# We Can Work It Out





What Is Possible

3 hospitals, over 25 locations and more than 2,000 doctors. Find one close to you.

# Hiding in Plain Sight....Part 2





## Medical History

- Normal pregnancy/birth
- Healthy through all newborn well check
   appointments
  - April 8 (time=0)
    - Reported to primary care provider with complaints of high fever (101-104°C)for two days
    - UTI ruled out
    - Tested positive for influenza B via rapid test
    - Decision made to treat symptoms



- Medical History
  - April 16 (time= 1 week)
    - Reported to primary care provider for routine 6month well baby check
    - No concerns
      - Fever had resolved



- Medical History
  - -May 4 (time = 4 weeks)
    - Reported to primary care provider with complaints of:
      - Two day history of sinus congestion, greenish nasal discharge, fever, fussiness, decreased appetite, vomiting
      - Temperature 102.9°F
    - Tested positive for influenza B
      - MD believes still positive from previous infection and current symptoms caused by other virus.



- Medical History
  - May 6
    - Reported to primary care provider for complaints of
      - Continued high fever
      - Temperature 102.9°F
      - Not eating well
      - Vomiting
      - Cold symptoms resolved
      - No signs of bacteria infection (otitis or pneumonia)



- Medical History
  - May 7
    - Reported to primary care provider for complaints of
      - Continued high fever
      - Lack of activity
      - "lazy eye", not looking directly at mother
      - Moaning
      - Diarrhea
      - Not eating well
    - Patient walked by provider to ED



### • ED evaluation (5/7)

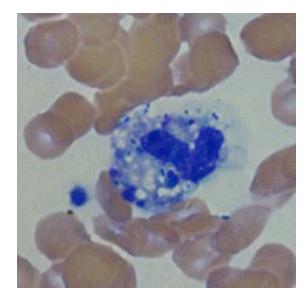
- Fever
- Altered mental status
- Not responsive to staff
- ED course
  - Bolus of fluids
  - Routine labs
  - Blood cultures
  - Head CT
    - No significant findings
  - Patient transferred to Children's Hospital of WI

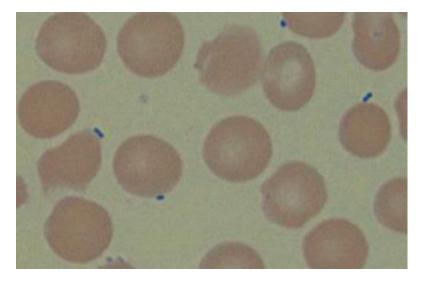


- Significant Laboratory results
  - UA
    - Not significant for infection
  - Chemistries
    - Sodium 130 (slightly low)
    - Bicarb 16 (low)
    - Creat 0.21 (low)
    - Other results within normal ranges
  - WBC 9.7
    - Unable to perform differential due to deteriorating cells
    - Bacteria observed on smear
    - Path review ordered
  - Pathology review of slide
    - Confirms presence of diplococci



- Significant Laboratory results
  - Intra and Extracellular bacteria present





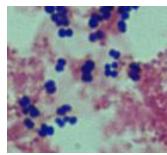


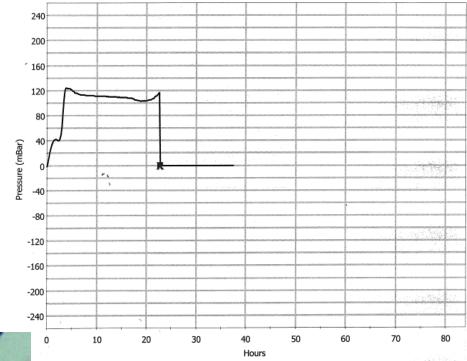
# Microbiology

course

- Blood culture positive at 22.9 hours
- Gram stain Gram positive cocci in clusters
- Verigene Strep pneumoniae







## Microbiology culture

 Subculture from positive blood culture bottle





## Microbiology culture

Subculture from original EDTA tube

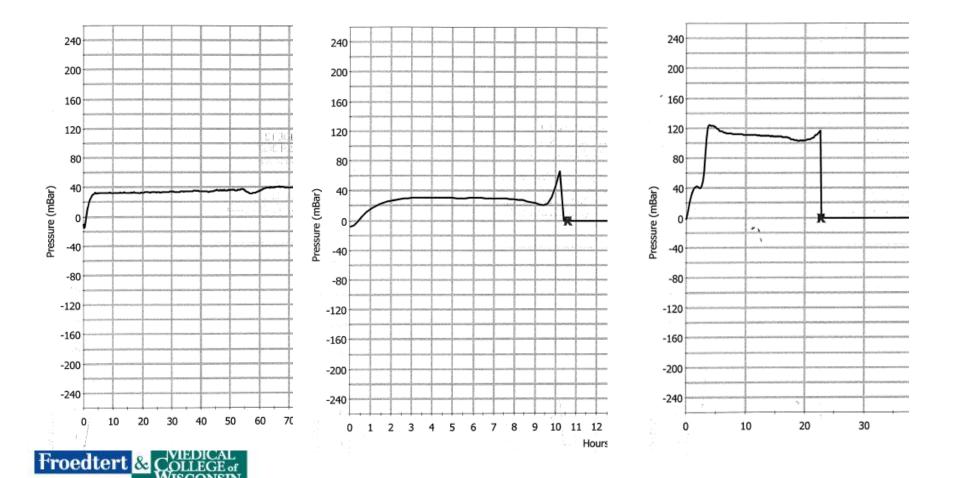




## Microbiology culture

# – Why was S pneumo not isolated from blood culture bottles?













#### A Review of Pneumococcal Vaccines: Current Polysaccharide Vaccine Recommendations and Future Protein Antigens

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## Pneumococci virulence

- Polysaccharide chains
  - Protect organism from phagocytosis
  - Highly immunogenic
  - 92 different subunits
    - Organisms express one



Table 1. Pneumococcal Vaccine Approval Dates, Serotypes, and General Effect on

Vaccine FDA Approval		Serotypes Contained in Vaccine*†
PPSV23	June 1983	1, 2,* 3, 4, 5, 6B, 7F, 8,* 9N,* 9V, 10A,* 11A,* 12F,* 14, 15B,* 17F,* 18C, 19A, 19F, 20,* 22F,* 23F, and 33F*

Recommendations

,

- All adults >65 y.o.
- At risk adults <65 y.o.</li>
- At risk children
  - Chronic heart or lung disease
  - Diabetes
  - CSF leaks
  - Cochlear implant
  - Asplenia
  - Immunocompromised

- Efficacy
  - Reduced infections with covered serotypes by 65%
  - Did not reduce carriage rate
  - Did not generate immune response in children <2 y.o.</li>



Vaccine	FDA Approval	Serotypes Contained in Vaccine*†	
PPSV23 June 1983		1, 2,* 3, 4, 5, 6B, 7F, 8,* 9N,* 9V, 10A,* 11A,* 12F,* 14, 15B,* 17F,* 18C, 19A, 19F, 20,* 22F,* 23F, and 33F*	

PCV7 February 2000 4, 6B, 9V, 14, 18C, 19F, and 23F

### Polysaccharide Conjugate Vaccine

- T-cell dependent antibody response
  - Effective in children <2
  - Generated higher antibody titers



PCV7 February 2000 4, 6B, 9V, 14, 18C, 19F, and 23F

#### Efficacy

- Reduced infection rate in vaccinated children
- Reduced infection rates in unvaccinated children via herd effect
- Hospitalizations due to pneumonia reduced in immunized children and elderly
- Reduced carriage rates in vaccinated and unvaccinated children and household members
- Did NOT reduce otitis media rate



# Smart bugs?

- After PCV7
  - 5 years post implementation
    - Serotype replacement noted
      - Increase in 19A
      - Infection rates returned to pre vaccine levels in high risk groups
    - Resistance rates increased
- 2010 PCV13

- 1, 3, 4, 5, 6A, 6B, 7F, 9V, 14, 18C, 19A, 19F, 23F



# Smart bugs?

- After PCV7
  - 5 years post introduction
    - Serotype replacement noted
    - Infection rates returned to pre vaccine levels in high risk groups
    - Resistance rates increased
- 2010 PCV13
  - 1, 3, 4, 5, 6A, 6B, 7F, 9V, 14, 18C, 19A, 19F, 23F
  - Included 6 most prevalent serotypes not included in PCV7
  - Post introduction Increase in only one serotype
    - 35B



Table 1. Pneumococcal Vaccine Approval Dates, Serotypes, and General Effect on Pneumococcal Disease

Vaccine	FDA Approval	Serotypes Contained in Vaccine*†	Pneumococcal Disease Effect From Vaccine Serotypes
PPSV23	June 1983	1, 2,* 3, 4, 5, 6B, 7F, 8,* 9N,* 9V, 10A,* 11A,* 12F,* 14, 15B,* 17F,* 18C, 19A, 19F, 20,* 22F,* 23F, and 33F*	<ul> <li>Reduced invasive disease</li> <li>No effect on carriage</li> </ul>
PCV7	February 2000	4, 6B, 9V, 14, 18C, 19F, and 23F	<ul> <li>Reduced invasive disease</li> <li>Reduced carriage</li> <li>Protective herd effect</li> <li>Increase in 19A infections</li> </ul>
PCV13	February 2010	1, 3, 4, 5, 6A,† 6B, 7F, 9V, 14, 18C, 19A, 19F, and 23F	<ul> <li>Reduced invasive disease</li> <li>Reduced carriage</li> <li>Increase in 35B infections</li> </ul>

PCV7, 7-valent pneumococcal conjugate vaccine; PCV13, 13-valent pneumococcal conjugate vaccine; PPSV23, 23-valent pneumococcal polysaccharide vaccine \*Serotypes in PPSV23 are unique to this vaccine. †Serotype in PCV13 is unique to this vaccine.



# **PCV13 Vaccine Impact**

- Reduction in incidence of invasive pneumococcal disease
  - -65% reduction in children <5 y.o.
  - Decrease in invasive disease in nonimmunized adults
  - Possible reduction in otitis media



# **Pneumococcal Vaccine Future**

## Concerns

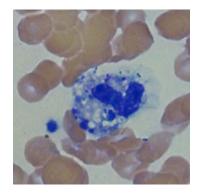
- Continued serotype replacement
- Continued increase in resistance in replacement strains
- Large diverse pool of capsular polysaccharide serotypes

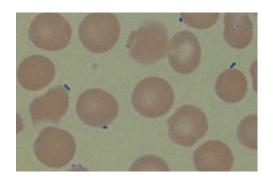
## • Future

 Search for more conserved protein-based antigen



• Intra and Extracellular bacteria present





- Received three doses of PCV13 vaccine
  (1, 3, 4, 5, 6A, 6B, 7F, 9V, 14, 18C, 19A, 19F, 23F)
- Clinical isolate serotyped as 33F/33A/37









