

Evaluation of the synbiotic Prodefen GG drops in pediatric patients with acute diarrhea of probable viral etiology

Submission date 17/07/2024	Recruitment status No longer recruiting	<input type="checkbox"/> Prospectively registered <input type="checkbox"/> Protocol
Registration date 28/08/2024	Overall study status Completed	<input type="checkbox"/> Statistical analysis plan <input type="checkbox"/> Results
Last Edited 17/07/2024	Condition category Infections and Infestations	<input type="checkbox"/> Individual participant data <input checked="" type="checkbox"/> Record updated in last year

Plain English summary of protocol

Background and study aims.

Acute viral gastroenteritis is a very common illness in healthy children worldwide. Little children and their caregivers must cope with several days of unpleasant diarrhea till full recovery. Recent studies have shown the efficacy of probiotics in the improvement of diarrhea.

This study objective is to assess clinical benefit of a new liquid formulation of a multistrain synbiotic (probiotics+prebiotics) over standard measures (diet and/or oral rehydration) for the management of acute diarrhea of probable viral origin in infants.

Who can participate?

Infants aged 2 years or younger, with acute diarrhea of less than 48 hours of evolution.

What does the study involve?

The present randomized controlled trial, PRODINFANT study, was designed to assess the effectiveness of a multistrain synbiotic compound in a new drops formulation for treating Acute Gastroenteritis of viral origin in a group of children in the first 2 years of life. Patients were randomized in 2 groups in order to receive a 7-day treatment of a 7-multistrain synbiotic drops in addition to supportive therapy (Synbiotic group) or supportive therapy (diet and/or oral rehydration) alone (Control group). The objective of the study is to evaluate the additional benefits of the synbiotic therapy to treat acute diarrhea in children over standard supportive measures.

What are the possible benefits and risks of participating?

There are risks associated with any treatment intervention; however regarding to probiotics, this risk is low. Randomized clinical trials have examined the safety of probiotics and have shown that they are well tolerated and associated with few adverse events in healthy individuals at doses higher than those used in the current study. Some of the adverse events reported in previous studies with probiotics included rash, nausea, gas, flatulence, abdominal bloating, abdominal pain, vomiting, increased phlegm, chest pain, constipation, taste alteration, and loss of appetite.

The microorganisms contained in Prodefen GG drops are considered probiotics and therefore

are usually part of the host's intestinal microbiota. For these reasons, we can consider it a safe product.

Probiotics are effective in the prevention and management of various diseases, especially those directly related to the human intestine, including acute diarrhea of possible viral etiology or antibiotic-associated diarrhea (AAD). Therefore, we can consider that the study is conducted according to usual clinical practice.

For the children in the treated group, a benefit is assumed, as a reduction in the duration of diarrhea is expected due to the action of the dietary supplement Prodefen GG drops.

For the control group receiving only the standard treatment of diet and oral rehydration, participation in this study will not represent any additional benefit.

Where is the study run from?

The study will be run from 10 pediatric primary care consultations or emergency services throughout Spain.

When is the study starting and how long is it expected to run for?

July 2018 to July 2022

Who is funding the study?

ITF RESEARCH PHARMA S.L.U. (Spain)

Who is the main contact?

Mr. Francisco Javier Hidalgo Bermejo
spain.medinfo@italfarmacogroup.com

Contact information

Type(s)

Public, Scientific, Principal Investigator

Contact name

Mr Francisco Javier Hidalgo Bermejo

ORCID ID

<http://orcid.org/0000-0002-4345-9829>

Contact details

Calle San Rafael 3
Alcobendas
Spain
28108
+34 916572323
spain.medinfo@italfarmacogroup.com

Additional identifiers

EudraCT/CTIS number

Nil known

IRAS number

ClinicalTrials.gov number

Nil known

Secondary identifying numbers

RES-6066-C1

Study information

Scientific Title

Effectiveness of a multistrain synbiotic product in children with acute viral diarrhea: a multicenter prospective randomized controlled study

Acronym

PRODINFANT

Study objectives

To assess the clinical benefit of a new liquid formulation of a multistrain synbiotic (probiotics+prebiotics) over standard measures (diet and/or oral rehydration) for the management of acute diarrhea of probable viral origin in infants.

Ethics approval required

Ethics approval required

Ethics approval(s)

Approved 05/11/2018, Clinical Research Ethics Committee (CEIC) of Hospital Universitario Puerta de Hierro (C/ Joaquín Rodrigo, 2, Madrid, 28222, Spain; +34 91 191 60 00; secreceic.hpth@salud.madrid.org), ref: 19.18

Study design

Multicenter prospective randomized open-label and controlled clinical trial

Primary study design

Interventional

Secondary study design

Randomised controlled trial

Study setting(s)

Childcare/pre-school, GP practice, Hospital

Study type(s)

Treatment, Safety, Efficacy

Participant information sheet

Not available in web format, please use the contact details to request a participant information sheet.

Health condition(s) or problem(s) studied

Acute diarrhea of probable viral etiology

Interventions

PRODINFANT study is designed as a multicenter, prospective, randomized, open-label and controlled clinical trial. A cohort of children presenting an acute diarrhea will be randomized between two groups, Synbiotic Group will receive a one-week treatment with a 7-multistrain synbiotic drops in addition to supportive therapy and Control Group will be assigned to supportive therapy (diet and/or oral rehydration) alone.

Investigators belong to pediatric primary care consultations or emergency services throughout Spain.

Children who met the inclusion criteria will be assigned a consecutive number at each study site according to the order of arrival, and then randomized to treatment with the synbiotic product (synbiotic group; odd number) or standard supportive measures (control group; even number). All participants regardless of their assigned group receive supportive standard treatment with diet and/or oral rehydration therapy. Children randomized to the synbiotic group receive a daily recommended dosage (10 drops) of the synbiotic product (Prodefen® drops, ITALFARMACO, S. A., Alcobendas, Madrid, Spain), with or without food and preferably in the morning for 7 consecutive days.

The study includes a 7-day treatment period with 3 visits (1 in-person visit and 2 visits by phone) and a final safety follow-up phone call at day 10. Children visiting the study center (visit 1) will be assigned to the treatment group, after inclusion criteria is confirmed, and signed informed consent is obtained. The first dose of the synbiotic product will be administered by the investigator at the study center (day 1) and parents are provided with a drop bottle of the product for treatment during the subsequent 6 days. Parents of all participants are given a patient diary card in which all information of the study variables have to be recorded on a daily basis. The parents are contacted by phone on days 4 and 7, so that the investigator can collect information on the study variables related to the course of the disease. Data of acceptability and tolerability of treatment are collected on day 7. In addition, on day 10, 3 days after the end of the treatment, adverse events occurring during that period will be recorded. If scheduled phones calls are foreseen during the weekend, a 2-day window period will be allowed.

Intervention Type

Supplement

Primary outcome measure

Efficacy is measured using the percentage of children who suffer diarrhea according to several definitions, the WHO criteria, the Bristol Stool Form Scale (BSFS) criteria, both WHO and BSFS criteria, and the parents' opinion regarding more stools per day or more fluid in consistency for at least 1, ≥ 2 or ≥ 3 consecutive days from day 1 to the end of the study

Secondary outcome measures

1. Number of days of diarrhea (three or more passage of stools per day) and duration (days) of diarrhea are measured until the end of the study
2. Recovery is measured by the percentage of children recovered from diarrhea (2 consecutive days without diarrhea) and the stool characteristics (frequency, consistency according to the BSFS, severity of diarrhea) throughout the study
3. Associated symptoms (vomiting, nausea, fever, mucus in stools and abdominal pain) are measured by the duration of symptoms and need of treatment throughout the study
4. Recovery are measured counting the number of visits to the primary care pediatrician or the emergency service as well as the need of concomitant medication throughout the study
5. Tolerability of the synbiotic product and satisfaction with treatment (acceptability) are

measured using a 5-point Likert scale, from “very good” to “very bad” and from “very satisfied” to “not at all satisfied”, respectively at week 8

6. The impact of treatment on the quality of life and daily activities of the parents will be assessed using a 5-point Likert scale from “not at all affected” to “extremely affected” at week 4 and 8

Overall study start date

19/07/2018

Completion date

08/07/2022

Eligibility

Key inclusion criteria

1. Children from both sex whose parents/guardians have given their informed consent to participate in the study
2. Children aged 2 years or younger who visited the pediatrician in the primary or the emergency healthcare center, presenting an acute diarrhea episode lasting less than 48 hours, with a probable viral origin
3. Children whose diarrhea will be treated only with diet and/or oral rehydration, on an outpatient basis
4. Children for whom the pediatrician considers treatment with the dietary supplement Prodefen GG drops as an additional treatment measure

Participant type(s)

Patient

Age group

Child

Lower age limit

0 Years

Upper age limit

2 Years

Sex

Both

Target number of participants

80

Total final enrolment

75

Key exclusion criteria

1. Children who have presence of any disease that may cause diarrhea, different from an infection, such as inflammatory bowel disease, food allergy, lactose intolerance, etc.
2. Immunocompromised children, children with valve implants, and/or those with a history of

infectious endocarditis, due to the risk of bacteremia

3. Children with severe dehydration or malnutrition

4. Children with visible blood in stools

5. Children with current pharmacological treatment for diarrhea

6. Children who have used antibiotics and/or probiotics the previous 7 days

7. Children allergic to milk protein (Prodefen GG drops may contain traces of milk, but do not affect those who are lactose intolerant)

8. Children currently participating in another study

9. Children allergic to any of the components of Prodefen GG drops

Date of first enrolment

01/09/2020

Date of final enrolment

26/11/2021

Locations

Countries of recruitment

Spain

Study participating centre

Centro medico privado-MEDInfant

Barcelona

Spain

08022

Study participating centre

Hospital Vigen del Mar

Madrid

Spain

28016

Study participating centre

Hospital Público Universitario del Sureste

Arganda del Rey

Spain

28500

Study participating centre

HBN 39 Especialidades Médicas

Madrid

Spain

28036

Study participating centre

Centro Medico Oporto

Madrid

Spain

28025

Study participating centre

Hospital Quiron A Coruña

A Coruña

Spain

15009

Study participating centre

Consultorio Pediátrico Dr. Hugo Daniel Ruiz Morena

San Sebastián de los Reyes, Madrid

Spain

28701

Study participating centre

Centro Médico Valdebernardo, S.L.P.

Madrid

Spain

28032

Study participating centre

Madrid Pediátrica Consultas

Madrid

Spain

28016

Study participating centre

Consultorio Privado
Madrid
Spain
28042

Sponsor information

Organisation

ITF RESEARCH PHARMA S.L.U.

Sponsor details

Calle San Rafael 3
Alcobendas
Spain
28108
+ 34 916572323
spain.medinfo@italfarmacogroup.com

Sponsor type

Industry

Funder(s)

Funder type

Industry

Funder Name

ITF RESEARCH PHARMA S.L.U.

Results and Publications

Publication and dissemination plan

Planned international conference presentations and publication of manuscript in a biomedical peer-reviewed journal.

Intention to publish date

15/09/2024

Individual participant data (IPD) sharing plan

The datasets generated during and/or analysed during the current study are not expected to be made available due to sponsor product patent requirements, as well as preserving participants sensitive data protection.

IPD sharing plan summary

Not expected to be made available