Hand, Foot and Mouth Disease

Hand, foot and mouth disease (HFMD) is a viral illness which commonly causes lesions involving the mouth, hands and feet. However, it may also affect other areas such as the buttocks and genitalia. The most common causes of HFMD are Coxsackievirus A16 (CA16) and enterovirus 71 (EV71). It is normally a mild, self-limiting illness but occasionally has serious complications, and fatalities have occurred in epidemics around the world.

Epidemiology

- HFMD occurs worldwide, with a peak incidence in the summer and autumn in temperate climates.
- It is most common among infants and children aged younger than 10 years but can occur at any age.
- Adults generally develop less severe symptoms than children although severe cases, particularly in immunocompromised adults, have been reported.
- It is moderately infectious so epidemics can occur. The risk of transmission is 10-30%.
- Epidemics have occurred in Southeast Asia in particular in recent years. A large outbreak occurred in China in 2009, with numbers of fatalities above 300. China has led the development of a vaccine. Those caused by enterovirus 71 are more likely to cause neurological complications and fatalities; however, CA16 has also been involved in epidemics and deaths.
- In the UK, outbreaks occur regularly in nurseries, schools and childcare centres. Most adults have developed immunity.
- It is not a notifiable illness.

Aetiology

HFMD is most often caused by the group A Coxsackieviruses (particularly CA16) or enterovirus 71 (EV71). These closely related viruses are of the Picornaviridae family, Enterovirus genus which includes echoviruses and polioviruses. Occasionally it can be caused by other group A and B Coxsackieviruses, particularly Coxsackieviruses A6 and A10.

Transmission is commonly by the faecal-oral route. Contact with skin lesions and oral secretions (including coughs and sneezes) can also allow transmission. An infected individual can continue to shed the virus in the stools for some weeks.

The incubation period is 3-5 days.

Presentation

- **Prodrome**: includes a low-grade fever, malaise and loss of appetite. There may be a sore mouth or throat, cough or abdominal pain. EV71 can occasionally cause vomiting.
- **Mouth lesions**: after the prodrome, lesions develop in the mouth. These may be on the buccal mucosa, tongue and hard palate. The uvula, gums, lips and skin around the mouth are sometimes involved. They begin as macular lesions that progress to vesicles which then erode. Mouth lesions are typically yellow ulcers surrounded by red haloes. They may be uncomfortable or painful. Children aged less than 5 years tend to have more severe symptoms than older children.
**Skin lesions**: most usually then also develop skin lesions. They are mainly on the palms, soles and between the fingers and toes. Lesions start as 2-5 mm erythematous macules but rapidly progress to grey vesicles with an erythematous base. Lesions may also appear on the trunk, thighs, buttocks and/or genitalia. These less typical lesions are mainly an erythematous maculopapular rash rather than the papulovesicular ones found on the hands and feet. The rash lasts about 3 to 6 days. Lesions are usually asymptomatic but can be itchy or painful.

**Differential diagnosis**[1]

- Herpangina (caused by similar Coxsackieviruses or echoviruses with lesions similar to HFMD but limited to the posterior oral cavity with no skin lesions).
- *Herpes simplex* and *herpes zoster* viruses.
- Chickenpox.
- Kawasaki disease.
- Erythema multiforme (Stevens-Johnson syndrome).
- Viral pharyngitis.
- Oral forms of lichen planus.
- Gingivitis.

**Investigations**[1, 2]

- Diagnosis is usually clinical.
- The virus can be isolated from throat, vesicle or rectal swabs (placed in viral transport medium) or from faecal culture. Viral shedding in faeces can be intermittent so more than one specimen may be required.
- Polymerase chain reaction (PCR) has made enteroviral subtyping possible and is increasingly the test of choice in specialist centres, although rarely used in routine clinical practice.
Management

There is no known treatment, so management is generally supportive:

- Parents may need to be reassured that it is unrelated to foot and mouth disease in animals.
- Encourage adequate fluid intake. If the mouth is uncomfortable, dehydration may result from poor fluid intake.
- Suggest a soft diet (soup, ice cream, mashed potato, etc).
- Antipyretic analgesics, such as paracetamol or ibuprofen, are usually all that is required.
- If the mouth is very painful, the following topical agents can be considered although there is no evidence of efficacy:
  - Lidocaine oral gel.
  - Benzydamine spray or mouthwash. (Spray only for those aged over 5 years, mouthwash for those aged over 12 years.)
  - Choline salicylate oral gel (not advised in children under the age of 16 years or in women who are pregnant or breast-feeding).
  - Mouth rinses with warm salty solutions if the person can be relied upon not to swallow it.

- Any cardiovascular or neurological complications need to be treated appropriately in secondary care (see below).

Indications to consider referral

- Signs of significant dehydration (particularly in a child). Admission to hospital is rarely required.
- Neurological signs or symptoms - eg, myoclonic jerks, persistent or severe headaches, signs of developing encephalitis.
- Persistent oral ulcers.

Advice about school and childcare settings

Public Health England does not recommend that children be kept away from school/nursery/childcare for any period of time. There is no need to isolate a child with HFMD and it is not notifiable.

Advise on general hygiene measures (see 'Prevention', below).

Advice for pregnant women

There are no known adverse consequences for the fetus if a pregnant woman is in contact with HFMD. Seek specialist advice if a woman develops HFMD within three weeks of expected delivery, as there may be a risk of passing the infection to the newborn. In rare cases this can lead to severe infection in the neonate, although usually illness is mild.

Complications

Complications are rare. They include the following:

- Secondary infection of skin that has been scratched.
- Painful stomatitis due to oral involvement. This can lead to dehydration.
- Neurological involvement and meningitis are more likely if EV71 is the causative organism. Neurological involvement can include aseptic meningitis, encephalitis, neurogenic pulmonary oedema and acute flaccid paralysis. Tremors, ataxia and cranial nerve palsies can occur.
- Cardiorespiratory failure can occur in severe cases.

Prognosis

This is generally excellent with a full recovery in the vast majority people.

Symptoms tend to improve within 3-6 days, usually with full resolution of skin and mouth lesions within 7-10 days.
Those cases caused by EV17 have a greater likelihood of developing neurological complications, although this is rare. Aseptic meningitis usually resolves without sequelae but encephalitis is more likely to have adverse outcomes. During epidemics, particularly in the Western Pacific region, there have been a number of fatalities, particularly in the outbreak in China in 2009.

**Prevention**[^1]

Good hygiene measures prevent spread within the family:

- Thorough handwashing after using the toilet. There is evidence that handwashing has a significant protective effect.[8]
- Covering of nose and mouth when coughing or sneezing.
- Hygienic disposal of used tissues and nappies.
- Avoidance of sharing of cups, eating utensils, towels, etc.
- The virus can be shed in faeces for some time after recovery, so good hygiene should be maintained.

China has led the way with development of, and clinical trials for, a vaccine to protect against HFMD. Plans for an immunisation strategy against the more potentially fatal EV71 are underway in China.[9,10] There is currently no vaccine against the Coxsackieviruses, although there is work being done in this area.[3]

**Further reading & references**

1. Hand foot and mouth disease; NICE CKS, March 2010 (UK access only)
2. AGuide to Clinical Management and Public Health Response for Hand, Foot and Mouth Disease (HFMD); World Health Organization (WHO) 2011
6. Guidance on infection control in schools and other childcare settings; Public Health England (September 2014)

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