances (in 119 patients, 68 men and 51 women), the incidence being 54.7/100 000. The mean age was 60.8 years (range 21-92 years). The etiology was gallstone disease, alcohol abuse and drugs (usually NSAIDS) in 45, 17 and 7%, respectively, while 8% of the cases were classified as idiopathic. Cholecystectomy and papillotomy had been performed earlier in 11% and 4% of the patients, while these procedures were done in 11% and 31%, respectively, during the hospital stay. A comparison between the patients with alcoholic and gallstone induced pancreatitis showed the former to be more men, younger, staying shorter time in hospital and with lower median values of serum amylase, bilirubin and amiontransferases. Seventeen patients (14%) had a history of recurrent pancreatitis, the cause being alcohol abuse and gallstone disease in 7 and 3, re- spectively. Pseudocysts or abscesses were seen in 9 patients (7.3%), while 7 patients (5.7%), 6 of them men, died. Their mean age was 79 years.

The incidence of acute pancreatitis in Buskerud county in southern Nor- way is apparently higher than in British and American populations, but lower than the figures from some Swedish studies. Regarding etiologies, sex- and age distribution and complications our results are comparable to earlier stud- ies.

**1882 Study of 136 Main Pancreatic Duct Retrograde Catheterisation Over 70 Years of Age**

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The authors evaluated during a 6 year period all E.R.C.P performed in patients over 70 years of age.

The aim of this study was to determine the morphological abnormalities of the pancreatic ductal system in elderly patients. 136 E.R.C.P were thus as- sessed. 31 cases showed an abnormality related to organic disease (22.8%): 13 pancreatic cancers (41.9%), 10 chronic calcifying pancreatitis (32.2%), 4 vaterians ampullomas (12.9%), 3 pancreas divisum (9.7%) and one case of oddits following biliary lithiasis (3.3%). The 105 remaining E.R.C.P showed a normal or dilated wirsung duct as follows: wirsung duct normal in 31.4% of patients – partially dilated in 23.8% (caphalic dilatation in 64% and caphalo- corporeal dilatation in 36% of patients). The wirsung duct was globally dilated in 44.8% of cases (10.6% of which were markedly dilated; in these cases, we have coined them as "megawirsung" of the elderly). The collateral ducts were dilated in 26.7% of cases and cysts were found in 14.2%. In conclusion this study shows that there are two types of wirsung duct in the elderly: (1) 20% of wirsung ducts are morphologically normal; (2) 80% of wirsung ducts are dilated, 22.8% of which can be ascribed to organic causes and 57.2% with out any cause. The authors discuss the mechanisms of main pancreatic duct dilatation.

**1883 Prospective Echoendoscopy Study of the Prevalence of Early Chronic Pancreatitis in 47 Non Cirrhotic Alcoholic Patients**


Echoendoscopy (EES) is increasingly used in the assessment of pancreatic disorders. EES has been proposed as a valuable screening test for early stages of chronic pancreatitis (ECP). The most commonly encountered signs are: the areolar aspect of the pancreatic parenchyma and the dilatation of the main pancreatic duct (MPD). The aim of our study was to determine the prevalence of these echoendoscopic signs.

Methods: 72 males and females, mean age 52 years, were investigated by EES. All patients underwent EES under neuroleptanalgesia with a EUM 3 Olympus endoscope. Meatal alcohol consumption was 273.7 g/day in group I and 13.6 g/day in group II.

Liver biopsy was performed in 41 patients (89%) and showed acute alco- holic hepatitis in 8 patients and fibroatosis without cirrhosis in 34 patients.

Results: (1) Group I: EES was normal in 26 patients (55.3%) and estab- lished the diagnosis of ECP in 21 patients (44.7%) based on the following find- ings: (a) parenchymal changes: areolar pattern in 18 cases; hyperemogenic spots in 6 cases; microcotyl in 1 case. (b) ductal changes: increased diameter of the MPD in 6 cases, hyperemogenic limits in 6 cases, distorted MPD in 4 cases. The MPD was normal in 7 cases. (2) Group II: EES was normal.

Conclusions: the prevalence of ECP in non cirrhotic chronic alcoholic pa- tients is 44.7%. This high prevalence should encourage the use of EES as a routine exam in the management of alcoholic patients undergoing with- drawal. Combined with clinical follow-up, EES may also become part of the monitoring of disease progression in order to detect chronic calcifying pan- creatitis in case of persistent alcohol addiction.

**1884 Selenium in Patients with Pancreatitis**


Acute pancreatitis is thought to be disease in which free oxygen radicals plays an important role. We have measured the serum level of selenium, one of the important free radicals scavengers. We used the atomic absorption spectrometry method. Medium selenium level in a healthy control group was 50.48 gl/ml.

We found a very low level of selenium with the patients with acute pan- creatitis. The medium was 17.13 gl/ml, the range was 3.4 gl/ml to 38.9 gl/ml (N 17). There was also lower level of selenium with the patients with chronic pancreatitis. The medium level was 34.7 gl/ml and vary from 4.7 to 77.9 gl/ml (N 33).

Conclusions. Our pilot study suggested the important role of selenium in acute and chronic pancreatitis. We concluded, that the low selenium level with patients with acute pancreatitis is caused by consumption of selenium as a scavenger of free oxygen radicals. We suppose that the treatment with selenium may play an important role in the course of the acute disease as well as in chronic pancreatitis.

**1885 The Rat Model is Invalid in Enzyme Secretion Studies of the Human Pancreas**


In the rat, removal of or inhibition of proteolytic activity in the duodenum caused a CCK mediated enhanced secretion of all pancreatic enzymes. This CCK stimulated secretion was shown to be atopine insensitive. The humoral regulation works through two independent trypsin-sensitive peptides both causing increased CCK release from the duodenal mucosa. The enhanced CCK release is assumed to be the trofic factor responsible for the pancreatic hypertrophy, hyperplasia and later cancer, in rats fed soy bean products containing trypsin inhibitors. The aim of the study was to see if intraduodenal instillation of various (i.e. soy bean) proteinase inhibitors in humans induced enhanced CCK release and enzyme secretion as in the rat.

Enzyme inhibition was measured by determination of residual enzyme ac- tivities and amounts of enzyme secreted as immunoreactive material. Com- pared with the rat, in which CCK levels increase 20-30 x basal value, only moderate (1-2 X) or no increase were obtained in the rats. In spite of this, a substantial increase (2-5 x) in enzyme secretion was obtained due to inhibitor instillations. However, depending upon the particular inhibitor used, selective inhibition and/or stimulated enzyme secretion could be obtained. After having established an elevated secretion rate (3-4 x) by inhibitor instilla- tion, i.e. atopine caused an immediate drop in enzyme output. Also increased secretion of pancreatic secretory trypsin inhibitor (PSTI) could be induced due to the particular inhibitor used, and also this enhanced secretion was atropine sensitive. No effects of atopine on the plasma CCK was seen. Inhibitor contain- ing soy bean products induce a neurally regulated pancreatic secretion and are consequently not assumed to be a nutritional hazard to man.
**1886** Prognosis of Pancreas Cancer Course by Tumor Markers Method

A.V. Kalinin, S.V. Skvortsov, B.N. Lytsar, E.A. Antokhin. The Main Military Hospital, Moscow, Russia

**Purpose of study:** to find out if it is possible to prognosticate the course of pancreatic cancer (PS) depending on the lever of tumor markers (TM) SA-19-9 and carcinoembryonic antigen (CEA) in blood serum.

**Method of study:** SA-19-9 and CEA count in blood serum of PS patients was determined by enzyme immunassay using sets produced by DIALplus, a Swiss-Russian joint venture company.

**Results:** the table below shows correlation between TM concentration in blood and length of survival of PC patients.

<table>
<thead>
<tr>
<th>Survival after TM test was made</th>
<th>Normal TM in blood</th>
<th>Increased TM count in blood</th>
<th>SA-19-9</th>
<th>CEA</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>moderate</td>
<td>high</td>
<td></td>
<td></td>
</tr>
<tr>
<td>&lt; 3 months</td>
<td>2</td>
<td>3</td>
<td>2</td>
<td></td>
</tr>
<tr>
<td>3-6 months</td>
<td>2</td>
<td>2</td>
<td>2</td>
<td>1</td>
</tr>
<tr>
<td>6-12 months</td>
<td>2</td>
<td>2</td>
<td>1</td>
<td></td>
</tr>
<tr>
<td>n &gt; 9</td>
<td>1</td>
<td></td>
<td>1</td>
<td>2</td>
</tr>
</tbody>
</table>

**Note:** increased TM count in blood: moderate – not more than by 2-3 times, high – over 10 times.

A distinct correlation was found out between the length of survival of PS patients and higher than normal TM count as well as TM level in blood at the moment of initial test. All 22 patients who died within 3 months had increased SA-19-9 and CEA count in their blood, with 15 (68.2%) of them having one time excess of normal content; 5 more (22.7%) had a considerable increase of SA-19-9 and CEA in their blood as against the normal. At the same time not one patient of the 4 who lived longer than a year showed a considerable increase of TM and simultaneous pathological count of SA-19-9 and CEA, and one case showed their normal level.

**Conclusion:** the length of survival of a PC patient depends directly on the level of SA-19-9 and CEA in blood at the initial testing. At the same time excess of SA-19-9 and CEA in blood against the norm is a poor prognosis for the length of survival of a PC patient.

**1887** Are There Simple Markers for the Prompt Diagnosis of Cholelithiasis in Patients with Acute Pancreatitis

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The purpose of this study was to investigate possible diagnostic markers for the prompt diagnosis of coexistent cholelithiasis in patients with acute pancreatitis.

**Subjects/methods:** 68 patients with acute pancreatitis, aged 45 ± 12 years were retrospectively distributed in two groups. The group I was consisted of 32 patients and the group II of 36 pts according to the coexistence or not of cholelithiasis respectively. Serum alanine aminotransferase (ALT), aspartate aminotransferase (AST), alkaline phosphatase (SAP) and bilirubin (Bil) levels were evaluated and a multivariant logistic regression analysis was performed between the two groups.

**Results:** Statistically significant differences (p < 0.01), were noted in female patients, and ALT, AST, SAP, Bil, values in group I patients as compared with group II patients. The following laboratory results (cut off), received on admission were of value in predicting whether a pancreatitis attack was associated with gallstones or not: ALT > 55 u/L (n.v.: 10-42), SAP > 140 u/L (n.v.: 40-120) Bil > 1.3 mg/dl (n.v.: 0.5-1.1).

It was estimated by regression analysis that the presence of more than 3 criteria predicted coexisted gallstones, while the absence of all five markers excluded this association.

**1888** The Influence of Naloxone in the Outcome of Experimental Acute Pancreatitis

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Naloxone, an opioid antagonist considered to inhibit the release of free oxygen radicals, was assessed on the progression and outcome of acute pancreatitis in rats. A model of experimental pancreatitis was induced by injection of 5% sodium taurocholate into a closed duodenal loop (group P). In a similar group naloxone was then administered intravenously (group P + N). As a comparison, two other groups were used, a sham-operation group (group S) and another one in which naloxone was given as previously (group S + N). Twenty four hours later serum amylase levels were determined and the pancreas of all groups was removed for histologic/semiquantitative morphometry and for biochemical determination of malondialdehyde levels, which account for the oxygen free radicals formation.

Amylase and malondialdehyde were statistically increased in the pancreatitis group (P) compared to groups S and S + N. Malondialdehyde levels though were statistically decreased in the naloxone [group P + N] treated rats [1618 ± 327 nmol/g] compared to pancreatitis group [P] [1978 ± 428 nmol/g]. Morphometric evaluation of pancreatitis findings were apparent in group P compared to less findings in naloxone group P + N [4.17 ± 1.2 versus 2.83 ± 1.83 grade].

Concluding, we may therefore imply that naloxone significantly inhibits the release of oxygen free radicals, suggesting that it may be used in delaying the progression of acute pancreatitis in rats.

**1889** Clinical Value of Dolichol Measurement in Pancreatic Cancer Control

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Pancreas contains a highest level of Dolichol (Dol) concentration in human tissues. Dol is responsible for the state of cell membranes and synthesis of N-glycoproteins in Dolichyl Phosphate Cycle (DPC). Urinary Dol have been reported to be 5 to 40 times the normal values, suggesting a metabolic abnormality of DPC in patients with cancer (Pullarkat et al., 1984). With focus on a tumor marker, the present study was carried out to estimate blood and urinary levels of Dol in patients with pancreatic cancer (PCA) and chronic pancreatitis (CHP).

The samples obtained from 29 patients with PCA (male, 38-67 years old) and 48 patients with CHP (male, 40-58 years old). Dol in blood and urine was assayed by HPLC method (Turpeinen, 1986).

Dol in healthy men’s blood and urine are 124 ± 8.5-ng/ml and 5.9 ± 0.6 µg/mmol creatinine respectively. In CHP Dol content in urine was much the same, but Dol content in blood showed an increase of 45-82%. Blood Dol concentration in patients with PCA increased at stage II up to 25%, at stage III up to 55%, making up 204 ± 12.9 ng/ml at stage IV. There was a significant difference between urinary Dol content in patients with CHP and that of cancer patients. Urinary Dol concentration increased at stage I up to 19-38%, making up 12.8 ± 2.8 µg/ml at stage II. At stage III the level of urinary Dol was 3-5-fold increased.

These findings suggest that Dol appeared in urinary excretion is one of the first manifestations of malignancy in pancreas. In this way CHP therapy should be carried out under Dol excretion control. The interest drawn to the employment of Dol as marker is explained by the fact that known PCA markers are glycoproteins (CA-50 and CA-19-9). There is reason to suggest that markers detected in PCA may evidence of the prime DPC cycle. Monitoring CHP patients with monthly urinary Dol determinations is a reliable method to diagnose a PCA.

**1890** Clinical Peculiarities of Alcoholic Pancreatitis

A.S. Loginov, V.M. Sadokov. Central Research Institute of Gastroenterology, Moscow, Russia

Examination of clinical peculiarities of alcoholic pancreatitis (AP) is the aim of this study. 170 patients with AP (167 male, 3 female, age 25-62 years) were observed for the period 2-12 years by special program.

The first clinical symptoms of pancreatitis appeared after 15 years of alcoholic abuse. The middle age of patients by the appearing of the first symptoms was 36 years. Dull pain in the upper part of abdomen preceded the first attack in 76% of these patients. We called this period as the stage of the early clinical symptoms. 93% of patients had the relapse stage, which began with the first attack. Duration of this stage depended on the development of complications and the degree of loss of the functional tissue in pancreas. Complications of AP appeared in 38% of patients. The complication stage may appear immediately after the first attack or a number of relapses may precede it during a few years. Exocrine pancreatic insufficiency was observed in 87% patients with AP its clinical signs were detected only in 12% patients on the late stages of the diseases. 21% of patients had diabetes mellitus.

The study of natural history of AP discovered that its multinominal clinical variants were due to the different stages of the disease. The stages of early clinical symptoms, relapses, complications and severe functional insufficiency of pancreas were the most distinct. The clinical stages reflect the severity of morphological disturbances in pancreas.
**1891** Success and Pitfalls in Ultrasound- and Endoscope-Guided Internal Drainage of Pancreatic Pseudocysts

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Purpose of the Study: Percutaneous ultrasound- and endoscope-guided internal drainage is a rather simple and fast method to cease pancreatic pseudocysts lying near the stomach. We performed this intervention in 72 cases, successfully in 65 ones.

Patients and Methods: Fifty-nine men and six women had been treated in this way in six years. All except for two suffered from alcoholic pancreatitis. There were pseudocysts treated with a diameter only of 3 cm or larger due to technical reasons. The intervention was followed by regular check-ups and the catheter was endoscopically removed after one year.

<table>
<thead>
<tr>
<th>Benign cysts</th>
<th>Malignancies</th>
<th>Recovery</th>
<th>Complication</th>
</tr>
</thead>
<tbody>
<tr>
<td>62</td>
<td>3</td>
<td>48 (77.5%)</td>
<td>14 (22.5%)</td>
</tr>
</tbody>
</table>

Results: The attempt to insert the catheter was unsuccessful in 7 cases (5 wrong guidances, 1 leakage, 1 vagotomy) and malignant in 3 cases (diagnosed 1-60 days after intervention). In 48 cases (77.5% of the treated 62 benign cysts) complication occurred (e.g. 8 abscess forming, 2 urinations). The amylase activities of the absceding cysts were under 10 000 IU.

Conclusions: In spite of the numerous benefits this method has some considerable drawbacks. Some of them are thought to be developing on physiological grounds (absceding of cysts with low enzyme content), others may be caused by the insufficiencies of the device (usurration, poor diagnostics of malignancies). Thus an improvement of the method and catheter seems to be necessary.

**1892** Expectant Management of Acute Biliary Pancreatitis (ABP). A Prospective Clinical Trial

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The optimum management of ABP still remains a matter of debate. Surgery during the attack is followed by unexpected high morbidity/mortality rates. Conservative management including endoscopic sphincterotomy for impacted stone removal is gaining widening popularity but only few studies have been published, mainly showing the immediate effectiveness of sphincterotomy, a procedure which could be avoided as being either unnecessary or harmful in certain cases.

In this prospective clinical trial, 39 noncasevous cases (19 males, m.a. 62 yrs & 20 females, m.a. 51 yrs) of ABP were submitted within the first 48 hrs from the onset of the attack to a strict conservative therapeutic regimen including: n.p.o., nasogastric drainage, TPN, broad spectrum antibiotics IV, gastric antisecretive medication and anticoagulants. On admission all patients had upper abdominal US and CT scans, full lower-pancreas-kidney and blood tests and close monitoring thereafter of their respiratory, renal, hepatic, pancreatic and cardiac/circulatory functions. Upper abdominal US and CT scans were repeated on the 7th day. The general condition and the seriousness of the attack were estimated independently both on admission and on the 7th day using the APACHE II scoring system and the Ranson’s scale respectively. All patients but one (38/39-97.5%) had a full clinical and biochemical recovery by the 7th, on average, day. It was found that: (a) although the seriousness of the attack is in accordance with the CT findings, it does not necessarily correspond with the general condition, (b) plasma and urine amylase levels dropped to normal by the 4th, on average, day, (c) in 38/39 patients (97.5%) there was neither respiratory nor renal function impairment, (d) in 10/39 patients (25%) the impaired liver function was normalized by the 7th day and (e) the overall morbidity rate was 15% and the mortality 2.5% (1/39 patients died of MOF). All patients underwent cholecystectomy in 8/39-21% the CBD had to be opened-searched for stone removal) 4-6 weeks after the attack had fully subsided.

The overall data in this series strongly suggest that the early start of conservative treatment in ABP can result in biological reverse, prevent further pathological pancreatic changes, diminish respiratory and renal complications and finally improve the overall outcome figures.
**1896** Ultrasonographic Abnormalities of the Pancreas in Insulin Dependent Diabetes Mellitus (IDDM) Patients

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We evaluated the pancreatic structure and the size in IDDM patients. A real time array sensor (3.5 MHz electronic transducer) was used for ultrasonographic measurements of the maximum diameters of the head, body and tail of the pancreas. The pancreatic head was measured anterior to inferior vena cava and the body of the pancreas anterior to the superior mesenteric artery. At the end of the tail was measured from its beginning near the left lateral border of the lumbar vertebral body. In this study, 25 IDDM patients (23 male, 2 female) and 27 healthy control subjects (25 male, 2 female) were studied. The diameters measured are shown in table.

<table>
<thead>
<tr>
<th></th>
<th>Pancreas</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Head (mm)</td>
<td>Body (mm)</td>
</tr>
<tr>
<td>IDDM patients</td>
<td>20.16 ± 0.78</td>
<td>9.50 ± 0.38</td>
</tr>
<tr>
<td>Control subjects</td>
<td>24.13 ± 0.12</td>
<td>12.31 ± 0.20</td>
</tr>
<tr>
<td>t Value</td>
<td>5.196</td>
<td>16.890</td>
</tr>
<tr>
<td>p Value</td>
<td>&lt;0.01</td>
<td>&lt;0.01</td>
</tr>
</tbody>
</table>

These results were not related to differences in age, sex or body size. Pancreatic image was predominantly hypoechogenic in 81% of IDDM patients. At the end of our study, we observed that in IDDM patients, pancreatic diameters were smaller than control subjects, probably due to acinar atrophy and fibrosis.

**1897** Acute and Chronic Pancreatitis Induced by Trinitrobenzen Sulfonic Acid (TNB)

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Animal models of chronic inflammatory lesions of the pancreas are scarce. TNB has been successfully employed to induce chronic colitis and cholangitis by direct infusion into the appropriate luminal segment. The mechanism of TNB chronic inflammation involves an immunological reaction to the haptenic TNB-protein complex. We hypothesized that TNB installation into the pancreatic ducts would also induce inflammatory lesions and ultimately produce a viable model of chronic pancreatic disease. Male S-D rats (300-350 gr) were ketamine anesthesized and the pancreatic duct cannulated through its duodenal opening. The duct was tied close to the liver and 0.4 ml of TNB 2% in PBS-10% ethanol (pH 8) was infused into the pancreas with the aid of a volumetric pump connected to a pressure transducer. Intrapancreatic pressure was never allowed to exceed 5 mmHg over basal. Control rats underwent the same procedure, but TNB was absent in the buffer. When infusion was completed, ligatures were released and bilio-pancreatic secretion allowed to drain for 30 min. The canula was then withdrawn and the duodenum and abdominal wall sutured. Evaluations were performed in the acute and in the chronic phase. For the acute phase rats were killed at 48 h. Serum amylase rose to 22.2 ± 1.8 U/ml at 24 h and to 7.9 ± 1.4 at 48 h, well above control levels of 5.1 ± 0.1, p < 0.05 for both time points. Pathology assessment was consistent with severe necrohemorragic pancreatitis with prominent intra and interpancreatic steatonecrosis. Spontaneous mortality rate during the acute phase was 50% (12 out of 24 rats). For the chronic phase rats were studied up to four weeks. At that end point serum amylase had returned to normal (4.7 ± 0.5) and pathology disclosed chronic changes in 43% of the surviving rats, consisting in various degrees of periductal fibrosis, patchy acute and chronic inflammatory infiltrates, common duct stenosis and signs of segmentary gland atrophy. Thus, TNB induces acute necrotizing pancreatitis that may progress towards development of chronic pancreatic disease, with features mimicking the human disease.

**1898** Study of Pancreatic Lesions in Liver Cirrhosis and Hepatocellular Carcinoma – Examination of Pancreatic Ducts

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We examined pancreatic lesions in autopsy of patients with hepatic failure caused by liver diseases. Subjects were 109 autopsies (25 patients with liver cirrhosis [LC] and 84 with hepatocellular carcinoma [HCC]). The average age of patients with LC and HCC was 55.7 ± 6.3 and 57.4 ± 7.1 years, respectively. The male-to-female ratio in the LC group was 6:19, while the ratio in the HCC was 16:68.

In LC, 10 with hepatic failure and 2 without it had pancreatic lesions: in HCC, pancreatic lesions were exhibited in 44 with hepatic failure and 5 without it. In both groups, patients with hepatic failure had frequent complications, and HCC group showed the higher frequency. There was no difference in the size of tumor and effects of treatment in HCC group. On the other hand, in patients with tumor embolus in the portal vein, pancreatic lesions were recognized very frequently (95%). Pancreatic lesions appeared more frequently with complications of portal hypertension and disseminated intravascular coagulation (DIC) than without complications. Details of pancreatic lesions: fat necrosis and hemorrhage were recognized more frequently in HCC than LC, and hemorrhage was often shown with complications of DIC. Complications of pancreatitis was recognized with portal hypertension or complications of DIC, and shown in HCC more than LC. Among patients with HCC, pancreatic lesions were often recognized with tumor embolus in the portal vein. Therefore, it is suggested that the change in portal circulation is a factor causing pancreatic lesions as well as change in the coagulating fibrinolysis system.

**1899** Does Serum Amylase Determination Give More Information Over the Diagnosis in Patients with Acute Biliary Pancreatitis (ABP)?

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**Background:** Although serum amylase determination is the main diagnostic examination in acute pancreatitis (AP), the data concerning its usefulness in the diagnosis of AP is inconsistent.

**Material and methods:** The diagnosis of AP was based on clinical picture, serum amylase levels (normal: <200 U) and positive pattern of pancreatitis in US and CT-scan.

280 patients with ABP underwent urgent (<24 h) duodenoscopy. Patients with stone impacted in the papilla of Vater were treated by endoscopic sphincterotomy (ES); (group I). Patients with normal papilla were treated by ES (group II) or managed conventionally (group III).

Serum amylase values were determined on admission and on following ten days (or longer if needed). Special attention was paid to initial amylase values and the duration of hyperamylasemia.

**Results:** In patients with stone impacted in the papilla of Vater (group I) initial amylase values (median: 1000 [IQ range: 831 to 1634]) were significantly higher than in remaining patients (median: 729 [IQ range: 491 to 1142]); p = 0.020. For levels above 1000 U, the ratio of stone impaction was 1.89 (95% CI from 1.3 to 2.7); p = 0.002. Predicted and clinical severity were not connected with any difference in admission serum amylase level.

Mean duration of hyperamylasemia was longer in patients with complicated than in uncomplicated course of pancreatitis; 4 days (IQ range: 3 to 5.5) vs. 3 days (IQ range: 2 to 4) respectively; (p = 0.0006). This was true for pancreatic (p = 0.009) as well as for extrapancreatic complications (p = 0.026).

The persistence of hyperamylasemia over 4 days was connected with 3.4 times greater risk of surgical treatment due to complications (95% CI from 1.6 to 7.1); p = 0.0005, and 3.2 times greater risk of death (95% CI from 1.2 to 8.3); p = 0.012.

The duration of hyperamylasemia also reflected the treatment used. Patients treated endoscopically (group I + III presented with significantly shorter hyperamylasemia than those managed conventionally (group III); p = 0.026.

**Conclusions:** (1) High admission serum amylase value (>5 times normal) could be useful in the prediction of stone impaction in patients with ABP.

(2) Prolonged hyperamylasemia seems to be connected with greater risk of complications and death in the course of ABP.

**1900** The Plasma Exchange as a New Therapeutic Mean in Acute Necrotising Pancreatitis (ANP)


Several experimental and clinical studies has shown that systemic complications related to ANP are caused by circulating toxic substances originating from the damaged pancreas and probably from the RES. Plasma exchange seems to be a reasonable method for diminishing the severity of Multisystem Organ Failure (MOF) being the most important cause of death in this disease.

In 8 patients went on surgery because of ANP and suffered from MOF the effect of plasma exchange was investigated in the postoperative period. The APACHE-II score, serum C-reactive protein, alpha-2-macroglobulin, and lactate-hydrogenase were determined before and after plasma exchange. There was not any complications related to plasma exchange. The APACHE-II score decreased (p <0.05) and CRP (p <0.05) level, LDH (p <0.05) activity lowered and AMG (p <0.05) level raised significantly after the procedure. We concluded that plasma exchange has probably a beneficial effect in the therapy of MOF in ANP, but further investigations are needed.
### 1901 Arginin-Induced Chronic Pancreatitis an Experimental Study


At present there is no simple model for chronic pancreatitis in the rat. Although the chronic pancreatic damage caused with L-arginine in Sprague-Dawley rats is a well recognized phenomenon, in Wistar rats, the same dose of the aminocid resulted in a surprisingly high early mortality in our preliminary study. 40 fasted male Wistar rats (b.w. 245-285 g) were used in the present study. 2 x 250 mg/100 g of L-arginine was given intraperitoneally in 1 hour interval as a 20% solution in 0.15 NaCl. The control rats received the same quantity of glycin. One glynin treated and one 250 mg/100 g arginine injected rat died. The remaining animals were killed at day 1 (8 rats), 2 (8 rats) and later at weeks 1 (8 rats), 2 (5 rats), 3 (5 rats) and pancreatic histology as well as serum amylase determinations were performed. Amylase elevations were maximal at day 1. Histology demonstrated edema, parenchymal necrosis and acute inflammatory infiltrate at days 1, 2, gradually developing to chronic subatotal atrophy with some ductular proliferation and fibrosis later on. Our modified pancreatitis model seems to be useful for examination of pathomechanism of acute and chronic pancreatitis in Wistar rats.

### 1902 Is Chronic Pancreatitis in the Aged Different?

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The existence of a form of chronic pancreatitis specific to the aged has been suggested. The aim of this work was to study the main characteristics of a population of aged patients and to compare them with a population of younger ones.

From January 1990 to December 1992, the 265 gastroenterologists working in 8 French departments have registered 596 new cases of chronic pancreatitis. The repartition of cases vs age was monophasic. So, the age of 65 (mean = 66,5) was chosen to define the aged group ("A"; n = 123) and compare it to the younger group ("Y", n = 473).

There was no significant difference for sex (male in 85.4% of group A and 85.8% of group Y), nor for the duration of symptoms (Median 7 vs 3 m respectively). In the A group there were less previous episodes of acute pancreatitis (25 vs 54%), pancreatic pain (44 vs 79%), amylase >3N (20 vs 50%), but more frequent diabetes (41 vs 27%) and steatorrhoea (51 vs 36%); pancreatic calcifications were more frequent (76 vs 62%); alcoholism was less often considered responsible for pancreatitis (58 vs 74%).

In conclusion, chronic pancreatitis diagnosed after 65 years of age is more often calcified, clinically silent except for diabetes and steatorrhoea, and less often considered of alcoholic etiology. The absence of a second incidence peak in the aged population suggest however that a specific senile pancreatitis does not exist (or is very rare), but that a more slowly progressive disease (perhaps due to a lesser daily alcoholic consumption) is predominant in this age group.

### 1903 Distribution of the Carbonyl Anhydrase Isoenzymes I, II and VI in Pancreatic Tumours. An Immunohistochemical Study

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The distribution of carbonyl anhydrase (CA) isoenzymes I, II and VI in normal and neoplastic pancreatic tissue was studied using polyclonal antisera and the immunoperoxidase technique. Samples were obtained from patients with well- (n = 4), moderately (n = 1), and poorly differentiated (n = 4) ductal adenocarcinomas, cystadenocarcinoma (n = 2), adenocystic carcinoma (n = 1), acinar adenocarcinoma (n = 1), gastrinoma (n = 3), insulinoma (n = 3) and glucagonoma (n = 1). The control specimens were from a patient with traumatic laceration of the pancreas.

The normal and malignant endocrine tissue showed intense positive staining for CA I that located in the cells expressing glucagon. In the exocrine pancreatic tissue, CA I was present in the normal and neoplastic ductal epithelium. No specific staining was detected with anti-CA VI serum neither in normal nor malignant tissue.

Our results suggest that CA isoenzyme I is abundantly expressed in the malignant exocrine pancreatic gland and to a lesser extent in the neoplastic pancreatic duct cells. These isoenzymes may thus serve as usable marker proteins for studying the origin, invasion and metastasis of pancreatic cancers.

### 1904 Results of Endoscopic Drainage in the Treatment of Chronic Pancreatitis


Abdominal pain due to chronic pancreatitis (CP) is partly attributed to poor drainage and subsequent dilatation of the main pancreatic duct (MPD). Endoscopic stenting appears to be useful at reducing MPD pressure. Required duration of drainage and long term efficacy are still uncertain. We describe the results of a standardized protocol for endoscopic management of CP.

Methods: 23 patients (pts) with abdominal pain due to alcoholic CP and stricture of the MPD were treated by endoscopy according to the following protocol: after balloon dilatation of the stricture a 17French stent was placed into the MPD and then exchanged every 2 months, the total duration of drainage being 6 months. This protocol was chosen to decrease the rate of stent occlusion and of iatrogenic ductular lesions. Pts were seen one year after stent removal (ultrasonography, wirsungography), the effect of the treatment on pain was analyzed. Specifically, we compared the difference in all pts, except in one who presented an infected pseudocyst attributed to stent occlusion. Rapidly resolving acute pancreatitis, was observed in 9 pts. Upon termination of drainage: Use of analgesics could be discontinued in 17 pts (74%). Abstinence of alcohol was observed in 20 pts, MPD diameter reduction in 19, and stricture resolution in 11. One year after stent removal: 12 pts (52%) did not use analgesics. Abstinence of alcohol was observed in 14 pts, MPD diameter reduction in 14, and stricture resolution in 8. Pain disappearance was significantly associated with reduction of MPD diameter and resolution of stricture but was not associated with abstinence of alcohol.

Conclusion: MPD stenting results in short-term clinical improvement in pts with CP and proximal MPD stricture. Persistence of advantageous clinical results is to be expected when the stricture has resolved and is observed in 50% of pts.

### 1905 Relationship Between Insulin and C-Peptide Secretion and Pancreas Morphology in Patients with Chronic Ethanol-Induced Pancreatitis

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In this study we have examined 30 patients with chronic ethanol-induced pancreatitis. The aim of this investigation has been to establish a high degree of function of endocrine pancreas dependent on chronic pancreatitis (CP) severity. Diagnosis of CP has been based on ultrasonography, ERCP, PABA-test as well as on determining of pancreatic enzymes in serum, urine and feces. According to obtained results, using by Cambridge international definition of pancreatography, all patients have been divided into three groups: one with focal CP with focal alterations (10 patients), second- CP with calcification (7 patients), third- CP with pancreatic calcification (7 patients). The fourth group have been consisted of 10 healthy control subjects. The function of beta-cells has been investigated by determination of basal values of C-peptide as well as oGTT and glucagon test. All this patients had normal basal values of insulin and C-peptide. Comparing the results of oGTT and glucagon tests no statistically significant difference has been found between control group and CP with focal alterations (p > 0.05). No statistically significant difference has been found between CP with diffuse alterations and with CP with focal alterations, too (p > 0.05).

No statistically significant difference has been found between control subjects and CP with focal alterations from the one side and CP with diffuse alterations and chronic calcific pancreatitis from the other side (p < 0.01). It can be concluded that the degree of residual beta-cells capacity depends of CP severity. The difference grows up at maximal stimulation of beta-cells secretion (glucagon test).

### 1906 Effect of Endotoxemia on the Severity of Experimental Acute Pancreatitis

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Introduction: Systemic endotoxemia is common in severe acute pancreatitis. The aim of the study was to determine changes in histology and mortality of rats with acute pancreatitis treated with different doses of endotoxin.

Methods: Male Wistar rats (mean body weight: 300 g) were used. Acute pancreatitis (AP) was induced by ligature of common pancreatic-biliary duct. Endotoxin (E) was given from E. Coli, was given intraperitoneally in 0.5 mg/kg body weight and 1 mg/100 g b w. Saline (Sa) treated and sham operated (S)-endotoxin treated animals served as control. Experimental animals were divided into five groups according to the treatments: 1. AP + saline, 2. AP + E 0.5 mg /kg, 3. AP + E 1.0 mg/kg, 4. E 0.5 mg /kg, 5. E 1.0 mg/kg. At 48 h after the treatment mortality rate was recorded and the pancreata
of the survivors were evaluated by light microscopy according to a grading scale from 0 to 4.

Results: There were no death in AP + saline, AP + E, 0.5, E + 1.0 groups, but in AP + E 1.0 group mortality was 60% (p < 0.05, chi-square test). The histologic (mean ± SD) parameters are summarized in the table.

<table>
<thead>
<tr>
<th>Groups</th>
<th>Necrosis</th>
<th>Inflammation</th>
<th>Hemorrhage</th>
<th>Edema</th>
</tr>
</thead>
<tbody>
<tr>
<td>AP + Sal</td>
<td>1.2 ± 0.2</td>
<td>1.6 ± 0.3</td>
<td>0.5 ± 0.3</td>
<td>1.8 ± 0.5</td>
</tr>
<tr>
<td>AP + E  0.5</td>
<td>1.8 ± 0.3</td>
<td>3.2 ± 0.0**</td>
<td>2.0 ± 0.5</td>
<td>2.6 ± 0.7*</td>
</tr>
<tr>
<td>AP + E  1.0</td>
<td>1.3 ± 0.3</td>
<td>2.6 ± 0.3*</td>
<td>2.3 ± 0.5**</td>
<td>2.5 ± 0.3</td>
</tr>
<tr>
<td>S + E 0.5</td>
<td>0.2 ± 0.3</td>
<td>0.6 ± 0.3</td>
<td>0.8 ± 0.7</td>
<td>0.6 ± 0.3</td>
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<tr>
<td>S + E 1.0</td>
<td>0.5 ± 0.3</td>
<td>0.6 ± 0.3</td>
<td>0.6 ± 0.3</td>
<td>1.2 ± 0.2</td>
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</table>

*p < 0.05, **p < 0.01 (Wilcoxon’s test). The numbers of animals are indicated in parenthesis.

Conclusion: AP is associated with an increased sensitivity to endotoxin. Endotoxemia may transfer acute pancreatitis into a more severe form of the disease.

1907 Non Typhic Salmonella Colitis May Induce Acute Reactional Pancreatitis

J.M. Reimund1, B. Duclos1, D. Schmitz1, M. Imler1, R. Baumann1

Infectious colitis is usually a self limited disease, when non typhic salmonella bacteria are the causative agents; however we report here five cases of acute pancreatitis (2 females, 3 males; mean age 39 years) occurring in the course of this pathological condition.

None of the 5 patients has been suffering from an other disease, none was taking any medication, and there was no history of alcohol abuse.

The diagnosis of pancreatitis was established by an increased level of serum enzymes; amylase: mean 143 U/l(range 69–249; N < 53); lipase: mean 856 U/l(range 225–1853; N < 190); and the urinary amylase excretion: mean 155 U/l(range 290–2336; N < 250). In all cases the aminotransferase and the alkaline phosphatase activities were within a normal range. Echomography and computed tomography were normal except in one case for which the pancreas was swollen with a necrotic area.

On a clinical point of view, in all cases, pancreatitis occurred as the diarrhoea began. The colitis was self limited and resolved in few days, whereas the pancreatitis had a prolonged course, and lasted several weeks. In all cases the colitis was associated with salmonella bacteria (S. typhimurium n = 2; S. braenderup n = 1; S. enteritides n = 1). Two facts deserve attention: in one case there was a concomitant arthritis; and 3 cases had the DR53 histocompatibility group.

Conclusions: Acute pancreatitis may occurred with self limited colitis secondary to salmonella bacteria. Their clinical and biological course is prolonged compared to the colitis. This pancreatitis may be secondary to immunogenous reactions as enlightened by the occurrence of arthritic reactions in some case and a possible HLA DR marker.

1908 Selective Regulation of Individual Exocrine Pancreatic Proteinases in Man


Pancreatic secretion of hydrolytic enzymes has been extensively studied, and amylase and proteinases are known to be secreted in a non-parallel fashion in man.

A more detailed examination of the feed-back regulation of different exocrine proteinases were performed using duodenal instillation of both sepa-rate and a combination of purified serine proteinases inhibitors. Activities and immunoreactive amounts of enzymes in the collected samples of duodenal juice were quantified.

Pancreatic secretion of different proteinases, amylase and lipase were demonstrated to be stimulated due to differences in inhibition profiles. Inhibi-tions of mainly trypsin activity by instillation of Köntz trypsin inhibitor (KI), resulted in an enhanced secretion of trypsin and lipase only. However, in-stillation of the more broad reacting Bowman-Birk inhibitor (BBI), resulted in enhanced secretion of trypsin, chymotrypsin and lipase together with pancreatic secretory trypsin inhibitor (PSTI). Secretion of elastase 1 and amylase were not stimulated by neither of the two inhibitors. A 100% inhibition of trypsin and a 80–85% reduction of total proteolytic activity by duodenal instillation of a mixture containing KTI, BBI and FOY-305 induced a slight increase in amylase as well. This non-parallell pancreatic secretion of exocrine enzymes might illustrate a safety system, allowing proper digestion of the specific food components stimulating pancreatic secretion and minimizing the loss of endogenous proteins.

1909 The Effects of Prophylactic Octreotide on ERCP-Induced Pancreatic Reactions

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Acute pancreatitis is the most serious complication after endoscopic cholangio-pancreatography (ERCP) and endoscopic sphincterotomy (EPT). Prevention of this complication has so far proven unsuccessful. The aim of this study was to evaluate the prophylactic effects of injecting patients due to undergo ERCP procedures with Octreotide. The study was carried out on 100 patients, randomly allocated in to two groups. The first group (50 pts.) received 0.1 mg of Octreotide s.c. one hour before the ERCP and two more doses on day 8 hours after the second group received placebo. Clinical signs of pancreatitis were noted and, before starting the ERCP, at 6, 24 and 36 hours after the end of the procedure serum amylase levels were determined.

Acute pancreatitis, in all cases mild, developed in 6% of all patients; in 8% of treated cases and in 4% of the controls (p = n.s.). 68 patients (38 controls and 30 from the treated group) underwent ERCP; acute pancreatitis was observed in 6.6% of treated patients (2/50) and in 2.6% of controls (1/38) (p = n.s.). EPT was performed on 32 patients (12 controls and 20 from the treated group); in Octreotide treated patients the incidence of acute pancreatitis was 10% (2/20) and in controls, it was 8.3% (1/12). In conclusion, the prophylactic Octreotide makes not reduce the incidence of acute pancreatitis following ERCP or EPT. More appropriate data may be obtained by studying a larger group of patients or by administering a higher dose of this drug.

1910 Differential Diagnostic Value of CA 19-9 Tumour Marker in Pancreatic Cancer

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The aim of our study was to get more information about the diagnostic value of tumour marker CA 19-9.

Methods: Serum level of CA 19-9 was determined by Immune Radiometric Assay (IRMA) in 194 patients with pancreatic malignancies or chronic pancreatitis. All patients were surgically treated. 70 patients had a final diagnosis of pancreatic cancer and 124 had chronic pancreatitis. The diagnosis was based on cytological, histology and operative macroscopic findings.

Results: In 55 of the 70 patients with pancreatic cancer CA 19-9 serum level was found elevated (above 40 U/ml), and in 15 patients it proved to be normal. In the group of patients with chronic pancreatitis normal level was found in 109 cases out of 124. The serum level was elevated in 15 patients. In the present analysis the determination of CA 19-9 provided 78.6% sensitivity and 87.9% specificity.

In this review we determined the diagnostic value of imaging techniques, too. The sensitivity and specificity provided by Ultrasonography was 78.6% and 90.3%, while CT yielded 81.6% and 83%, ERCP showed 90.7% and 95.8% respectively.

Conclusion: From these data we can conclude, that the determination of CA 19-9 tumour marker should be used in combination with imaging techniques in order to be a reliable method in the differentiation between chronic pancreatitis and pancreatic malignancy.

1911 The 14C-Xylose Breath Test in Chronic Pancreatitis: Evidence for Small Intestinal Bacterial Overgrowth

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The 1 gm 14C-xylose breath test has been found to be a highly sensitive and specific test in detecting the presence of small intestinal bacterial overgrowth. To test the hypothesis that chronic pancreatitis (CP) with exocrine insufficiency may lead to bacterial over growth we performed the 14C-xylose breath test in CP patients with and without steatorrhoea.

Methods: 10 CP patients with steatorrhoea (> 7 g fat/d), 15 CP patients without steatorrhoea, and in 14 healthy volunteers without steatorrhoea participated in the study. Breath excretion of 14CO2 after oral administration of 5 g 14C-xylose and an oral meal of unlabelled xylose in 200 ml of water was determined at 30 min intervals during 240 minutes. All subjects were studied after an overnight fast.

Results: The cumulative breath 14CO2 excretion (expressed as percentage × 10^-3) of ingested 14C-xylose expired as breath 14CO2 over 240 minutes, mean ± SEM) in controls was 522 ± 36, in CP patients without steatorrhoea 578 ± 46 (not statistically different when compared to controls), and in CP patients with steatorrhoea 863 ± 101 (p < 0.01 compared with controls and with CP patients without steatorrhoea).

Conclusion: These results suggest that patients with pancreatic insufficiency secondary to chronic pancreatitis are prone to develop small intestinal
Idiopathic Acute Pancreatitis (IAP) A Diagnostic Review

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The IAP is that one with unknown cause or etiology. Its frequency ranges between 2% and 30% in all AP. 148 AP have been diagnosed and treated prospectively from April 1987, to April 1992. At first, 18 cases were classified as IAP; their evolution was controlled from 2 to 48 months (x: 22) and their medical histories were evaluated a second time. Three non alcoholic patients had an associated to alcohol large intake, prior to the attack. Thus, they were classified as alcoholic (3/18: 16%). An asthmatic patient was having prednisone and was classified as drug-induced (1/18: 5%). Four patients had recurrences, in three of them gall-bladder microthiasis was found and the individual underwent surgery. Eleven patients (11/18: 61%) were considered as IAP (11/14: 7.4%) in a new group, but seven of them had undergone a previous cholecystectomy one or two years prior to the first attack; only four (4/18: 22%)/(4/14: 2.7%) had an unknown history causing AP. Sensibility at first classification was 61%. IAP is more severe (18%) than the gallstone pancreatitis (10%) but without mortality. In conclusion: a critical review after an initial idiopathic attack in necessary in the controlled series. The first case (unknown?) of IAP is gallbladder microthiasis, and must be meticu- lously looked for by iterates ultrasound, duodenal bile analysis and ERCP the results of this will be the right choice of the treatment to follow and the prevention of recurrences.

Acute Gallstone Pancreatitis (AGP) in Association with Acute Cholecystitis (AC)

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The association between AGP and AC needs a diagnostic and therapeutic approach because AC with hyperamylasemia is frequently seen. We studied a prospective series of 133 patients with AGP being 120 patients operated at the first admission. AGP and AC diagnosis were performed by clinical history, hyperamylasemia and/or lipasemia, ultrasound, intraoperative and pathological findings. We compared this series with other series of patients who had acute biliary abdomen without AGP: n = 64 (true negatives). We had 4 false negative (AGP without hyperamylasemia) (Sensitivity 97%) and 6 false positive (Pseudo AGP) (Specificity 94%). Finally we found 18 cases of AGP associated to AC (13%: 18/120).

Most of the patients received medical treatment for a week (7.7 ± 4.7 days) and then they were operated. Morbidity was 12% (2/16) and mortality was 0%.

AGP had more severity prognosis in 3 cases (18%) for persistent abdominal pain and leukocytosis higher than 15,000. In these cases surgery was performed between the 3rd and 4th day after admission.

In conclusion: association between AGP and AC is not exceptional (13% of certificate cases). AGP prognosis may be more severe in 18% and an early surgery may be necessary. Post-operative morbidity is similar in comparison with AGP alone (12% and 9%; NS). Our criterion of conservative treatment is safe (none mortality) for the patient with AGP and AC is the same admission.

Atypical Pattern of Main Pancreatic Duct’s Side Branches without Pathological Substrates on Anatomical and Clinical Pancreatograms

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When presented on pancreatograms, the main pancreatic duct’s branches are of great importance in diagnosing the minute pathological changes. Even so, some variations of these branches may give misleading in radiographic evaluation of the pancreas, and should be noted. Our study was based on 97 anatomical pancreatograms of human pancreas from autopsies, and 103 en- doscopic retrograde pancreatograms, both without pathological substrate. In only 3 cases (1.5%) atypical side branches were detected. The first one presented serpiginous branches, as chronic pancreatitis, localized to an upper head-body segment of pancreas. The second case involved a ductal bridge over the main pancreatic duct, as seen in pancreatitis with stenosis of the main duct. In the third case there were two vertical upward branches for the uncinate process, which resembles the picture of pathological displacement.

Conclusion: Though rare as a finding, atypical patterns of main pancreatic side branches might produce misinterpretations of otherwise normal clinical pancreatograms.

A Useful Method for the Diagnosis of Infected Pancreatic Necrosis: The Tc99m-HM-PAO-Leukocyte Scintigraphy

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Numerous attempts have been made to achieve a reliable prognostic ass- sessment in the early phase of acute pancreatitis. However, the results of these tests (clinical features, laboratory parameters, imaging techniques in several variations) have been disappointing in prospective studies. The aim of our work was to evaluate the prognostic role of Tc99m-HM-PAO-leukocyte scintigraphy in the early phase of acute pancreatitis and those with compli- cations.

Patients and Methods: 31 leukocyte scintigraphies were performed. Mixed leukocytes were labeled in vitro with Tc99m-HM-PAO. Images were taken after the biliary voids 30 minutes, 4 and 8 hours following the reinsertion of the labeled cells. In group 1, 20 consecutive patients with acute pancreatitis (at least threefold elevation of serum amylase level as well as typical findings in ultrasound and/or computed tomography) were tested. Scintigraphy was performed in all patients during the first week of hospitalisation (mean: 4.3 days). In group 2, 11 patients were tested with pancreatic pseudocyst following acute pancreatitis. Clinical features, laboratory parameters and outcome were registered during the hospitalisation.

Results: In group 1, scintigraphy showed an abnormal accumulation of leukocytes in the pancreatic area in 7 patients. Median duration of hospitalisation of these patients was significantly longer (mean: 25/16–48/48 days) than those with negative scintigram (mean: 13/5–24/48 days). 4 patients with positive leukocyte scan had an infected pancreatic necrosis confirmed by surgery and bacterial culturing, the other 3 patients were successfully treated with antibi- otics. Scintigraphy was normal in 13 patients with favorable outcome without antibiotic treatment. In group 2, among the 11 patients with pancreatic pseudo- cysts there were 4 cases with positive leukocyte scan. Pancreatic abscess was found in 3 of them during surgery. 2 out of 7 patients with normal scinti- gram, surgery revealed a noninfected pancreatic pseudocyst.

Don't Miss Eosinophilic Pancreatitis

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Among 5 cases already published and 2 new cases we observed, a preoper- ative diagnosis of eosinophilic pancreatitis (EP) was evoked only in one. The others were treated surgically as if they were cancer (3 Whipple’s procedures, 2 splenopancreatectomies, 1 palliative bypass). Simulating cancer were pic- tures of duodenal compression and stiffness (3 cases) or of irregular bile duct stenoses (2 cases) and above all the operative discovery of a very hard and en- larged pancreas. The pancreatic enlargement was total in 3 cases, left sided in 2, right sided in 1, non precised in 1.

EP is characterized by a homogenous hypoechogenic hypertyrophy on U.S., hypodense on X-Rays with regular narrowing of the pancreatic ducts and without calcification or cyst which make it quite different from the usual chronic pancreatitis. As for eosinophilic gastroenteritis of which EP is a particular extension, the diagnosis can rely neither on demography (from 21 to 70 years of age) nor on an allergic history (3 cases only) nor on blood eosinophilia, which is variable and non specific. It can rely only on histology. In 6 of the cases the massive pseudotumoral eosinophilic infiltration has been demonstrated in some place (s) of the digestive tract walls. But it is often irregularly strown among the 3 layers. Among 5 cases of EP which have had endoscopic biopsies only 2 had a positive result. That emphasises the need of multiple biopsies.

A correct diagnosis would avoid needless exereses and lead to a useful steroid therapy.

Surgical Treatment of Pancreatic Cancer. Possibilities and Pitfalls in Diagnosis

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Between January 1989 and December 1993 645 patients out of 727 with pancreatic carcinoma were treated surgically at the 1st Surgical Dept. of the Sem- melweis University of Medicine. Radical operation was performed in 16% of the cases including partial or total pancreateoduodenectomy and distal re-
section. In the rest of the patients only different types of palliative procedures could be performed.

Overall morbidity rate amounted to 6%, while hospital mortality could be kept as low as 3.4%. The retrospective analysis covered the diagnostic procedures – ultrasonography, computed tomography, ERCP, tumor marker CA 19-9 and fine needle aspiration biopsy – pre- and postoperative staging. 85% of the patients in the nonresected group had stage III and IV pancreatic cancer, 15% were classified as stage II, and were considered unresectable because of local spread. In the resected group it was surprising to find upon pathologic examination, that in spite of the resectability of the tumor 8% of the patients were classified having stage III; 35% stage II and 57% stage I cancer, respectively. Overall mean survival of the resected group was 16 months. It seems that early pancreatic cancer is a rare clinical finding. This might be due to the lack of characteristic symptoms and the uncertainty of the diagnostic procedures in our experience.

1918 Endoscopic Treatment of Pancreatic Diseases

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Introduction: Strictures, stones and pseudocyst impair the normal outflow of pancreatic juice in chronic pancreatitis (CP) and may cause pain and/or recurrent attacks of acute pancreatitis (AP). Patients with these problems must be treated. One of the possibilities is a surgical intervention a second is endoscopic treatment: pancreatic papillosphincterotomy (PPS) stenoses dilatation, stones extraction, lithotrypsy and stenting. In this non-randomised single centre study evaluated the endoscopic management of these problems.

Materials and methods: In the Clinic of Gastroenterology 40 PPS have been done. There were 27 men and 13 women with a mean age 56.6 years (range 25-85). Indication for PPS was CP 34 times (from this PPS of papilla duodenii minor was done 4 times). Stenosis of pancreatic duct with major prestenotic dilatation was seen in 16 patients with CP. After PPS was done extraction of stones or drainage (pancreatic duct or pseudocyst). The attempt of drainage of pancreatic duct was done 16 times and was successful 12 times.

Results: In 10 patients with CP after pancreatic duct drainage the pain significantly decreased in 1 patients only mild change of the pain could be seen and in 1 patient the pain was very strong and did not decrease after the drainage. Therefore he was indicated for surgery.

Conclusion: Our experiences with a small number of patients with painful CP demonstrated good results after pancreatic duct drainage. Recurrent attacks of AP after the drainage were not seen in 83.3% of our patients and their pain decreased significantly. In some patients could be seen dilatations of pancreatic three after removing of pancreatic stent.

1919 Protonene: Preconditions for its Usage in Treatment of Chronic Pancreatitis

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The cytoprotective action of prostaglandins (PG) in inflammatory lesions of pancreas, along with their ability to modulate exocrine pancreatic secretion prompted us to examine the effects of Protonene (P), PGE2 derivates, in chronic pancreatitis (CP).

52 patients with mild manifestations of CP recurrence and 19 healthy volunteers were studied. The effects of intravenous P (Tallinn pharm. factory: 0.08 mg/kg/min for 40 min) on basal and Enzaprost F-stimulated (PGF2α: 0.4 mg/kg/min) gastric HCl secretion, pancreatic enzymes’ activities, Na bicarbonate output, pancreatic juice (PJ) volume, and cyclic nucleotides (CN) levels in PJ were tested.

Administration of P in volunteers resulted in quick (in 20 min) but short-term (for 40 min) increase in Na bicarbonate production and PJ volume (p < 0.05). Enzymes activities, CN contents in PJ and HCl secretion did not change. Pretreatment with Enzaprost F caused growth of enzymatic activity in PJ and HCl secretion in both groups. This effect was abolished by P (P < 0.05), which, however, did not suppress Enzaprost-stimulated rise in PJ output and bicarbonate production. P was also found to relieve pain. Analgesia started 20–30 min earlier than the beginning of the dilatation, lasted for up to 1–2 days and was most likely related to relaxation of smooth muscles verified by ultrasonographic examination.

Thus, P appears to stimulate directly (without CN involvement) H2O and bicarbonate secretion, relieve pain and provide a “functional relaxation” for the acinar cells under stimulation. These properties of the drug may be helpful in treatment of CP patients.

1920 Vagal Tone and Antral Size in Duodenal Ulcer and Functional Dyspepsia Patients

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Patients with functional dyspepsia (FD) are characterized by low vagal tone and wide gastric antrum compared to healthy controls (C) (Scand J Gastroenter 1992; 27: 427-432, Psychosomatic Medicine 1993; 55: 12-22). The aim of the present study was to investigate vagal tone and antral size in patients with duodenal ulcer (DU). Methods: Sixteen consecutive patients with endoscopically verified DU and Helicobacter pylori infection were investigated the day after endoscopy. No patients received any ulcer treatment. All subjects were investigated fasting in the morning. The antral width was measured ultrasonographically in a standardized vertical section, and measurements of respiratory sinus arrhythmia (RSA) was calculated to index vagal tone. The results were compared to those in 25 patients with FD and 25 C similarly examined previously. Results: Vagal tone was significantly higher in DU (7.2) and C (5.7) compared to FD (1.9) (p < 0.001). DU patients tended to have higher vagal tone than C (p = 0.07). Fastig antral area was significantly wider in DU (4.8 cm²) and FD (4.6 cm²) than in C (3.4 cm²) (p = 0.01). Conclusions: DU patients are characterized by high vagal tone compared to functional dyspepsia patients. Both groups have a wide gastric antrum; possibly due to a functional disorder in FD and to a mechanical outlet obstruction in DU.

1376 Does Systemic Absorption of 5-Aminosalicylic Acid from Olsalazine (Dipentum®) and Mesalazine (Asacol® and Pentasa®) Differ Significantly in Ulcerative Colitis?

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5-Aminosalicylic acid (5-ASA), the active moiety of sulphasalazine, formulated as the azo-bonded analogue olsalazine (Dipentum®), or as delayed release or slow release mesalazine (Asacol® and Pentasa®), is effective in prevention of relapse in ulcerative colitis. However, significant systemic absorption of 5-ASA may make it less effective and may increase adverse effects.

Method: We compared the systemic absorption of D, A and P at steady state. 15 patients (age 21-69 y, 6 female) with ulcerative colitis in remission were given in a randomised block design 7-day courses of D 1.0 g/d, A 1.2 g/d, and P 1.5 g/d. Plasma and urine were collected on days 6 and 7 of each course and analyzed for 5-ASA and its main metabolite, acetylated-5-ASA, by HPLC.

**Results:**

| Mean (SEM) plasma and urine 5-ASA and Acetyl-5-ASA in 15 patients |
|-----------------|-----------------|-----------------|
| **Plasma**      | **Dipentum**    | **Asacol**      | **Pentasa**    |
| 5-ASA (µmol/L)  | 1.48 (0.35)     | 3.76 (1.27)     | 1.35 (0.21)    |
| Ac-5-ASA (µmol/L)| 3.51 (0.57)     | 7.90 (1.39)     | 5.29 (1.26)    |
| Urine 5-ASA     | 53.1 (22.8)     | 149 (40.0)      | 252 (65.7)     |
| 5-ASA (µmol/24 h)| 1180 (136)      | 1730 (132)      | 3190 (266)     |
| Total 5-ASA     | 1230 (151)      | 1880 (155)      | 3450 (272)     |
| Equimolar 5-ASA | 21.2 (2.60)     | 24.0 (1.98)     | 35.1 (2.77)    |

* p < 0.01 compared with A, p < 0.001 compared with P

**Conclusions:** 5-ASA and acetyl-5-ASA in plasma and urine during olsalazine (Dipentum) treatment were significantly lower than with the two mesalazine formulations (A and P). At therapeutic doses olsalazine appears to have the least systemic absorption at steady state.