Management of chronic pancreatitis

Liver update meeting 2024 Dr Neo Seabi



Outline

- Definition
- Aetiology
- Diagnosis
- Management options
 - Pharmacotherapy
 - Endoscopic
 - Surgical

Definition

- CP
 - Progressive disorder
 - Recurrent episodes of inflammation
 - Replacement of pancreatic parenchyma with fibrous tissue
- Consequences
 - Pain
 - Exocrine and endocrine insufficiency
 - Malignancy

CLASSIFICATION SYSTEMS OF AETIOLOGIES

MANNHEIM

M indicates multiple risk factors including:

Alcohol consumption: excessive (>80 g/d), increased (20-80 g/d), moderate (<20 g/d)

Nicotine consumption

Nutritional factors: high caloric proportion of fat and protein, hyperlipidemia

Hereditary factors: hereditary, familial, idiopathic (early onset, late onset), tropical

Efferent duct factors: pancreas divisum, annular pancreas and other congenital abnormalities of the pancreas, pancreatic duct obstruction (eg, tumors), posttraumatic pancreatic duct scars, sphincter of Oddi dysfunction

Immunological factors: autoimmune pancreatitis

Miscellaneous and rare metabolic disorders: hypercalcemia, hyperparathyroidism, chronic renal failure, drugs, toxins

TIGAR-O

Toxic-metabolic: alcohol, tobacco smoking, hypercalcemia, hyperlipidemia, chronic renal failure, medications, toxins

Idiopathic: early onset, late onset, tropical

Genetic mutations: PRSS1, CFTR, SPINK1, others

Autoimmune: isolated, syndromic

Recurrent and severe AP-associated CP: postnecrotic (severe AP), vascular disease/ischemic,

postirradiation

Obstructive: pancreas divisum, sphincter of Oddi disorders, duct obstruction (eg, tumor),

posttraumatic pancreatic duct scars

M-ANNHEIM and TIGAR-O

Diagnostic criteria for aetiology, clinical and diagnostic stage

Provides a checklist for recording risk factors

Dig Dis Sci 2018

Am J G 2020

Disease burden

- Epidemiology
 - Rare
 - Estimated prevalence of 35-50 per 100000 adults in US
 - Incidence 5 PER 100 000 patient years
 - SA: Madela et al
 - Looked at pathology and outcomes of pancreatic surgeries
 - 126 surgeries for cancer and CP: 77 PD
 - Prevalence of CP was 29.9% and 55.9% in PDAC
- Poor quality of life
 - Physical and metal health



Progression

Latent

Pain
Intact endocrine and
exocrine function

Compensatory

Pain under control.

Transitional

Acute on chronic Pancreatitis

Decompensated

PEEI are the main features

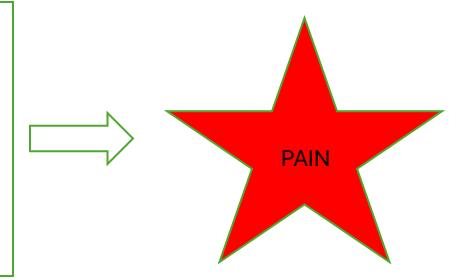
Sequelae of pancreatic tissue destruction

Pain

Pancreatic ductal hypertension

Stones Strictures Oxidative stress from schaemia

Altered nociception Neuropathy



High BP does not equate to pain

Exocrine insufficiency

- Results in maldigestion and malabsorption
 - Steatorrhoea
 - Fat soluble vitamins
 - Abdominal distension
 - Sarcopenia
 - Reduced QOL

Endocrine

- Pancreatogenic DM (Type 3c)
 - Glycaemic control often poor with wild fluctuations
 - 40% point prevalence
 - Islet cells replaced with fibrosis
 - Insulin resistance, ?impaired incretin hormone response

- Metabolic bone disease
 - Chronic pancreatitis osteophathy
 - Increased fracture risk

Malignancy

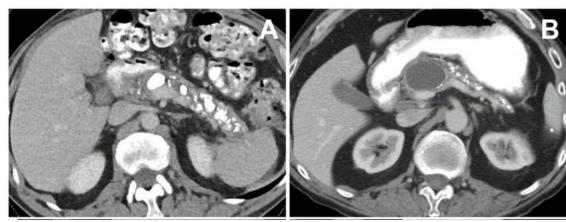
More common in CP from hereditary causes Less in AIP

Chronic Pancreatitis Is a Risk Factor for Pancreatic Cancer, and Incidence Increases With Duration of Disease: A Systematic Review and Meta-analysis

Sonal Gandhi, MBBS¹, Jaime de la Fuente, MD², Mohammad Hassan Murad, MD³ and Shounak Majumder, MD²

Diagnosis

- Symptoms
- Imaging: Early vs late CP
 - X-Ray, US, CT, MRI (MRCP or secretin-stimulated MRCP), EUS (+/-TE) and ERCP
- Laboratory: PEI
 - Quantitative fecal fat estimation >7g/day: gold standard for steatorrhea
 - Fecal elastase-1: good alternative
 - Secretin stimulated pancreatic function
- J gastroenterol 2022
- AMJG 2020



Conventional criteria Rosemont criteria

Parenchymal criteria Major criteria A

Hyperechoic foci (>2 mm in length/width with

Hyperechoic strands shadowing)

Hypoechoic lobules, Major duct calculi (echogenic structure[s] within the

foci or areas MPD with acoustic shadowing)

Cyst

Major criteria B

Lobularity (≥3 contiguous lobules = 'honeycombing')

Duct criteria

Irregular duct contour Minor criteria

Visible side branches Cyst (anechoic, round/elliptical with or without

Hyperechoic duct septations)*

margin

Stone

Dilated duct (≥3.5 mm in body or >1.5 mm in tail)*

and ectatic course)

Dilated side branch (>3 tubular anechoic structures each measuring ≥1 mm in width, budding from the

MPD)*

Hyperechoic duct wall (echogenic, distinct structure

>50% of entire MPD in the body and tail)

Hyperechoic strands (≥3 mm in at least 2 different directions with respect to the imaged plane)

Hyperechoic foci (>2 mm in length/width that are

nonshadowing)*

Lobularity (>5 mm, noncontiguous lobules)

Diagnostic criteria

Can J Gastro 2011, GIE 2009, J Gastro 2022

- EUS
 - Hyperechoic foci (non-shadowing) or strands
 - Lobularity
 - Hyperechoic MPD margins
 - Dilated side branches

^{*}If any of these minor criteria are present, the patient cannot be classified as 'normal'. MPD Main pancreatic duct. Data from references 14 and 15

Management

- Focused on:
 - Lifestyle changes
 - Alcohol and smoking reduces exocrine insufficiency, abdo pain and local complications

P Goltl et al. Alcohol 2023

- Pain control
- PERT/ Nutrition
- Diabetes control

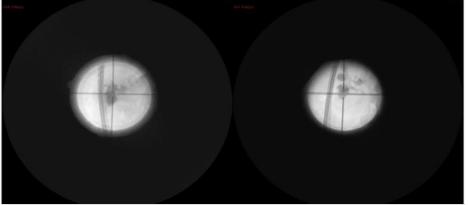
PAIN

- NSAIDs
- Weak then strong opioids
- Other drugs e.g pregabalin, antioxidants
- Fat restricted diet may be effective
- Extra-corporeal shock wave lithotripsy
- Endoscopy
- Surgery
- IR: pseudoaneurysm or haemosuccus

Endoscopic management

- Stone management
 - ERCP with or without panctreatoscopy and lithotripsy
 - The stones can be rigid and refractory
 - ESWL
 - Stenting: can be initial therapy
- ERCP: Strictures
 - Stent
 - If dominant stricture
 - Sphincterotomy: if papillary stenosis
 - Plastic vs fcSEMS
- EUS: case by case
 - Coeliac plexus: Uni or bilateral, direct or indirect
 - Collections







Pancreatic Exocrine Insufficiency

- Pancreatic enzyme replacement therapy
- Nutrition without fat restriction
- A,D,E,K etc. supplementation

- CP associated osteopathy: BMD and high fracture risk
- Sarcopenia- high risk of hospitalization and mortality
 - Nutritional therapy and PERT

PERT/ Nutrition

Clinical nutrition 2024

- Enteric-coated microspheres
- Released at pH >5.5
- Recommended dose 20-50000 PhU with main meals: adjust
- Spread capsules out over a meal
- Colonic strictures with high dose, esp children
- PERT improves fat absorption and GI symptoms
- PPI can be added if incomplete symptomatic response
- High fiber not recommended
- If malnourished, high energy, high protein meals 5-6 per day





Role of surgery

- Usually reserved for refractory cases related to anatomic changes
 - Pancreatic, duodenal or biliary obstruction
 - Refractory pain
- Types of interventions
 - Drainage procedures
 - Partial pancreatectomy with/without duodenal resection and pyloric preservation
 - Frey, Berge and Berne most commonly done
 - Total pancreatectomy and islet autotransplantation
 - Survival of the graft sometimes poor: mesenchymal contransplantation, etarnecept, HCQ tried
- Different but often comparable outcomes wrt: Pain improvement, LOS, blood T/F, weight gain
- Organ sparing procedures widely used

Ther adv gastroenterol 2024

Malignancy screening

Key concept

There is a lack of evidence to suggest that performing screening examinations on patients with CP to detect pancreatic malignancy is beneficial.



Incidence and risk of pancreatic cancer in patients with chronic pancreatitis: defining the optimal subgroup for surveillance

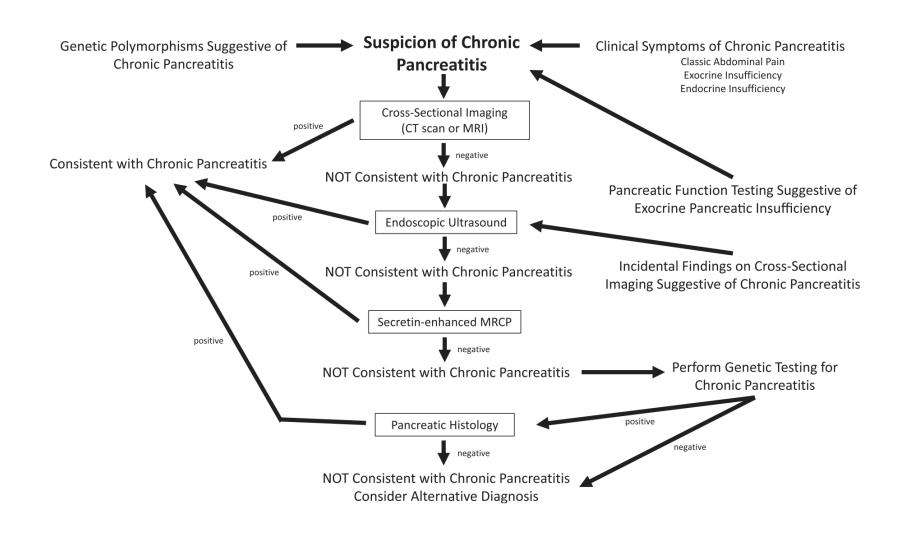
Hyo Suk Kim¹, Tae-Geun Gweon¹, Sang Hi Park², Tae Ho Kim¹, Chang Whan Kim¹ & Jae Hyuck Chang^{1,3⊠}

New therapies

- Neuromodulation
 - Cervical transcutaneous nerve stimulation
- Cognitive behavioural therapy
 - Pain self-management
- Energy therapy (Pranic healing)

Diagnostic approach

Gardner et al AJ Gast 2020



Conclusion

- Individualized therapies
- Improve quality of life
- Malignancy screening for a select group