



Case Studies: Perplexing Infectious Disease Presentations

Each of us has encountered a number of infectious disease cases that just do not fit the classic textbook description. Using a case studies format, examples of perplexing cases will be presented. There will be a discussion of the evaluation, differential diagnosis, and treatment for each case presented. (Participants can submit cases for consideration.)

- Discuss the differential diagnosis based on the cases presented.
- Describe the appropriate diagnostic work-up and management for each case.

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Baffling Bugs: Mysterious Infectious Disease Cases

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Case: 28 y/o male

Presents with 6 day hx nasal congestion, fever
dry cough, aches, sinus pain

Previously **healthy**

No HIV risk factors

T **38.2°** Pulse 96 BP **115/70** Resp 16

NAD, clear rhinorrhea, **TMs** clear, OP clear

Chest clear

Antibiotics for URIs

Survey of 1529 **MDs** - 20,707 pt. visits

Antibiotics prescribed for:

51% of colds

52% of **URIs**

66% of bronchitis

12,000,000 Antibiotic Rxs for URI /bronchitis
- 21% of all antibiotic prescriptions

Gonzales R, et al. *JAMA* 1997;278:901.

Antibiotics - Patient Expectation

Survey of 113 URI pts. and their **MDs** in clinic.

65% patients expected Abr prior to visit

Physicians were incorrect or uncertain
whether pts. expected Abx 51% of the time

Patient satisfaction not correlated with Rx.
but with perception of time spent by MD

(Hamm et al. *J Fam Pract* 1996;43:56)

Cold Preparations

- Decongestants
 - pseudoephedrine, phenylpropanolamine
- *Antihistamines
- . **Antitussives/Expectorants**
- Vitamin C
- Zinc
- Echinacea

OTC Cold Products

Review of 51 studies 1950-1991:

only 27 met minimal methodology criteria

No **efficacy** of drugs for children < 5

Antihistamines: **4/10** studies showed benefit

Decongestants **4/5** studies showed **benefit**

Expectorants: 1 study failed to show **benefit**

Antitussives: No studies adequate to evaluate

Smith MB, Feldman W. *JAMA* 1993; 269: 2258-63.

Case – 28 y/o male returns

1 week after initial ER visit

Persistent fever, increased sinus pain, purulent nasal discharge

T 38.4° Pulse 92 BP 110/70 Resp 18

NAD, purulent rhinorrhea, sinus tenderness, TMs clear, OP clear, teeth normal

Chest clear

Sinusitis with Colds

90% of viral URIs have associated sinus disease

"Rhinosinusitis"

One billion cases a year in U.S.

~ 1% seek medical care

Estimated that 0.5-2% get bacterial infection

5-20 million cases a year

~ 10% seek medical care

No simple "gold standard" for diagnosis

Sinusitis – Clinical Features

Usual URI symptoms (rhinorrhea, facial pressure, headache, sneezing)

Fever

Purulent nasal discharge

Pain increased when bending forward

Erythema over sinuses

Maxillary toothache

Poor response to decongestants

Timing!!! - > 7 days after URI

Sinusitis - Bacteriology

Strep. Pneumoniae

H. Influenzae

Anaerobes

M. Catarrhalis

Staph. Aureus

Streptococci

Gram-negatives

Sinusitis - Antibiotics

First-line agents:

Amoxicillin

TMP/SMX

Second-line agents

Amox/clav

Azithromycin

2nd/3rd ceph

Clarithromycin

Sinusitis – Ancillary Drugs

Decongestants

Oral preferred over topical

No proven **benefit**, but makes sense

Intranasal Steroids

No **benefit** in clinical trials

Antihistamines – 1st generation

? **Benefit** from anticholinergic effects

Guaifenesin

? benefit

Case – 28 y/o male returns (again!)

3 months after initial ER visit
 Recurrent episodes of sinusitis
 Multiple courses of antibiotics
 TMP/SMX, **Azithromycin**, **Amox/clav**
 Now with fever, facial swelling and pain getting worse after 10 days **Amox/clav**

Case – 36 y/o woman

Abdominal pain for 5 days, fever for 1 day
 Vomited X 2, no diarrhea, no urinary **sx**
 Previously healthy, no prior surgeries
 T **38.6°** Pulse 106 BP **100/70** Resp 20
 Mild distress, HEENT nl, Chest clear
 Abdomen – **Diffuse** tenderness, greatest RLQ
 + Guarding, no rebound, negative **psoas**
 Pelvic – No discharge, +/- CMT. + R **adnexal** tend

Case – 36 y/o woman

CBC: WBC 16.4 **79%P**, 13% L, 6% M
 UA: Dip + ket, 3 WBC, 0 RBC
 Urine pregnancy test negative

Appendicitis - Presentation

M&-analysis of 10 studies, **>4000** pts

Finding	Sens.	Spec. (%)
RLQ pain	61	53
Migration	64	62
Psoas	16	95
Rectal Tend.	41	77
Anorexia	68	36

Appendicitis - CBC

Raftery '76 - (175 pts.)
 WBC **>10**: 92% **sens** 75% **spec**
Sasso '70 - (525 pts.)
 79% of appy pts had WBC **>10**
 62% had elevated PMN count
 96% had one or the other

Appendicitis - CBC

Sensitivity for appendicitis:

	WBC	PMN%	Either
Lau '69	61%		91%
Raftery '76	92%	92%	96%
Sasso '70	79%	62%	96%
Bower '81	07%	92%	96%
Doraiswamy '79	42%	96%	
Lansden '63	64%	60%	

Appendicitis - Ultrasound

Noncompressible appendix
 ≥ 7 mm diameter

Appendicolith

Periappendiceal fluid

Necrosis - loss of wall continuity

Appendicitis - Ultrasound

Sensitivity **76% - 96%**

Specificity 85% - 90%

Accuracy **83% - 96%**

Ultrasound - Other Diagnoses

Ovarian cyst

Ovarian torsion

TOA

Ectopic pregnancy

Cholecystitis

Urolithiasis

Diverticulitis

Intussusception

Ileocectitis

Malignancy

Meckel's diverticulum

Mesenteric adenitis

Ultrasound - Limitations

Operator Dependent

Availability

Indeterminate if poorly visualized

Difficult to exclude appendicitis

Perforation decreases sensitivity

Does not replace repeat exams

Appy: CT vs. UTZ

100 pts., 54 had appendicitis

	CT	UTZ (%)
Sensitivity	96	76
specificity	89	91
Accuracy	94	83
PPV	96	95
NPV	95	76

Discordant **results** in 20:

CT correct in 17, UTZ correct in 3

Balthazar EJ et al. *Radiology* 1994;190:31

Dog and Cat Bite Infections: Organisms

Cat		Dog	
<i>Pasteurella</i>	78%	<i>Pasteurella</i>	54%
<i>strep</i>	52%	<i>strep</i>	46%
Staph	33%	Staph	36%
<i>Fusobacterium</i>	30%	<i>Fusobacterium</i>	36%
<i>Bacteroides</i>	22%	<i>Bacteroides</i>	32%
<i>Porphyromon.</i>	22%	<i>Porphyromon.</i>	32%
<i>Prevotella</i>	22%	<i>Peptostrep.</i>	23%

Talan DA, NEJM 1999;340:85

Pasteurella -Antibiotic susceptibility

What Works

MIC₉₀ (μg/ml)

Pen G	0.006-0.25
Amoxicillin	0.25
Ciprofloxacin	0.008-0.03
Trovaflaxacin	0.03
Cefuroxime	0.06
Azithromycin	0.125-0.5

Goldstein EJC *Antimicrob Agents Chemother* 1998;41:391

Pasteurella -Antibiotic susceptibility

What Doesn't Work

MIC₉₀ (μg/ml)

Cephalexin	1-16
Erythromycin	4-8
Clarithromycin	1-4
Dicloxacillin	8-16
Nafcillin	4
Clindamycin	32
Gentamicin	2

Goldstein EJC *Antimicrob Agents Chemother* 1997;41:1552.

Dog and Cat Bite Wound Infections: Empirical Antimicrobials

	Pasteur.	Staph	Anaerobes
Amoxicillin/clav.	+	+	+
Ampicillin/sulbactam	+	+	+
Cefotetan/cefox.	+	+	+
Azithromycin	+	+	+/-
Trovaflaxacin	+	+	+
Pen/cephalexin	+	+	+/-
Clindamycin/cipro	+	+	+

Selected Reading

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Baffling Bugs: Mysterious ID Cases

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Case: Double Trouble

4 m/o male with **cough** and vomiting

HPI

- 3 days non-productive cough
- 1 day tactile fever and post-tussive emesis
- No blood or bile in emesis
- No diarrhea
- No travel, ill contacts, hemoptysis

Past Medical History

- Previously-healthy
- Full term, NSVD
- No medications
- No allergies
- No immunizations

Physical Exam

VS -- T 37.2, HR 120, RR 32, Pulse Ox 99%
Gen -- NAD, vomited "yellow stomach contents"
HEENT -- AFOF, TMs and OP normal, MMM
Neck -- supple, no adenopathy
Lungs -- Clear, no retractions

Physical Exam

CV -- RRR without murmur
Abd -- soft, ND/NT, no HSM or mass
GU -- normal circumcised male
Ext -- no cyanosis or edema, CR < 2 secs
Neuro -- awake, alert, interactive, nonfocal
Skin -- no rash

Initial Management

- Dx viral URI
- Post-tussive emesis
- Discharge **home**
- Follow-up PMD within 1 week

The Plot Thickens

- Mom calls PMD due to more vomiting and coughing
- Coughing continuously over the phone
- Returns to ER -- Temp 38.7. HR 166
- General physical exam unchanged
- Rectal exam -- maroon stools, heme +

Lab Testing

WBC 19,100
Diff -- 45% PMN, 10% Bands, 34% Lymphs
Hct 36%
PLT 495,000
Chcm-7 normal
UA normal

Hospital Course

- Abdominal series -- normal
- BE -- ileocolic intussusception (reduced)
- NP swabs for *B. pertussis* DFA and culture
- Admitted for observation and Erythromycin
- Respiratory isolation

Hospital Course

HD 4

- first NP culture negative
- D/C home on p.o. ERY x10 days
- Respiratory isolation for 5 days (total)

7 days after D/C

- NP swab positive for 1 colony *B. pertussis*

Pertussis -- Overview

- Whooping cough
- *Bordetella pertussis* -- Gram neg bacterium
- Household attack rates >90%
- Often unrecognized and unreported
- & 3 months 5 years (peak 6-11 months)
- Most are not fully immunized
- Older children and adults are reservoirs

Pathophysiology

- Fimbriae attach to respiratory epithelium
- Toxin production
 - Pertussis toxin
 - Adenylate cyclase toxin
 - Cilio-tracheal toxin
 - Dermonecrotic toxin
- Loss of ciliary function
- Debris and mucous accumulation
- Pneumonia and bronchiectasis

Clinical Presentation

Catarrhal stage

- Follows incubation period 6-20 days
- Lasts 1-2 weeks
- Rhinorrhea, lacrimation, mild cough
- Fclet usually <101F

Clinical Presentation

Paroxysmal stage

- Lasts 2-4 weeks
- Paroxysmal cough (5-10)
- Inspiratory "whoop" (not all patients)
- Cyanosis, plethora, diaphoresis, tongue protrusion
- Highest morbidity and mortality

Clinical Presentation

Convalescent stage

- Lasts 1-4 weeks
- Symptoms gradually improve
- Cough may persist up to 6 months

Complications

- Pneumonia most common (14% of admits)
- CNS complications from hypoxia and toxins
- Seizures (3%)
- Encephalitis (1%)
 - » 1/3 fatal
 - » 1/3 serious sequelae
 - » 1/3 no sequelae
- Rectal prolapse, hernia, PTX, hemorrhage

Ancillary Data

- initial diagnosis is clinical
- WBC > 15,000 with lymphocytosis
- Less common age < 6 months
- NP culture on Bordet-Gengou media

Culture Sensitivity

Time	Sensitivity
Shortly after exposure	67 - 81%
Week 3	25%
Week 4	14%
Week 5	0%

Other Tests

- . DFA -- less sensitive, much less specific
- . Antibody titer to virulence factors -- most sensitive
- . PCR -- high sensitivity and specificity, fast, less available

Bordetella in Adults

Wright, et al. *JAMA* 1995

- 75 ED patients with cough > 2 weeks
- 21% met serologic criteria for pertussis
- . Symptoms/lymphocyte count not helpful

- Benign disease in adults

Nenning, et al. *JAMA* 1996

- Incidence in adults similar to PUD

Management

- Antibiotics to eradicate organism
- Onl> ameliorates disease in catarrhal stage
- ERY 50 mg/kg/d x 14 days
- Isolate patients for 5 days after starting ERY
- Alternatives --other macrolides. TMP/SMX
- Household and close contact PEP

Patient Disposition

- Admit age < 6 months and clinical penussis
- Discharge others unless complications

Pearls and Pitfalls

- Easy to miss if not on differential
- Pay attention to immunization status
- Less typical in age < 6 months and adults
- Disease most serious in age < 6 months
- Always send culture with antibody tests
- Fever > 101F uncommon

Intussusception -- Overview

- Common abdominal emergency in infants
- Prolapse of intestine into adjacent lumen
- Spontaneous reduction is rare
- Survival rate 99% if timely treatment

Pathophysiology

- Intussusceptum = invaginating portion
- Intussusciens = receiving portion
- Lead point proximal to ileocecal valve
- Ileocolic invagination most common

Pathophysiology

- Mesentery dragged with intussusceptum
- . Venous return obstructed
 - Mucosal edema and bleeding ("currant jelly" stools)
- Arterial occlusion
- Intestinal gangrene and perforation

Etiology

- Male:Female = 2:1
- Ages 3 months - 5 years
- 60% age < 1 year
- Peaks incidence spring and fall
- Age < 2 years -- Peyer's patches (<10% pathologic)
- Age > 5 years -- 75% underlying lesion
 - Polyp, ileal duplication, lymphoma, inverted Meckel's, HSP hematoma

Rotavirus Vaccine (RRV-TV)

- . 9/98 - 7/99: 15 cases of intussusception
 - . Ages 2-11 months
 - . 87% after first dose (RotaShield)
 - . 80% within 1 week of vaccine
 - . 8/15 required surgical resection
 - . Lymphoid hyperplasia
 - . NCKP/Minnesota data suggest causation
- MMWR July 1999*

Diagnostic Findings

- Classic Triad (20-40%) -- colicky abdominal pain, vomiting, bloody stool
- Abdominal pain -- 50-90%, usually first symptom, 4-5 minute episodes, knees to chest, 10-20 minute rest periods
- vomiting -- 60-90%, often bilious
- Bloody stools -- 20-60%, usually after 12 hours of pain

Other Exam Findings

- Elongated RUQ mass
- Absent bowel sounds in RLQ (Dance's sign)
- May palpate intussusceptum in rectum
- Fever to 103°F may develop
- May present as lethargy, pallor, AMS

Ancillary Tests

- Labs -- nonspecific
- Abdominal films -- normal early in disease. later may see obstruction. soft tissue mass. li-cc air
- Barium enema -- diagnostic and therapeutic, "cervix-like mass", "coiled-spring" appear once of distal ileum after reduction
- Air enema -- increasing in popularity use depends upon radiologist's preference

Ultrasound

- Detect intussusception / confirm reduction
- "Target sign"
 - Hypoechoic ring / hyperechoic center
- "Pseudokidney sign"
 - Hypoechoic walls / hyperechoic mucosal layers
- One study of 65 patients
 - Sensitivity 100% (95% CI: 83-100%)
 - Specificity 93% (95% CI: 81-99%)

Bhisitkul DM, et al, *J Peds* 1992

Management

- General -- IV fluids, electrolytes (K+), nasogastric tube
- Surgical consultation
- Barium enema -- success rate 50-90%
- Analgesics prn with BE

Pearls and Pitfalls

- If no diarrhea, no "gastroenteritis"
- Vomiting without diarrhea age < 1 year = rectal exam
- Bilious vomiting (yellow/green) is not normal in young children
- Most do not have classic triad of colicky pain, vomiting, and bloody stools

Pearls and Pitfalls

- Pitfalls may appear well between episodes
- BE contraindicated in perforation or shock
- Pathologic lead point very common in older children (age > 5 years)
- Consider if recent RRV-1 V and symptoms

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