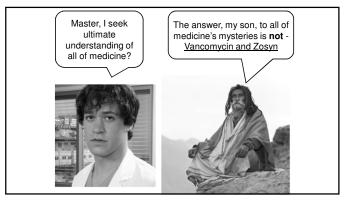
### Paul Carson, M.D. Sanford Health NDSU Dept. of Public Health

1

Problems that Should Raise
Your Sphincter Tone

ว



ID - 1st Principles

Fever or Increased WBC alone ≠ Emergency

Fever ≠ Antibiotics ≠ Antipyretics

3

# When Are Antibiotics Really Necessary? After appropriate cultures expeditiously obtained... > You have a target > Patient is "septic, shocky, toxic" > Patient is immunosuppressed • Transplant recipient • Neutropenic • AIDS with low CD4 • High dose steroids or "mabs" • Splenectomized

SIRS w or w/o Infection w Sepsis Septic Shock MODS infection or w/o SIRS

> SOFA - PaO2/FiO2, Glasgow coma scale, MAP, Bilirubin, Platelets, Cr
2 or more points = "Sepsis"

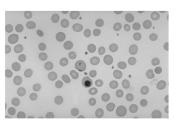
> gSOFA - 1 point each for: SBP < 100, RR > 22, Altered mentation
2 or more points associated with poor outcome

5

"The prescription comes too late when the disease has gained strength by long delays"

> Ovid (43 BC - 18 AD) Remedia Anoris

24 y.o presents to ER with 24 hrs fever. No other sx's or signs. Past hx remarkable for MVA 5 yrs ago with expl. laparotomy. Lab tech notes findings below incidentally on CBC. Why should you be worried??



"Create her child of spleen; that it may live"

William Shakespeare
The Tragedy of King Lear

#### **Post-Splenectomy Sepsis**

Overall incidence is lowChildren: 0.13-8.1%Adults: 0.28-1.9%

8

- Higher if ass'd with hemoglobinopathy or malignancy, lower if trauma
- > 50-70% occur in 1st 2 years after splenectomy
- > Risk increase 6x if prior episode of PSS

9 10

#### **OPSI - When to Think of It**

Fever plus......

7

- > Known past splenectomy
- > Functional Hyposplensism
  - Sickle Cell Dz, Hemoglobinopathies
  - Amyloidosis
  - SLE and RA
  - Acute alcohol use
- > Howell-Jolly bodies or nucleated rbc's on blood smear

**OPSI - Organisms** 

- ➤ Encapsulated bacteria (≅ 75%) "S-N-K-H-S-C-P"
- Pneumococcus over 50%
- > Capnocytophaga canimorsus
- > Babesiosis and Malaria
- > Bordatella holmesii
- > Other bacteria
- > Empiric Rx: High dose Ceftriaxone and Vanco

11 12

#### **Antibiotic Prophylaxis**

> Controversial

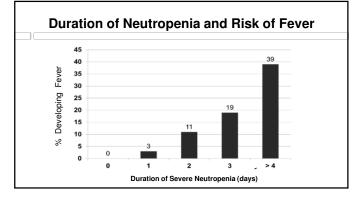
13

- > Children < 15 in 1st 3-5 years post-splenectomy
- Patient-initiated therapy for fever in adults followed by prompt evaluation
  - Amoxicillin/clavulanate 875mg po
  - Cefuroxime 1 gm po
  - Levofloxacin 750mg po
- Consider lifelong prophx in survivors of PSS

### **Case History**

14

45 y.o. male recently diagnosed with non-Hodgkins lymphoma. Started 1st cycle of CHOP chemoRx 10 days ago. Now presents with fever, chills, sore mouth, diarrhea, nausea/vomiting. Exam shows T-102 degrees and mild abdominal distension and tenderness. What is your initial impression and management?



Severity of Neutropenia and **Episodes of Severe Infection** 40 35 30 25 20 18 Infection 15 10 < 100 101-500 501-1000 > 1000 **Absolute Neutrophil Count** 

15 16

#### Source of Infection in Febrile Neutropenia

- ➤ Infectious source only found in 30% of cases
- > Bacteremia documented in 25% of patients
- > 80% of infections arise from patient's endogenous flora

#### **Common Sites of Neutropenic Infection**

Site	Percent
Mouth/Pharynx Respiratory tract Skin/Soft tissue/IV sites Perineal region Urinary tract Nose/Sinuses GI	25 25 15 10 8 5

Note: Exam very important. Classic findings of inflammatory response will be missing. Pay close attention to pain, redness or induration

17 18

#### Positive BC's in Febrile **Granulocytopenic Patients** % GNB %GPB 1980's 60 25 1995 22 62 2000 14 76

#### **Treatment of Febrile Neutropenia**

- > Targeted therapy if clinically apparent
- Empiric Therapy
  - Monotherapy (Cefepime, Imepenem, Meropenem, Pip-Tazo). Use <u>full</u> <u>dosing anti-pseudomonal dosing</u>
- > Vancomycin
- If suspected line infection, SSTI, or OP source
- Gram positives growing Still febrile at 72 hrs
- > Antifungals add if still febrile after 5 days

19 20

#### **Case History:**

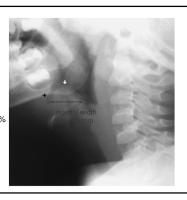
28 yo female has 1 week history of sore throat and low grade fever. Over last 2 days, pain is severe, trouble eating and drinking. Exam shows toxic looking woman with a T-103, drooling, difficulty opening mouth wide. OP appears modestly reddened but nothing else.

Differential diagnosis? Next Step?

Fever and stridor or inability to handle secretions are warning signs

Thumb sign on soft-tissue lateral neck

Sensitivity of plain X-ray: 43-70%



22 21

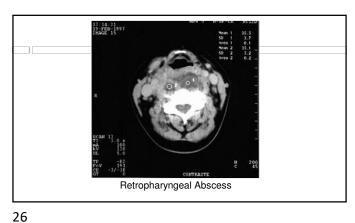
#### **Epiglottitis**

- ➤ Usually caused by H. influenza, S. pneumo, S. aureus (MRSA)
- Past: children > adults. Since Hib vaccine incidence equivalent or higher in adults with *S. pneumo* predominating
- > Dx best made with laryngoscopy in a controlled setting
- Ready for urgent intubation and surgical airway if needed
- > Airway control takes precedence over diagnostic evaluation
- > Treatment: 2nd or 3rd generation CP and Vancomycin

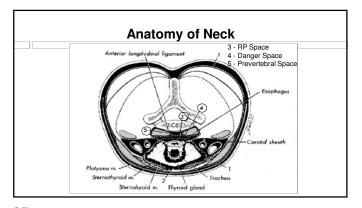


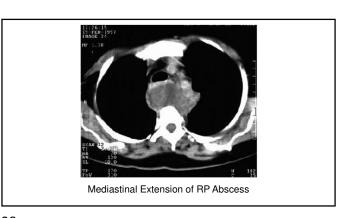
#### **Retropharyngeal Space Infection**

- > Usually due to mixed aerobic and anaerobic oral flora
- > Risk of entering "Danger Space"



25





27 28

#### **Retropharyngeal Space Infection**

- > Diagnosis: CT scan of the neck
- > X-ray of soft-tissue lateral neck
  - RP space is normally < 1/2 adjacent vertebral body
- Management
- Abx: ampicillin sulbactam, clindamycin, or carbapenem
- Involve ENT surgeon, possibly CV surgeon

#### **Case History**

22 yo female college student presents to U.C. with 3 day history of temps. to 103, arthralgias, myalgias, headache, and malaise. In the last 24 hours she has noticed an unusual rash. PMH is unremarkable. Exam is only noteworthy for fever and a purple blotchy rash.

29 30



#### Fever and Purpura/Petechial Rash

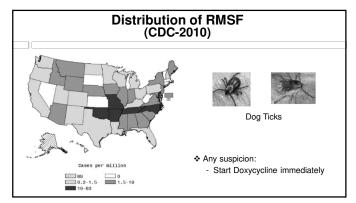
- >Life threatening infection until proven otherwise
  - Meningococcal sepsis +/- meningitis
  - Rocky Mountain Spotted Fever

#### ➤Other diagnoses:

- Vasculitis
- DIC
- TTP

32

- Febrile illness with thrombocytopenia



#### **Meningococcal Sepsis - Management**

- > Patient should be placed in respiratory isolation
- > Blood cultures obtained immediately
- > Gram stain of buffy coat
- Empiric penicillin G or CNS penetrating cephalosporin while awaiting cultures
- > Lumbar puncture

34

33

#### **Case History**

45 y.o. construction worker fell off of scaffolding hit left thigh on the ground. No obvious trauma or break in skin. Over next 48 hours pain becomes worse in leg and he then spikes a temp of 104 degrees with chills. He presents to the ER and looks toxic. Temp is 103, P-110, BP- 98/40. Exam reveals following appearance to the extremity......

Differential diagnosis? Next Step?

35 36

"Beauty is only skin deep, but ugly goes all the way to the bone"

Anonymous

#### When to Suspect Necrotizing Fasciitis

- > Exquisite pain
- > Patient looks toxic
- > "Woody" induration
- > Bullae
- > Fixed dusky purple/grayish discoloration

37 38

#### **Prediction Rule for Nec Fasc** Number of **Clinical Parameter Points** CRP ≥ 150 mg/dL WBC 15-25k > 25k 2 Hgb 11-13.5 g/dL < 11 g/dL Serum Sodium < 135 meq/L Serum Creat > 1.6 mg/dL Serum glucose > 180 mg/dL 80% probability of Nec Fasc Possible Nec Fasc 7-10% cases of Nec Fasc Score > 8 Score 6-8 Score < 6

#### **Necrotizing Fasciitis - Management**

- > Diagnosis rests on clinical exam and ultimately surgical exploration
- > Involve surgeon promptly
- Diagnosis +/- aided by:
- MRI, CT, bedside incision and probing, frozen biopsy
- Antibiotics: Clindamycin + PCN? IV immunoglobulin

39 40





41 42

60 yo Chinese male who works for John Deere and travels regularly to and from China and India presents to WIC with fever, chills, headache 6 weeks after return from a recent purchasing trip to southern India. History is also significant for hiking in Itasca Park in MN 3 weeks before becoming ill. Hematology techs note abnormality on blood smear.

Diagnosis?

Next Step?

43

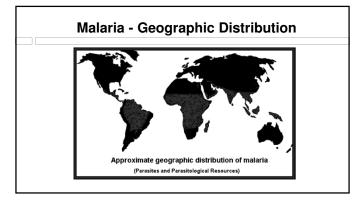
#### Falciparum Malaria - Diagnosis

- > High index of suspicion
- > Anemia and thrombocytopenia common
- Dx usually can be made with 2-3 sets of thick and thin blood smears over 48 hr period
- > Thick smears are sensitive for presence of malaria
- > Thin smears identify morphology and degree of parasitemia

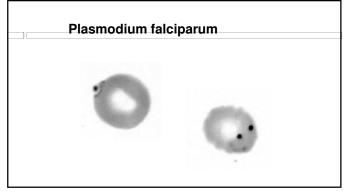
44

#### **Suspected Malaria - Management**

- Febrile patient from endemic area in last 6 weeks without obvious source.... P. falciparum until proven otherwise
- > Potentially life-threatening
- > Admit for observation while smears being obtained
- > Once diagnosis made, immediately begin:
  - Atovaquone/proguanil or Artemether-lumefantrine or quinine+doxycycline
  - Chloroquine in limited areas of Caribbean and Central America



45 46

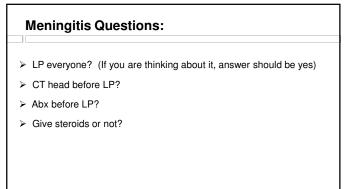


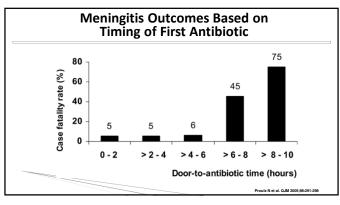
Plasmodium falciparum

Plasmodium falciparum

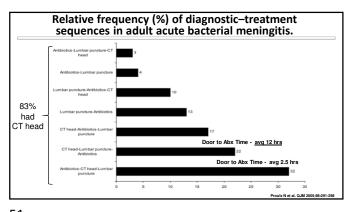
(original image provided by Steve Aley)

47 48





49 50



#### 

NEJM 2001;345:1727-33.

51 52

# Treatment of Meningitis - General Principles > Delay in therapy increases morbidity and mortality > Try to obtain LP 1st, but don't delay therapy beyond 30' after presentation > Use maximum doses of abx > Get BCs before the abx if LP must be delayed

Steroids in Meningitis

Clear ↓ morbidity in children with *H. flu* meningitis, conflicting results in pneumococcal infxn

Benefit also unclear with pneumococcus in adults, however, one blinded prospective study showed clear benefit with pneumococcus

Dexamethasone 0.15 mg/kg q6h x4d should be started just before or concomitant with antibiotics in both children and adults

Don't give if antibiotics were already given prior

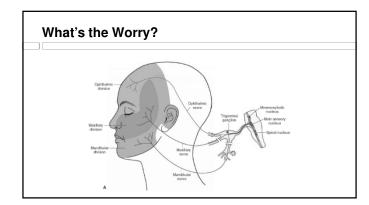
May ↓ penetration of vancomycin

Discontinue in adults if NOT pneumococcus

53 54

78 y.o. female presents with 2 days of tingling and burning pain in her R upper face. She then notices these lesions on her nose. Why should you be concerned?





55 56



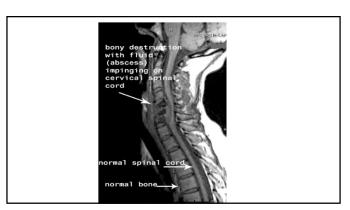


57 58

### **Case History**

55 y.o. male admitted to the hospital with 8 weeks of progressively severe upper neck pain. Has been seen in WIC and ER multiple time in past 2 mos with dx of MS neck pain. Pain became unbearable and now he is having LUE and LLE weakness of the last 6 hrs.

DDx? Next Step?



59 60



#### **Epidural Abscess - Clinical Manifestations**

- Classic triad: exquisite localized back pain, progressive neurologic deficit, and fever
- > Fever may only be present in 50%
- $\succ$  Typical course: Pain and radicular sx's 1st 3d  $\rightarrow$  weakness over 36h  $\rightarrow$  paralysis over next 24h

61 62

#### **Epidural Abscess-Diagnosis**

- ➤ Think of it!
- Note: Most patients will try to ascribe pain to some inciting event, no matter how trivial
- > ESR universally elevated
- > MRI diagnostic modality of choice

#### **Epidural Abscess - Management**

- > Identify organism:
  - BC's x 2-3 before antibiotics
  - Aspirate of abscess
  - PPD if suspicious
- > Empiric antibiotics
  - Vancomycin ± Ceftriaxone or Ciprofloxacin
  - Should be started ASAP
- > Neurosurgical consultation ASAP

63 64

#### **Epidural Abcess-Outcome**

- > Overall mortality 18%, up to 38% for cervical abscess
- Paralysis
  - Less frequent with lumbar disease
  - Some probability for recovery if treated within 24h or incomplete cord lesion manifested by sensory sparing
  - No recovery if Rx >36h after onset

### Emergent Infectious Diseases of the Nervous System

- >Acute bacterial meningitis
- ➤ Encephalitis (HSV)
- ➤ Epidural abscess
- ➤ Brain abscess

65 66

Acute Bacterial Meningitis - Presentation				
Finding	Frequency %			
Fever, headache, meningusmus*	85			
Altered mental status*	50-80			
Kernig's or Brudzinski's sign	50			
Focal cerebral signs	10-20			
Seizures	30			
Rash with meningococcemia	50			
*Meningusmus may be subtle, esp. in eld Confusion more prominent in elderly.	lerly.			

Lumbar Puncture

➤ If you think of it, it should probably be done

➤ Should be done ASAP

➤ Image head first IF:

■ papilledema present

■ focal neurologic signs

■ suspected ↑ intracranial pressure (coma, bradycardia, hypertension, CN III palsy)

67 68

## Possible Tests to Order on CSF? > Gram Stain, routine culture > AFB smear and culture > Cryptococcal antigen > Latex agglutination tests > Fungal culture > PCR for HSV, Enterovirus, CMV, EBV, VZV, JC virus, Lyme, Toxoplasma > VDRL, Lyme titer, viral cultures > C-Reactive Protein

69 70

> Oligoclonal bands, Myelin Basic Protein

## Tests to Order on CSF If cell count, protein, or glucose abnormal, consider in the appropriate clinical context: • AFB smear and culture • Cryptococcal antigen • Fungal culture • PCR for HSV, Enterovirus, CMV, EBV, VZV, JC virus, Lyme • VDRL, Lyme titer, viral cultures • C-Reactive Protein

Empiric Therapy for ABM				
	Predisposing Factor	Antibiotic(s)		
	Age 18-50 yrs	3rd Gen CP +/- vanco		
	Age > 50 yrs	3rd Gen CP + Amp +/- vanco		
	Immunocompromised	Vanco + ceftazidime + Amp		
	Head trauma, neuro- surgery, CSF shunt	Vanco + ceftazidime		

71 72

#### Steroids in Meningitis

- Adult benefit only with S. pneumoniae in the developed world (decreased mortality from sepsis and shock). Greatest benefit on the sickest patients (Glasgow coma scale of 8-11 on presentation).
- >Must be given prior to or along with 1st dose Abx
- ➤ Dose Dexamethasone is 0.15 mg/kg q 6 hrs x 4 days
- >Stop early if not pneumococcal meningitis

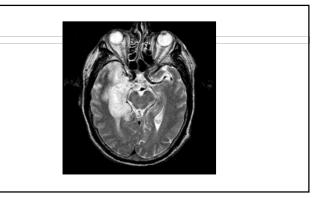
#### **Case History**

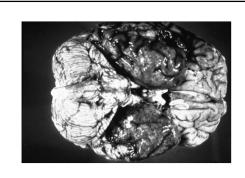
60 y.o. male presents to E.D. with 3 day hx of progressive headache and fever to 102 degrees. He had complained of some mild neck stiffness. Over the last 24 hours he has become progressively confused, somnolent, and at times combative. His wife has noticed he is not using his left arm much.

DDx?

Next Step?

73 74





75 76

#### **HSV Encephalitis**

- > Was most common of viral encephalitides (now WNV)
- > 10-20% of all viral encephalitides
- > Frequency in U.S. ~ 1/250-500,000 persons/yr
- > HSV-1 principal agent
- > Occurs in all ages throughout the year

#### **HSV Encephalitis – Clinical Presentation**

- > Altered mentation
- > Decreasing level of consciousness
- > Focal neurologic findings: dysphasia, weakness, paresthesias
- > Fever and personality changes uniformly present
- ➤ Seizures in up to 2/3

77 78

## Diagnosis of HSV Encephalitis >EEG • Sensitivity 84% • Characteristic spike-and-slow wave activity and PLED's focally >CT head • Mass effect in temporal lobe in 50-75% at some time >MRI • v. sensitive, asymmetric early white matter changes >HSV PCR for CSF • diagnostic test of choice • highly sensitive • replaced brain biopsy

HSV Encephalitis Treatment

Acyclovir 10 mg/kg q8h x 10-14d

Consider empiric therapy in any suspected encephalitis pending EEG/MRI/PCR studies

79 80

HSV Encephalitis-Outcome			
	Treated	Untreated	
Mortality	70-80%	30%	
Neurologic sequelae	80-90%	50-60%	