

# CHRONIC DIARRHOEA OF INFANCY

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## Introduction

Not long ago adjectives such as chronic or persistent or protracted or intractable have been equivalently used to define diarrhoea lasting for more than 2 weeks, in an infant with less than 3 months of age, who is losing or not gaining weight, and with stools that are negative for infective pathogens. This definition has been coined by Avery almost 40 years ago (1968) for a severe diarrhoeal illness of often undefined etiology, with a high mortality approaching 15% - 30%, mainly due to infection and malnutrition. However, in the last couple of decades the clinical picture and etiology have changed substantially. Therefore, the aim of this extended summary is to review the etiology and the pathogenesis of chronic diarrhoea starting in the first 6 months of life. Also, the up-to-date therapeutic approaches will be presented.

## Definitions & etiology

A broad spectrum of different diarrhoeal diseases, initiated within the first 6 months of life, has recently been divided into two major categories: i. protracted / persistent and ii. intractable diarrhoea. In the first subtype are diseases which subside despite the initial severity, following an appropriate nutritional treatment, such as postenteritis syndrome, food protein sensitization, enzyme deficiencies, immune deficiencies, early coeliac disease, etc. In the second group diarrhoea persists for years, despite different modes of nutritional and drug treatment, and therefore is today named as intractable diarrhoea of infancy. This second category is further divided according to the presence of small intestinal mucosal lesion into two subtypes, the first one with mucosal atrophy caused by disease such as autoimmune enteropathy, epithelial dysplasia and microvillous inclusion disease, and the second one with normal mucosal findings embracing conditions such as transport defects and micronutrient deficiencies. In the table 1 different causes of protracted and intractable diarrhoea are presented.

## Intractable diarrhoea: pathogenesis & treatment

Diseases presently known under the name of «intractable diarrhoea» are developed either due to genetic defects in ion transport (Na/H & glucosa/galactosa malabsorption..) or in cell structure and metabolism (microvillous atrophy, epithelial dysplasia) or are due to immune hyperactivity and autoimmunity (autoimmune diarrhoea, IPEX syndrome..). Diarrhoea commonly develops soon after birth or in the first few months of life, and by pathophysiology is either secretory or a combination of secretory & osmotic. Patients with autoimmune pathogenesis may present with extradiigestive

disorders such as diabetes, arthritis, thyroiditis, etc. Disorders are mostly life-threatening, have progressive course, and patients present with intestinal failure, requiring prolonged parenteral nutrition or are even candidates for intestinal transplantation. Those with autoimmune background may respond to steroids and/or massive immunosuppressive medications.

### **Protracted / persistent diarrhoea: pathogenesis & treatment**

Protracted / persistent diarrhoea, defined by WHO in 1985 as diarrhoeal episode lasting for more than 2 weeks, and following mostly an infection-induced illness, has a complex pathogenesis. Despite initial acute episode, persistent diarrhoea is not caused by persistence of the micro-organism but is due to alteration in host factor(s) that render the children more susceptible to bacterial overgrowth and to other causes of malabsorption. Although causes may be very different a vicious cycle ensues, whereby malnutrition, immune deficiencies, food sensitization and bacterial overgrowth are common mechanisms, irrespective of the initiating factor. However, the final common pathway probably involves self-perpetuating injury to the small intestinal mucosa, while the predisposing conditions are previous diarrhoeal episodes, malnutrition, poverty, formula and / or diluted cow's milk feeding practices. Concerning specific pathogens associated with persistent diarrhoea, the following are most commonly listed: enteropathogenic E.coli, enteroaggregating E. coli, Clostridium difficile, Giardia lamblia, Cryptosporidium parvum, Salmonella, Entamoeba histolytica. Protracted diarrhoea is primarily a nutritional disorder and, therefore, an appropriately chosen nutritional treatment is the mainstay of therapeutic approach. The current algorithm for nutritional treatment of infants with protracted diarrhoea is presented.

### **Recommended literature**

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