

## **وبینار چالش های تشخیصی و درمانی کووید در کودکان**

دانشگاه علوم پزشکی اراک- اسفندماه ۱40۰



# MIS-c Management

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## • تعریف: درگیری التهابی سیستم های مختلف بدن (بیش از دو سیستم) در کودکان ناشی از بیماری کووید

- بدنبال کووید علامت دار و یا اثبات شده
  - بدنبال كوويد بدون علامت
    - گرفتاری اطرافیان
- کشف اتفاقی کووید در کودک بدنبال ازمایش PCR
- پیدا کردن ردپای کووید رد کودک مبتلا به MIS-c در سرولوژی



## Criteria

### **CDC**

All 4 criteria must be met

بحمن روماتولوژی کودکان ایران

- 1. Age < 21 yrs
- Clinical presentation consisting of MISC including all:
  - Fever
  - Lab evidence of inflammation
  - Multisystem involvement (2 or more organs)
  - Severe illness requiring hospitalization
- 3. No alternate diagnosis
- 4. Recent or current SARS-COV2 infection or exposure
  - PCR (+)
  - Serology (+)
  - Antigen test (+)
  - COVID19 exposure within 4 weeks prior to onset of symptoms



### **WHO**

- 1. Age < 19 yrs
- 2. Fever > 3 days
- 3. Clinical signs of multisystem involvement (at least 2 or more)
- 4. Elevated inflammatory markers
- 5. No alternate explanation
- 6. Evidence of SARS-COV2 infection
  - PCR (+)
  - Serology (+)
  - Antigen test (+)
  - Contact with an individual with COVID19



## Organ Inolvments



- Cardiac involvement
- Liver Dysfunction
- Brain
- Pancreas
- Pulmonary
- Renal involvement

- Skin
- Eye
- Mucosal
- GI
- Joint
- Hematologic





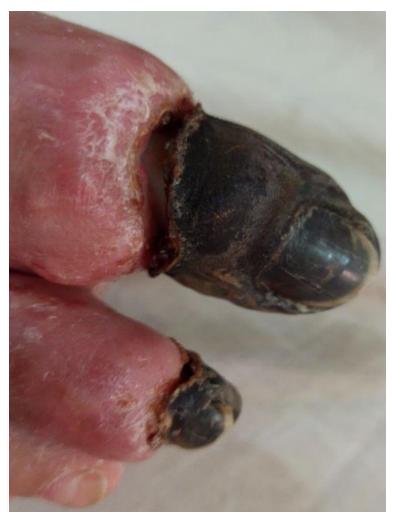








## Skin Manfestation









## Classification

- Based on severity:
  - Mild
  - Moderate to severe
- Based on Age:
  - In neonate: MIS-n
  - In Children: MIS-c
  - In Adult: MIS-a





## Management

- Mild case can be followed outpatient.
- ☐ Most children with MIS-C need to be treated in a hospital.
- □ Some need **treatment** in a pediatric intensive care unit.
- □ Treatment usually involves supportive care and measures to reduce inflammation in any affected vital organs to protect them from permanent damage.





### **Tehran Children's Medical Center Protocol**

Iran J Pediatr. 2020 October; 30(5):e108617.

doi: 10.5812/ijp.108617.

Published online 2020 September 25.

**Methods Article** 



## An Algorithmic Approach to Multisystem Inflammatory Syndrome in Children with COVID-19: Tehran Children's Medical Center Protocol

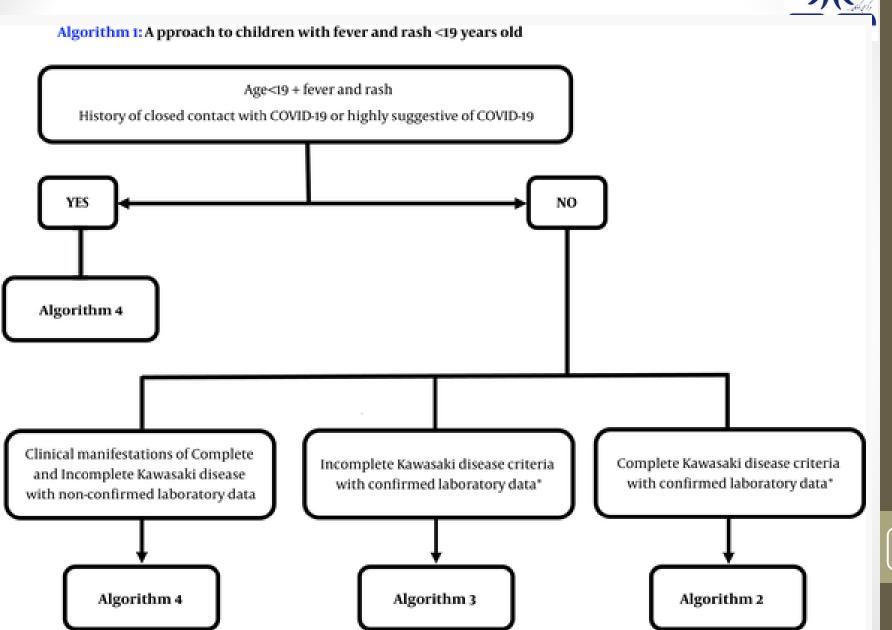
Vahid Ziaee (1,2,3,\*, Raheleh Assari (1,2), Setarh Mamishi (1,2), Aliakbar Zeinaloo (1,2,6), Masoud Mohammadpour (1,2,7), Iran Malekzadeh (2,8) and MIS-C COVID-19 Collaborators Groups (1,3,4,6,7,8)



Iranian Journal of Pediatrics:
Published Online: September 25,
2020

Article Type: Methods Article Accepted: August 22, 2020





Laboratory data

Response to

treatment

(fever resolved)

### Fever more than 4 days and 4 of five criteria:

- · Polymorphous skin rashes
- Bilateral non-purulent conjunctival injection
- Mucosal changes: cracked lips, strawberry tongue, mucosal erythema
- Extremity changes: hands and feet edema or palmar erythema
- Unilateracervical lymphadenopathy>1.5 cm

#### Confirmed laboratory data:

- ESR>40
- CRP>30
- WBC>15000
- Platelet in upper limit or thrombocytosis
- ALT elevation
- Normal ferritin or <450</li>

No other organ involvements explained in algorithm 4

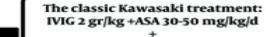
#### Treatment with ASA 3-5mg/kg/d continued + Echocardiography/after 10-14

Echocardiography (after 10-14 day and 2 months later)

Abnormal Echocardiography

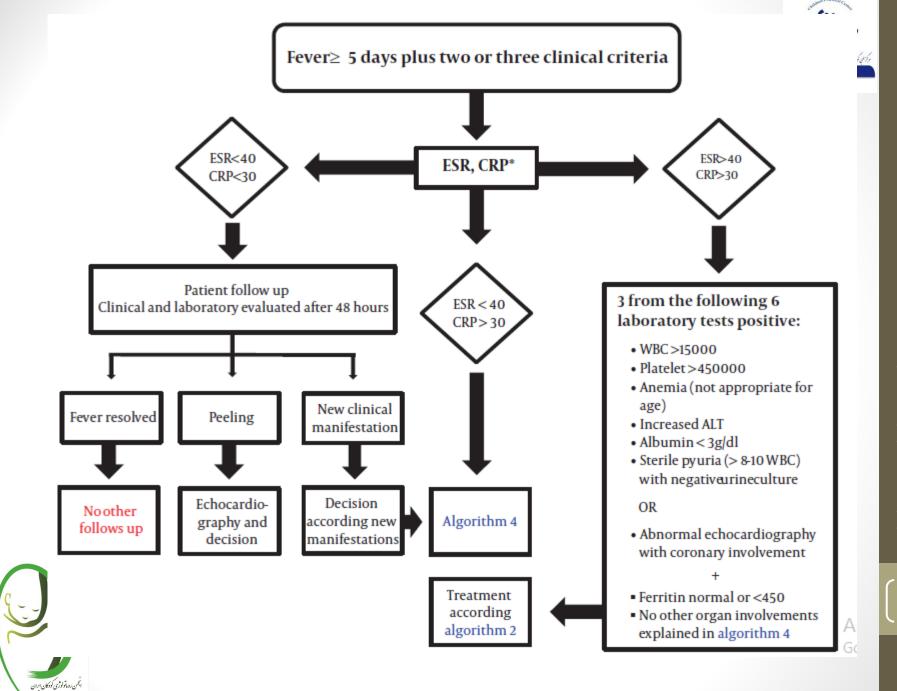
Referred to Pediatric Rheumatologist

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Echocardiography

No response to treatment (Fever not resolved)





### BOX 1: Para-clinic evaluation for all suspicious patients:

- CBC, ESR, CRP, B/C, U/A, U/C
- Liver function test, PT, PTT, INR, serum albumin and serum Ferritin
- D-Dimer, BNP, IL6, Procalcitonin (if available)
- Pancreatic and Cardiac Enzymes (in selective patients)
- Abdominal ultrasonography
- Echocardiography
- COVID-19 confirmation\*:
  - o SARS-COV2 RT-PCR

OR

- o Serology
- Diagnostic complementary evaluations (for patients suspicious to other infectious disease)
   based on pediatric infectious disease consultation
- \* If negative, recheck (both PCR and serology) after 7-10 days



and

Closed contact with COVID-19 or high suspicio us to COVID-19

#### Box 2

Unusual clinical manifestations in KD:

- Shock or low blood pressure
- Cardiac failure, Carditis
- Acute abdominal presentations
- o Acute gastroenteritis
- o Ascites or edema of intestinal wall
- Hepatic failure
- Pancreas involvement
- Splenomegaly
- Clinical icterus
- Clinical manifestations of coagulopathy (thrombosis, bleeding, ...)
- CNS involvements (seizure, loss of consciousness, cranial nerve involvement)

#### Box 3

Unusual laboratory results in KD:

- Hematologic involvement:
- o Leukopenia (WBC < 4000)
- o Thrombocytopenia (platelets <150000)
- o Lymphopenia (according to age)
- Acute phase reactants:
- o ESR <30 with highly elevated CRP
- o Ferritin >450 ng/ml
- o IL-6 >100 pg/mL
- o Procalcitonin >1 ng/mL
- o BNP>100 pg/mL
- Hypercoagulopathy state:
- o Increased PT, PTT
- o Decreased fibrinogen <150
- o D-Dimer>1000
- Hepatic dysfunction:
- o Increased ALT > 2 times

#### BOX 4

Imaging evaluations in CT scan and ultrasonography

- Pulmonary involvements according to COVID-19
- Free fluid in abdominal cavity
- Edema of intestinal wall
- Thrombosis suspicions in imaging
- Ejection fraction <45%



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#### **Box 5: Diagnostic criteria**

■ Suspicious to KD according to clinical criteria (algorithm 2, 3)

Plus

- Two of the 3 following criteria:
  - o At least One clinical criterion (box 2)
  - o At least two laboratory criteria (box 3)
  - o One imaging criteria (box 4)
- \* In shock state, fever for one day would be accepted





### Algorithm 5: Therapeutic approach to Kawasaki-like syndrome or multisystem inflammatory syndrome in children (MIS-C) by SARS-COV2

#### **Box 6: Treatment of cytokine storm\*:**

- Treatment with methylprednisolone pulses (20-30 mg/kg/day) for 3 consecutive days.

+

- Treatment with IVIG 2gr/kg single dose after the first MTP (If coronary arteries involved)

+

- Enaxaparin in critical patients (admitted in ICU) 0.5 mg/kg/dose every 12 hours

+

- Aspirin 3-5 mg/kg IVIg should be infused with blood pressure and pulse rate monitoring in a prolongated time of about 18-24 hours with complete hydration

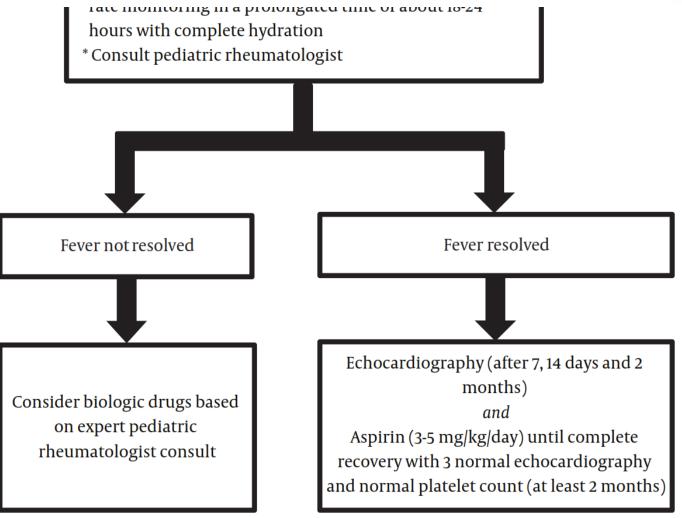
\* Consult pediatric rheumatologist

#### **Box 7: Suspicious to infectious disease\***

- Antibiotic therapy (if necessary)
- Anti viral therapy (if necessary)
- Hydroxychloroquine treatment
- Treatment of other symptoms
- \* The above mentioned approaches should be performed parallel cytokine storm treatment with pediatric infectious disease consultation

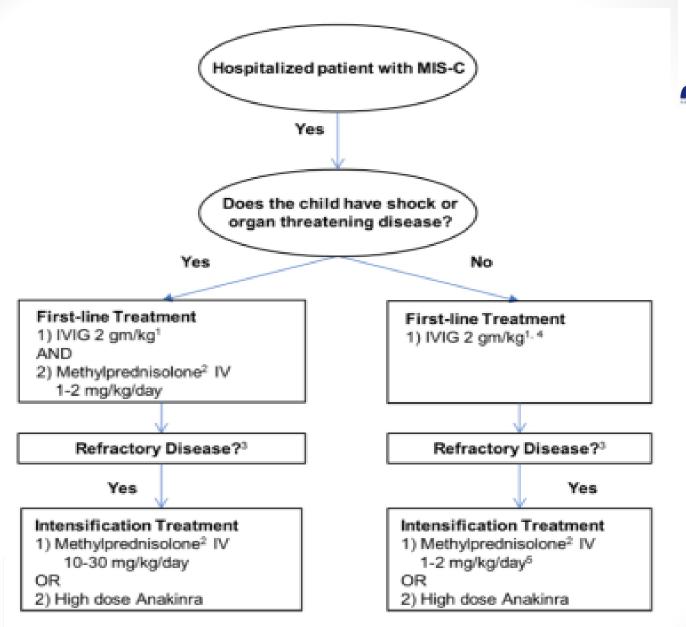












Clinical Guidance for Pediatric Patients with MIS-C Associated with SARS-CoV-2 and Hyperinflammatin in COVID-19. approved by the ACR Board of Directors on June 17, 2020

ا بجمن رو بالوفرري كودكان ايران Pediatric Rheumatology Society of Iran

## Final outcome



### COVID infection:

Total patients: 1215

Mortality: 28

Mortality rate: 2.3%

### MIS-c:

Total patients: 320

Mortality: 5

Mortality rate: 1.5%



## Evaluation for Moderate to



### severe cases

- Lab studies:
  - CBC, CRP, ESR
  - Ferritin
  - LDH
  - Liver function tests
  - Serum electrolytes and renal function tests
  - Urinalysis
  - Coagulation studies (PT, INR, PTT, D-dimer, fibrinogen)
  - Troponin, BNP or N-terminal pro-BNP



# Printed Control of Declinary

## **Evaluation for Mild Cases**

- Lab studies:
  - CBC
  - CRP
  - Serum electrolytes and renal function tests
- If these results are abnormal, additional testing is performed (listed above).
- Following cardiac complication (at least 1 echocardiography afetr 2 weeks).



