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Learning objectives: By the end of this lecture the students should know::

- What is acute diarrhea?
- Why the children may get acute diarrhea? `Causes.``
- Clinical scenario of common infectious causes of acute diarrhea
- What are the Complications may associated with diarrhea?.
- How can we Prevent acute diarrhea?.

Def.: Diarrhea is best defined as excessive loss of fluid and electrolytes in the stool (frequent loss of loose stool). **Or defined as** an increase in stool frequency to twice the usual number per day in infants or three or more loose or watery stools per day in older children.

Diarrheal diseases are one of the leading causes of morbidity and mortality in children world wide.

Causes of acute diarrhea:-

- 1. Gastroenteritis.
- 2. Systemic infection (like pneumonia, UTI, etc.)
- 3. Antibiotic associated (penicillin, macrolid, etc)
- 4. Others: food poisoning

Mechanism of diarrhea

- 1. <u>Secretary</u>: diarrhea results from defective absorption or increase secretion of fluid and electrolytes, producing watery diarrhea of normal stool osmolality. e.g cholera,
- 2. <u>Osmotic</u>: result from maldigestion or ingestion of unabsorbable solute produce watery diarrhea of high stool osmolality (lactase deficiency, glucose-galactose malabsorption)
- 3. <u>Increase motility:</u> result from decrease transit time. e.g. irritable bowel syndrome
- 4. <u>Decrease motility</u>: stasis lead to bacterial over growth (pseudoobstruction, blind loop)
- 5. <u>Reduce surface area</u>: decrease functional absorptive capacity. e.g. short bowel syndrome, celiac disease, Rota virus enteritis.
- 6. *Mucosal invasion*: inflammation and ulceration of mucosa. e.g. amoebiasis

Gastroenteritis:

The term *gastroenteritis* refers to infections of the GIT caused by bacterial, viral, or parasitic pathogens. Many of these infections are food-borne illnesses. The most common manifestations are diarrhea and vomiting, which may also be associated with systemic features such as abdominal pain and fever.

Enteric pathogen varies by geographical location, children in developing countries become infected with different group of bacterial and parasitic pathogens, whereas both developed and developing countries children acquire Rota virus (and other viruses), and Giardia during their first 5 years of life.

Risk factors for gastroenteritis

- 1. Major risks include environmental contamination.
- 2. lack of exclusive or predominant breast-feeding.
- 3. Malnutrition increases several fold the risk of diarrhea and associated mortality The risks are higher with vitamin A deficiency, and Zinc deficiency
- 4. Ingestion of raw or undercooked food, improper food handling.
- 5. Additional risks include young age, immune deficiency and measles,

Others: like blood group O, vitamin A deficiency in Vibrio cholerae O1 and O139

Clinical Signs Associated With Dehydration

MINIMAL OF NO

SYMPTOM	DEHYDRATION	SOME DEHYDRATION	SEVERE DEHYDRATION
Mental status	Well; alert	Normal, fatigued or restless, irritable	Apathetic, lethargic, unconscious
Thirst	Drinks normally; might refuse liquids	Thirsty; eager to drink	Drinks poorly; unable to drink
Heart rate	Normal	Normal to increased	Tachycardia, with bradycardia in most
1			severe cases
Quality of pulses	Normal	Normal to decreased	Weak, thready, or impalpable
Breathing	Normal	Normal; fast	Deep
Eyes	Normal	Slightly sunken	Deeply sunken
Tears	Present	Decreased	Absent
Mouth and tongue	Moist	Dry	Parched
Skinfold	Instant recoil	Recoil in <2 sec	Recoil in >2 sec
Capillary refill	Normal	Prolonged	Prolonged; minimal
Extremities	Warm	Cool	Cold; mottled; cyanotic
Urine output	Normal to decreased	Decreased	Minimal

Modified from Duggan C, Santosham M, Glass RI: The management of acute diarrhea in children:

Viral-gastroenteritis:

Rotavirus:-is the most frequent cause of diarrhea during winter months.

Rout: Fecally contaminated foods. Ready-to-eat foods touched by infected food workers (salads, fruits) **Aerosol transmission of rotavirus may be possible**

Nearly all infants and children worldwide were infected by 2 yr of age before vaccine introduction. Primary infection with rotavirus causes moderate to severe diseases, while reinfection in adolescent usually produce mild illness. The virus invades the epithelium of the upper S.I., and in sever cases may extend throughout the S.I. and colon causing villous damage and sometime secondary transient dissaccaridase deficiency. the patient have vomiting which may last for 3-4 days with diarrhea which usually persist for 7-10 days.

Lab finding: Isotonic dehydration with acidosis is the most common finding in children with severe viral enteritis.

- GSE: The stools are free of RBCs and WBCs.
- CBC count: the white blood cell count may be moderately elevated secondary to stress, the marked left shift seen with invasive bacterial enteritis is absent

The diagnosis is confirmed by **ELISAs**, which offer >90% specificity and sensitivity, are available for detection of group A rotavirus and enteric adenovirus in stool samples.

Treatment: 1- Avoiding and treating dehydration are the main goals in treatment of viral enteritis. The rehydration may require admission for I.V. fluid therapy.

- 2- **Oral zinc** for 10–14 days during and after diarrhea (10 mg/day for infants <6 mo of age and 20 mg/day for those >6 mo)
- 3-Therapy with probiotic organisms such as Lactobacillus species has been shown to be helpful only in mild cases and not in dehydrating disease

Bacterial gastro-enteritis:

Many bacterial pathogens may produce diarrhea, the infection usually acquired by **feco-oral** route:

*Salmonella:- It transmitted by ingestion of contaminated food or by contact with infected animals .it produce diarrhea by invading the intestinal mucosa causing nausea, vomiting, crampy abdominal pain followed by loose watery stool and fever.

Diagnosis: General Stool Examination **GSE**: The stool typically contains RBCs and pus cells.

• stool culture. And Blood culture in young infants.

Treatment: -Rehydration and Oral zinc

Antibiotics: No antibiotics for uncomplicated gastroenteritis in normal hosts caused by non-typhoidal species. **Treatment indicated** in infants <3 mo, and patients with malignancy, chronic GI disease, severe colitis hemoglobinopathies, or HIV infection, and other immuneincompetent patients. **Cefotaxime** 100-200 mg/kg/day every 6-8 hr for 5-14 days

Or **Ceftriaxone** 75 mg/kg/day once daily for 7 days

Or **Ampicillin** 100 mg/kg/day every 6-8 hr for 7 days

Or **Cefixime** 15 mg/kg/day for 7-10 days

*Shigella: Usually affect children < 2 years age.

it transmitted by direct person to person feco-oral contact, it penetrate the epithelium produce inflammation, ulceration and bleeding .it can also produce toxins resulting in watery diarrhea, crampy abdominal pain and high grade fever >40 c.

Shigellosis may associated with seizure .

Diagnosis: GSE and Stool culture.

Treatment: 1- Rehydration.

- 2-A single large dose of vitamin A (200,000 IU) lessens severity of shigellosis in settings where vitamin A deficiency is common.
- 3- Oral zinc
- 4- Antibiotics: Currently, in most developed and developing countries, Shigella strains are often resistant to ampicillin and TMP-SMX; therefore, these drugs should not be used for empirical treatment.

Ceftriaxone (50 mg/kg as a single daily dose IV or IM) can be used for empirical therapy, **Oral cefixime** (8 mg/kg/24 hr divided every 12-24 hr).

Azithromycin (12 mg/kg orally for the 1st day, followed by 6 mg/kg/day for the next 4 days).

Ciprofloxacin (20-30 mg/kg/24 hr divided into 2 doses) used to be a back-up drug to treat shigellosis but is now the drug of choice recommended by the WHO for all patients with bloody diarrhea, irrespective of their ages.

A child who has typical dysentery and who responds to initial empirical antibiotic treatment should be continued on that drug for a full 5-day course even if the stool culture is negative.

- E. coli:- may produce diarrhea by different mechanism:-
 - *Both *Entero-pathogenic E.coli EPEC* and *Entero-toxogenic E.coli* produce toxin which increase secretion and reduce absorption.
 - *Entero-invasive E.coli (EIEC)cause mucosal damage.
- *Entero-haemorrhagic E.coli cause haemorrhagic colitis +/-Hemolytic uremic syndrome. Specific antimicrobial therapy of diarrheagenic E. coli is improving with accurate, rapid molecular diagnostic panels using direct fecal samples. Treatment is complicated by resistant to antibiotics because of their previous exposure to inappropriate antibiotic therapy. If rapid molecular panel tests are not available, EIEC infections may be treated before culture results are available because the clinician suspects shigellosis and has begun empirical therapy. If the organisms prove to be susceptible, TMP-SMX is an appropriate choice, treatment of EPEC infection with TMP-SMX intravenously or orally for 5 days may be effective in speeding resolution,

*Campylobacter jejuni: the infection is spread by person to person contact and by contaminated food and water: Raw and undercooked poultry, unpasteurized milk, contaminated water .it invade the mucosa causing enterocolitis most pt have mild disease recover spontaneously before reaching the diagnosis.

Treatment: Rehydration, Oral zinc

Antibiotics are recommended for patients with severe course, and for children who are immunosuppressed or have underlying diseases. Most Campylobacter isolates are susceptible to macrolides, fluoroquinolones, aminoglycosides, chloramphenicol, tetracyclines, and clindamycin

*Yersinia enterocolitica:-Usually transmitted by pets and foods causing self-limiting disease (3days-3weeks). Appendicitis-like symptoms (diarrhea and vomiting, fever, and abdominal pain) occur primarily in older children and young adults.

Treatment Enterocolitis in an immunocompetent patient is a self-limiting disease, and no benefit from antibiotic therapy is established. For patients with systemic illness and very young children with either of:- TMP-SMX aminoglycosides, third-generation cephalosporins, and quinolones.

*Vibriosis (cholera):

Cholera is acute watery diarrhea caused by group of toxins produced by <u>V. cholerae</u>, the bacteria is transmitted by feco-oral route, the affected pt gets watery diarrhea &vomiting with low grade fever. in sever cases the pt have profuse painless watery rice-water diarrhea with fishy-odor &rapidly develop sever dehydration +/-shock & renal failure if untreated

Diagnosis: GSE and Stool Culture:

transport media :Sea salt , and Culture media: TCBS, Or TTGA ELISA testing for toxins .

<u>Treatment:</u> 1-REHYDRATION: Vomiting is not a contraindication to ORS. Severely dehydrated patients require intravenous fluid, ideally with lactated Ringer solution.

- 2- Oral Zinc should be given as soon as vomiting stops. usually within 4-6 hr after initiation of rehydration therapy)
- 3- Antibiotics: Antibiotics should only be given in cases with moderately severe to severe dehydration

Adults :Doxycycline 300 mg PO given as a single dose OR **Ciprofloxacin** 1g PO single dose Or **Azithromycin** 1g PO single dose

Children: Erythromycin 12.5 mg/kg/dose4 times a day \times 3 days (up to 500 mg per dose \times 3 days) or Azithromycin, 20 mg/kg as a single dose (up to 1 g)

Or Ciprofloxacin 20 mg/kg PO as a single dose

Or **Doxycycline** 2-4 mg/kg PO as a single dose

Parasitic gastroenteritis:

- 1. Entameba histolytica
- 2. Giardia lamblia
- 3. Cryptospordium spp.
- 4. Blastocystis hominis

Amebiasis: protozoal infection caused by *Entameba histolytica*, it transmit by any uncooked food or food contaminated by an ill food handler after cooking; drinking water, or by person to person contact, it can cause invasive disease with flask shape ulcer in the intestinal mucosa. The infected individual is mostly asymptomatic. 2-8 % of pt develop diarrhea with blood and mucus, fever, abdominal pain headache and chills, untreated pt may have perforation and subsequent peritonitis Ameba may disseminated to involve extra-intestinal organs, the liver is the most common organs affected (amebic liver abscess).

Diagnosis: detection of E. histolytica antigens in stool. GSE, 3 samples may needed. **Treatment**: 1- rehydration. 2- Oral zinc.

3- Nitroimidazole such as **Metronidazol** for amebic Colitis or liver abscess: 35-50 mg/kg/day in 3 divided doses for 7-10 days or **Tinidazol tab.** for amebic Colitis 50 mg/kg/day once daily for 3 day Then followed by **Diloxanide furoate** 20 mg/kg/day in 3 divided doses for 7 days.

<u>Giardiasis:</u> infection with <u>G lamblia</u> is world wide distribution affects humans, dogs, and other wild animals. <u>Route</u>: feco-oral route. outbreak may occurs in day care center. the pt may be asymptomatic, or may have sudden explosive watery foul-smelling stool, flatulence, abdominal distension, nausea, and vomiting.some time the pt present with chronic diarrhea.

- **Diagnosis OF Giardiasis**: GSE(3 sample)
 - o Examination of duodenal aspirate.
 - o ELISA.

<u>Treatment</u>: Rehydration., Oral zinc. <u>Tinidazole tab >3 yr: 50 mg/kg once,</u>

or **Metronidazol**(15 mg/kg/day tds for 5-7 days)

Complications of acute diarrhea:

Most of the complications associated with gastroenteritis are related to delays in diagnosis and delays in the institution of appropriate therapy

- 1-dehydration, pre-renal renal failure, shock, acidosis
- 2-Electrolytes disturbance: Hypokalemia may cause ileus., Hypo/hypernatremia, and Hypocalcemia may cause convulsions.
- 3-Disseminated infections.
- 4-Hypoglycemia.
- 5-Secondary dissaccharidase deficiency... chronic diarrhea... Malabsorption and malnutrition.
- 6-Convulsion:
- 7- Extraintestinal Manifestations of Enteric Infections:
- a) Reactive arthritis: (Salmonella, Shigella, Yersinia, Campylobacter)
- b) Guillain-Barré syndrome: (Campylobacter)
- c) Glomerulonephritis: (Shigella, Campylobacter, Yersinia)
- d) Erythema nodosum: (Yersinia, Campylobacter, Salmonella)
- e) Hemolytic uremia syndrome: (Shigella dysenteriae 1, Escherichia coli 0157: H7)
- f) Hemolytic anemia: (Campylobacter, Yersinia).

Prevention of diarrhea:

- **1.Promotion Of Exclusive Breast-Feeding:** Exclusive breast-feeding protects very young infants from diarrheal disease through the promotion of passive immunity and through reduction in the intake of potentially contaminated food and water.
- **2. Improved Complementary Feeding Practices** Complementary foods should be introduced at 6 mo of age while breast-feeding should continue for up to 1 yr (longer period for developing countries).

Vitamin A supplementation reduces childhood mortality by 34%; improved vitamin A status reduces the frequency of severe diarrhea

- 3. Vaccination: Rotavirus, vaccines against Shigella and ETEC., cholera vaccine
- 4 .Improved water and sanitary facilities and promotion of personal hygiene.
 - 5. **Improved management of diarrhea** The ORS and zinc supplementation for the management of diarrhea, coupled with selective and appropriate use of antibiotics, have the potential to reduce the number of diarrheal deaths among children.

References:

- Nelson Textbook of Pediatrics, 21 edition.
- Nelson essentials Textbook of Pediatrics, 7th edition.
- Illustrated textbook of pediatrics.5th edition