North Carolina Society of Gastroenterology 2024 Annual Meeting



Updates in the Management of Acute Pancreatitis

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Joint Providership



American Society for Gastrointestinal Endoscopy

Disclosures: none



Learning Objectives

- Appreciate the burden of acute pancreatitis in the US
- Discuss the evidence behind the mainstays of treatment in acute pancreatitis
- Define pancreatic fluid collections and review management strategies
 for severe acute pancreatitis with necrosis
- Recognize the quality indicators for patients admitted with acute pancreatitis

Acute Pancreatitis in the US



Peery et al. Gastroenterology 2019

Revised Atlanta Criteria

Diagnosis requires 2 of the following 3 criteria:

- 1. Epigastric abdominal pain
- 2. Lipase > 3 times normal
- 3. Acute pancreatitis on cross-sectional imaging*

Banks et al. Gut 2013



Drug Induced Pancreatitis

• RARE!

Summary of drug-induced acute pancreatitis based on drug class

Class Ia	Class Ib	Class II	Class III	Class IV	
Azodisalicylate	Amiodarone	Chlorothiazide	Atorvastatin	Ampicillin	
Bezafibrate	Azathioprine	Clozapine	Carbamazepine	Bendroflumethiazide	
Cannabis	Clomiphene	Didanosine	Captopril	Benazepril	
Carbimazole	Dexamethasone	Erythromycin	Ceftriaxone	Betamethasone	
Codeine	Ifosfamide	Estrogen	Chlorthalidone	Capecitabine	
Cytosine	Lamivudine	L-asparaginase	Cimetidine	Cisplatin	
Arabinoside	Losartan	Pegaspargase	Clarithromycin	Colchicine	
Dapsone	Lynestrenol/methoxyethinylestradiol	Propofol	Cyclosporin	Cyclophosphamide	
Enalapril	6-mercaptopurine	Tamoxifen	Gold	Cyproheptadine	
Furosemide	Meglumine		Hydrochlorothiazide	Danazol	
Isoniazid	Methimazole		Indomethacin	Diazoxide	
Mesalamine	Nelfinavir		Interferon/ribavirin	Diclofenac	
Metronidazole	Norethindronate/mestranol		Irbesartan	Diphenoxylate	
Pentamidine	Omeprazole		Isotretinoin	Doxorubicin	
Pravastatin	Premarin		Ketorolac	Ethacrynic acid	
Procainamide	Trimethoprimsulfamethazole		Lisinopril	Famciclovir	
Pyritonol			Metolazone	Finasteride	
Simvastatin			Metformin	5-fluorouracil	
Stibogluconate			Minocycline	Fluvastatin	
Sulfamethoxazole			Mirtazapine	Gemfibrozil	
Sulindac			Naproxen	Interleukin-2	
Tetracycline			Paclitaxel	Ketoprofen	
Valproic acid			Ponatinib	Lovastatin	
			Prednisone	Mefenamic acid	
			Prednisolone	Nitrofurantoin	
				Octreotide	
				Oxyphenbutazone	
				Penicillin	

What happens to our patients with Acute Pancreatitis?



Adapted from Forsmark et al. NEJM 2016

IAP/APA evidence-based guidelines for the management of acute pancreatitis

Wor		
^a Interna ^b Ameria	Gastroenterology 2018;154:1096–1101	201
	AGA SECTION	3
	American Gastroenterological Association Institute Guideline on Initial Management of Acute Pancreatitis	
An	Seth D. Crockett, ¹ Sachin Wani, ² Timothy B. Gardner, ³ Yngve Falck-Ytter, ^{4,5} and Alan N. Barkun ⁶ ; on behalf of American Gastroenterological Association Institute Clinical Guidelines Committee	201
Ma		
Scott 7	Cenner, MD, MPH, FACG ¹ , John Baillie, MB, ChB, FRCP, FACG ² , John DeWitt, MD, FACG ³ and Santhi Swaroop Vege, MD), FACG ⁴

-9-,

Severity Assessment

1040015 11+10	4000	At a decision and at 40 hours
APACHE-II*10	1989	At admission and at 48 hours
		Temperature, MAP, heart rate, respiratory rate, Pao ₂ , arterial pH, HCO ₃ , sodium, potassium, creatinine,
		hematocrit, WBC, Glasgow Coma Score, age, chronic health points
BISAP ¹⁶	2008	At admission and at 48 hours
		BUN (>25 mg/dL), impaired mental status (Glasgow Coma Score <15), SIRS (≥2), age (>60 y), pleural effusion
Glasgow ⁸	1984	At admission and at 48 hours
		Age (>55 y), WBC (>15,000/mL), glucose (>180 mg/dL), BUN (>45 mg/dL), Pao ₂ (<60 mm Hg), calcium (<8 g/dL), albumin (<3.2 g/dL), LDH (>600 IU/L)
HAPS ²¹	2009	At admission and at 48 hours
		Abdominal tenderness, hematocrit (>43 mg/dL for men or >39.6 mg/dL for women), creatinine (>2 mg/dL)
JSS ²⁰	2009	At admission and at 48 hours
		Base excess (≤3 mEq/L), Pao ₂ (≤60 mm Hg or respiratory failure), BUN (≥40 mg/dL) or Cr (≥2 mg/ dL), LDH (≥2× upper limit of normal), platelet (≤100,000/mm ³), calcium (≤7.5 mg/dL), CRP (≥15 mg/dL), SIRS (≥3), age (≥70 y)
Panc 314	2007	At admission and at 48 hours
		Hematocrit (>44 mg/dL), BMI (>30 kg/m ²), pleural effusion
POP ^{15a}	2007	At admission and at 48 hours
		Age, MAP, Pao ₂ :Fio ₂ , arterial pH, BUN, calcium
Ranson ^{4,5}	1974	At admission: age (>55 y), WBC (>16,000/mL), glucose (>200 mg/dL), LDH (>350 IU/mL), AST (>250 IU/mL)
		At 48 hours: hematocrit (decrease >10%), BUN (increase >5 mg/dL), calcium (<8 mg/dL), Pao ₂ (<60 mm Hg), base deficit (>4 mEq/L), fluid sequestration (>6 L)
SIRS ¹²	2006	At admission and at 48 hours
		Temperature (<36°C or >38°C), heart rate (>90/min), respiratory rate (>20/min or $PaCo_2 <32 \text{ mm}$ Hg), WBC (<4000/mm ³ , >12,000/mm ³ or >10% bands)

Severity Index

- BUN > 25
- MPAIRED MENTAL STATUS
- SIRS PRESENT
- AGE > 65
- PLEURAL EFFUSIONS

- BISAP = 3 → → → pancreatitis? mortality

Does trending lipase

help prognosticate outcomes in acute

• BISAP = $5 \rightarrow 22\%$ mortality

BISAP > 3: consider a setting with close clinical monitoring (i.e. ICU)

Wu et al. Gut 2008

Resuscitation Choices:

TABLE 7-2. COMPOSITION OF CRYSTALLOIDS						
	NA	CL	K	CA	LACTATE	
FLUID	(mmol/L)	(mmol/L)	(mmol/L)	(mg/dL)	(mmol/L)	PH
Normal saline solution	154	154	_	_	_	6.0
Lactated Ringer's solution	130	109	4	3	28	6.5

 Recent data suggest that LR may be better in critically ill patients* and equivocal to NS in noncritical ill patients

> Semler et al. NEJM 2018 Self et al. NEJM 2018

Fluid Management

Lactated Ringer's reduces pancreatic acidosis and tryspin activation



Lactated Ringers = Normal Saline*

Wu et al. Clin Gastroenterol Hepatol 2011

Balance is Key

Circulatory collapse

Pancreatic microischemia







Hydration Strategies in Acute Pancreatitis

Close monitoring using clinical parameters (Vitals, BUN and Hct) improves outcomes

ORIGINAL ARTICLE

Aggressive or Moderate Fluid Resuscitation in Acute Pancreatitis

E. de-Madaria, J.L. Buxbaum, P. Maisonneuve, A. García García de Paredes, P. Zapater, L. Guilabert, A. Vaillo-Rocamora, M.Á. Rodríguez-Gandía,
J. Donate-Ortega, E.E. Lozada-Hernández, A.J.R. Collazo Moreno, A. Lira-Aguilar, L.P. Llovet, R. Mehta, R. Tandel, P. Navarro, A.M. Sánchez-Pardo,
C. Sánchez-Marin, M. Cobreros, I. Fernández-Cabrera, F. Casals-Seoane,
D. Casas Deza, E. Lauret-Braña, E. Martí-Marqués, L.M. Camacho-Montaño, V. Ubieto, M. Ganuza, and F. Bolado, for the ERICA Consortium*



- Start with 10cc/kg bolus if hypovolemic (no bolus if normovolumic) → 1.5cc/kg/hr
- Careful Monitoring to maintain EUVOLUMIA
- Diuresis is OK in first 72 hours to support euvolumia











Maintain Euvolemia!

Early Management Strategies in Acute Pancreatitis

Drain the fluid?

Risk Factor Modification?

Imaging?

To Eat or Not to Eat?

Annals of Internal Medicine



Early Versus Delayed Feeding in Patients With Acute Pancreatitis Shorter LOS in mild/mod severe AP; no diff in complications Equal LOS in severe AP; limited data

Early versus On-Demand Nasoenteric Tube Feeding in Acute Pancreatitis

No difference in infective complications between groups in patients with SAP Is post-pyloric feeding really necessary?

NG = NJ feeding EQUAL OUTCOMES!



Baron et al. Gastroenterology 2020



*TPN reserved if not receiving adequate enteral feeding or if gut is compromised



What should I do if they have recurrent pain?

Starting low fat solid diet upfront <u>REDUCES</u> hospital length of stay (-1.5d, p < 0.05) without recurrence of abdominal pain

> Jacobsen et al. CGH 2007 Moraes et al. J Clin Gastroenterol. 2010



Low Fat Diet ASAP

Maintain Euvolemia!

Early Management Strategies in Acute Pancreatitis

Drain the fluid?

Risk Factor Modification?

Imaging?

What additional work-up is needed?

WHILE ADMITTED

- Rule out gallstones
- Medication Reconciliation
- Substance Abuse Counselling
- LFTs, calcium and triglycerides

- AFTER DISCHARGE
- If initial work up negative and/or age > 40, proceed with MRI/MRCP
- If Family history positive for pancreatitis or concern for CF, will consider genetic testing

When is imaging necessary?

Imaging Study	Utility in Assessment in the Acute Setting	
Ultrasound		\bigstar
СТ		
MRCP	_	
EUS		
ERCP		

Risk Modification

Recurrent Acute Pancreatitis



Lankisch et al. Am J Gastroenterol 2009 Yadav et al. Am J Gastroenterol 2012



Take out the gallbladder during this index admission!

Da Costa et al. Lancet 2015 Vege et al. Gastroenterology 2018

The Recurrence of Acute Alcohol-Associated Pancreatitis Can Be Reduced: A Randomized Controlled Trial

ISTO NORDBACK, HANNA PELLI, RIITTA LAPPALAINEN-LEHTO, SATU JÄRVINEN, SARI RÄTY, and JUHANI SAND

Department of Gastroenterology and Alimentary Tract Surgery, Division of Surgery, Gastroenterology and Oncology, Tampere University

- 120 patients randomized to either one intervention during index admission or repeated interventions every 6 months for 2 years:
- 1. Information on toxicity of ETOH
- 2. Need to reduce ETOH intake
- 3. Evaluation of contributing factors
- 36 patients were lost to follow-up

iver		Initial intervention Only	Repeated interventions at 6-month intervals	
		(n = 61)	(n = 59)	Р
	Total hospital admissions			
	Abdominal complaints			
	Number of patients, N (%)	16 (26)	7 (12)	.038
	Number of admissions	30	15	.004
	Recurrent AP ^a			
	Number of patients, N (%)	13 (21)	5 (8)	.042
	First recurrence			
	<6 mo, N (%)	5 (8)	4 (7)	NS
	>6 mo, N (%)	8 (13)	1 (2)	.018
	Number of episodes			
	Overall	20	9	.012
	<6 mo	5	4	NS
	>6 to 24 mo	15	5	.009
	Admissions because of abdominal			
	complaints not fulfilling			
	the criteria of recurrent			
	AP			
	Number of patients (%)	3 (5)	2 (3)	NS
	Number of admissions	10	6	NS

GASTROENTEROLOGY 2009;136:848-855



Low Fat Diet ASAP

Maintain Euvolemia!

Early Management Strategies in Acute Pancreatitis

Drain the fluid?

CCY in Index AdmissionRule out gallstones 1stETOH CounsellingMalignancy after d/c

	Interstitial edematous pancreatitis	Necrotizing pancreatitis		
	Acute (peri)panreatic fluid collection	Acute necrotic collection		
< 4 weeks	Homogenous fluid adjacent to pancreas without a recognizable wall	Intra and/or extra pancreatic necrotic collection without a well-defined wall		

Indications for Drainage

Indications for Antibiotics

 Radiographic evidence of
infection
 Clinically appearing infected

- Infected pseudocyst or necrosis (preferably after 4 weeks)
- Ongoing organ failure despite optimal several weeks of medical therapy
- Sterile fluid collection with persistent symptoms (after 4 weeks)
- Disconnected duct from gland necrosis

Disconnected Duct



Pancreatic Fluid Collection Drainage

Transmural drainage (i.e.

cystgastrostomy and necrosectomy)

Transampullary drainage (i.e. disconnected duct syndrome)

Percutaneous (IR) drainage

Surgical drainage (i.e.

cystgastrostomy or cystjejunostomy)

Varadarajulu et al. Gastroenterology 2013 Van Brunschot et al. Lancet 2018

Warning...





No longer than 4 weeks!

Garcia-Alonso et al. Endoscopy 2018 Bang et al. Gut 2017 Gastroenterology 2020;158:67-75

Check for

AGA CLINICAL PRACTICE UPDATE: EXPERT REVIEW

American Gastroenterological Association Clinical Practice Update: Management of Pancreatic Necrosis

Intra-abdominal hypertension common;

unusual

clinical abdominal compartment syndrome

Management of Severe Acute Pancreatitis with Necrosis



patient with persistent nutritional failure or organ dysfunction Approach to debridement Dependent on pattern of necrosis and institutional expertise

- Central retrogastric collection:
 - Endoscopic transgastric or laparoscopic transgastric approach
- Retrogastric with paracolic gutter extension:
 - Percutaneous drainage with step-up to videoscopic retroperitoneal debridement or endoscopic debridement with addition of percutaneous drain as needed
- Retrogastric collection with extension to the right of the mesenteric vessels:
 - Endoscopic or laparoscopic transgastric; may need open debridement

Important: MULTIDISCIPLINARY APPROACH!

Adapted from Baron et al. Gastroenterology 2020



Low Fat Diet ASAP

Maintain Euvolemia!

Early Management Strategies in Acute Pancreatitis

Only if symptomatic. No prophylactic antibiotics!

CCY in Index AdmissionRule out gallstones 1stETOH CounsellingMalignancy after d/c

Adherence to Guidelines



Enforcement of Guideline Based Care



DiMagno et al. Am J Gastroenterol 2014





Vivian et al. Am J Gastroenterol 2019 Mosko et al. Clin Gastroenterol Hepatol 2020

How is Duke doing with this?

 <u>Population</u>: Retrospective cohort of 309 adults with a diagnosis of acute pancreatitis admitted to DUH, DRH, or DRaH from 10/7/2019-10/7/2020



Hein A et al. Am J Gastro 2021 Namasingh N et al. Am J Gastro 2021

How can we be better?

- Acute Pancreatitis Order Set
- Consult Surgery when suspected Gallstone Pancreatitis!!



Nost patients with acute pancreatitis do not need to be made strictly NPO. If NPO, reassess daily.
 Clear liquid diet probably of low utility and most patients who can tolerate oral intake can be placed on low fat diet.
 If unable to tolerate oral nutrition, strongly consider placement of nasoenteral feeding tube and initiation of tube feeding by hospital day #3.

Diet low fat low cholesterol - Consistency Advanced (Mech Soft)

Diet clear liquid

Diet NPO except for sips and ice chips

Diet NPO Until Specified



Thank you for your attention!

Questions?

Darshan.Kothari@duke.edu

CME/MOC Question:

A 47 year old female with hypertension presents to the ED with acute onset epigastric pain with nausea and vomiting and is found to have a lipase of 4000 (ULN 70) and imaging findings consistent with acute pancreatitis. She only takes HCTZ. Her social history of notable for tobacco use and 1-2 drinks per week. Her LFTs are normal and her ultrasound shows gallstones. She receives IV fluids, analgesia and is initially bowel rest. By hospital day 1, her pain improves and her diet is advanced to a low fat solid diet. Prior to discharge, which of the following should be done?

- A. Stop HCTZ
- B. Consult social work for alcohol cessation resources
- C. Surgical consultation for cholecystectomy
- D. MRCP to rule out choledocholithiasis
- E. Send IgG4 subclasses

Joint Providership





CME/MOC Question:

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Joint Providership





CME/MOC Answer: Correct Choice: C

Most common cause of acute pancreatitis is gallstones and with documented cholelithiasis, patients should undergo cholescystectomy during this index admission.

Joint Providership



