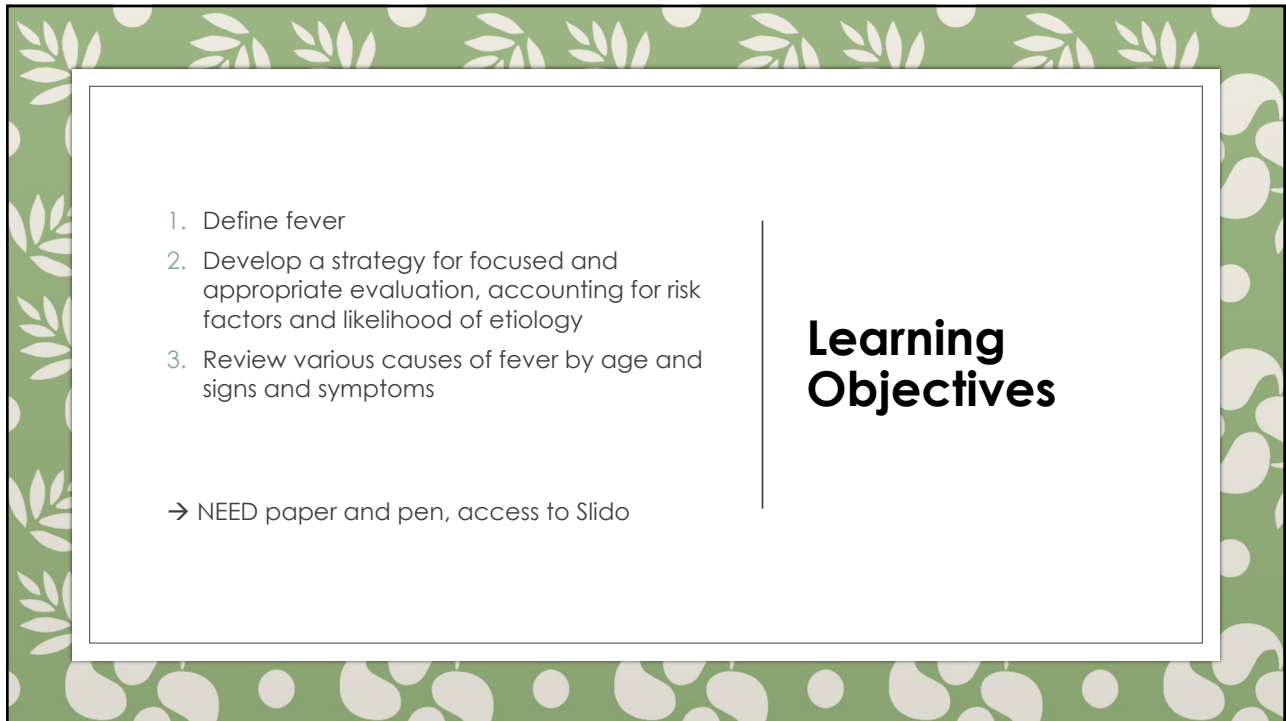


1

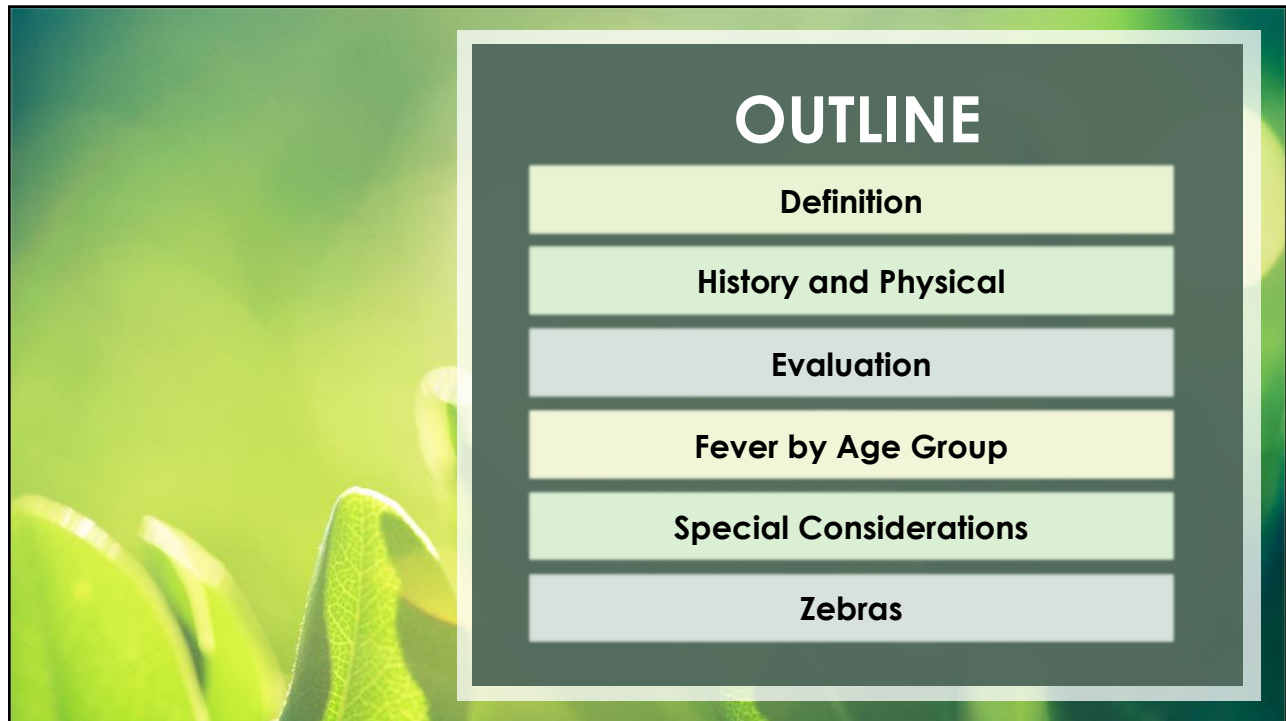


1. Define fever
2. Develop a strategy for focused and appropriate evaluation, accounting for risk factors and likelihood of etiology
3. Review various causes of fever by age and signs and symptoms

→ NEED paper and pen, access to Slido

Learning Objectives

4

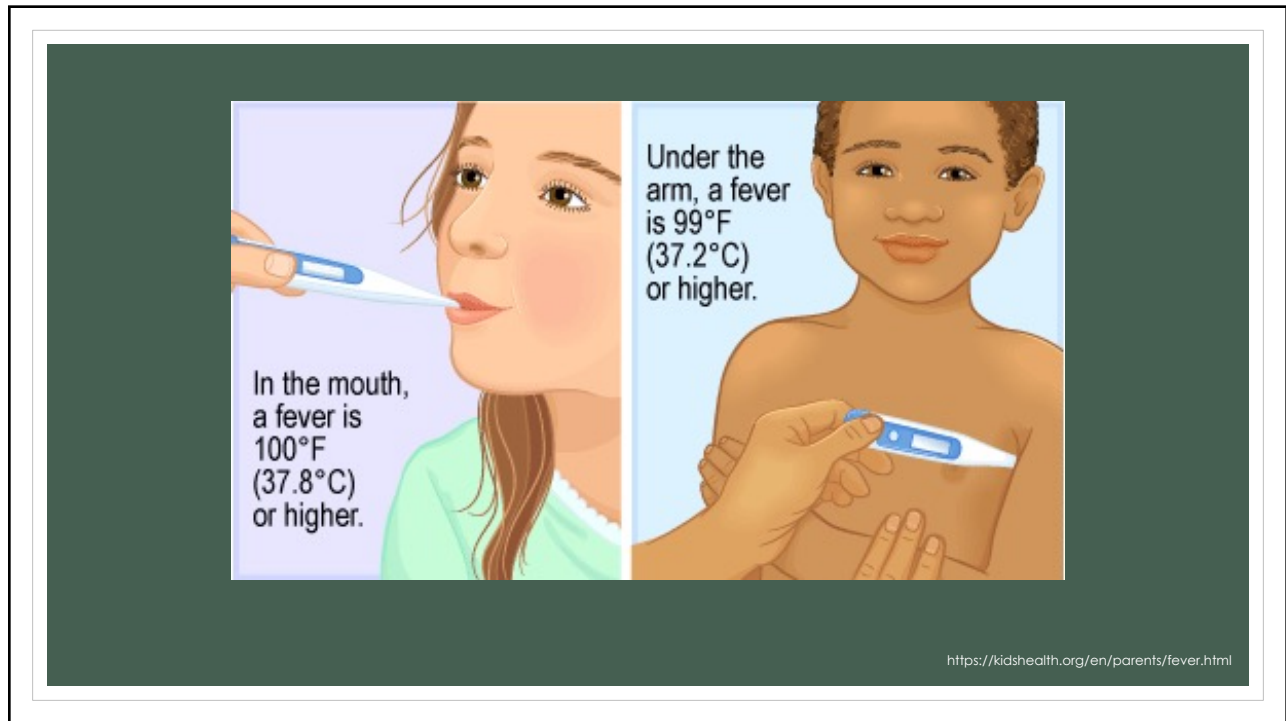


5

What is a fever?

- **It depends!**
- AAP: ≥ 38.0 rectal, > 37.2 oral
- SickKids/About Kids Health: ≥ 38.0
- Merck Manual: > 38.2 rectal, > 37.8 oral
- Harrison's Principles of IM: > 37.2 oral in the AM and > 37.7 in the PM
- CDC: ≥ 38.0 or feels warm to touch or feels feverish

8



9

How to best measure temperature?

- **It is confusing for parents!**
- SickKids/About Kids Health: Rectal if < 3 years, oral if ≥ 3 years
Axillary and TM less reliable
No forehead or pacifier thermometer
- AAP: Rectal if < 4 years, oral if ≥ 4 years
Temporal artery and TM okay
Axillary not accurate

10

- Fever duration, height, pattern, prodrome
- Chronology
- Associated symptoms
- Events, injuries
- **ROS:** respiratory, GI, pain
- **PMHx:** co-morbidities, recent surgery, immunizations
- **FamHx:** rheumatologic
- **SocHx:** exposures, animals, travel, bites, pica

History

11

- **Vital signs:** relative bradycardia
- **Skin:** rash, petechiae, ECM, erythema nodosum, malar erythema, salmon pink rash, diaphoresis or lack thereof
- **HEEN:** conjunctivitis (palpebral, bulbar, limbic-sparing), hemorrhage, sinus tenderness/drainage
- **OP:** erythema, ulcers, dentition, gingival changes
- **Lymph Nodes:** local, diffuse, tender
- **Chest/♥:** crackles, wheeze, new murmur
- **Abd:** tenderness, HSM
- **MSK:** tender bones, joints, muscles
- **GU:** perianal abscess/tag, ulcers, discharge, tenderness

Physical Exam

12

- **CBC with differential**
 - WBC ↑ . . . neutrophils, lymphocytes, eosinophils
 - WBC ↓
 - WBC morphology . . . atypical, toxic granulation
 - Hemoglobin ↓
 - Platelets ↑
 - Platelets ↓
 - Pancytopenia
- ESR/CRP
- Electrolytes/transaminases (Na, AST/ALT)
- Blood culture
- UA/urine culture
- CXR
- TB skin test

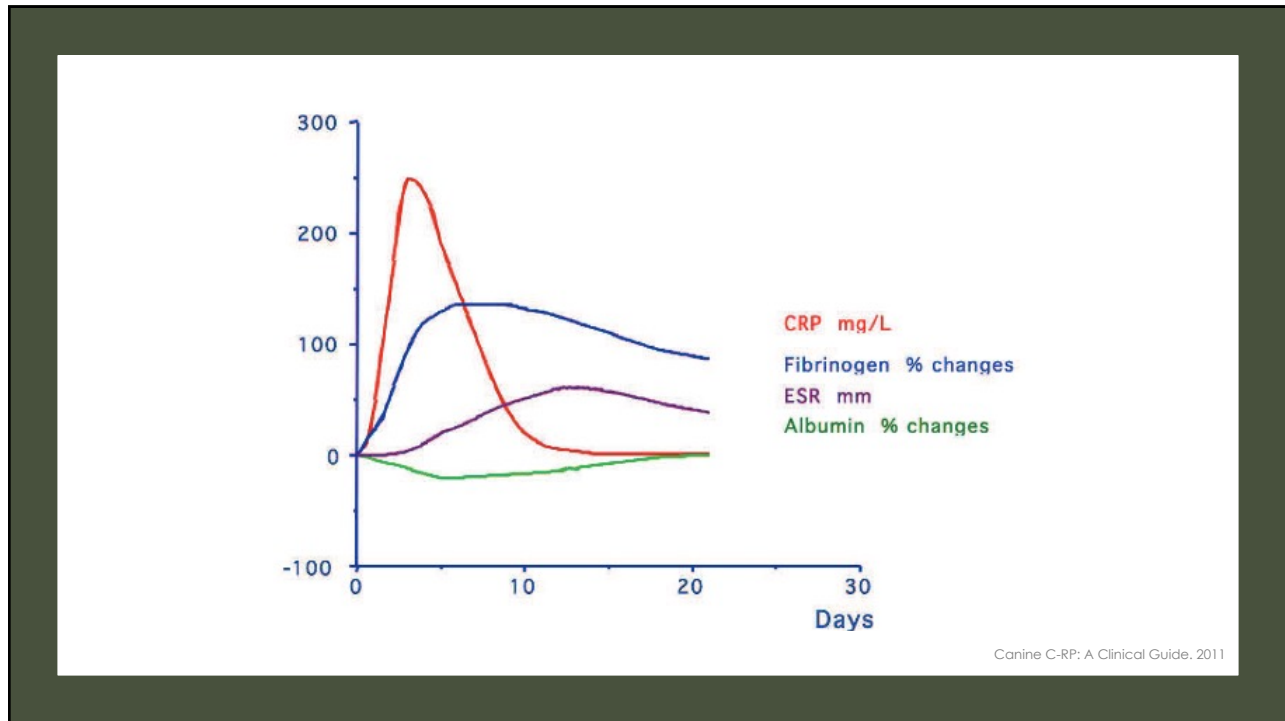
Evaluation

13

CRP vs. ESR

CRP	ESR
<ul style="list-style-type: none"> • Direct measurement of protein • No variations based on age, gender, temperature, other conditions • More sensitive to clinical changes • Rapid rise and fall 	<ul style="list-style-type: none"> • Indirect test of RBC behaviour • Varies with age, pregnancy, anemia, plasma proteins, certain drugs • Slower response to changes • Slower rise and fall

14



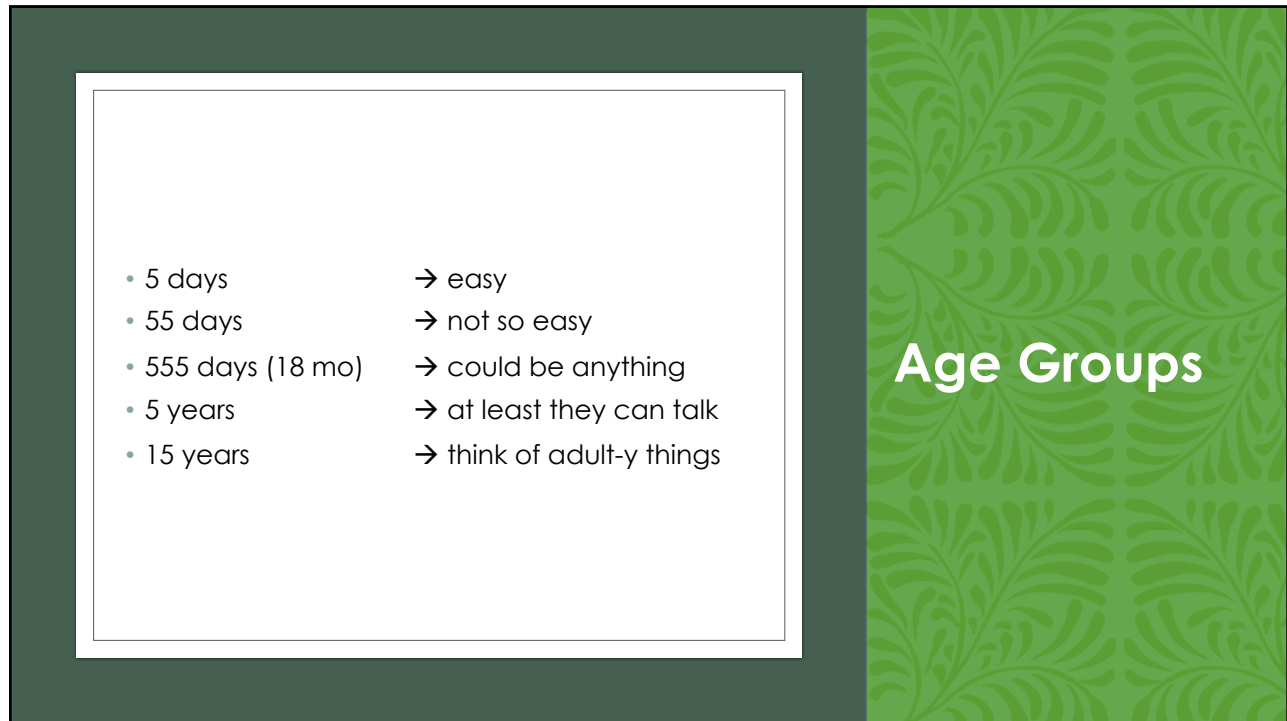
15

CRP vs. ESR

CRP	ESR
<ul style="list-style-type: none"> • Direct measurement of protein • No variations based on age, gender, temperature, other conditions • More sensitive to clinical changes • Rapid rise and fall 	<ul style="list-style-type: none"> • Indirect test of RBC behaviour • Varies with age, pregnancy, anemia, plasma proteins, certain drugs • Slower response to changes • Slower rise and fall

Procalcitonin: helpful in bacterial infection, sepsis

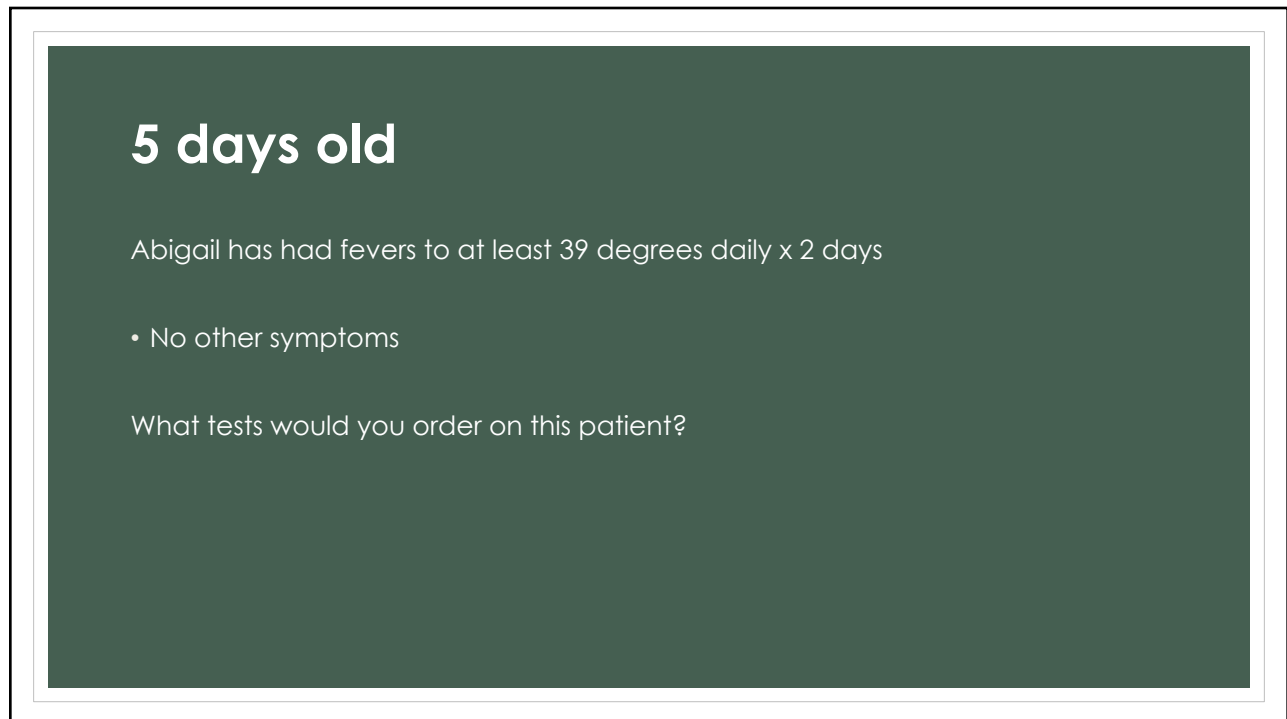
16



• 5 days	→ easy
• 55 days	→ not so easy
• 555 days (18 mo)	→ could be anything
• 5 years	→ at least they can talk
• 15 years	→ think of adult-y things

Age Groups

17



5 days old

Abigail has had fevers to at least 39 degrees daily x 2 days

- No other symptoms

What tests would you order on this patient?

18

5 days old

Easy!

- CBC/Blood culture
- UA/Urine culture
- LP
- Anything else?
 - CXR, viral swabs, HSV

→ Empiric antibiotics and admit

19

55 days old

Brayden has had fevers to at least 39 degrees daily x 2 days

- No other symptoms

→ Not so easy . . .

20

CLINICAL PRACTICE GUIDELINE

American Academy of Pediatrics



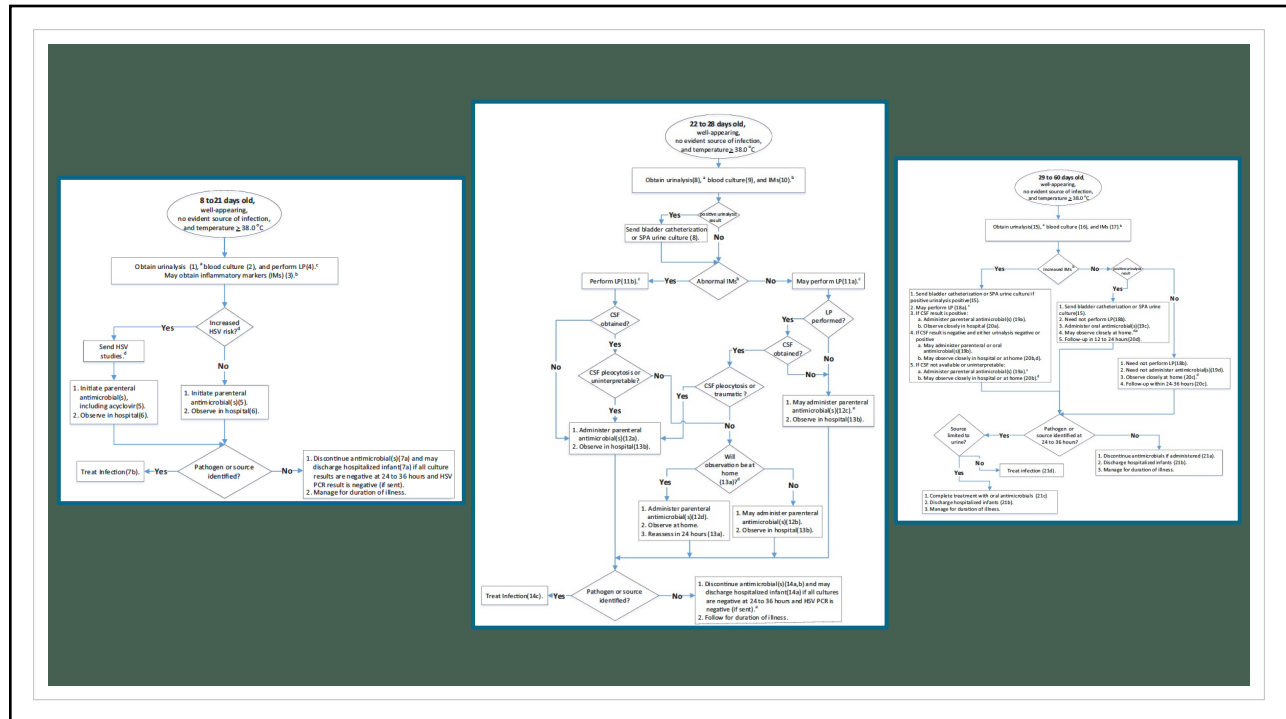
DEDICATED TO THE HEALTH OF ALL CHILDREN™

Evaluation and Management of Well-Appearing Febrile Infants 8 to 60 Days Old

Robert H. Pantell, MD, FAAP,^a Kenneth B. Roberts, MD, FAAP,^b William G. Adams, MD, FAAP,^c Benard P. Dreyer, MD, FAAP,^d Nathan Kuppermann, MD, MPH, FAAP, FACEP,^e Sean T. O'Leary, MD, MPH, FAAP,^f Kyrnika Okechukwu, MPA,^g Charles R. Woods Jr, MD, MS, FAAP^h SUBCOMMITTEE ON FEBRILE INFANTS

PEDIATRICS Volume 148, number 2, August 2021

21



22

Chronological Age

8-21 days

- Blood culture
 - Urinalysis +/- Ucx
 - LP
 - (HSV risk/test/ACV)
 - (IMs: ANC, CRP, ProC)
- Abx, admit
→ Discharge 24-36 hours

22-28 days

- Blood culture
 - Urinalysis +/- Ucx
 - IMs: ANC, CRP, ProC
 - If IMs abN, LP
 - (HSV risk/test/ACV)
- Abx, admit
→ No abx, admit
→ Abx, home, f/u 24 hours

29-60 days

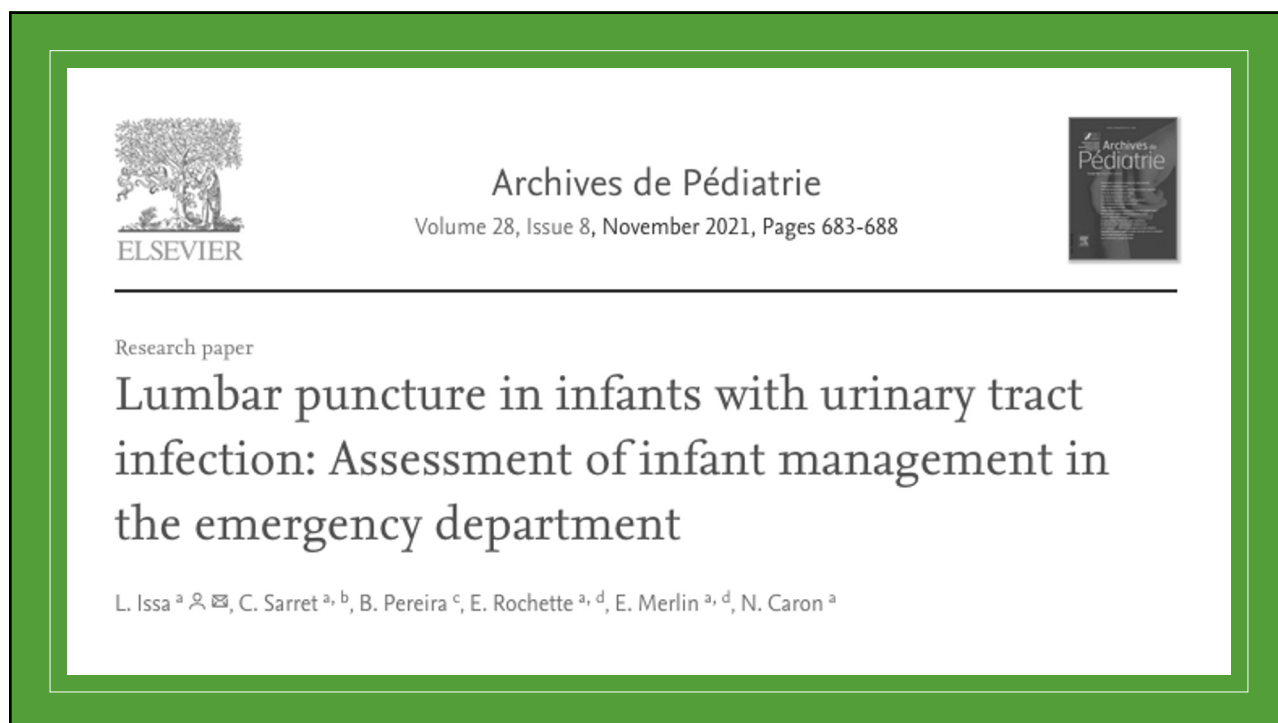
- Blood culture
 - Urinalysis +/- Ucx
 - IMs: ANC, CRP, Procalcitonin
 - If IMs abN, consider LP
 - LP+ : abx, admit
 - LP- : parenteral/oral abx, admit or home
 - LP? : parenteral abx, admit or home
- UA+: home, PO abx, f/u 12-24 h
→ All neg : home, no abx, f/u 24-36 h

23

TABLE 3 Initial Empirical Antibacterial Therapy for Well-Appearing Febrile Infants 7 to 60 Days Old

Suspected Source of Infection	8–21 d Old	22–28 d Old	29–60 d Old
UTI ^a	Ampicillin IV or IM (150 mg/kg per d divided every 8 h) and either ceftazidime IV or IM (150 mg/kg per d divided every 8 h) or gentamicin IV or IM (4 mg/kg per dose every 24 h)	Ceftriaxone IV or IM (50 mg/kg per dose every 24 h)	Ceftriaxone IV or IM (50 mg/kg/dose every 24 h). Oral medications for infants older than 28 d. ^b Cephalexin 50–100 mg/kg per d in 4 doses or cefixime 8 mg/kg per d in 1 dose
No focus identified ^c	Ampicillin IV or IM (150 mg/kg per d divided every 8 h) and either ceftazidime IV or IM (150 mg/kg per d divided every 8 h) or gentamicin IV or IM (4 mg/kg per dose every 24 h) ^d	Ceftriaxone IV or IM (50 mg/kg per dose every 24 h)	Ceftriaxone IV or IM (50 mg/kg/dose every 24 h)
Bacterial meningitis ^e	Ampicillin IV or IM (300 mg/kg per d divided every 6 h) and ceftazidime IV or IM (150 mg/kg per d divided every 8 h)	Ampicillin IV or IM (300 mg/kg per d divided every 6 h) and ceftazidime IV or IM (150 mg/kg per d divided every 8 h)	Ceftriaxone IV (100 mg/kg or d once daily or divided every 12 h) or Cefazidime IV (150 mg/kg or d divided every 6 h) and vancomycin ^f IV (60 mg/kg or d divided every 8 h)

26



27

55 days old

WHAT IF FEVER AND . . .

- Exam shows bulging fontanel
- Exam shows hepatosplenomegaly
- Exam shows swelling and tenderness to palpation over multiple extremities

28

55 days old

WHAT IF FEVER AND . . .

- Exam shows bulging fontanel → meningitis, roseola
- Exam shows hepatosplenomegaly → EBV, CMV, HLH, malignancy
- Exam shows swelling and tenderness to palpation over multiple extremities → infantile cortical hyperostosis (Caffey disease)

29

555 days old = 18 months

Charlotte has had fevers to at least 39 degrees daily x 2 days

- Evaluation dependent on other symptoms/findings
- Consider screening investigations (e.g., UA/urine culture in girls)

WHAT IF:

Charlotte has had fevers to at least 39 degrees daily x 5 days

- Most would consider screening investigations
- Some viral illnesses consistently have fever 5+ days

30

555 days old = 18 months

- Pallor
- Fatigued
- Fussy
- Doesn't want to walk
- Gums sensitive when brushing teeth

→ Acute lymphocytic leukemia

31

555 days old = 18 months

WHAT IF . . .

- Known diagnosis of ALL and recent chemotherapy
- Recent return from month in India
- Exam showed warm and tender knee joint
- Exam showed conjunctivitis and red cracked lips
- Exam showed abnormal appearance of teeth and absence of sweating

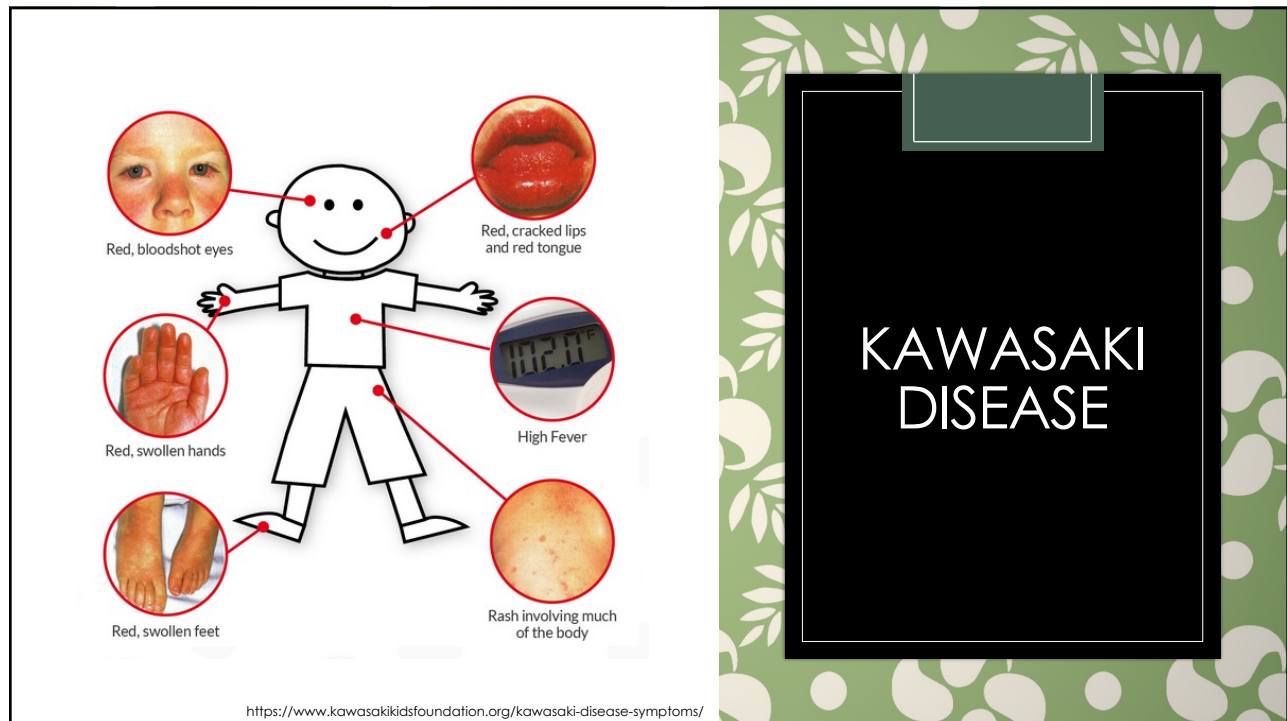
32

555 days old = 18 months

WHAT IF FEVER AND . . .

- Known diagnosis of ALL and recent chemotherapy → febrile neutropenia
- Recent return from month in India → malaria, dengue, leptosporosis
- Exam showed warm and tender knee joint → septic arthritis, osteo, JIA
- Exam showed conjunctivitis and red cracked lips → KD, SJS, TSS, MIS-C, adeno
- Exam showed abnormal appearance of teeth and absence of sweating → ectodermal dysplasia

33



34

CRASH AND BURN

Conjunctivitis (non-purulent, limbic-sparing)

Rash (polymorphous, maculopapular, diffuse)

Adenopathy (cervical lymph node >1.5 cm)

Strawberry tongue/mucous membrane changes

Hand and feet changes (edema, erythema)

Burn (fever at least 38.5 for at least five days)

KAWASAKI DISEASE

35

KAWASAKI DISEASE

```

graph TD
    Start[Child with a fever for ≥5 days without an alternative explanation] --> Q1[How many clinical diagnostic criteria are met?]
    Q1 -- "0 or 1 clinical criteria" --> Q2[Is the child ≥6 months old with a fever ≥7 days?]
    Q1 -- "2 or 3 clinical criteria and a strong clinical suspicion of KD" --> Q3[Assess CRP/ESR]
    Q1 -- "≥4 clinical criteria" --> Q4[Consistent with KD]
    
    Q2 -- Yes --> Q5[Possible KD]
    Q2 -- No --> Q6[KD unlikely]
    
    Q3 -- "CRP <3.0 mg/dL and/or ESR <40 mm/hour" --> Q6
    Q3 -- "CRP ≥3.0 mg/dL and/or ESR ≥40 mm/hour" --> Q7[Assess for supplemental laboratory criteria and obtain echocardiogram]
    
    Q7 -- "≥3 supplemental laboratory criteria OR positive echocardiogram" --> Q8[Incomplete KD]
    Q7 -- "≤2 supplemental laboratory criteria and negative echocardiogram" --> Q6
    
    Q4 --> Q9[Start treatment and obtain baseline echocardiogram]
    
    Q5 --> Q10[Obtain baseline echocardiogram]
    Q10 -- Positive --> Q11[Incomplete KD]
    Q10 -- Negative --> Q12[Suspected incomplete KD]
    
    Q12 --> Q13[Assess for supplemental laboratory criteria in addition to CRP/ESR and monitor closely including serial echocardiograms until KD is diagnosed and treatment started or until patient is afebrile and otherwise well for 48 hours]
    
    Q6 --> Q14[Perform serial clinical and laboratory re-evaluation if fever persists and obtain echocardiogram if typical percutaneous discoloration develops even if fever has resolved]
    
    Q8 --> Q15[Start treatment]
    
    Q9 --> Q15
    
```

Adapted from: McCrindle BW, Rowley AH, Newburger JW, et al. Diagnosis, treatment, and long-term management of Kawasaki disease: A scientific statement for health professionals from the American Heart Association. *Circulation* 2017; 135:e927.

36

5 years old

David has had fevers to at least 39 degrees daily x 5 days

- Bilateral leg pain
- Refuses to walk
- Exam: swollen knees and ankles with limited ROM secondary to pain
- Exam: diffuse macular rash
- WDWTK?

37



38

- Fever, arthralgia/arthritis, lymphadenopathy, rash, edema of hands/feet
- Type 3 hypersensitivity reaction to drug
 - Primary: 6-21 days after exposure
 - Recurrent: 1-4 days after exposure to same antigen
- Common triggers include: amox, cefaclor, septr, penicillin
- Clinical diagnosis
- Self-limited
- NSAIDs, antihistamines, steroids
- Labs if ill-appearing

SSLR

39

5 years old

WHAT IF FEVER AND . . .

- Joint pain is migratory, affecting one joint at a time
- Bone pain
- Recurrent fevers every few weeks associated with mouth ulcers
- Myalgias to calf muscles, toe-walking, tender to gastrocnemius squeeze
- Recent COVID infection one month ago
- Night sweats, arthralgias, recent travel to South America
- Cervical lymphadenopathy, splenomegaly

40

5 years old

WHAT IF FEVER AND . . .

- Joint pain is migratory, affecting one joint at a time → ARF, JIA
- Bone pain → leukemia, osteomyelitis, brucellosis
- Recurrent fevers every few weeks associated with mouth ulcers → PFAPA
- Myalgias to calf muscles, toe-walking, tender to gastrocnemius squeeze → influenza
- Recent COVID infection one month ago → MIS-C
- Night sweats, arthralgias, recent travel to South America → brucellosis, malaria
- Cervical lymphadenopathy, splenomegaly → EBV, Kikuchi-Fujimoto disease

41

15 years old

Emily has had fevers to at least 39 degrees daily **x 5 days**

- Intermittent history of RUQ pain
- Worsening pain in the last week
- Looks unwell on exam
- Abdomen tense, tender RUQ and lateral to umbilicus with guarding
- Murphy's sign positive

42

15 years old

WHAT IF FEVER AND . . .

- History of recent weight loss and hematochezia
- Recent immigrant from African country
- Recent appendectomy and abdominal pain
- Headaches and hypernatremia
- Jaundice and recent travel to Mexico
- Petechiae
- Dysphagia, cervical LAD, abdominal pain and tenderness BUQ
- Abdominal pain and tenderness lower abdomen, vaginal discharge

43

15 years old

WHAT IF FEVER AND . . .

- History of recent weight loss and hematochezia → IBD
- Recent immigrant from African country → Sickle Cell Disease
- Recent appendectomy and abdominal pain → intra-abdominal pyogenic abscess
- Headaches and hypernatremia → pituitary tumor
- Jaundice and recent travel to Mexico → Hepatitis A
- Petechiae → meningococcus
- Dysphagia, cervical LAD, abdominal pain and tenderness BUQ → mono
- Abdominal pain and tenderness lower abdomen, vaginal discharge → PID

44

- Recent chemotherapy
- Sickle cell disease
- Immigrant/returning traveler
- Periodic/recurrent fever
- Prolonged fever

Special Considerations

45

Special Considerations

- **Chemotherapy**
 - Febrile neutropenia → increased risk for opportunistic infections
 - Precautions
 - Management
 - Blood cultures line + peripheral, broad-spectrum antibiotics
 - Admit
- **Sickle cell disease**
 - Functionally asplenic → increased risk for encapsulated, invasive infections
 - Management
 - Urgent antibiotics: ceftriaxone or levofloxacin +/- vancomycin
 - CBC, blood culture, swab for flu/COVID

46

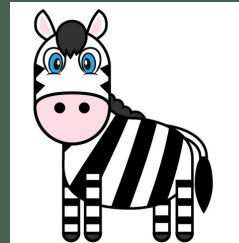
Special Considerations

- **Recent travel**
 - Malaria
 - Dengue fever
 - Salmonella/typhoid
 - Tuberculosis
 - Hepatitis A
- **Periodic fever**
 - PFAPA
 - Muckle-Wells syndrome

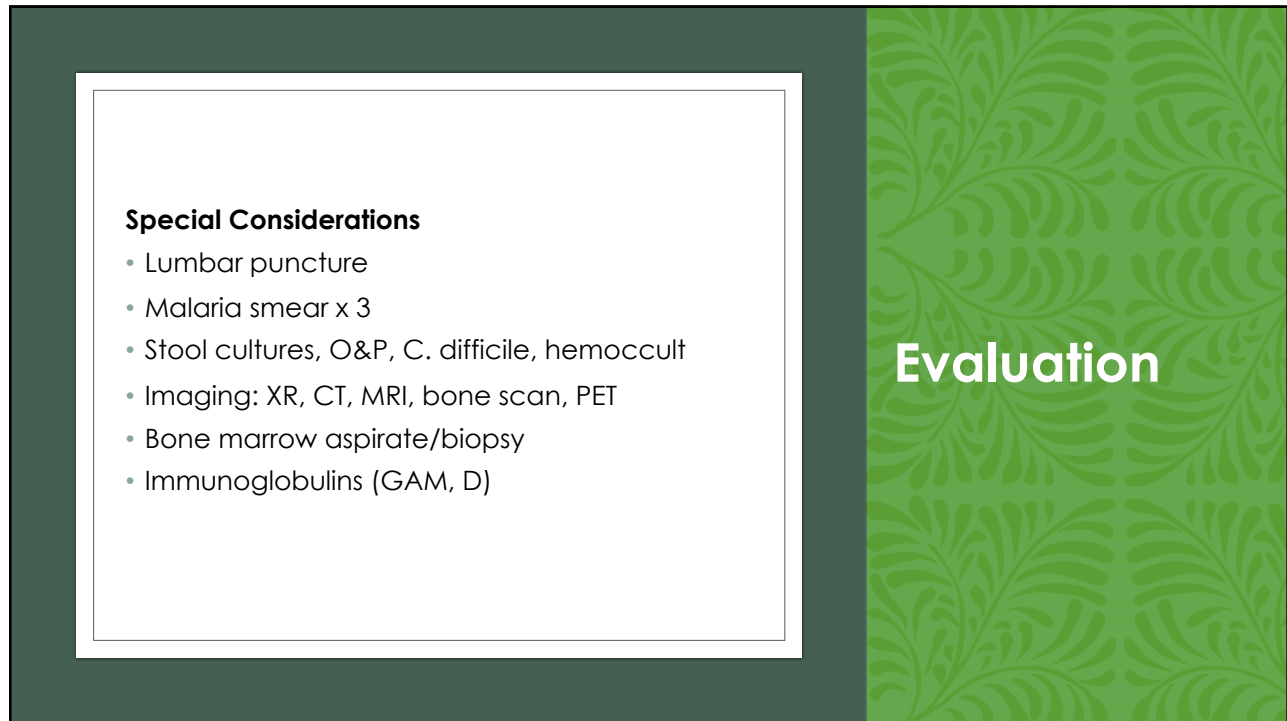
47

Special Considerations

- **Prolonged fever (> 1 week)**
 - Infections 50% - prolonged or sequential
 - Connective tissue disorder
 - Malignancy
 - Kawasaki Disease
 - Zebras
 - Drug fever
 - Central/CNS dysregulation
 - Factitious
 - HLH
 - IBD
 - And more - hyperthyroid, ectodermal dysplasia, immunodeficiencies, familial dysautonomia, Kikuchi-Fujimoto disease



48



Evaluation

Special Considerations

- Lumbar puncture
- Malaria smear x 3
- Stool cultures, O&P, C. difficile, hemocult
- Imaging: XR, CT, MRI, bone scan, PET
- Bone marrow aspirate/biopsy
- Immunoglobulins (GAM, D)

49



SUMMARY

- Fever can be a sign of a large variety of conditions
- Tailor evaluation to appearance, signs, symptoms, and risk factors (including age)
- Special circumstances require increased clinical vigilance
- Phone a friend when you need one

50

