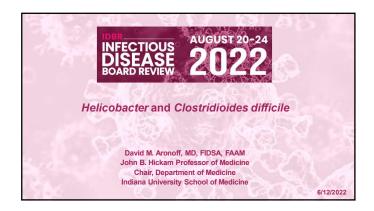
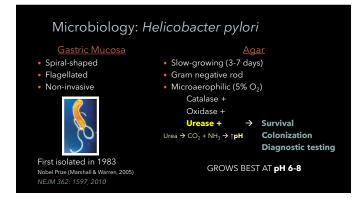
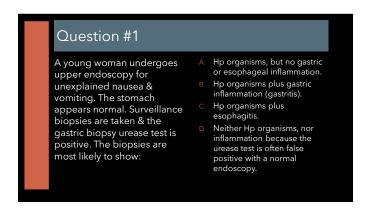
Speaker: David Aronoff, MD









Answer #1 A young woman undergoes Hp organisms, but no gastric upper endoscopy for or esophageal inflammation. Hp organisms plus gastric inflammation (gastritis). unexplained nausea & vomiting. The stomach Hp organisms plus appears normal. Surveillance esophagitis. biopsies are taken & the Neither Hp organisms, nor gastric biopsy urease test is inflammation because the positive. The biopsies are urease test is often false positive with a normal most likely to show: endoscopy.

Question #2

What is the most likely source for humans to acquire H. pylori infection?

A. Perinatally from mother
B. Ingestion of raw vegetables
C. Ingestion of undercooked meat
D. Ingested tap water from a municipal source
E. Contact with infected secretions from another human

Speaker: David Aronoff, MD



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### Helicobacter pylori: Key Points

- Humans are the only natural Hp host
- Infects > 50% of the world's population
- US ~20-40%\*
- A leading chronic infection in humans
- Similar to dental caries
- Majority are asymptomatic but <u>all</u> have chronic active gastritis
- Severity of gastritis varies depending on the Hp strain & the host

 ${}^\star\!At\,greater\,risk:indigenous\,Americans,Black/AA,Hispanic,\&\,immigrants\,from\,high\_cancer-risk$ countries like Japan, Korea, Taiwan & China

### Helicobacter pylori: Key Points

Hp is a carcinogen that causes an inflammationdriven cancer

- 1-3% of infected individuals will develop cancer
- Hp causes 15% of the total cancer burden globally
- Up to 89% of all gastric cancer is attributable to Hp



Lee Y, et al. Annu Rev Med (2022) Shah SC, et al Gastroenterology (2021)

### Transmission of H. pylori

- Exact route of transmission is not known
- Likely **fecal-oral** or **oral-oral**
- Intrafamilial spread (person-to-person, esp. mother-tochild but not during pregnancy)
- Low socioeconomic status, poor sanitation, crowding associated with *†*transmission

### Disease Paths for Helicobacter pylori Infection

Asymptomatic gastritis

85-90%

• Peptic ulcer (DU, GU)

MALT lymphoma

1-17%

• Gastric cancer

0.1-3% <0.01%

DU, duodenal ulcer

GU, gastric ulcer MALT, mucosal-associated lymphoid tissue

### H. pylori: Disease Associations

• #1 cause of chronic gastritis

• PUD: 90% of DU, 80% of GU

• Gastric Cancer (60 - 90%)

• MALT lymphomas (72 - 98%)

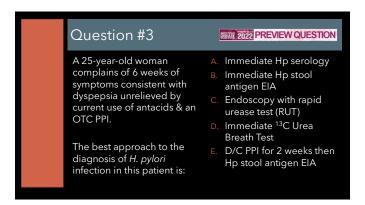
Hр

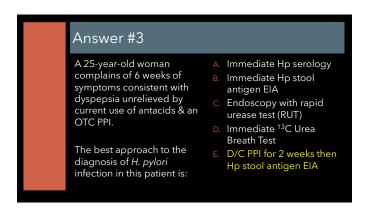
H. pylori is a World Health Organization-designated carcinogen & the strongest nown risk factor for non-cardia

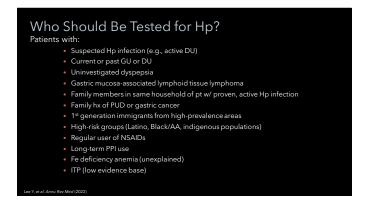
- Iron deficiency anemia, B12 deficiency, ITP
- Eradication Hp neither causes nor exacerbates GERD
- Hp poss. reduces risk for Barrett's esophagus/esophageal CA

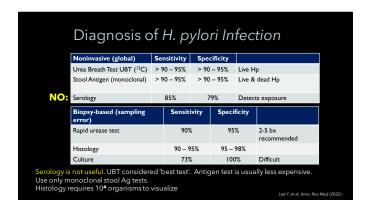
HP is classified by WHO as a Class 1 carcinogen. MALT = mucosal-associated lymphoid tissue

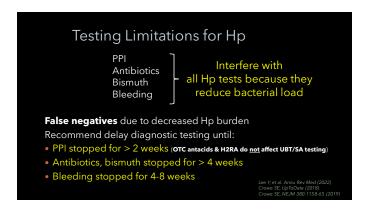
Speaker: David Aronoff, MD

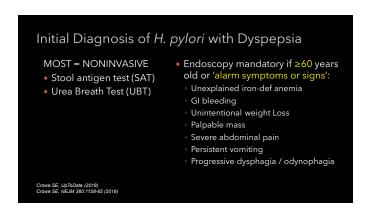










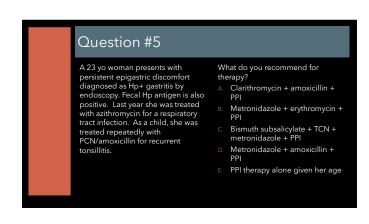


Speaker: David Aronoff, MD

# Which of the following is the most appropriate next step for evaluating a 29-year-old previously healthy but overweight male patient with typical retrosternal heartburn symptoms? A. Stool antigen test for H. pylori B. Urea breath test for H. pylori C. No testing for H. pylori D. Serological testing for H. pylori E. Empiric therapy for H. pylori regardless of testing

# Answer #4 • Which of the following is the most appropriate next step for evaluating a 29-year-old previously healthy but overweight male patient with typical retrosternal heartburn symptoms? • A. Stool antigen test for H. pylori • B. Urea breath test for H. pylori • C. No testing for H. pylori • D. Serological testing for H. pylori • E. Empiric therapy for H. pylori regardless of testing

# Explanation for Q#4 H. pylori is not implicated as an etiological factor in gastroesophageal reflux disease (GERD) Treatment for (eradication of H. pylori) can increase the risk for Barrett's esophagus and esophageal adenocarcinoma Serology is **not** a recommended test for H. pylori



A 23 yo woman presents with persistent epigastric discomfort diagnosed as Hp+ gastritis by endoscopy. Fecal Hp antigen is also positive. Last year she was treated with azithromycin for a respiratory tract infection. As a child, she was treated repeatedly with PCN/amoxicillin for recurrent tonsillitis.

What do you recommend for therapy?

A. Clarithromycin + amoxicillin + PPI

B. Metronidazole + erythromycin + pPI

C. Bismuth subsalicylate + TCN + metronidazole + PPI

D. Metronidazole + amoxicillin + PPI

E. PPI therapy alone given her age

Who should be treated for *H. pylori* infection?

Houston Consensus Conference on Testing for *Helicobacter pylori* Infection in the United States

Hashern B. El-Serag. \*\*John Y. Kao. \*\*Fasiha Kanwal. \*\*L. Mark Gilger. \*\*Frank LoVecotio.\*\*
Storen F. Mosa. \*\*Johna Crows. \*\*Johna Elfant. \*\*Thornas Haas. \*\*Johna Hapise. \*\*Adam Elfant. \*\*Thornas Haas. \*\*Johna Hapise. \*\*Johna

Speaker: David Aronoff, MD

### Takeaways about Treatment of Hp

- Cure rates of most Hp therapies are **relatively low** (e.g., 80% or lower).
- Antibiotic resistance is a HUGE challenge, even with 3-drug therapies, provoking newer quadruple therapies
- Ask about prior antibiotic exposure hx (especially clarithromycin &
- Discuss the critical importance of adherence to treatment
- Use high dose PPI (BID dose; increase gastric pH>4-5)
- Hp grows optimally at pH 6-8
  Acidity hinders stability & activity of macrolides, amoxicillin a lot
  Fast metabolizers of PPIs (CYP2C19 genotypes) reduce levels of omeprazole/lansoprazole
  Vonoprazan is a new potassium-competitive acid blocker that appears very promising

### Takeaways about Treatment of Hp

- Combination therapy is essential
- The optimal duration of Hp therapy is 14 days
- Consider antibiotic susceptibility testing after multiple relapses Culture-based and non-culture-based (NGS) techniques can determine resistance
- Success should always be confirmed by a **test of cure** after treatment of every patient (e.g., UBT performed 4 or more weeks after therapy)

Lee YC. Annu Rev Med (2022)

### Eradication of Helicobacter pylori

- Triple therapy with a PPI, clarithromycin, & amoxicillin or metronidazole is not favored due to increased prevalence of macrolide resistance (but might still be an option on boards!) Clarithromycin resistance in the US now ≥ 15%
- Use a bismuth-based quadruple therapy for 14 days as 1stline therapy:
  - Bismuth subsalicylate or subcitrate
- Tetracycline (not doxycycline: results are inferior)
- Metronidazole

### RIFABUTIN-Based Combinations

- 2020: The FDA approved of omeprazole, amoxicillin & rifabutin (Talicia) for *Hp* treatment in adults
- Omeprazole 10 mg, amoxicillin 250 mg, & rifabutin 12.5 mg
  - The recommended dosage is 4 capsules (with food) every 8 hours for 14 days.
- Summary: Omeprazole/Amoxicillin/Rifabutin (Talicia) A fixed-dose, rifabutin-based, 3-drug combination FDA-approved for treatment of Helicobacter pylori infection.
- First rifabutin-based product to be approved for treatment of H. pylori infection.
- Rifabutin-based triple therapy has been used for years as a salvage regimen for treatment-refractory *H. pylori* infection.
- Approval was based on the results of two trials in treatment-naive patients; H. pylori was eradicated in about 80% of those treated with the combination.
- How the efficacy of Talicia compares to that of other regimens used for first-line treatment of H. pylori infection is unknown.
- Rates of *H. pylori* resistance to rifabutin have been low; whether more widespread use as part of a first-line regimen would result in higher rates of resistance remains to be established.
- Common adverse effects include diarrhea, headache, rash, and

Has the potential to interact with many other drugs

# Eradication of Helicobacter pylori

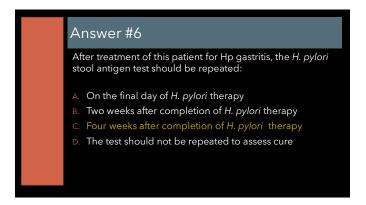
- Fluoroguinolone resistance is common now (>50%)
  - They are not recommended in 1st-line treatment regimens
- Resistance to amoxicillin, tetracycline & rifabutin is uncommon
- Clinical significance of resistance to metronidazole not straightforward

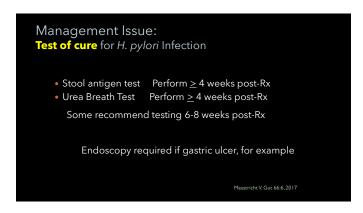
### Question #6

After treatment of this patient for Hp gastritis, the H. pylori stool antigen test should be repeated:

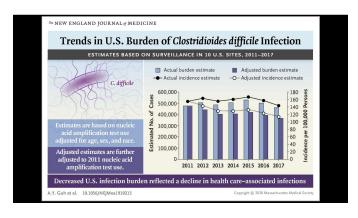
- A. On the final day of *H. pylori* therapy
- B. Two weeks after completion of H. pylori therapy
- Four weeks after completion of H. pylori therapy
- The test should not be repeated to assess cure

Speaker: David Aronoff, MD

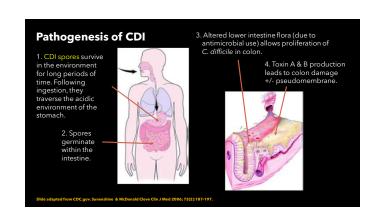




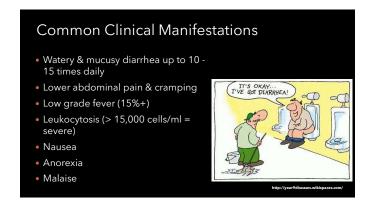




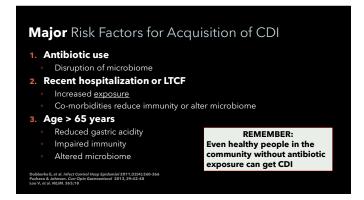
# Antibiotic-Associated Diarrhea (AAD) Common In 5-25% of antibiotic treatment courses especially with > 3 days of Abx but one dose is sufficient 10-40% of AAD is associated with *C. difficile* infection (CDI) but nearly all AA colitis is CDI Disruption of colon microbiome & bile acid physiology are key mechanisms

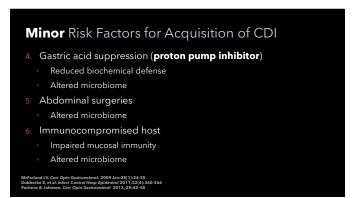


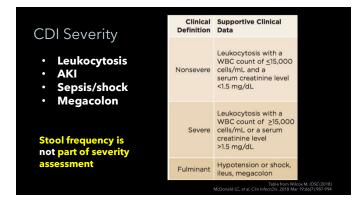
Speaker: David Aronoff, MD

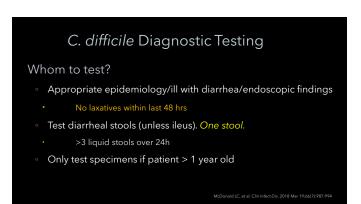




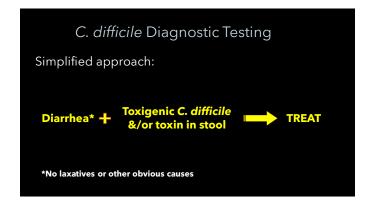


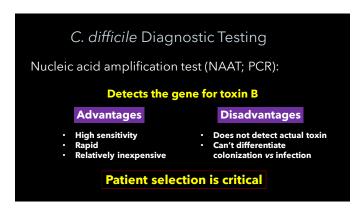


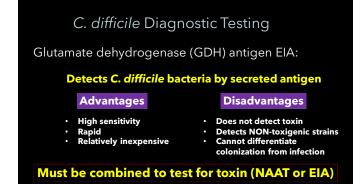


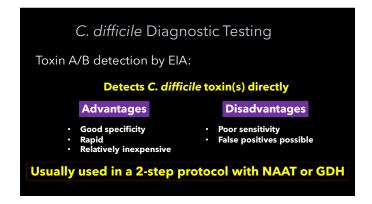


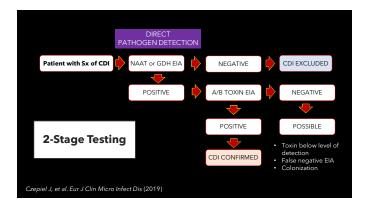
Speaker: David Aronoff, MD

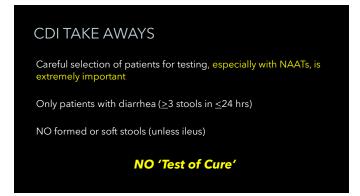






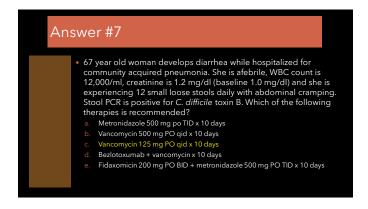


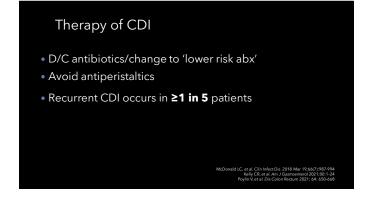


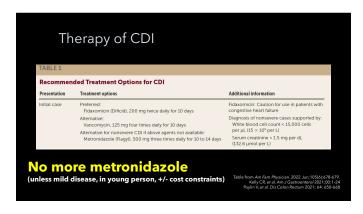


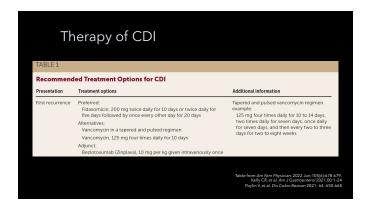
Speaker: David Aronoff, MD

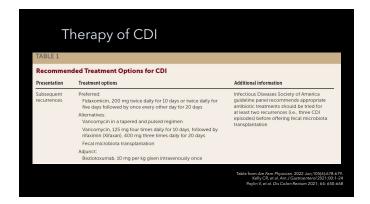
# • 67 year old woman develops diarrhea while hospitalized for community acquired pneumonia. She is afebrile, WBC count is 12,000/ml, creatinine is 1.2 mg/dl (baseline 1.0 mg/dl) and she is experiencing 12 small loose stools daily with abdominal cramping. Stool PCR is positive for C. difficile toxin B. Which of the following therapies is recommended? a. Metronidazole 500 mg po TID x 10 days b. Vancomycin 500 mg PO qid x 10 days c. Vancomycin 125 mg PO qid x 10 days d. Bezlotoxumab + vancomycin x 10 days e. Fidaxomicin 200 mg PO BID + metronidazole 500 mg PO TID x 10 days



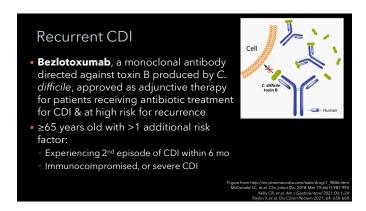


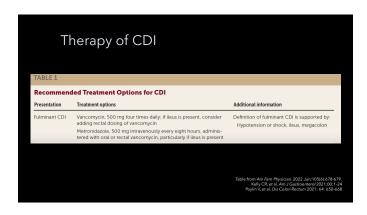






Speaker: David Aronoff, MD





Prevention of *C. difficile* Disease (HCW & visitors)

Contact precautions for patient care.

Gloves, gowns while diarrhea persists.

Single rooms

Handwashing with SOAP & WATER

Alcohol gel rubs do not kill *Cd* spores

Sporocidal solutions for hospital cleaning.

(eg. hypochlorite solutions)

Antibiotic restriction policies

(Antimicrobial stewardship programs).

Lancet ID 17:194, 2017 Scotland Lancet ID 17:411, 2017 England

