

EARLY CHRONIC PANCREATITIS – ARE YOU MISSING IT?

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FINANCIAL DISCLOSURES

- MEDICAL ADVISORY BOARD MEMBER COOK MEDICAL
- AUSTRALASIAN PANCREATIC CLUB RECEIVES FINANCIAL SUPPORT FROM MYLAN

ROAD MAP...

- DEFINITION AND SPECTRUM OF CHRONIC PANCREATITIS
- SIGNS AND SYMPTOMS
- WHO IS AT RISK?
- HOW CAN WE IDENTIFY PATIENTS WITH ECP?
- CONSEQUENCES
 - PANCREATIC EXOCRINE INSUFFICIENCY
 - PANCREATIC MALIGNANCY
- OPTIMAL MANAGEMENT: LIFESTYLE CHANGES AND PHARMACOLOGICAL THERAPIES
- FUTURE DIRECTIONS – DIAGNOSTIC TOOL FOR GENERAL PRACTICE (APC STUDY)

DEFINITION: CHRONIC PANCREATITIS

Chronic pancreatitis is a pathologic fibro-inflammatory syndrome of the pancreas in individuals with genetic, environmental and/or other risk factors who develop persistent pathologic responses to parenchymal injury or stress

- PANCREATIC ATROPHY
- FIBROSIS
- DUCT DISTORTION AND STRICTURES
- CALCIFICATIONS
- PANCREATIC EXOCRINE DYSFUNCTION
- PANCREATIC ENDOCRINE DYSFUNCTION
- DYSPLASIA

TRADITIONAL TEACHING

- RARE - 0.5% OF POPULATION
- ALCOHOLIC OR 'CLOSET ALCOHOLIC'
- PAINFUL, STEATORRHOEA, WEIGHT LOSS
- DIAGNOSIS WITH CT, 72 HOUR FAECAL FATS



AETIOLOGY OF CHRONIC PANCREATITIS



NAPS2: A HIGHER THAN EXPECTED NUMBER OF CP PATIENTS (55%)
WERE CLASSIFIED WITH NON-ALCOHOLIC AETIOLOGY

North American Pancreatitis Study 2 (NAPS2)

Chen G, et al. Gut Gastro & Hepatol. 2011;6:66-71.

HOW COMMON IS CHRONIC PANCREATITIS?

There is evidence to suggest that pancreatic disease may exist undiagnosed in medical practice¹⁻⁴



AUTOPSY STUDY

• 1978¹ (DENMARK)



- 394 AUTOPSIES WERE CONDUCTED, WHEREBY THE PANCREAS OF EACH CADAVER UNDERWENT HISTOLOGICAL EXAMINATION
- POST MORTEM RESULTS SHOWED THAT 52 OF THE 394 (13%) EXAMINED BODIES SHOWED EVIDENCE OF CHRONIC PANCREATIC INFLAMMATION
- OF THE BODIES SHOWING EVIDENCE OF CHRONIC PANCREATIC INFLAMMATION, ONLY 2 HAD BEEN DIAGNOSED WITH CHRONIC PANCREATITIS



SCREENING ASYMPTOMATIC PATIENTS 1

• 1991² (FRANCE)



- 180 PATIENTS THOUGHT TO BE FREE OF ANY ABDOMINAL OR PANCREATIC DISEASE AGED FROM 16-83 YEARS OLD HAD THEIR DUODENAL JUICE COLLECTED
- OF THE STUDY GROUP ANALYSED, THE OLDER PATIENTS WERE FOUND TO HAVE A LOWER FUNCTIONING PANCREAS (SUGGESTING PANCREATIC FUNCTION DECREASES WITH AGE) AND 6% OF PATIENTS WERE FOUND TO HAVE PANCREATIC EXOCRINE INSUFFICIENCY (PEI)



SCREENING ASYMPTOMATIC PATIENTS 2

• 2005³ (GERMANY)



- A TRIAL TOOK STOOL SAMPLES FROM 914 PARTICIPANTS AGED BETWEEN 50-75 YEARS OLD
- THE LEVELS OF ELASTASE-1 (AN ENZYME SECRETED BY THE PANCREAS) WERE MEASURED TO DETECT PANCREATIC FUNCTION
- RESULTS SHOWED THAT 105 (12%) OF PARTICIPANTS DISPLAYED SIGNS OF PANCREATIC EXOCRINE INSUFFICIENCY (PEI) AND 47 (5%) DISPLAYED SIGNS OF SEVERE PEI
- THE FINDINGS FROM THIS STUDY SUGGEST THAT THE PREVALENCE OF PEI INCREASES WITH AGE



SCREENING IBS SYMPTOMS

• 2014⁴ (USA)



- MEDICAL RECORDS OF 2256 PATIENTS WITH IRRITABLE BOWEL SYNDROME (IBS) WERE ANALYSED FOR ABNORMAL FAECAL BIOMARKERS
- APPROXIMATELY 7% OF PATIENTS ASSESSED WERE FOUND TO HAVE LOW FAECAL ELASTASE LEVELS WHICH INDICATES PANCREATIC EXOCRINE INSUFFICIENCY (PEI)
- THE FINDINGS OF THIS STUDY SUGGEST THAT SOME PATIENTS DIAGNOSED WITH IBS MAY HAVE A MORE SINISTER, UNDERLYING CONDITION



SCREENING SYMPTOMATIC PATIENTS

• 'FUNCTIONAL DYSPEPSIA'

- 22% OF SMOKERS WITH EPIGASTRIC PAIN NOT RESPONSIVE TO PROTON PUMP INHIBITORS
- CHRONIC PANCREATITIS CHANGES ON EUS

• 'TYPE 1,2 DIABETICS'

- CLINICAL RECORDS OF 1,922 PATIENTS WERE RETROSPECTIVELY EXAMINED FOR FEATURES OF T1/T2 DM, OR PE/CP ON IMAGING
- 157 (8%) WERE RECLASSIFIED AS TYPE 3C DIABETES

Dominguez Muñoz, Personal data, Harth Diabetes Care 2008

EVIDENCE FOR UNDIAGNOSED PANCREATIC EXOCRINE INSUFFICIENCY IN THE GENERAL POPULATION

Study author	Patient characteristics	Country	Patient numbers	Method of investigation	Result
Olsen T, 1979 ¹	Post mortem	Denmark	394 autopsies	Histology	13% pancreatic inflammation
Laugier R, et al. 1991 ²	Healthy persons aged 18 to 83 years	France	180 prospective	Direct secretion, CCK test	6% PEI
Rotherkugel D, et al. 2003 ³	General population aged 50-75 years	Germany	214 prospective	Fa-1-200mg/gg	11.5% PEI
Herzig K, et al. 2011 ⁴	Persons aged 60-80 years	Germany	108 prospective	Fa-1c 200mg/gg	21.7% PEI
Leeds J, et al. 2010 ⁵	IBS-D	United Kingdom	214 prospective	Fa-1-1000mg/gg + CT scan	6.1% severe PEI 21% CP
Goepel J, et al. 2014 ⁶	IBS related symptoms	USA	256 retrospective	Fa-1-200mg/gg	7.1% PEI
Comford J, et al. 2014 ⁷	Diagnosis, abdominal pain, weight loss	United Kingdom	1027 retrospective	Fa-1-1000mg/gg CT, MRI, US scan	11.4% PEI 33.1% CP or PC

1. Olsen T. Acta Path Microbiol Scand. 1979; Sect A. 86:341-366. 2. Laugier R, et al. Digestion. 1991; 52:200-213.3. Rotherkugel D, et al. Scandinavian J Gastroenterol. 2003; 38:609-614. 4. Herzig K, et al. BMC Gastroenterol. 2011; 11: 3. 5. Leeds J, et al. Clin Gastroenterology and Hepatology. 2010; 8:834-838. 6. Goepel J, et al. Scand J Gastroenterol. 2014; 49:19-25. 7. Comford J, et al. Gut. 2014; 63 A35.

DOES IT MATTER?

THE CONSEQUENCES OF UNDIAGNOSED PANCREATIC DISEASE



1. Schneider D, et al. Am J Gastroenterology. 2005; 100:1930-1935. 2. Comford J, et al. Clin Gastroenterology. 2014; 63:1201-1203. 3. O'Brien J, et al. Pancreas. 2010; 39:82-84. 4. Durrig V, et al. Am J Gastroenterology. 2006; 101:1540-1544. 5. Tiel P, et al. Gastroenterology. 2007; 132:1847-1852. 6. Comford J, et al. Gastroenterology. 2014; 136:1212-1217. 7. Durrig V, et al. Clin Gastroenterology and Hepatology. 2007; 5:1272-1276. 8. Comford J, et al. Gut. 2014; 63:1201-1203. 9. Comford J, et al. Gut. 2014; 63:1201-1203. 10. Comford J, et al. Gut. 2014; 63:1201-1203. 11. Parola A, et al. Cancer. 2014; 115:202-209.

WHY MISSED:

- CLINICAL SYMPTOMS AT EARLY STAGE ARE NON-SPECIFIC
- MAINSTAY OF INVESTIGATIONS POORLY SENSITIVE IN EARLY DISEASE



Need a high index of suspicion

HISTORY

- ALCOHOL
- SMOKING
- GENETICS
 - CF
 - HEREDITARY PANCREATITIS
 - PRSS1
 - SPINK1
 - CTRC
- DUCTAL OBSTRUCTION
- SYSTEMIC DISEASES
 - SLE
 - HYPERTRIGLYCERIDAEMIA
 - HYPERCALCAEMIA
 - COELIAC
- AUTOIMMUNE
- TROPICAL

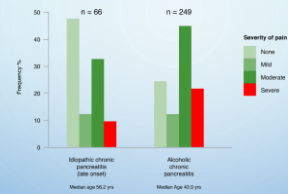
SIGNS AND SYMPTOMS IN CHRONIC PANCREATITIS

- ABDOMINAL PAIN
- PANCREATIC INSUFFICIENCY
 - MALABSORPTION
 - WEIGHT LOSS
 - DIARRHOEA
 - NUTRIENT DEFICIENCIES – PARTICULARLY FAT-SOLUBLE VITAMINS (ADEK)
- TYPE IIIC DIABETES

PAIN

- USUALLY UNRELATED TO MEALS
- EARLY – PAIN IN DISCRETE ATTACKS; CAN BE MISTAKEN FOR ACUTE PANCREATITIS, NUD
- LATE – CONTINUOUS, CLASSIC
- DOES NOT IMPROVE OVER TIME
 - AMMANN, GASTROENTEROLOGY, 1999, MULLADY GUT 211

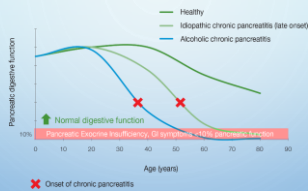
WHY DOES CHRONIC PANCREATITIS EXIST UNDETECTED?



- 90% OF PATIENTS WITH IDIOPATHIC CHRONIC PANCREATITIS PRESENTS WITH MILD, MODERATE OR NO PAIN SYMPTOMS

Leppä P. et al. Gastroenterology, 1988, 107:1481-1485

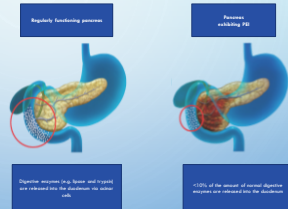
PEI AND CHRONIC PANCREATITIS



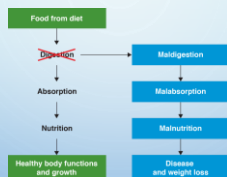
Adapted from Leppä P. et al. Digestion 1987; 50:202-211, Srinivasan S. et al. Scandinavian Journal of Gastroenterology 2000; 41:827-834, Miller R. et al. International Journal of Pancreatology 1989; 2:437-450

WHAT IS PANCREATIC EXOCRINE INSUFFICIENCY (PEI)?

- PROGRESSIVE LOSS OF ACINAR CELLS
- INSUFFICIENT SECRETION OF DIGESTIVE ENZYMES INTO THE DUODENUM¹
 - FUNCTIONALITY OF THE PANCREAS DROPS BELOW 10%¹⁻³
- FOOD IS NOT DIGESTED PROPERLY AND THE NUTRIENTS THAT THE BODY REQUIRES ARE NOT ABSORBED – THIS CAN LEAD TO MALNUTRITION AND OTHER COMPLICATIONS (SUCH AS OSTEOPOROSIS)⁴



CLINICAL CONSEQUENCES OF PEI



SYMPTOMS OF PEI

- A COMBINATION OF ANY OF THE FOLLOWING SYMPTOMS IS AN INDICATION THAT A PATIENT COULD HAVE PEI⁴
 - GAS AND BLOATING
 - FOOD NOT BEING DIGESTED PROPERLY CAN LEAD TO BLOATING IN THE ABDOMEN
 - ABDOMINAL PAIN
 - GAS PRODUCED BY FOOD NOT BEING DIGESTED FULLY CAN LEAD TO ABDOMINAL PAIN
 - FREQUENT DIARRHOEA
 - CAUSED BY UNDIGESTED FOOD MOVING TOO QUICKLY THROUGH THE DIGESTIVE TRACT
 - WEIGHT LOSS
 - DUE TO THE IMPAIRED DIGESTION AND ABSORPTION OF FATS, PROTEINS AND CARBOHYDRATES, WEIGHT LOSS OFTEN OCCURS IN INDIVIDUALS WITH PEI
 - STEATORRHOEA
 - OILY, FOULING AND VERY FOUL SMELLING STOOLS THAT ARE DIFFICULT TO FLUSH – THIS IS DUE TO EXCESS FAT BEING EXCRETED IN THE FECES DUE TO POOR FAT DIGESTION AND ABSORPTION
 - MUSCLE WEARINESS
 - THE LACK OF PROTEIN BEING PROPERLY DIGESTED AND ABSORBED MEANS THAT THE AN INDIVIDUAL WITH PEI CAN HAVE WEAK MUSCLES

INVESTIGATIONS


RECOMMENDATIONS TO INVESTIGATE FOR PANCREATIC DISEASE AND PEI IN PATIENTS PRESENTING WITH CHRONIC DIARRHOEA SYMPTOMS

Britain 2003	USA 2012	Australasia 2015
<p>British Society of Gastroenterology¹ Guidelines for Investigation of chronic diarrhoea, 2nd Edition</p> <p>"Diarrhoea may result from: a) Bile salt malabsorption (intestine); b) Small bowel inflammation; c) Small bowel malabsorption; d) Malabsorption due to pancreatic insufficiency or primary disorders, and it can be difficult to separate these on clinical grounds."</p>	<p>Mayo Clinic, Rochester, USA² Evaluating the Patient With Diarrhea: A Case-Based Approach</p> <p>"When fatty diarrhea is identified, the initial goal is to distinguish malabsorption from malabsorption." The evaluation focuses on looking for a structural problem resulting in the small intestine or pancreas."</p>	<p>Australasian Pancreatic Club³ Australasian treatment guidelines for the management of pancreatic exocrine insufficiency</p> <p>"PEI may occur in patients with exocrine-pancreatic insufficiency (EPI). Treatment with PEI may reduce diarrhoea and abdominal pain (Level 2b) -"</p>


1. Thomas G, et al. Gut 2005; 54(Suppl 1):i1-i15. 2. Pancreatology 2012; 12(4):461-467. 3. Smeaton S, Mayo Clin Proc 2015; 90(5):585-592

IS EARLY DETECTION OF CHRONIC PANCREATITIS POSSIBLE?


Endoscopic ultrasound (EUS)



MRI



Computed tomography



Lermon A et al. Cancer Research; 74 (13); 1-9, 2014

AMERICAN PANCREATIC ASSOCIATION GUIDELINES FOR DIAGNOSIS OF CHRONIC PANCREATITIS

If inconclusive or non diagnostic Proceed to next step

- Step 1: Clinical signs and symptoms of chronic pancreatitis (abdominal pain, weight loss, malabsorption, diabetes mellitus)
- Step 2: CT scan (Calcification in combination with atrophy and/or dilated duct)
- Step 3: MR/MRCP with secretin enhancement (Carriage Class III dilated duct, atrophy of gland, filling defects in duct suggestive of stones)
- Step 4: Endoscopic Ultrasound (EUS) (≤ 5 EUS CP criteria)
- Step 5: Pancreas function test (with secretin) (Fecal elastase-1 < 80 mcg/g)

Endoscopic Retrograde Cholangiopancreatography (ERCP) (Carriage II, dilated main pancreatic duct and ≥ 2 dilated side branch)

Chronic pancreatitis

Adapted from Cornwell D, et al. Pancreas 2014; 33(1):151-152

HOW IS PEI DETECTED?

DIRECT TEST:

- SECRETIN-CAERULEIN TUBULAR TEST

INDIRECT TESTS:

- 3 DAY FAECAL FAT TEST (GOLD STANDARD)
- FAECAL ELASTASE-1 STOOL TEST**
- 13C MIXED TRIGLYCERIDE BREATH TEST

FAECAL ELASTASE-1 STOOL TEST

- FAECAL ELASTASE-1 TEST IS BECOMING MORE COMMON IN CLINICAL PRACTICE¹
- IN 2010, IT WAS REPORTED TO BE THE MOST POPULAR TEST USED TO EVALUATE PEI²
- REQUIRES A SINGLE FORMED STOOL SAMPLE²
- MEASURES THE ELASTASE CONCENTRATION IN THE STOOL^{1,2}
- SPECIFICITY: APPROXIMATELY 93%¹

>200 µg/g stool: normal value²

<200 µg/g stool: mild PEI¹

<100 µg/g stool: severe PEI¹

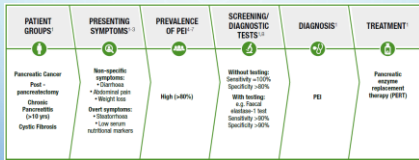
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1. Todd J, et al. MAJ 2010; 2. Smeaton S, et al. Best Pract Res Clin Gastroenterol 2010; 24(5):537-547. 3. Loner C, et al. Gut 1996; 39: 386-388

PEI DIAGNOSTIC PATHWAY

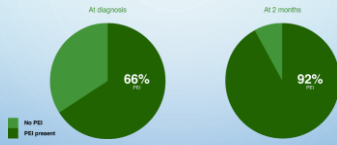


1. Todd J. et al. *Gut* 2010; 59:461-467. 2. Thomas P. et al. *Gut* 2009; 58:1661-1665. 3. Larkin J. et al. *Pancreatology* 2010; 10: 360-365. 4. *Journal of Pancreatic Research and Practice* 2010; 14: 101-104. 5. *Journal of Gastroenterology and Hepatology* 2010; 25: 101-104. 6. *Journal of Gastroenterology and Hepatology* 2010; 25: 101-104.

A FEW WORDS ABOUT PEI AND PANCREATIC CANCER...

PEI AND PANCREATIC CANCER

PEI PREVALENCE IN UNRESECTABLE PANCREATIC CANCER



Prospective study in 32 patients with newly diagnosed unresectable cancer of pancreatic head region recruited between 2010 and 2012, followed up for at least 6 months. PEI determined using faecal elastase-1 stool test <200 mcg/g.

Sharma S. et al. *J Clin Oncol* 2014; 32: 1052-1056.

QUALITATIVE RESEARCH: IMPACT OF PEI ON QOL IN UNRESECTABLE PANCREATIC CANCER

Participants identified that their priority cancer supportive care need was their difficulty in managing gastrointestinal problems, diet and digestion. They expressed strong feelings of frustration and anger related to struggling with symptoms of PEI.

"I found this very confronting, her not wanting to eat. I (crying) try to force food in. She gets upset, it makes it worse." - Carer, Male, 60yrs

"I just can't get enough nutrition you know..." - Patient, Female

"You've got this lack of food intake, I can't eat as much of what I used to and there are things that I can't eat." - Patient, Female

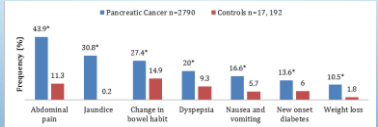
Qualitative research conducted by the NSW Cancer Council, in 61 people including patients diagnosed with pancreatic cancer, carers, family members, and bereaved participants.

Geelan H. & White K. *Support Care Cancer* 2012.

A SUBSTANTIAL TIME LAG OCCURS BETWEEN SYMPTOM PRESENTATION TO DIAGNOSIS OF PANCREATIC CANCER

UK case control study involving 2773 patients with pancreatic ductal adenocarcinoma (PDAC)
 Mean number of visits to GP = 18
 Mean number of "alarm" (not specific) symptoms = 11

Symptom presentation to primary care in UK, within 2 years prior to diagnosis of pancreatic cancer



Adapted from Keane et al. *BMJ* 2014
 * Pancreatic cancer versus controls, P<0.001
 # Symptoms with greatest frequency (>10%) shown

Keane M. et al. *BMJ* 2014; 349: g6570.

MANAGEMENT

LIFESTYLE MODIFICATION

- AVOID EXPOSURE TO OBVIOUS RISK FACTORS
 - ALCOHOL
 - SMOKING
- ENZYME SUPPLEMENTATION (PERT)
- SMALL MEALS FREQUENT MEALS

ADDITIONAL MANAGEMENT

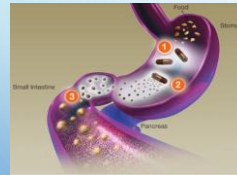
- ANALGESIA
 - SIMPLE
 - NEUROLEPTICS EG TCA, GABAPENTOIDS
- DECOMPRESSION
 - ENDOSCOPIC
 - SURGICAL
 - ESWL

PERT TREATMENT GOALS¹

- ELIMINATE MALDIGESTION
- ELIMINATE MALABSORPTION
- MAINTAIN ADEQUATE NUTRITION

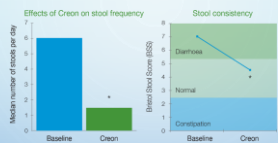
¹ Troost et al. *Ann Intern Med*. 2012; 156:480-487

DELIVERY OF PANCREATIC ENZYMES USING A MODERN ORAL FORMULATION



1. The capsule containing pancreatic enzymes enters the stomach along with the food.
2. The outer capsule dissolves rapidly to release enteric coated mini-microspheres
3. The mini-microspheres pass through the pylorus together with the chyme.
4. The active digestive enzymes are released in the duodenum to digest nutrients.

EFFECTS OF PERT ON GI SYMPTOMS (PEI IN CHRONIC PANCREATITIS)



¹ $p < 0.002$ vs. 888-C patients with normal fat levels
 Open label study, $n = 19$, to 12 weeks, all patients with PEI: FE-1 < 100 $\mu\text{g/g}$ stool.
 Dose: Creon 30,000 units three times daily.

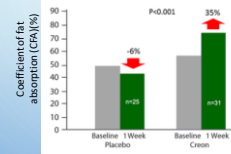
Patients on pancreatic enzyme therapy (Creon) showed a clinically significant response.

- Stool frequency reduced from 6 to 1.5 per day ($p < 0.002$)
- Stool consistency changed from "diarrhoea" to "normal" ($p < 0.002$)

Observational study involving 19 patients identified with PEI using the faecal elastase-1 stool test. Patients were treated with Creon 30,000 units three times daily for 12 weeks. Patient responses were measured using Bristol Stool scale and the number of stools per day.

EFFECT OF PERT ON FAT MALABSORPTION IN PATIENTS WITH PANCREATIC EXOCRINE INSUFFICIENCY POST PANCREATIC SURGERY

A RANDOMISED, DOUBLE BLIND, PLACEBO CONTROLLED STUDY OVER 7 DAYS

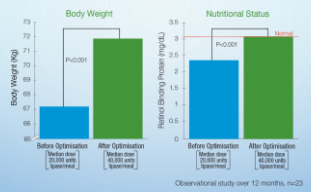


58 patients with pancreatic exocrine insufficiency after pancreatic resection due to malignancy or chronic pancreatitis.

PERT SIGNIFICANTLY IMPROVES FAT ABSORPTION BY 35% AFTER ONE WEEK.

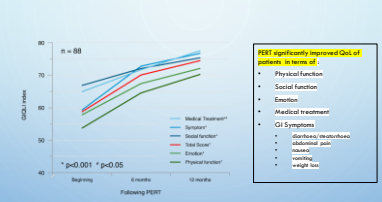
Seller C, et al. *Aliment Pharm & Ther*. 2013

EFFECTS OF PERT ON BODY WEIGHT AND NUTRITIONAL STATUS



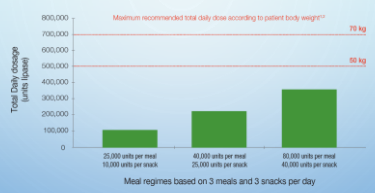
Observational uncontrolled study involving 20 patients with pancreatic exocrine insufficiency associated chronic pancreatitis. At baseline, these patients were treated with pancreatic enzyme replacement therapy with the dosage titrated to avoid the symptoms of weight loss and diarrhoea. The dosage was increased and titrated to achieve normal fat absorption as determined by 13C-mixed triglyceride breath test. The treatment was continued for one year.

EFFECTS OF PERT ON THE QUALITY OF LIFE (QOL) OF PATIENTS WITH CHRONIC PANCREATITIS



Observational prospective, multicentre study which compared response and quality of life over one year to patients with chronic pancreatitis treated with chronic enzyme therapy, without prior pancreatic resection. Quality of life was measured using questionnaire quality of life index (QOL)

PERT ADULT DOSE RANGE



Maximum dose recommended by:
 1. Cystic Fibrosis CFQ Guidelines Committee, Cystic Fibrosis Foundation
 2. Australian Guidelines for Management of Pancreatic Exocrine Insufficiency (Eckardt, et al. 2002, 1999, 1997-1991)

MANAGEMENT OF PEI USING PERT

- PERT:**
- INITIALLY – 25,000 to 40,000 UNITS LIPASE WITH EACH MEAL
 - ENCOURAGE PATIENTS TO EAT 6 SMALLER MEALS PER DAY RATHER THAN 3 LARGE MEALS
 - IF REQUIRED, INCREASE DOSE UP 80,000 LIPASE UNITS WITH EACH MEAL
- OTHER CONSIDERATIONS:**
- PATIENT COMPLIANCE
 - MANY PATIENTS HAVE ACIDIC INTESTINAL PH WHICH DECREASE ENZYME RELEASE FROM PREPARATIONS WHICH HAVE PH SENSITIVE ENTERIC COATING
 - ACID SUPPRESSING AGENTS MAY BE REQUIRED IN SOME PATIENTS
 - LACK OF WEIGHT GAIN DUE TO INADEQUATE FAT INTAKE
 - FAT RESTRICTION NOT REQUIRED WITH PERT

SUMMARY

- COMMON
- SIGNIFICANTLY UNDER-DIAGNOSED
- UNRECOGNISED DISEASE CAN LEAD TO POOR QOL, HEALTH COMPLICATIONS AND REDUCED SURVIVAL
- SYMPTOMS CAN BE NON-SPECIFIC
- NEED TO THINK OF IT!!!

FUTURE DIRECTIONS

- APC INITIATED DEVELOPMENT OF A DIAGNOSTIC TOOL
- BASED ON WORK DONE IN UK FOR PANCREATIC CANCER
- BROADENED TO ALSO INCLUDE
 - CHRONIC PANCREATITIS
 - PANCREATIC EXOCRINE INSUFFICIENCY
- PHASE 1 DEVELOPMENT PHASE (CURRENT)
 - LITERATURE REVIEW
 - DELPHI PROTOCOL
- PHASE 2 - VALIDATION PHASE
 - GENERAL PRACTICES TO UTILIZE PROSPECTIVELY