Frightening Flare-Ups: Tackling Chronic Pancreatitis in Primary Care

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Disclosures

- I have no financial disclosures
- I am a "burnt out" Primary Care Physician



Objectives

•Explore medical strategies for managing pain in Chronic Pancreatitis.

- •Examine surgical and endoscopic treatment options for Chronic Pancreatitis.
- Review the diagnosis and management of Exocrine Pancreatic Insufficiency (EPI).



What is Chronic Pancreatitis?

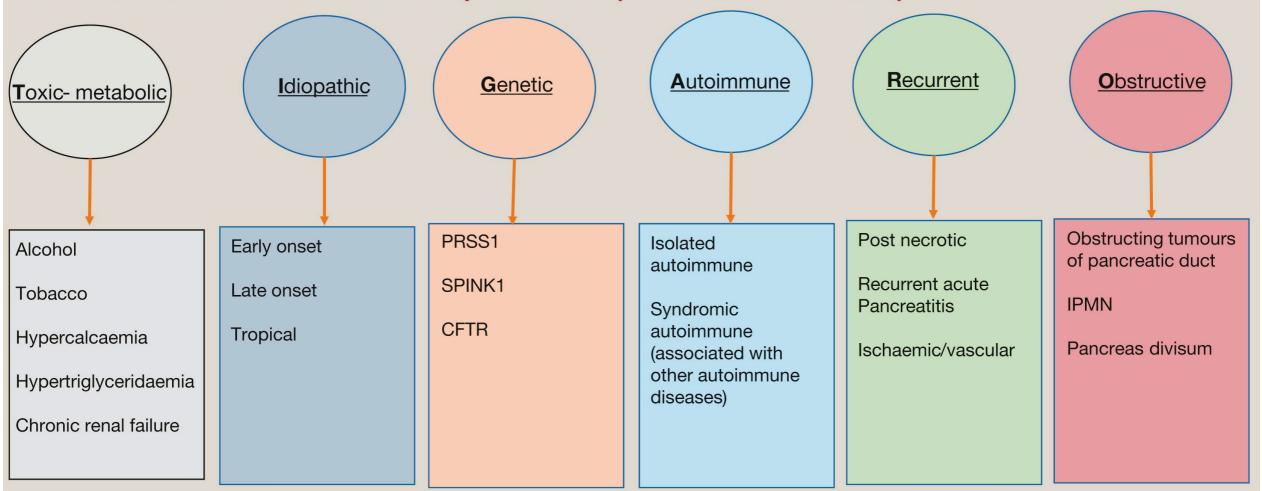
Chronic pancreatitis is a pathologic fibro-inflammatory syndrome of the pancreas in individuals with genetic, environmental and/or other risk factors who develop persistent pathologic responses to parenchymal injury or stress



Epidemiology

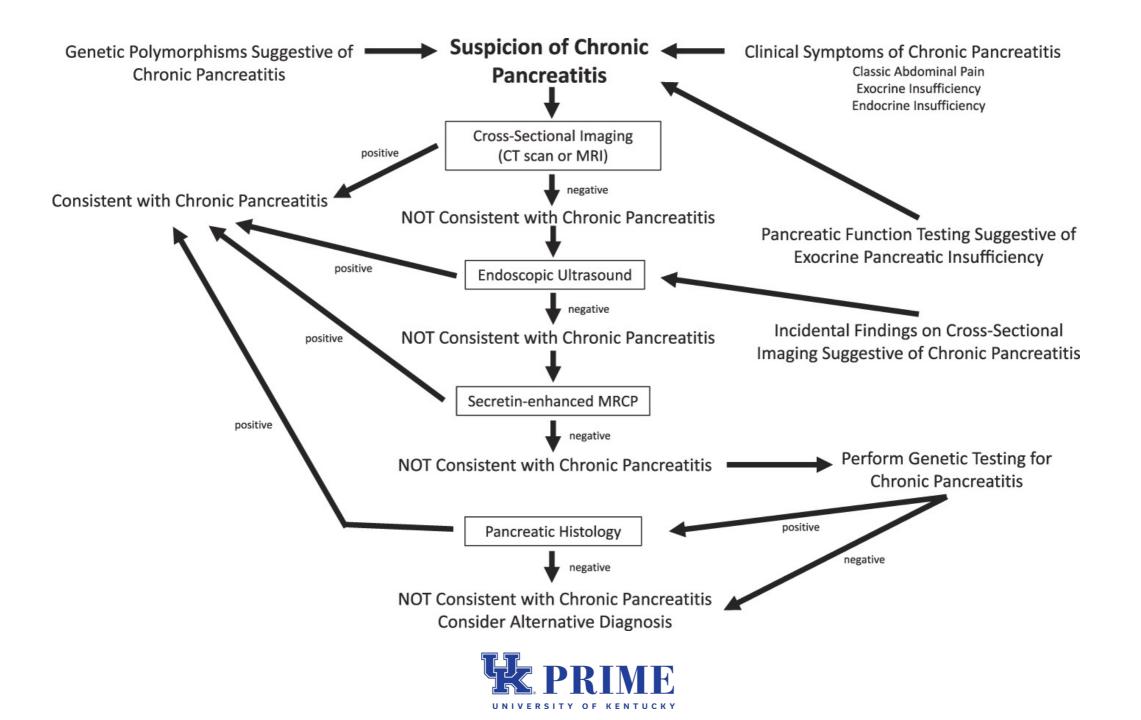
- The incidence of chronic pancreatitis is approximately 8/100,000 with a prevalence of about 50/100,000.
- Male Predominance
- Middle age around 40

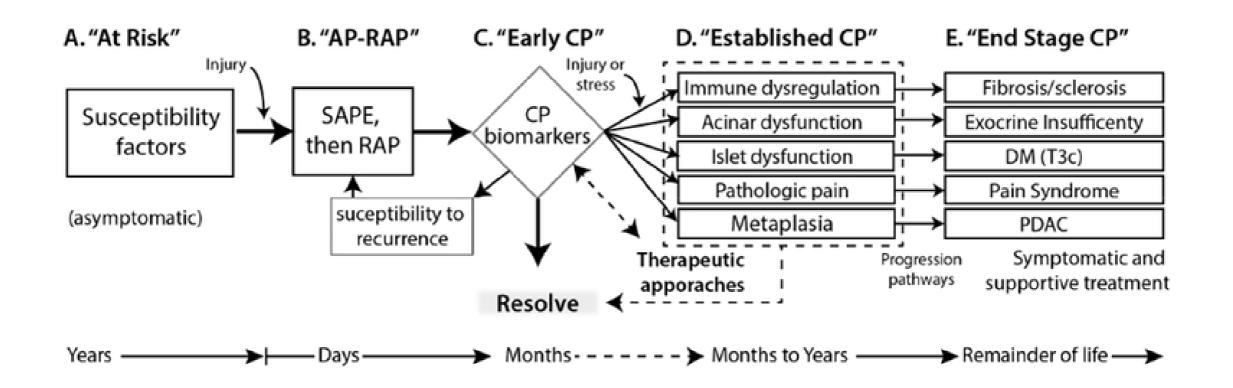


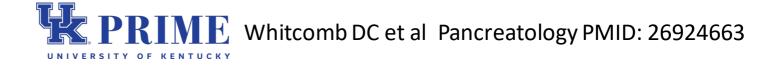


Risk factors associated with chronic pancreatitis (TIGAR-O classification)









Management of Chronic Pancreatitis

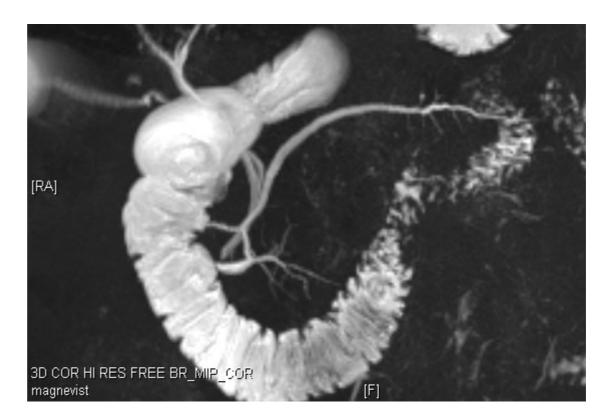


Patient A

45 yr old female

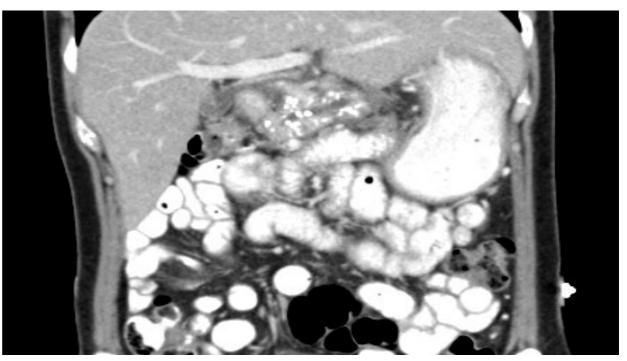
Chronic abdominal pain

On opiates



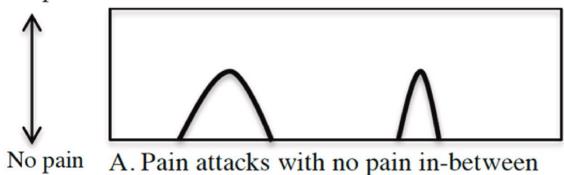


40 year old male Chronic Fluctuating abdominal pain

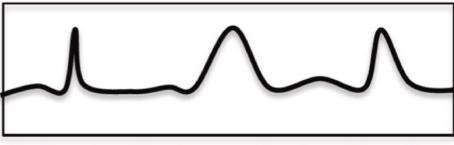




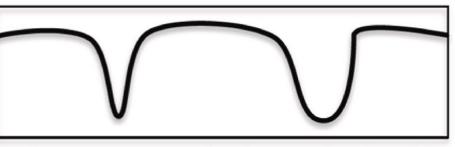
Worst pain



B. Constant pain daily



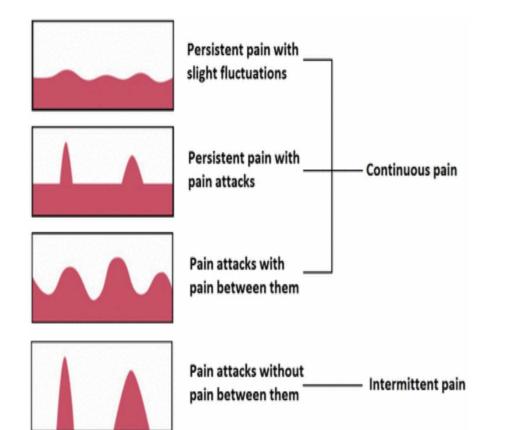
C. Constant background pain with pain attacks



D. Severe constant background pain with reduced pain periods in-between



Tuck et al Pancreatology 2022

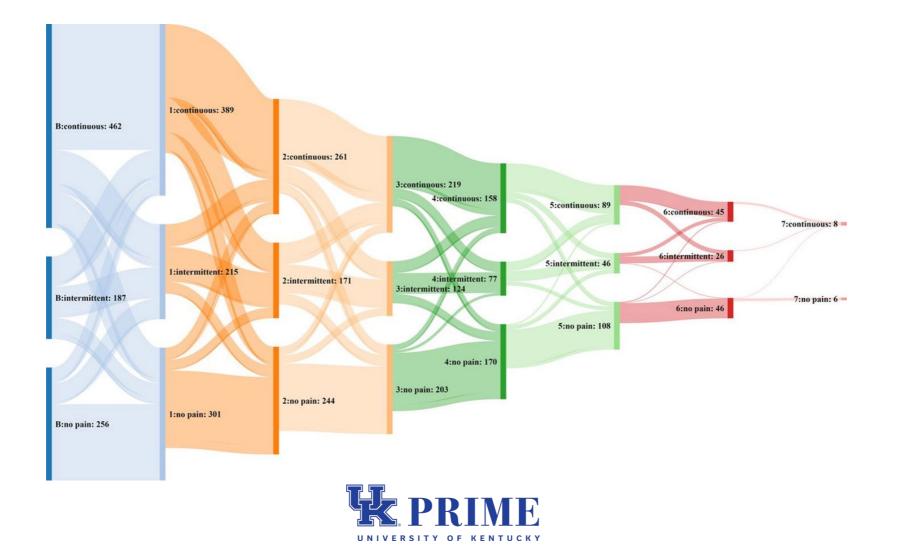


Continuous and intermittent pain patterns in chronic pancreatitis do not seem to be the result of distinctly different pathophysiological entities. The subjectively reported character of pain is not related to imaging findings or disease duration. Pain patterns often change over time and are merely a feature of how severity of pain is experienced.



Kempeneers at al Gut 2021

Pain Patterns change over time



One in ten patients with CP have primary painless disease.

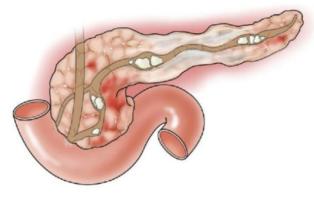
Majority of painless chronic pancreatitis attributable to an idiopathic/genetic etiology.



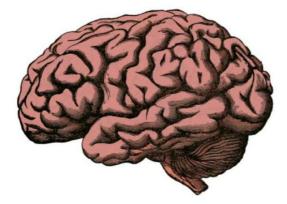
Bhullar et al. Pancreatology 2022

Overlap and cumulative effects of pancreatic duct obstruction, abnormal pain processing and psychological distress on patient-reported outcomes in chronic pancreatitis

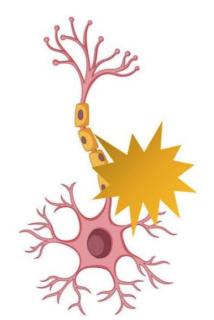
Pancreatic duct obstruction



Psychological distress

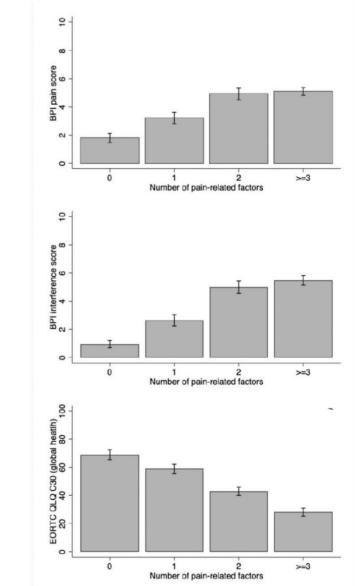


Neuronal sensitization

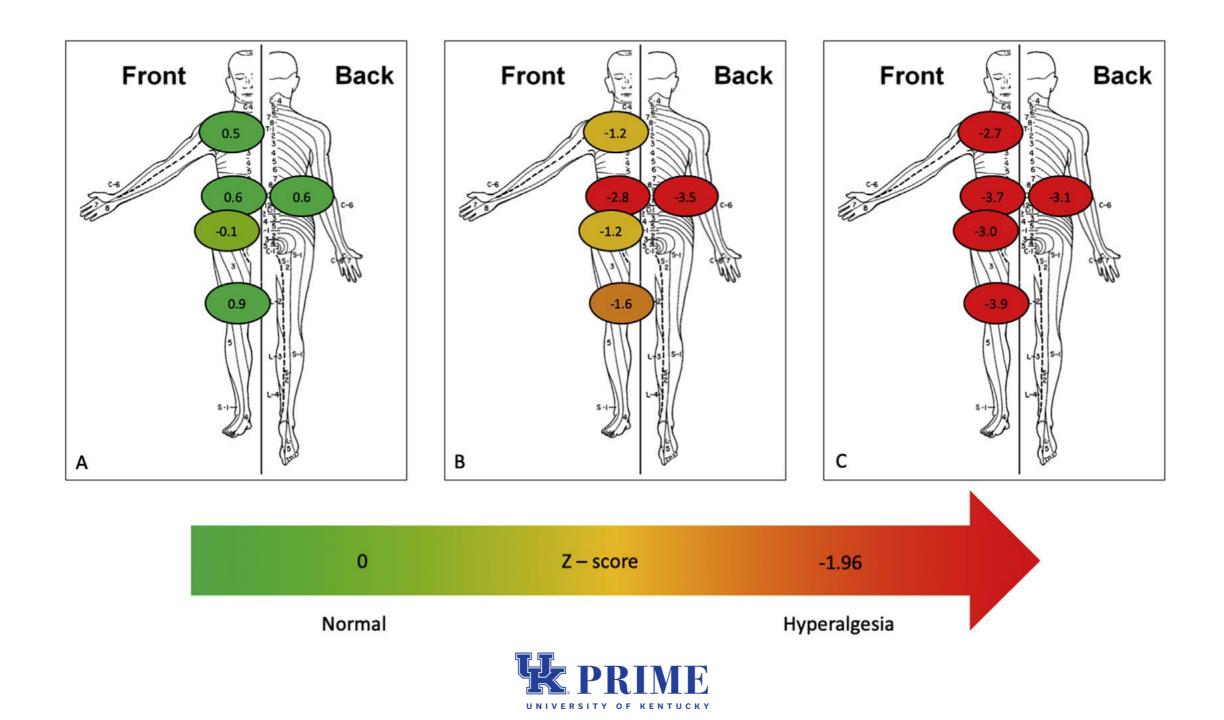


- Anxiety
- Depression
- Pain catastrophising

Olesen SS, et al. Gut 2021; doi: 10.1136/gutinl-2021-325855



Gut



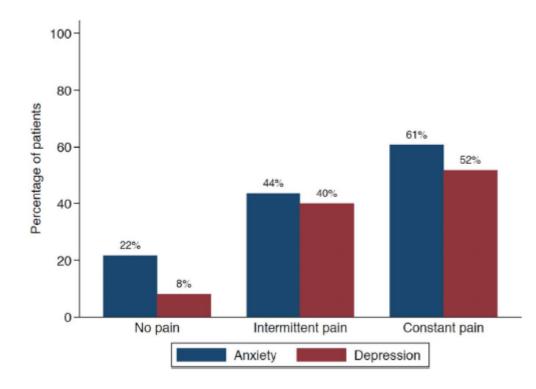
High prevalence of central sensitization in painful CP

2.A 2.B 100 100 -80 80 Percentage of patients Percentage of patients 60 60 40 40 20 20 0-0 Painful CP No current pain Painless CP No Hyperalgesia Segmental Hyperalgesia Widespread Hyperalgesia No pain Intermittent pain Constant pain No Hyperalgesia Segmental Hyperalgesia Widespread Hyperalgesia



Faghih at al Clinical Gastro Hep 2022.

Prevalence of Anxiety and Depression in CP



High Prevalence

- Anxiety : 46.8%
- Depression: 38.65
- Both: 29%

Chronic pancreatitis patients with symptoms of anxiety and depression are more likely to report

- Pain
- Pain of higher severity
- More pain interference with their lives



Philips at al Am J Gastro, 2020

Analgesic Ladder

Chronic non-cancer pain (CNCP)

Adjuvant analgesic STEP 3 Minimal invasive intervention +Non-opioid analgesics +Adjuvant analgesic +Adjuvant analgesic +Adjuvant analgesic +Adjuvant analgesic +Adjuvant analgesic +Adjuvant analgesic

STEP 1

Non-opioid analgesic + Adjuvant analgesic



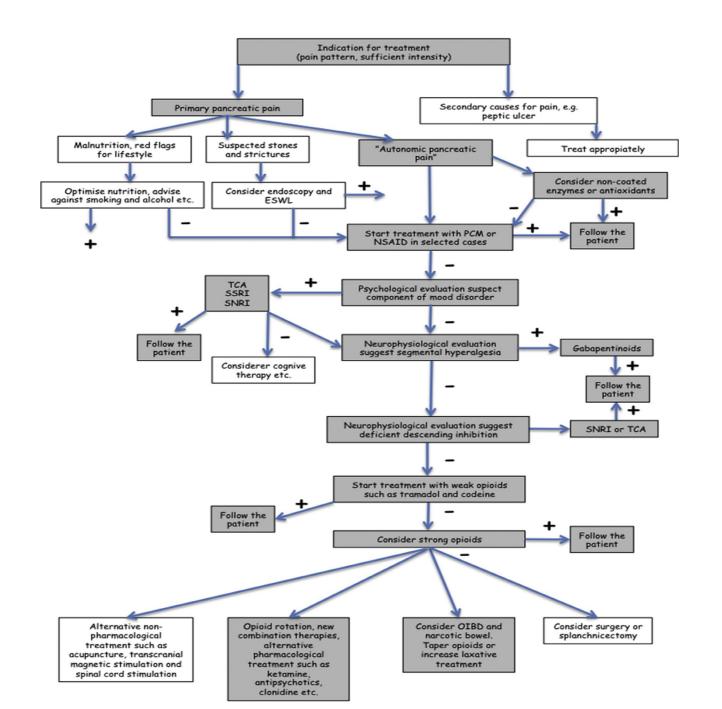




STEP 4

Strong-opioid analgesic

+Non-opioid analgesic

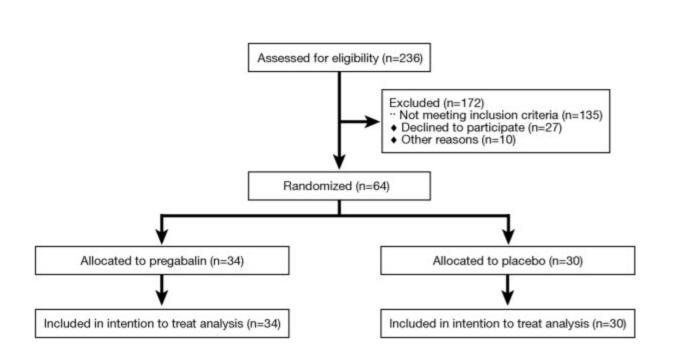


Adjunctive Therapies

- Psychiatric therapy for concomitant disorders
- Sleep aid for insomnia
- PERT for EPI
- GABA-ergic medications
- Antioxidants
- SNRI
- SSRI
- Procedural pain interventions



Philips AE, pancreas 2023



Pregabalin



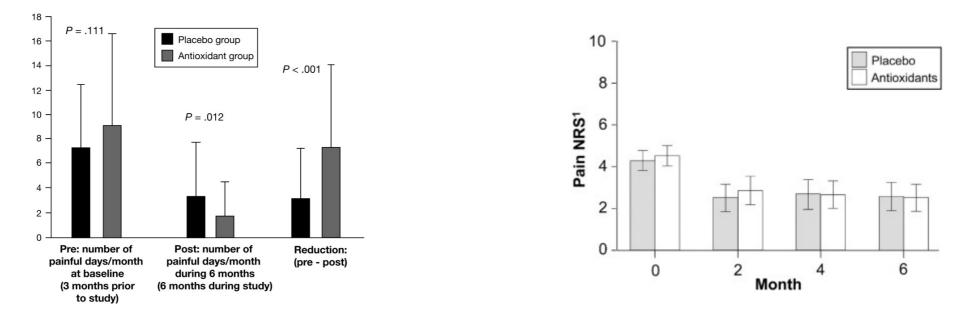
- Over three weeks pregabalin reduced daily average pain score by 36% compared to 24% in placebo group.
- Most patient were also on opiates to the effect of pregabalin was independent
- Side effects were common



Olesen et al, Gastroenterology 2011

Antioxidants show mixed results in CP

- Consist of Vitamin A,C, E, selenium and methionine
- Benefits were seen in younger patient with a non alcohol etiology

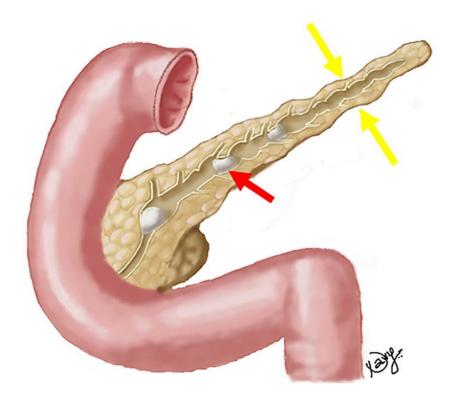


Bhardwaj et al Gastroenterology 2009 **The DRIME** Siriwardena at al Gastroenterology 2012

UNIVERSITY OF KENTUCKY

Pain Management with endoscopy

• Remove obstruction (stone, stricture) in the pancreatic duct.





Pain Management with endoscopy

 EUS and ERCP interventions are mostly based on observational data Biliary/pancreatic sphincterotomy Pancreatic duct stents Stone extraction Lithotripsy (ESWL vs intraductal) Celiac plexus block (weak data)



Pain management : surgery

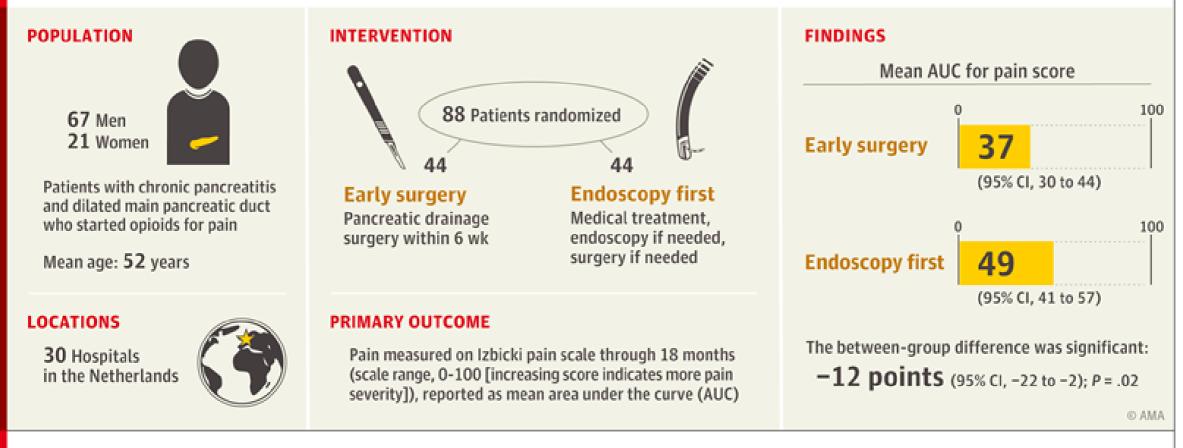
- Goal : relieve obstruction in pancreatic duct, common bile duct or gastric outlet obstruction
- Factors influencing surgical choice
 - Diffuse vs local disease
 - Ductal obstruction
 - Surgical expertise



🕺 JAMA Network⁻

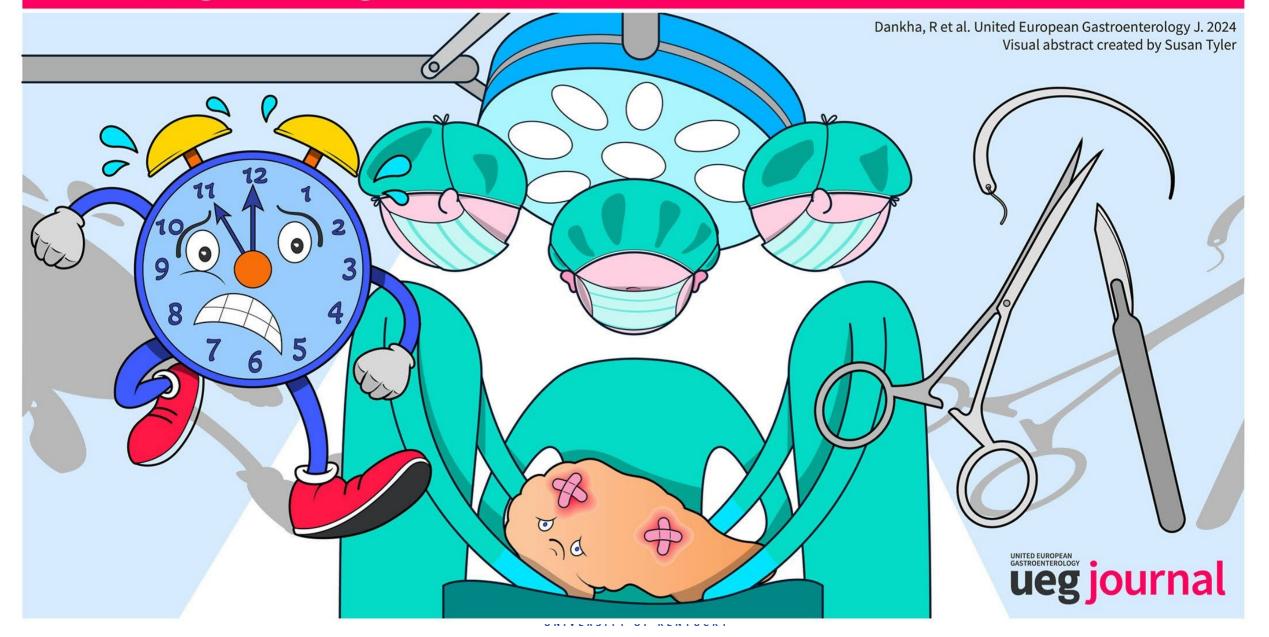
QUESTION For patients with painful chronic pancreatitis, is early surgery more effective than the endoscopy-first approach in addressing pain reduction?

CONCLUSION This randomized clinical trial found that early surgery resulted in less pain over 18 months.



Issa Y, Kempeneers MA, Bruno MJ, et al; the Dutch Pancreatitis Study Group. Effect of early surgery vs endoscopy-first approach on pain in patients with chronic pancreatitis: the ESCAPE randomized clinical trial [published January 21, 2020]. JAMA. doi:10.1001/jama.2019.20967

Surgical Management of Chronic Pancreatitis: A Narrative Review

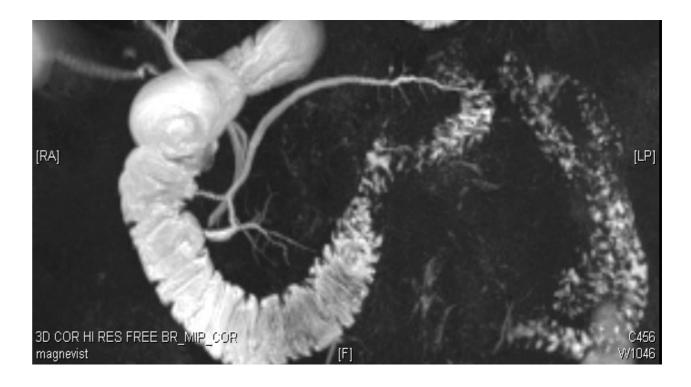


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Chronic abdominal pain

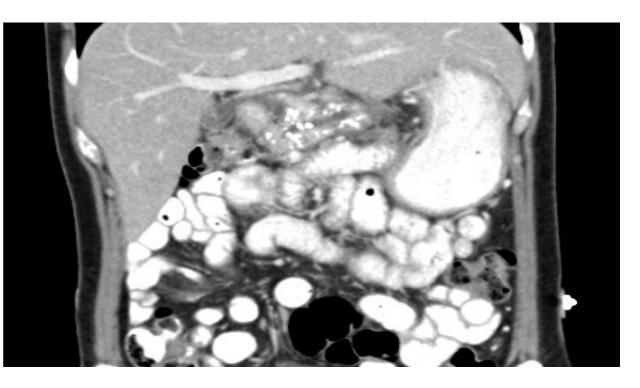
On opiates



I need an electrician !!



40 year old male Chronic Fluctuating abdominal pain



I need a Plumber !!



Chronic pancreatitis related Diabetes

- CP related DM is the most common type of pancreatogenic DM.
- ADA recommends annual diabetes screening.



Exocrine Pancreatic insufficiency

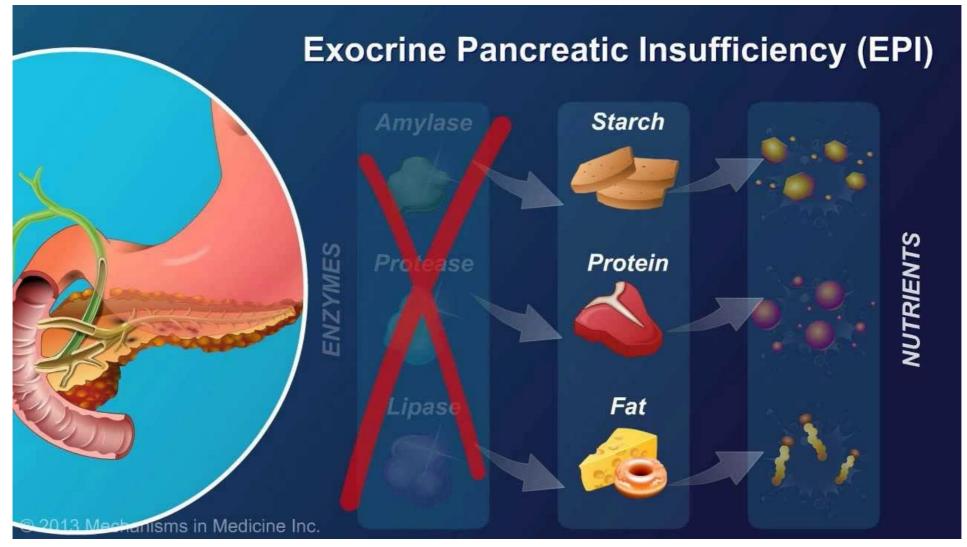
- Pancreatic enzyme secretion varies with content and volume of meal ingested
- Average lipase units per meal = 720,000 lipase U
- Exocrine pancreatic insufficiency occurs with 90% of enzyme output is diminished
- EPI is a condition caused by inadequate production, deliver and activity of pancreatic enzymes needed for normal digestion



Associated Medical Conditions

- Chronic Pancreatitis
- Occasionally after severe acute pancreatitis
- Tumors obstructing the main pancreatic duct
- Pancreatic resection
- Surgeries like distal gastrectomy or gastric byass surgery
- Systemic conditions
 - Cystic Fibrosis
 - IBD
 - Celaic Disease
 - Giardiasis
 - Gastrinoma
 - VHL
 - DM







EPI : Signs and Symptoms

- Gas
- Bloating
- Abdominal Pain
- "Diarrhea"
- Unexplained weight loss
- Steatorrhea
- Foul smelling stool
- Nocturnal stools
- Clinically relevant maldigestion occurs earlier than appearance of overt symptoms











Picture Credit : Dr Andres Gelrud

Making the diagnosis

- Fecal elastase
- Clinical response to PERT
- Suspect EPI when you see following labs : Low Hb, albumin, abnormal fat soluble vitamins
- 72 hour fecal fat



Fecal elastase

- Stool must be solid or semi solid
- Ok to test patient taking Pancreatic enzyme supplementation

Elastase, stool Collected on November 17, 2023 8:18 AM

Lab tests - Stool

Results

PANCREATIC ELASTASE-1 FECES View trends

mcg/g

Value >500

(NOTE)

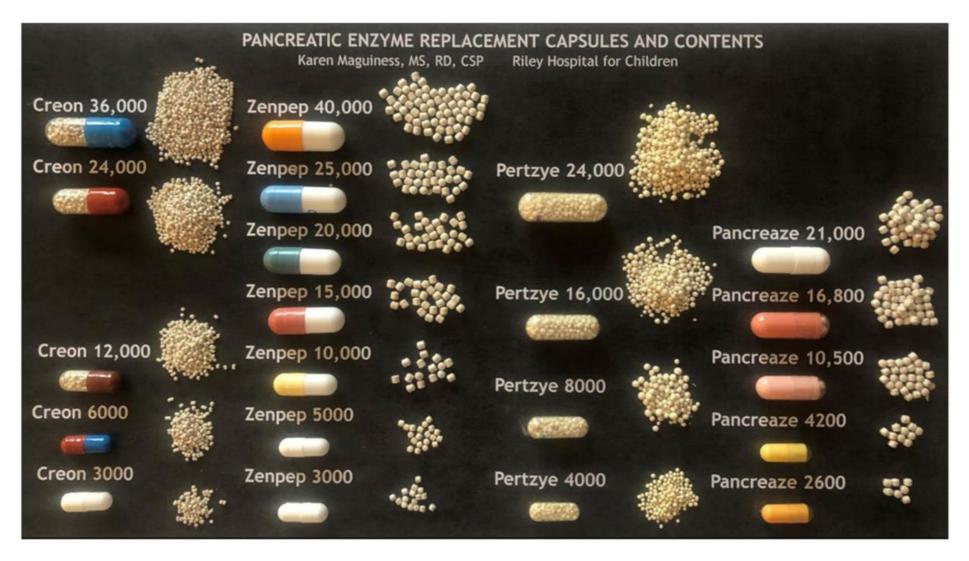
Adult and Pediatric Reference Ranges for Pancreatic Elastase-1:

Normal: >200 mcg/g Moderate Pancreatic Insufficiency: 100-200 mcg/g Severe Pancreatic Insufficiency: <100 mcg/g

Elastase-1 (E-1) assay results are expressed in mcg/g, which represent mcg E1/g feces.

It is not necessary to interrupt enzyme substitution therapy.







What is the best time to take the enzymes?

- Highest efficacy \rightarrow enzymes consumed throughout the meal
- Second Highest efficacy-> just after meals
- Lowest Efficacy-> just before meals



Pancreatic Enzyme replacement therapy

- How Much? Recommended starting dose is 25-50,000 USP units of lipase/meals
- Take 50% dose with snacks



Suboptimal Response

- Non compliance → assess affordability
- Underdosing → start low and increase approach
- Inactivation by gastric acid \rightarrow add PPI or H2A
- Improper timing of administration
- Concurrent GI diseases like SIBO, celiac, IBD, giardiasis







Metabolic Bone disease

 High prevalence of metabolic bone disease in CP Osteopenia -40%

Osteoporosis-25%

Increased risk to low trauma fractures

CP induces an inflammatory state, which contributes to bone loss.

Offer DEXA for osteoporosis screening.



Risk of Pancreatic Cancer

- Increased risk of pancreatic cancer vs general population
- Cumulative lifetime incidence is < 5%
- Insufficient data to recommend routine pancreatic cancer screening
- Screening should be offered to
 - Hereditary Chronic pancreatitis
 - Patients with new onset diabetes
 - Tropical pancreatitis (South East Asian patients with SPINK1)



Isolated Gastric Varices

- In setting of Splenic vein thrombosis
- In case of bleeding offer splenectomy

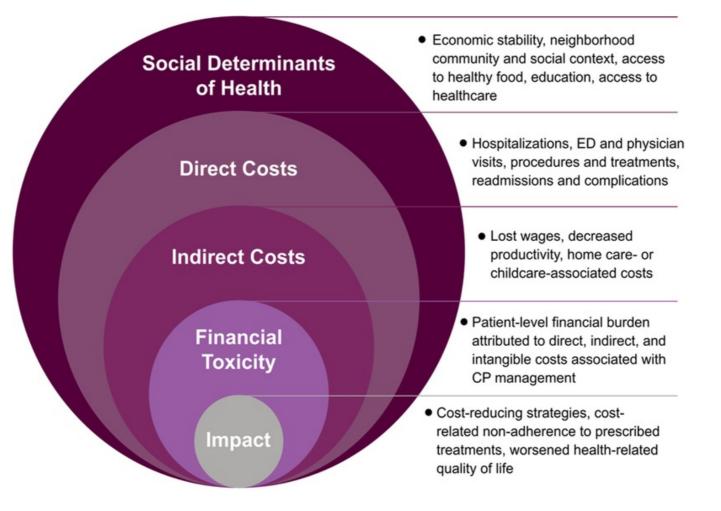


Nutritional Deficiencies

- Fat soluble Vitamin Deficiency ADEK
- Micronutrient deficiency- Magnesium and Zinc
- Sarcopenia
- Low BMI



Financial Toxicity





Choate et al 2024, Pancreas









Pancreatic Pain Consortium





Behavior and Development

IMPACT-2 STUDY

Do you have pain from recurrent acute or chronic pancreatitis?



About the Study

Researchers at Seattle Children's Research Institute want to learn more about the usefulness of an online pain self-management program for adults with pain from chronic pancreatitis

Eligibility

- () 18 years or older
- Diagnosed with recurrent acute or chronic pancreatitis and experience pain
- Access to, or willing to borrow, a web-enabled device

For more information, please contact our study team:

Participation Involves

Ĩ mood, and health at 3 time points over 9 months



Being randomly assigned to use one of two online programs for 8 weeks



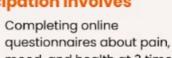
Optional participation in a telephone interview about the online program

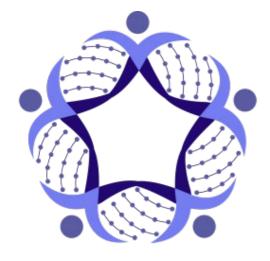
Earning up to \$225 in gift cards

https://redcap.link/impact2-info € 206-884-1559

ppsi@seattlechildrens.org







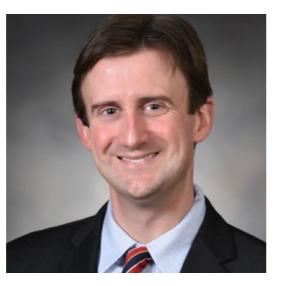
NATIONAL PANCREAS FOUNDATION CENTERS OF EXCELLENCE



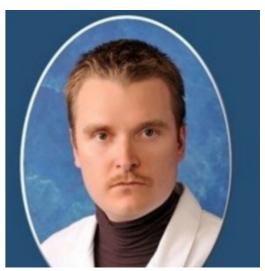
















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