

Antibiotic as an adjunct to treatment of severe acute malnutrition in infants less than six months of age. A protocol for the systematic review

List of Exclusions and Reasons for Exclusions.

Abate 2020 ¹	Inappropriate study design and appropriate control was not available
Adam 1954 ²	Inappropriate Study Design
Adem 2020 ³	Inappropriate Patient Population. Study was a cohort study and only included children 6-59 months of age
Ahmed 1999 ⁴	Inappropriate Patient Population. Even though all the children were severely malnourished, all of them had diarrhea. Also, the study was not a randomized study.
Ahmed 2001 ⁵	Inappropriate Study Design. Authors discuss a protocol for treatment of severe acute malnutrition. The study was a descriptive review
Akbani 1977 ⁶	Inappropriate study population. Children with SAM who were infected with tuberculosis.
AmorelliGonzaga 1989 ⁷	Inappropriate study population. Study investigated miomycin and included both well-nourished and malnourished children.
Amza 2013 ⁸	Inappropriate study population. Study investigated mass azithromycin distribution and included children irrespective of the nutritional status.
Amza 2014 ⁹	Inappropriate study population. Study investigated mass azithromycin distribution and included children irrespective of the nutritional status.
Angelakis 2014 ¹⁰	Inappropriate study population. Study included patients with Q fever endocarditis patients treated with doxycycline and hydroxychloroquine
Ashorn 2018 ¹¹	Inappropriate study population. Study included children with diarrhea, dysentery and malnutrition.
Ashraf 2012 ¹²	Inappropriate Comparator. Children aged 2-59 months having severe pneumonia with SAM were randomized to day-care or hospital-care. All the children received antibiotics.
Autret 1989 ¹³	Inappropriate study Design. Study to assess the pharmacokinetics of gentamicin.
Ayalign 2018 ¹⁴	Inappropriate study population. Children with urinary tract infections.
Ayiya 1987 ¹⁵	Inappropriate study Design. The study was a systematic review
Aziz 2015 ¹⁶	Inappropriate Intervention. Study investigated vitamin K supplementation.
Berkley 2009 ¹⁷	Wrong comparator
Berkley 2016 ¹⁸	Inappropriate Comparator. Both the intervention and comparison group were treated according to the WHO

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	protocol and then randomized to Co-trimoxazole vs. placebo.
Berkowitz 1992 ¹⁹	Inappropriate Study Design. Review article.
Bhatnagar 1992 ²⁰	Inappropriate patient population. All the patients had diarrhea. All of them were given gentamicin
Bhatnagar 1996 ²¹	Inappropriate Study Design. Observation study in children with persistent diarrhea
Birindwa 2020 ²²	Inappropriate Study Design. Observational study in children with severe pneumonia
Bleakly 2014 ²³	Inappropriate Study Design. Before and after study to assess the effectiveness of a protocol to treat SAM.
Boesen 1953 ²⁴	Inappropriate Indication. Antibiotics were given for treatment of gastroesophageal reflux.
Bolme 1980 ²⁵	Inappropriate Study Design: Pharmacokinetic study for Penicillin in children
Bolme 1988 ²⁶	Inappropriate Study Design: Pharmacokinetic study for streptomycin in children with Tuberculosis.
Bolme 1995 ²⁷	Inappropriate Study Design. Pharmacokinetic study in malnourished children for Penicillin.
Bravo 1982 ²⁸	Inappropriate study design: Study to assess the pharmacokinetics
Bravo 1984 ²⁹	Inappropriate study design: Study to assess the pharmacokinetics
Bredow 1994 ³⁰	Inappropriate Study Design. Observational study
Brodwall 2016 ³¹	Inappropriate Study Design. Letter to editor
Bruhn 2016 ³²	Inappropriate Study Design. Observational study
Brunozi 2019 ³³	Inappropriate Study Design. Observational study
Buchanan 1977 ³⁴	Inappropriate Study Design. Letter to editor
Buchanan 1978 ³⁵	Inappropriate Study Design. Observational study
Buchanan 1979 ³⁶	Inappropriate Study Design. Observational study
Bunn 2009 ³⁷	Inappropriate Study Design. Commentary
Cakir 2012 ³⁸	Inappropriate Patient Population. Children studied leukemia
Caksen 2000 ³⁹	Inappropriate Study Design. Observational study
Caksen 2000 ⁴⁰	Inappropriate Study Design. Observational study
Cedrato 1974 ⁴¹	Inappropriate Study Design. Observational study
Chisti 2015 ⁴²	Inappropriate Patient Population. Children with respiratory symptoms only
Church 2015 ⁴³	Inappropriate Study Design. Observational study
Cochlovius 1953 ⁴⁴	Inappropriate intervention. Observational study
Dicko 2016 ⁴⁵	Inappropriate study population. Intervention was given to all the children irrespective of nutritional status at the time of chemoprophylaxis for malaria.
Dieng 2014 ⁴⁶	Inappropriate Study Design. Observational study
Dubray 2008 ⁴⁷	Inappropriate study population. Both the study groups received antibiotics and these were given to children 6-59 months of age
Duffau 1990 ⁴⁸	Inappropriate study population. Children with diarrhea

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Dutta 2006 ⁴⁹	Inappropriate study design: Observational study
Eriksson 1983 ⁵⁰	Inappropriate Patient Population. Also, observational study
Eriksson 1988 ⁵¹	Inappropriate Patient Population. Also, observational study
Ford 1976 ⁵²	Inappropriate Study Design. Observational study
Friedrich 2016 ⁵³	Inappropriate Study Design. Commentary
Garcia 1974 ⁵⁴	Inappropriate Study Design. Observational study
Gardner 2011 ⁵⁵	Inappropriate Study Design. Commentary
Gennaro 2017 ⁵⁶	Inappropriate Study Design. Commentary
Gensch 1957 ⁵⁷	Inappropriate Study Design. Observational study
Gernaat 1998 ⁵⁸	Inappropriate Study Design. Observational study
Ghosh 1995 ⁵⁹	Inappropriate Patient Population. Children with persistent diarrhea
Girma 2018 ⁶⁰	Inappropriate Study Design. Observational study
Gore Langton 2020 ⁶¹	Inappropriate Patient Population. The antibiotics were given irrespective of nutritional status and study included children beyond six months of age. The comparison group was not appropriate.
Hecht 2015 ⁶²	Inappropriate Patient Population. The study included children beyond six months of age up to 18 years.
Heikens 1993 ⁶³	Inappropriate Patient Population, mean age of children at 1.2years.
Heikens 1993 ⁶⁴	Inappropriate Comparator, antibiotics compared with receiving health care from community healthcare aids
Hirschhorn 1971 ⁶⁵	Inappropriate Study Design, paper is a commentary on the current state of research.
Howard 1967 ⁶⁶	Inappropriate Study Design, all infants studied had acute diarrhea
Isaack 1992 ⁶⁷	Inappropriate Study Design, investigates nosocomial infections.
Isanaka 2020 ⁶⁸	Inappropriate Patient Population. All patients are above 6 months of age.
Islam 2021 ⁶⁹	Inappropriate Outcomes. Antibiotic usage was outcome not treatment group. Also all patients have diarrhea.
ISRCTN 2017 ⁷⁰	Wrong patient population. Patient age ranged from 2 months to 13 years, unable to obtain subgroup data for under 6 months.
Kabalo 2017 ⁷¹	Inappropriate Patient Population. Patients aged over 6 months.
Lares-Asseff 1999 ⁷²	Inappropriate Study Design. Study evaluated blood levels for antibiotic not mortality outcome.
Lares-Asseff 2016 ⁷³	Inappropriate Comparator. Compared different doses of antibiotics to look for toxicity, not mortality.
Lattes 1954 ⁷⁴	Inappropriate Study Design and lack of full text.
Lebl 2001 ⁷⁵	Inappropriate Patient Population, age range 2 years to 18 years.

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Lelijveld 2021 ⁷⁶	Inappropriate Patient Population, age range 6 months to 59 months.
Lepage 1984 ⁷⁷	Inappropriate Study Design, patients all have bacterial infection treated with antibiotics.
Lewis 1956 ⁷⁸	Inappropriate Patient Population, patients all over 1 year old.
Maataoui 2020 ⁷⁹	Inappropriate Patient Population. Patients all over 6 months of age.
Macdougall 1957 ⁸⁰	Inappropriate Patient Population. Average age is 2 years.
MacLean 1980 ⁸¹	Inappropriate Comparator. Antibiotic usage compared to enteral feeding.
Maitland 2006 ⁸²	Inappropriate Patient Population. Median age of 25 months.
Martischinig 1952 ⁸³	Inappropriate Intervention and lack of full text.
Mathew 2016 ⁸⁴	Wrong Patient Population. Patient all aged over 6months.
Mathew 2016 ⁸⁵	Wrong Patient Population. Patient all aged over 6months.
Melaku 1999 ⁸⁶	Inappropriate Patient Population. Patient median age 5.9 years.
Moschovis 2014 ⁸⁷	Inappropriate Study Design. Patients treated for pneumonia.
Mtango 1986 ⁸⁸	Inappropriate Intervention. Patients given antibiotics to treat acute respiratory infections.
Muhammad 2020 ⁸⁹	Inappropriate Comparator. Compares nutritional treatment with antibiotics usage
Mulholland 1995 ⁹⁰	Inappropriate Patient Population. All patients have pneumonia to treat.
Muller 1953 ⁹¹	Inappropriate Study Design and lack of full text.
Murce 1955 ⁹²	Lack of Full Text.
Muwanguzi 2021 ⁹³	Inappropriate Patient Population. Children aged 1 year and above.
Nalwanga 2020 ⁹⁴	Inappropriate Patient Population. Mean age 14 months.
Nalwanga 2020 ⁹⁵	Inappropriate Patient Population. Mean age 14 months.
Nansumba 2018 ⁹⁶	Inappropriate Study Design. observational study
Nantanda 2008 ⁹⁷	Inappropriate patient population. paediatric patients with pneumonia
NCT 2009 ⁹⁸	Inappropriate patient population. paediatric patients with pneumonia
NCT 2009 ⁹⁹	Inappropriate Patient Population. children age 6 months to 5 years
NCT 2020 ¹⁰⁰	Inappropriate outcome. studied the effect of prophylactic azithromycin has on the microbiome
NCT 2015 ¹⁰¹	Inappropriate patient population . hospitalized patients (not necessarily patients with diarrheal illness) receiving prophylactic course of azithromycin (not antibiotics to treat diarrheal illness) at the time of discharge, outcomes of malnourished patients not published
NCT 2012 ¹⁰²	Inappropriate patient population . age 6 months to 5 years
Nuzhat 2005 ¹⁰³	Inappropriate study design . observational study

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	Inappropriate population . children with a skin disease
O'Brien 2021 ¹⁰⁴	Inappropriate Patient Population . children age 6 to 59 months
O'Brien 2020 ¹⁰⁵	Inappropriate intervention . studied the effect on biannual prophylactic azithromycin (not antibiotics to treat a diarrheal illness) on the mortality rate of malnourished children
O'Brien 2020 ¹⁰⁶	Inappropriate study design . observational study Inappropriate intervention . biannual prophylactic azithromycin (not antibiotics for diarrheal illness)
Page 2013 ¹⁰⁷	Inappropriate Patient Population . children age 6 to 59 months
Pai 1985 ¹⁰⁸	Inappropriate Study Design
Parpia 2020 ¹⁰⁹	Inappropriate Study Design . antibiotics were given prophylactically to the children in the study (not in the treatment of diarrheal illness) Inappropriate patient population: children did not start receiving the prophylactic antibiotics until after 6 months of age
Pinto 2012 ¹¹⁰	Inappropriate Setting . PICU Inappropriate study design . observational study
Polster 1954 ¹¹¹	Lack of Full Text
Pombo 2017 ¹¹²	Inappropriate patient population . patients were hospitalized with pneumonia, not diarrheal illness Inappropriate study design . observational study
Prentice 2013 ¹¹³	Inappropriate Study Design . this paper is a review of various studies on the management of malnutrition in children
Rasul 2006 ¹¹⁴	Inappropriate Study Design . randomized control trial but no placebo control, controls were admitted to the hospital prior to the study and thus did not receive the WHO protocol Inappropriate patient population . malnourished children age 1 month to 5 years, included hospitalizations for many reasons not just diarrheal illness Inappropriate intervention . compared WHO protocol vs no WHO protocol, so patients in both groups may have received antibiotics and the difference between the groups includes many interventions other than antibiotics
Rawson 2016 ¹¹⁵	Inappropriate Study Design . letter to the editor
Reed 1996 ¹¹⁶	Inappropriate Intervention . desired outcome was the difference in incidence of bacteremia and the difference in mortality rate among bacteremic patients between malnourished and adequately nourished children Inappropriate study design . observational study Inappropriate patient population . age 0 to 5 years
Roboz 1955 ¹¹⁷	Lack of Full Text
Rogawski 2015 ¹¹⁸	Inappropriate Study design . observational study

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Rokstad 2013 ¹¹⁹	Inappropriate Patient Population . this paper is a brief review of a study done by Trehan in 2013, which studied a population of children age 6 to 59 months
Rosenberg 1974 ¹²⁰	Inappropriate Study Design . review/opinion paper
Roy 2010 ¹²¹	Inappropriate Patient Population . children age 5 to 12 years Inappropriate indication . desired outcome as whether there was a difference in the pharmacokinetics of isoniazid in malnourished children vs adequately nourished children
Sala 1955 ¹²²	Inappropriate Intervention and lack of full text
Samotra 1985 ¹²³	Inappropriate Indication . desired outcome was whether or not there was a difference in the pharmacokinetics of gentamicin in malnourished children vs adequately nourished children
Samotra 1986 ¹²⁴	Inappropriate Indication . desired outcome was whether there was a difference in the pharmacokinetics of chloramphenicol in malnourished children vs adequately nourished children
Sanogo 2018 ¹²⁵	Inappropriate Indication . desired outcome was the level of adherence among healthcare providers to WHO guidelines regarding management (antibiotics, oral rehydration therapy, and zinc) of paediatric diarrheal illness in Bamako, Mali.
Santoro 2002 ¹²⁶	Inappropriate Study Design . observational study. Inappropriate indication . desired outcome was to see how many children admitted to the PICU for lower respiratory tract infections needed ICU level care
Schapira 1971 ¹²⁷	Lack of Full Text
Sepehr 2009 ¹²⁸	Inappropriate Patient population . age 2 to 94 years Inappropriate study design . observational study Inappropriate indication . desired outcome was optimal length of post-operative antibiotic prophylaxis as well as risk factors associated with post-operative infection
Shahira 2002 ¹²⁹	Inappropriate Patient Population . population was all pediatric patients at a teaching hospital Inappropriate study design . observational study Inappropriate indication . desired outcome was risk factors for nosocomial infections
Soheir 1981 ¹³⁰	Inappropriate Indication . desired outcome was the appropriate dose of chloramphenicol for malnourished children, not mortality
Standing 2018 ¹³¹	Inappropriate indication . studied the proper dose of IV ceftriaxone and PO metronidazole needed to achieve a therapeutic level in malnourished children
Tan 2020 ¹³²	Inappropriate Indication . studied the use of anthropometric measurements as opposed to clinical signs to identify malnourished children
Taubenslag 1950 ¹³³	Inappropriate Study Design .
Thame 2001 ¹³⁴	Inappropriate Study Design . observational study

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Thorson 1989 ¹³⁵	Inappropriate Indication .
Tilg 2013 ¹³⁶	Inappropriate Study Design . this paper is a review of a few different studies. Inappropriate patient population . Of interest, it discusses the results from Trehan 2013, which has a patient population of children age 6 months to 59 months
Tornberg-Belanger 2017 ¹³⁷	Inappropriate Study Design . observational study Inappropriate topic . studied adherence to antibiotic guidelines of various hospitals
Trehan 2010 ¹³⁸	Inappropriate Patient Population . was children age 6 to 59 months Inappropriate study design . observational study
Trehan 2013 ¹³⁹	Inappropriate Patient Population . children age 6 to 59 months
Trehan 2016 ¹⁴⁰	Inappropriate Study Design . same exact study as trehan 2013 Inappropriate patient population : children age 6 to 59 months
Trehan 2016 ¹⁴¹	Inappropriate Patient Population . letter to editor
Usman 2019 ¹⁴²	Inappropriate Study Design . observational study
Uzan-Yulzari 2021 ¹⁴³	Wrong patient population. Age of patients range from birth to 2 years.
Vather 2018 ¹⁴⁴	Inappropriate Study Design . observational study
Verani 2019 ¹⁴⁵	Inappropriate Study Design . observational study.
Viso Gurovich 2003 ¹⁴⁶	Inappropriate study design . observational study. Inappropriate Intervention . studied the outcomes of paediatric patients who received antibiotics to treat various infections, most commonly bronchopneumonia, complicated pneumonia, and conjunctivitis.
Vygen 2013 ¹⁴⁷	Inappropriate Study Design . case series (descriptive study design) looking at the outcomes of malnourished infants who received care at a nutritional rehab center
Walsh 2018 ¹⁴⁸	Inappropriate Intervention . tested the effectiveness of a lactose free legume based nutritional feed to promote the resolution of diarrhea
Weingaertner 1961 ¹⁴⁹	Lack of Full Text
Wittmann 1967 ¹⁵⁰	Inappropriate Patient Population. age range was 2 months to 2 years and not all were malnourished
Woodd-Walker 1972 ¹⁵¹	Inappropriate Study Design, inappropriate patient population . age range was from 1 year to 4 years
Zecca 1951 ¹⁵²	Lack of Full Text and study is from 1951
BMJ 2013 ¹⁵³	Wrong study design. Research news article.
Langdon 2018 ¹⁵⁴	Wrong patient population. Patient ages ranged from 6 weeks to 60 weeks with no subgroup data available for under 6 months.
Journal of Agricultural and Food Chemistry 1954 ¹⁵⁵	Wrong study design. News report article.
Akush 1951 ¹⁵⁶	Wrong indication. Patients all treated for pneumonia.

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Alam 2020 ¹⁵⁷	Wrong study design. All patients treated for diarrhea.
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1. Abate BB, Tilahun BD, Kassie AM, Kassaw MW. Treatment outcome of Severe Acute Malnutrition and associated factors among under-five children in outpatient therapeutics unit in Gubalafto Wereda, North Wollo Zone, Ethiopia, 2019. *PLoS One* 2020;15(9):e0238231. DOI: 10.1371/journal.pone.0238231.
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