Diagnosis and Therapy of Acute Pancreatitis

Early Phase \leq 4weeks

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Radboudumc

Disclosure



















U NOVARTIS

Leerdoelen

- 1. Voeding en acute pancreatitis
- 2. I.V. vloeistof therapie
- 3. Wanneer een CT-scan?

- 4. Wanneer stuur je iemand naar een expert center?
- 5. Wanneer draineer je vloeistofcollecties?

Gebasseerd op

- NL/Europese IAP/APA 2013 richtlijn
- AGA 2018 richtlijn

Incidence / prognosis AP

- I= 5-30/ 100.000
- 20% moderate-severe course: necrosis and/or multi organ failure
- Mortality rate 5-8% (30% in severe AP)
- 8-10% develops chronic pancreatitis
- 17% recurrent attacks => 26% develops chronic pancreatitis

Pancreas 2018;47: 653-666

Clin Gastroenterol Hepatol. 2016 May;14(5):738-46.

Am J Gastroenterol 2006; 101:2379-400.

2013 AIP/APA guidelines

Etiology

Biliary (eg, gallstones, microlithiasis)

60-80%

Alcohol

Other causes:

- Anatomic variants
- Ampullary/ ductal obstructions
- Metabolic (eg, hypercalcemia, hypertriglyceridemia)
- Drugs/Toxins, Trauma, Ischemia, Hypothermia
- Infections
- Autoimmune
- Genetic (familial, sporadic)
- Idiopathic:

12-25%

- 1. Diagnosis
- 2. Etiology
- 3. Severity prediction
- 4. Severity assesment
- 5. CT/MRI in the early fase
- 6. Treatment

1. Diagnosis AP

Definition of acute pancreatitis: '2 out of 3' of the following criteria:

- 1. Clinical (upper abdominal pain),
- 2. Laboratory (serum amylase or lipase >3x upper limit of normal)
- 3. Imaging (CT, MRI, ultrasonography) criteria.

IAP/APA evidence-based guidelines for the management of acute pancreatitis: Pancreatology 13 (2013) e1-15.

Classification of acute pancreatitis—2012: revision of the Atlanta classification and definitions by international consensus: Gut 2013;62:102–111.

2. Etiology of Acute Pancreatitis

Goal: potential treatment / prevention recurrent attack

On admission:

- 1. Personal (i.e. previous acute pancreatitis, known gallstone disease, alcohol intake, medication and drug intake, known hyperlipidemia, trauma, recent invasive procedures such as ERCP)
- 2. Family history of pancreatic disease,
- 3. Physical examination,
- 4. Laboratory serum tests (i.e. liver enzymes, calcium, triglycerides),
- 5. Imaging (i.e. right upper quadrant ultrasonography).

IAP/APA evidence-based guidelines for the management of acute pancreatitis: Pancreatology 13 (2013) e1-15.

2. Etiology of Acute Pancreatitis

High probability of a biliary etiology (at least 1):

- 1) Gallstones or biliary sludge on imaging (any type),
- 2) Dilated CBD: > 8 mm in patients \leq 75 y or > 10 mm in patients > 75 y,
- $3) \quad ALAT > 2x ULN$

3. Severity prediction

3-dimension approach predict outcome of acute pancreatitis:

- 1. host risk factors (e.g. age, co-morbidity, body mass index),
- 2. clinical risk stratification (e.g. persistent SIRS)
- 3. monitoring response to initial therapy (e.g. persistent SIRS, blood urea nitrogen, creatinine).

3. Severity prediction (DPSG)

Within 24 h after presentation:

- **1**. APACHE II score ≥ 8
- 2. modified Glasgow score \geq 3
- 3. CRP > 150 mg per liter.

4. Cross-sectional Imaging

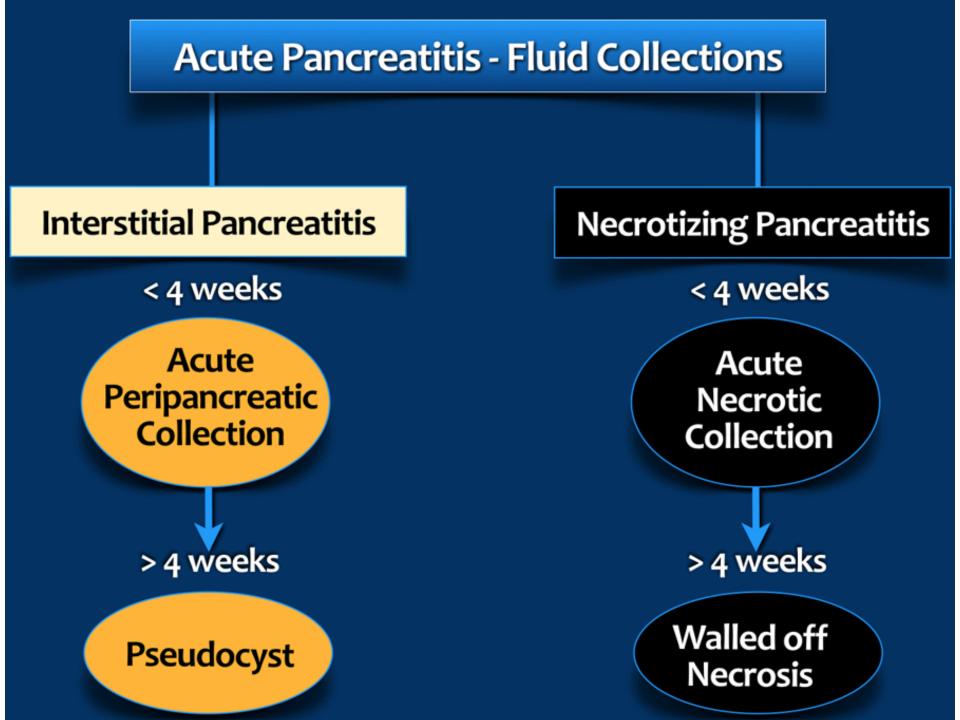
Indication for initial CT in acute pancreatitis:

- 1. Diagnostic uncertainty
- 2. Confirmation of severity based on clinical predictors of severe AP
- **3**. Failure to respond to conservative treatment or in the setting of clinical deterioration.

Optimal timing: At least 72-96 h after onset of symptoms.

IAP/APA evidence-based guidelines for the management of acute pancreatitis: Pancreatology 13 (2013) e1-15.

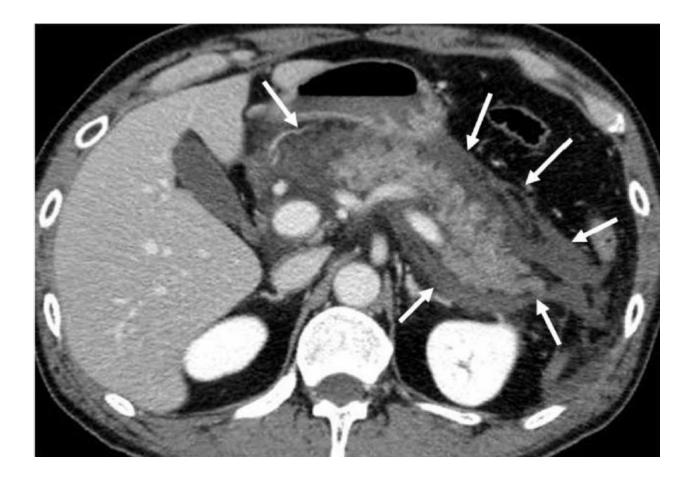
AGA Institute Guideline on Initial Management of Acute PancreatitisGastroenterology 2018;154:1096–1101



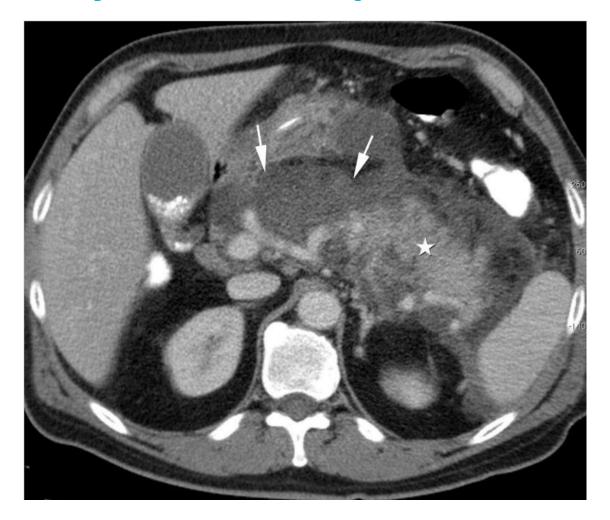
CT 3 days: Interstitial pancreatitis



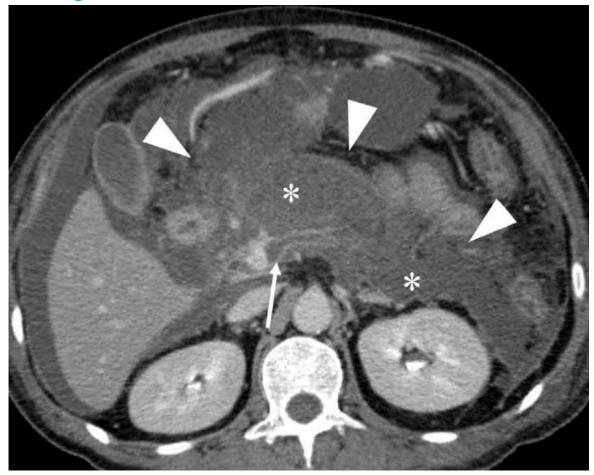
CT day 3: + fluid collections



CT 4 days: necrotic pancreatitis



CT 4 days: necrotic fluid collections



5. Severity assesment

2012: revised Atlanta criteria

- A. Mild AP
 - No organ failure
 - No local or systemic complications
- B. Moderately severe AP
 - OF resolves within 48h and/or:
 - Local/systemic complications without OF
- C. Severe AP
 - Persistant OF (>48h): single or multiple OF

5. Severity assesment

Cullen's sign: after 24-48h: Methemalbumin subcutaneously

- Increased mortility
- Also in: ectopic pregnancy, trauma, ruptured aortic aneurysm



5. Severity assesment

• Grey-Turner's & Fox's sign: 24-48h



6. Treatment AP

Treatment:

- A. Pain control
- B. Fluid rescussitation
- C. Role of Antibiotics
- D. Role of Nutrition
- E. Early ERCP in acute biliary pancreatitis
- F. Early drainage of infected fluid-collections
- G. Referal to expert center
- H. Future perspectives

A. Treatment: Pain control

Standard step-up approach:

- 1. PCM/NSAIDS
- 2. Opiods
- **3**. Patient Controlled Analgesia (PCA)
- 4. Epiduraal

High on drugs: lessons from opiates in pancreatitis

Animal research:

- 1. more ileus/bacterial translocation,
- 2. Temporal delay in the reparative inflammatory response
- 3. Prevents pancreatic regeneration

Gut 2018;67:600-602

Pancreas 2017;46: 858-866)

Gut 2018;67:719-27.

B. Fluid rescuscitation

- Ringer's lactate is recommended for initial fluid resuscitation
- Goal directed therapy with 5-10 ml/kg/h until **resuscitation goals**

Goals (1 or more):

- 1. Non-invasive clinical targets:
 - heart rate < 120/min,
 - mean arterial pressure between 65-85 mmHg, and
 - urinary output > 0.5-1ml/kg/h,
- 2. Invasive clinical targets: stroke volume variation, intrathoracic blood volume
- 3. Biochemical targets of hematocrit 35-44%.

IAP/APA evidence-based guidelines for the management of acute pancreatitis: Pancreatology 13 (2013) e1-15.

C. Role of AB

Use of prophylactic antibiotics is not recommended.

Technical review (n=10 RCT's):

- 1. AB: less infected necrosis: OR=0.56 (0.36-0.86)
- 2. Sub-group analysis (studies >2002): OR=0.81 (0.44-1.49)

D. Nutrition: 2018 AGA Guideline

- 1. Early vs delayed feeding: reduced risk
 - interventions for necrosis: OR, 2.47 (1.41-4.35)
- 2. Enteral vs parenteral nutrition: reduced risk:
 - infected peripancreatic necrosis (OR, 0.28; 95% CI, 0.150.51),
 - single organ failure (OR, 0.25; 95% CI, 0.10-0.62), and
 - multiple organ failure (OR, 0.41; 95% CI, 0.27-0.63).
- 3. Nasogatric vs nasoduodenal: ns mortality

D. Nutrition

Step-up approach:

- 1. Early (within 24 hours) oral feeding as tolerated
- 2. Inability to feed orally: enteral rather than parenteral nutrition: nasogastral or nasal enteral.
- 3. Parenteral nutrition can be administered as second-line therapy if nasojejunal tube feeding is not tolerated and nutritional support is required

AGA and IAP/APA guidelines

Lancet 2015;386:85-96.

JPEN J Parenter Enteral Nutr. 2018 Sep;42(7):1139-1147.

E. Role of ERCP in biliary pancreatitis

- Not in predicted mild cases
- **Probably not** in predicted severe cases ٠
- In cholangitis: indicated



Early biliary decompression versus conservative treatment in acute biliary pancreatitis (APEC trial): study protocol for a randomized controlled trial



Trials

F. Early drainage infected collecions?

- Interventions should be delayed to 4 weeks before intervention
- Mechanical or infectious complications





G. Referal to expert center

Severe acute pancreatitis: that may need:

- interventional radiologic,
- endoscopic,
- surgical intervention.

A specialist center:

- a high volume center with up-to-date intensive care facilities

- access to interventional radiology, interventional endoscopy with EUS and ERCP assistance as well as surgical expertise in managing necrotizing pancreatitis.

Future perspectives

- Epidural
- Pentoxifyllin: Mayoclinics
- Fluid trials: SVI netwerk US
- Fish-oil: PWN: PLANCTON-trial
 - Early intravenous omega-3 fatty acids
 - Predicted severe acute pancreatitis (Besseling and van Goor)



Questions?





