FEVER: SYNOPSIS

OBJECTIVES

- To determine which patients are at high risk of developing sepsis.
- To assess patient with fever.
- To initiate empiric therapy.

WHICH PATIENTS ARE HIGH-RISK FOR SEPSIS

- Neonates
- Transplant recipients
 - □ Bone marrow
 - Solid organ
- Oncology patients
 - □ Undergoing therapy, mucositis, central line
 - □ Most chemotherapy: nadir ~ 10 days after rx
- Asplenic patients, including sickle cell

DEFINITION OF FEVER

- **38.0**
 - □ Neonates (< 12 months)
 - □ Any immunocompromised patient
 - Including transplant patients, patients with immunodeficiencies, oncology patients (sustained ≥38 x 1 hour)
- **38.5**
 - □ All other patients
- These are general guidelines, individual patients/services may have different parameters

WHAT ETIOLOGIES CAUSE FEVER?

- Infectious
- Inflammatory
- Oncologic
- Other: CNS dysfunction, drug fever
- Life-threatening conditions

INFECTIOUS

- Systemic
 - □ Bacteremia, sepsis, meningitis, endocarditis
- Respiratory
 - URI, sinusitis, otitis media, pharyngitis, pneumonia, bronchiolitis
- Abdominal
 - □ Urinary tract infection, abscess (liver, kidney, pelvis)
- Bone/joint infection
- Hardware infection
 - □ Central line, VP shunt, G-tube

INFLAMMATORY

- Kawasaki disease
- Juvenile inflammatory arthritis
- Lupus
- Inflammatory bowel disease
- Henoch-Schonlein purpura

ONCOLOGIC

- Leukemia
- Lymphoma
- Neuroblastoma
- Sarcoma

OTHERS

- CNS dysfunction
- Drug fever

LIFE-THREATENING CONDITIONS

- Sepsis, febrile neutropenia
 - □ Vital sign instability, poor-perfusion, may have altered mental status, disseminated intravascular coagulation
- Hemophagocytic lymphohistiocytosis
 - □ Splenomegaly, bicytopenia, elevated ferritin, elevated triglycerides, low fibrinogen, hemophagocytosis, low/absent NK cell function, elevated soluble IL2 receptor
- Malignant hyperthermia
 - ☐ Following administration of inhaled anesthetics or depolarizing neuromuscular blockers (succinylcholine), at-risk patients include those with myopathy
 - □ Muscle rigidity, rhabdomyolysis, acidosis, tachycardia

ASSESSMENT

- Vital signs
- Repeat physical exam
 - □ Overall appearance (sick, toxic)
 - □ Central/peripheral lines
 - □ Incisions/wounds
 - □ VP shunt/tracheostomy/gastrostomy tube
 - □ Oral mucosa/perineal area for neutropenic patients
 - Perfusion
- Call for help if concerning vital signs/exam
 - □ Fellow or attending
 - □ Rapid response team (RRT)/PICU

LABORATORY EVALUATION

- What would you do if the patient has hardware (VP shunt, tracheostomy, gastrostomy tube) or central line?
 - CBC with differential
 - □ Blood culture
 - □ CSF (tap VP shunt)

- What would you do if the patient has a high risk for sepsis?
 - □ Immunocompromised
 - □ Transplant recipient
 - □ Oncology patient
 - □ CBC with differential
 - □ Blood culture
 - □ Urinalysis and urine culture
- What would you do for an infant < 2 months of age?</p>
 - □ CBC with differential
 - □ Blood culture
 - □ Catheterized urinalysis and urine culture
 - □ Lumbar puncture
- Who needs a urinalysis and urine culture?
 - □ Circumcised males < 6 months
 - □ Uncircumcised males < 1 year
 - □ Females < 2 years
 - □ Immunocompromised patients
 - □ Patients with history of UTI/pyelonephritis
- Who needs a lumbar puncture?
 - \Box Neonates ≤ 2 months
 - □ III-appearing
 - □ Altered mental status
- What tests do you send?
 - □ Gram stain and culture
 - □ Cell count and differential
 - □ Protein and glucose
 - □ Extra tube for additional studies
 - Enteroviral PCR, HSV PCR, CA encephalitis project
- Consider CRP, ESR
- Consider PT/PTT, fibrinogen
- Consider chest x-ray
- Consider nasopharyngeal DFA
- For immunosuppressed patients consider:
 - □ Viral PCR studies (ie CMV, EBV, HHV6)
 - □ Additional imaging (ie ultrasound, CT scan)

TREATMENT FOR NON-HIGH RISK PATIENTS

- May not need empiric antibiotics
- Consider the following issues:
 - □ Is patient clinically stable?
 - □ Are the screening laboratory studies suggestive of infection?

TREATMENT FOR PATIENTS WITH CENTRAL LINES

- Ceftriaxone
- Vancomycin

TREATMENT FOR NEONATES ≤ 2 MONTHS

- If < 28 days old</p>
 - □ Ampicillin AND cefotaxime OR
 - Ampicillin **AND** gentamicin
 - Consider acyclovir
- If 29-60 days old
 - □ Ceftriaxone ± Ampicillin **OR** Vancomycin
 - □ Until CSF results are known (cell count, protein, glucose), initiate therapy with meningitic dosing regimen

TREATMENT FOR FEBRILE NEUTROPENIA

- Broad-spectrum antibiotics with *Pseudomonas coverage*
 - □ Ex: use ceftazidime or piperacillin-tazobactam
- Consider double coverage for possible resistant *Pseudomonas*
 Ex: add amikacin or tobramycin
- Consider gram-positive coverage (central line, skin infections)

 Ex: add vancomycin
- Consider anaerobic coverage (mucositis, typhlitis)
 Ex: use piperacillin-tazobactam or add clindamycin

TAKE-HOME POINTS

- Infections are the most common cause of fever in children
- During assessment of a child with fever, pay close attention to vital sign changes, overall appearance, and potential sites of infection
- Closely monitor for clinical decompensation after antibiotic administration, particularly in patients at high-risk of developing sepsis

REFERENCES

Baraff LJ. Management of fever without source in infants and children. *Ann Emerg Med.* 2000. 36:602-14.

Meckler G, Lindemulder S. Fever and neutropenia in pediatric patients with cancer. *Emerg Med Clin N Am.* 2009. 27:525-44.

Palazzi EL. Approach to the child with fever of unknown origin. UpToDate. 2011

Palazzi DL. Etiologies of fever of unknown origin. UpToDate. 2011.

Tolan R. Fever of unknown origin: A diagnostic approach to this vexing problem. *Clin Pediatr*. 2010;49:207-13.