PANTHERYX



Colostrum Scientific Overview

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PanTheryx

Introduction to Colostrum





More Than Just A Mother's Love...

A Mother's Colostrum Prepares Her Baby For The World Around

- Transfers immunity from mother to child
- Helps in the development of the immune and gut systems
- Strengthens the intestinal barrier to help keep pathogens out and improve digestion
- Supports healthy gut bacteria (probiotics) and reduces bad bacteria and viruses (pathogens)



The same holds true for all mammals, including dairy cows



Harnessing The Benefits of Colostrum for All Ages



Colostrum from dairy cows enables us to harness the benefit of colostrum across the lifespan





Completely natural product, produced by nature and meant to nurture





Calf receives all the colostrum it needs





Excess colostrum is collected and dried into a powder





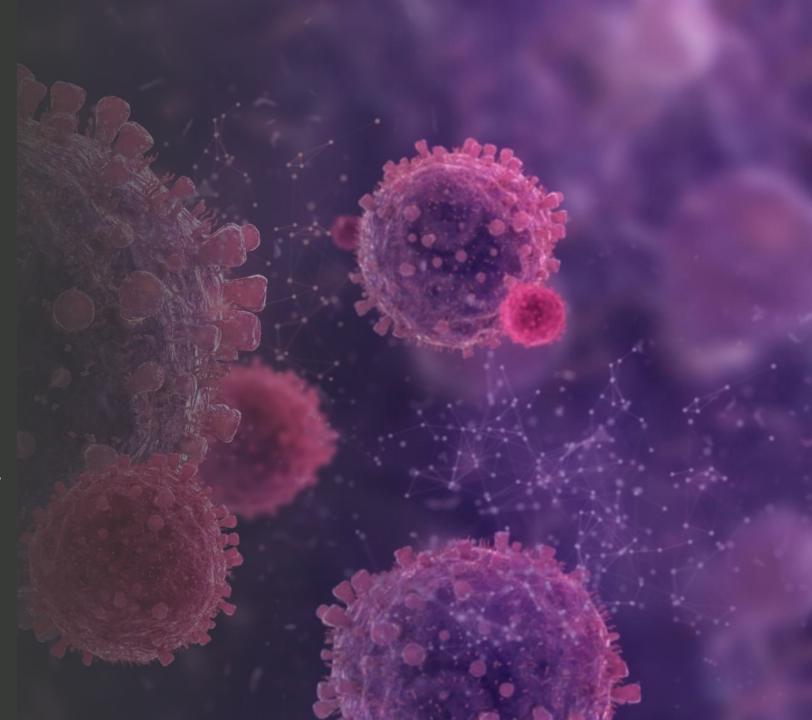
Many bioactive components to provide an array of benefits



Supports immune and digestive health, sports nutrition and more



Activity of Bioactive Components





Dairy Colostrum is a Powerhouse of Components

Unlike traditional prebiotics and probiotics....

Dairy Colostrum contains over

250

Characterized
Functional
Beneficial
Bioactive Components

Including immunoglobulins, growth factors, oligosaccharides, immune factors, and more



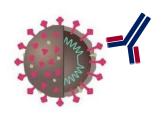


Activity for Immunity

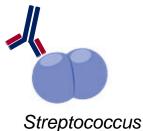
Immunoglobulins

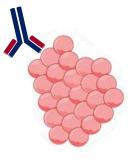
Bovine immunoglobulins have shown binding activity against a wide range of pathogens









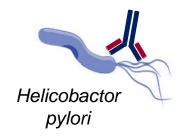


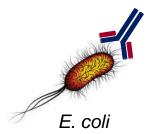
Coronaviruses

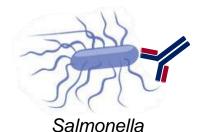
Rotavirus

Streptococcus pneumoniae

Staphylococcus







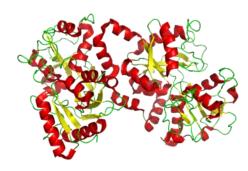
Candida albicans, Campylobacter, Cryptosporidium, Pseudomonas, and Enterobacter

And others including:



Activity for Immunity

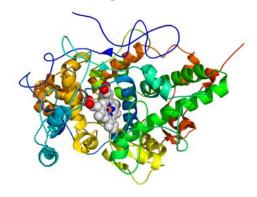
Lactoferrin



- Glycoprotein, antiviral and antibacterial
- Inhibits growth of E. coli, Salmonella, Listeria, Streptococcus, Giardia
- Blocks viruses from entering cells via binding cellular receptors or direct binding to viruses
- Antiviral against Rotavirus, RSV, herpes, Hepatitis B and C, HPV, HIV, and others

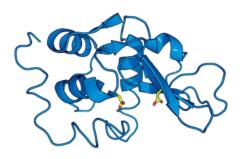
Immune Factors

Lactoperoxidase



- Enzyme with antibacterial functions
- Toxic to a wide array of bacteria via production of reactive oxygen species including hydrogen peroxide
- Inhibits bacterial metabolism

Lysozyme



- Enzyme with antibacterial functions
- Utilizes bacterial cell wall as an enzymatic substrate causing bacterial cell lysis and death



The Health of Your Gut Impacts Your Entire Body

Immune – more than 70% of the immune system is in the gut which directly and indirectly impacts immunity outside the gut (whole body)

Brain – the gut communicates directly and indirectly with the brain and can affect brain inflammation, mood, and more

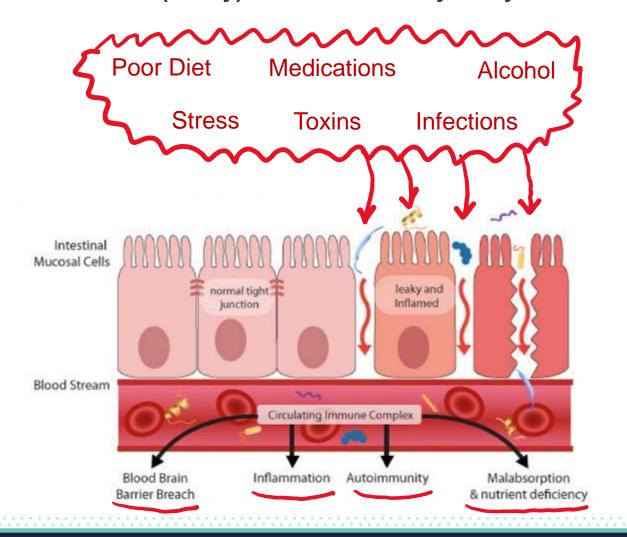
Nervous system – the gut contains 100-500 million neurons (specialized communication cells), more than the spinal cord, some call the gut a "second brain"

Heart – beneficial microbiota can affect things like good cholesterol (HDL) levels



Colostrum Growth Factors Can Help Heal The Gut

A Permeable (Leaky) Gut is Caused By Many Factors



Colostrum Contains Growth Factors

Can help protect and heal the intestinal barrier



No

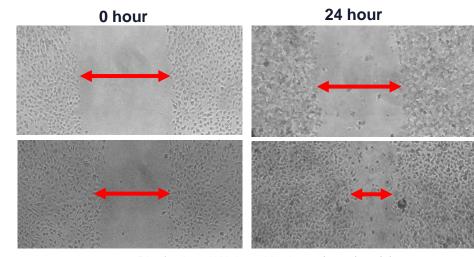
Colostrum

+ Colostrum

EGF – Epidermal Growth Factor

IGF – Insulin-Like Growth Factor and more...

Study: Colostrum from dairy cows speeds up closure of a wound in an intestinal cell model



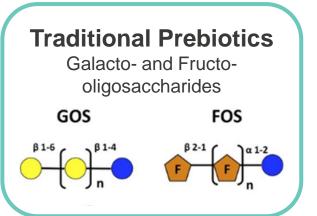
Playford and Weiser. Nutrients (2021).13(1):265



Colostrum Oligosaccharides Can Act as Prebiotics

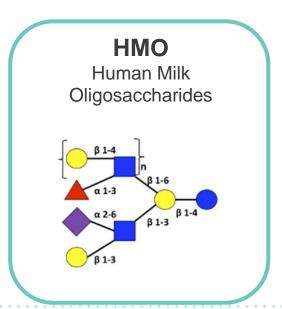
Traditional Prebiotics

Simple structure, have health benefits, but can have side effects (discomfort)



Human Milk Oligosaccharides

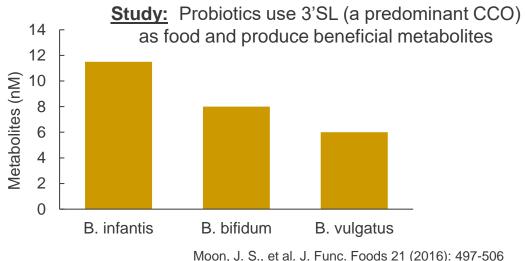
Complex structures that feed beneficial microbes in the infant gut (like Bifidobacterium infantis)



CCO Cow Colostrum Oligosaccharides α 2-6

Cow Colostrum **Oligosaccharides**

Complex oligosaccharides that have similarities to Human Milk Oligosaccharides (HMO)

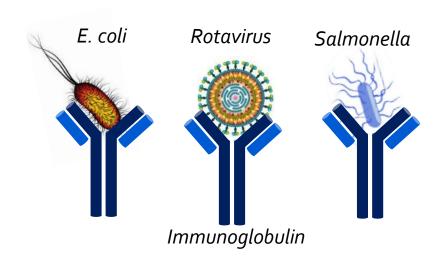




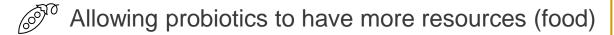
Colostrum Immunoglobulins Improve Intestine Environment

Colostrum Immunoglobulins

Can bind to pathogens in the intestine





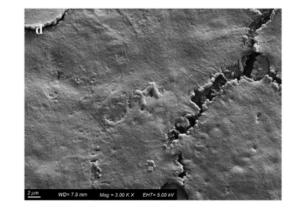


Helping to establish a good balance of bacteria

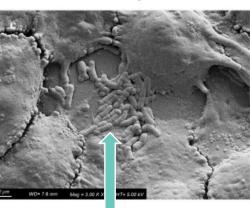
Colostrum Immunoglobulins

Can help probiotics adhere to the intestinal wall

Control



+ Immunoglobulin



Probiotic attached to intestinal cells



They likely support more successful colonization of probiotics

Morrin ST, et al. AMB Express, 2020: 10(1), 1-10.



Colostrum Can Support Probiotics

Immunoglobulins Growth Factors



Help create ideal intestinal environment

C-1

Study: Colostrum from dairy cows increases the growth of many different types of probiotic strains

	Col	
Strain	OD_{max}	†
Pediococcus acidilactici RBL39	1	_
Pediococcus acidilactici R1001	↑	4
Lactobacillus lactis RBL22	1	
Lactobacillus helveticus R0052	1	
Lactobacillus helveticus R0389	1	
Bacillus subtilis R0179	NE	
Enterococcus faecium R0026	↑	
Lactobacillus rhamnosus R0011	↑	
Lactobacillus rhamnosus GG	↑	
Lactobacillus rhamnosus RW-9595M	1	
Bifidobacterium lactis BB12	1	
Bifidobacterium longum R0175	1	

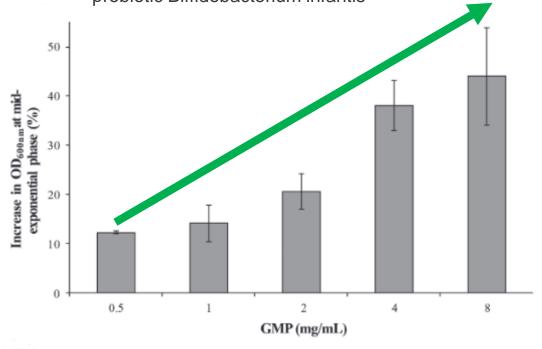
Champagne, C. P., et al. Can. J. Microbiol. 60.5 (2014): 287-295.

Oligosaccharides Glycoproteins



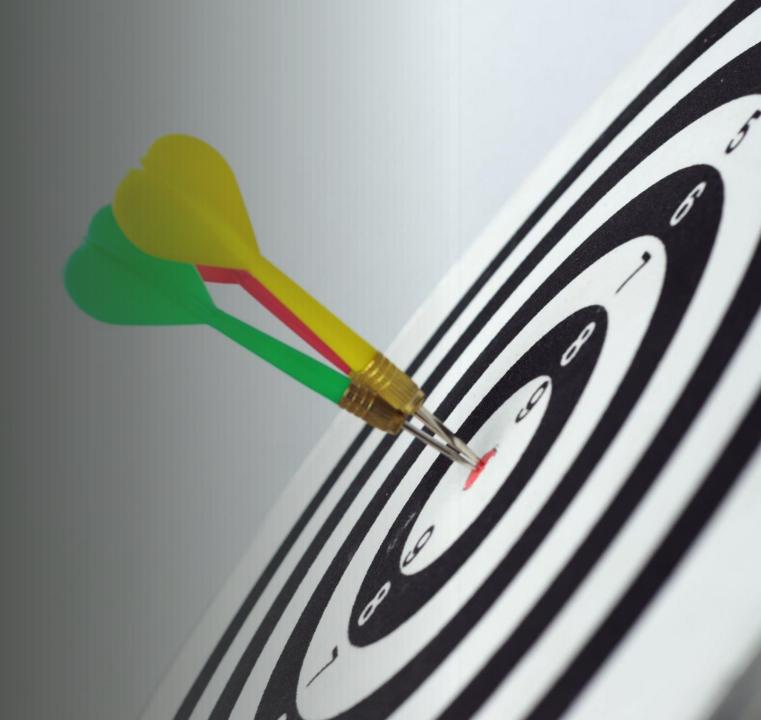
Help to feed and promote probiotic growth

<u>Study:</u> Colostrum glycoprotein (GMP) promotes the growth of probiotic Biffidobacterium infantis



O'Riordan, Noelle, et al. Journal of dairy science 101.8 (2018): 6730-6741

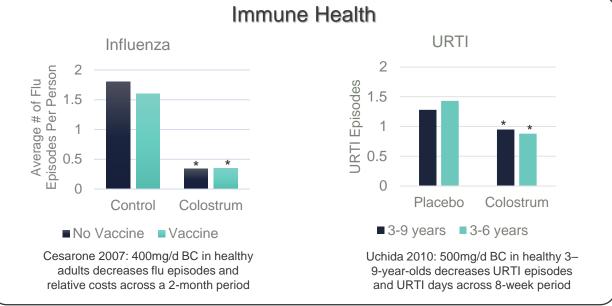
Clinical Overview Summary

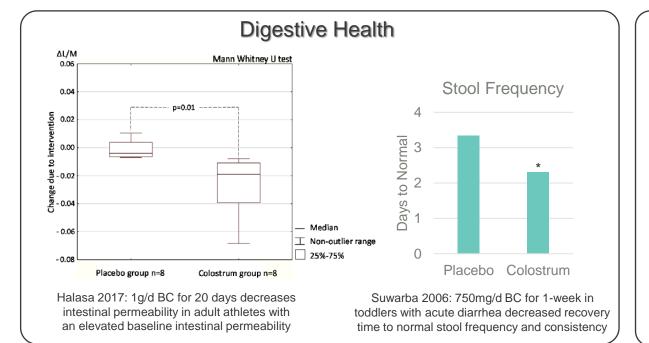


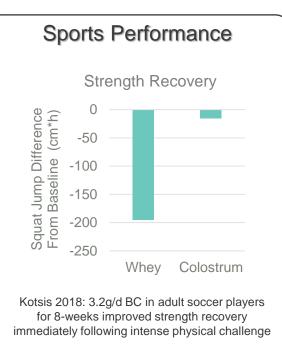


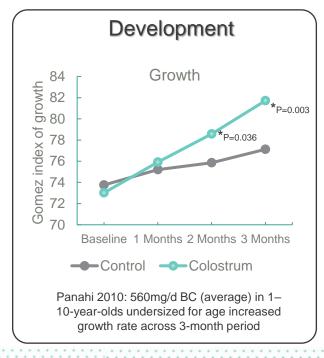
Clinical Dose Examples

Health Benefit	Adults	Children
Digestive Health	1 g	750 mg
Immune Health	400 mg	500 mg
Development (growth)	N/A	560 mg
Sports performance	3.2 g	N/A









Immune Health





Upper Respiratory Tract Infections (URTI)

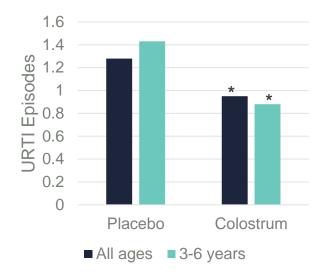
Study

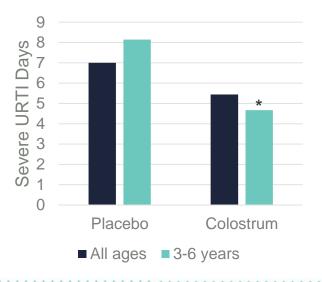
- Children 3 year to 9 years old
- 195 participants
- Normal healthy children
- Double-blind, placebo controlled randomized trial
- Milk powder placebo
- Received 500 mg bovine colostrum per day for 8 weeks

Results

Compared to placebo controls:

- After 8 weeks consuming bovine colostrum
 - Lower mean URTI episodes
 - Fewer days with severe URTI symptoms





^{*} Significant difference P<0.05 vs placebo



Upper Respiratory Tract Infections (URTI) and Diarrhea

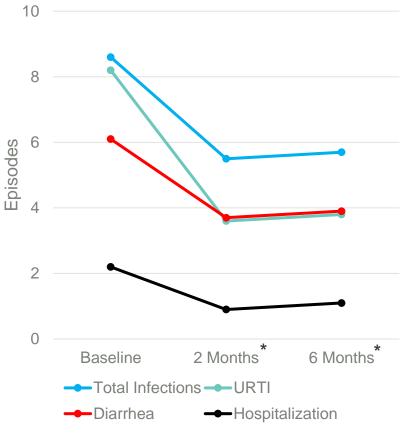
Study

- Children 1 year to 6 years old
- 160 participants
- Had recurrent URTI and/or diarrhea
- Open label, multicenter
- Received bovine colostrum 3g (<2 y) or 6g (>2 y) per day for 6 months

Results

Compared to intra-individual baseline:

- After 2 months consuming bovine colostrum
 - Lower total infections
 - Lower URTI episodes
 - Lower diarrhea episodes
 - Less hospitalizations
- After 6 months consuming bovine colostrum
 - All measures still significantly lower No regression to baseline levels



^{*} Significant difference P<0.05 vs baseline



Upper Respiratory Tract Infections (URTI) and Diarrhea

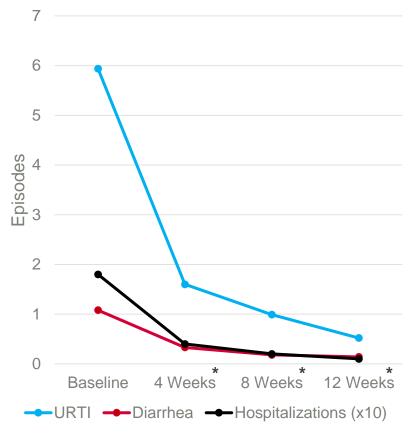
<u>Study</u>

- Children 1 year to 8 years old
- 605 participants
- Had recurrent URTI and/or diarrhea
- Open label, multicenter
- Received 3g bovine colostrum per day for 12 weeks

Results

Compared to intra-individual baseline:

- Reduction in number of URTI episodes
 - 73% by week 4
 - 83% by week 8
 - 92% by week 12
- Reduction in number of diarrhea episodes
 - 70% by week 4
 - 83% by week 8
 - 87% by week 12
- Reduction in number of hospitalizations
 - 76% by week 4
 - 89% by week 8
 - 92% by week 12



^{*} Significant difference P<0.05 vs baseline



Influenza

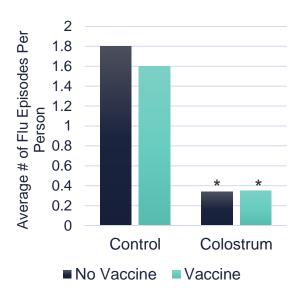
Study

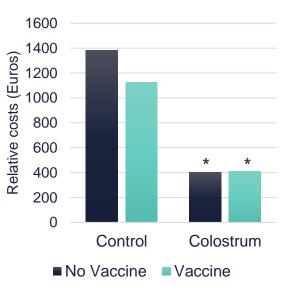
- Adults 30 to 80 years old
- 144 participants
- Normal healthy adults
- Participants placed into four groups:
 - · no vaccine and no colostrum
 - · no vaccine plus colostrum
 - vaccine but no colostrum
 - vaccine plus colostrum
- The colostrum groups consumed 400mg of bovine colostrum per day for 2 months

Results

Compared to control group

- After 2 months consuming bovine colostrum
 - Reduced number of flu episodes
 - Reduced relative costs
- Bovine colostrum was more effective than the influenza vaccine







RESEARCH ARTICLE

Open Access

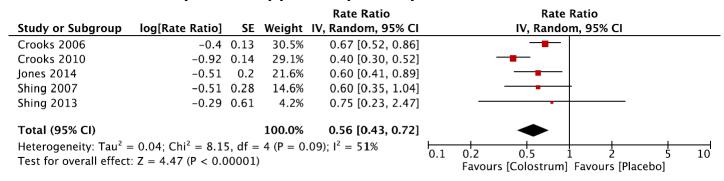


Bovine colostrum supplementation and upper respiratory symptoms during exercise training: a systematic review and meta-analysis of randomised controlled trials

Arwel W. Jones^{1*}, Daniel S. March², Ffion Curtis¹ and Christopher Bridle¹

- Meta-analysis of studies with adult participants undergoing exercise training
- Strict inclusion requirements, only 5 out of 22 eligible studies made the cut
- Exercise training included running, swimming, cycling, and general cardiovascular and weight training
- Dosing ranged from 10-20 grams of bovine colostrum, and length ranged 8-12 weeks long
- Pooled analyses revealed a significant decrease in days with URTI symptoms (P<0.00001) and number of URTI episodes (P<0.05)

Days with Upper Respiratory Tract Infection



Upper Respiratory Tract Infection Episodes

				Rate Ratio	Rate Ratio
Study or Subgroup	log[Rate Ratio]	SE	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Crooks 2006	-0.29	0.35	44.6%	0.75 [0.38, 1.49]	
Jones 2014	-0.56	0.37	39.9%	0.57 [0.28, 1.18]	
Shing 2007	-0.91	0.68	11.8%	0.40 [0.11, 1.53]	
Shing 2013	-0.29	1.22	3.7%	0.75 [0.07, 8.18]	
Total (95% CI)			100.0%	0.62 [0.39, 0.99]	•
Heterogeneity: Tau ² =	0.00; Chi ² = 0.70	6, df =	3 (P = 0)	.86); $I^2 = 0\%$	
Test for overall effect:	Z = 2.02 (P = 0.0)	04)			0.01 0.1 1 10 100 Favours [Colostrum] Favours [Placebo]

Jones et al. BMC Sports Science, Medicine and Rehabilitation (2016) 8:21

Digestive and Gut Health





Clinical Evidence: Gut Health

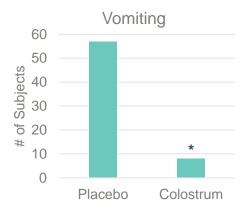
Acute Diarrhea

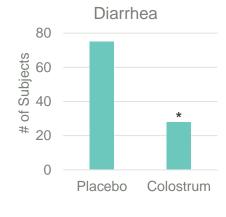
Study

- Infants and toddlers age 6 to 24 months
- 160 total subjects, 80 per group
- Double blind, randomized controlled trial
- Diagnosed with a case of acute diarrhea
- All received 3g per day of bovine colostrum for 1 week

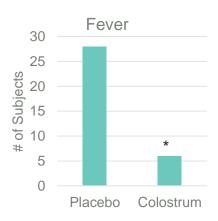
Results

- After 48 hours of bovine colostrum, the number of individuals with vomiting, diarrhea, and/or fever were significantly lower than the placebo group
- Over the 1-week trial, bovine colostrum significantly reduced the time to disappearance of all symptoms









*P<0.001 vs placebo

Barakat et al (2020). Journal of Tropical Pediatrics, 66(1):46-55.



Clinical Evidence: Gut Health

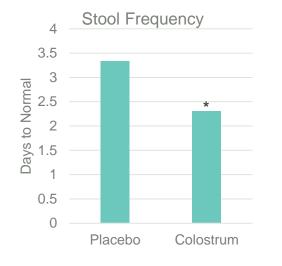
Acute Diarrhea

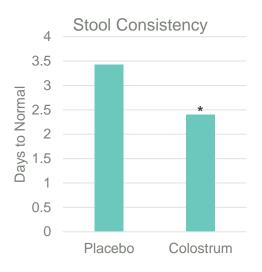
<u>Study</u>

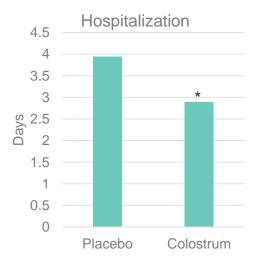
- Infants and toddlers age 6 to 24 months (mean 10.9)
- 70 total subjects, 35 per group
- Diagnosed with a case of acute mild-intermediate diarrhea
- · Double blind, randomized controlled trial
- All received 750 mg per day of bovine colostrum for 1 week

Results

- Infants and toddlers on bovine colostrum recovered significantly faster from acute diarrhea as measured by stool frequency and stool consistency compared to the placebo group
- Consumption of bovine colostrum significantly reduced the overall hospital stay length compared to the placebo group







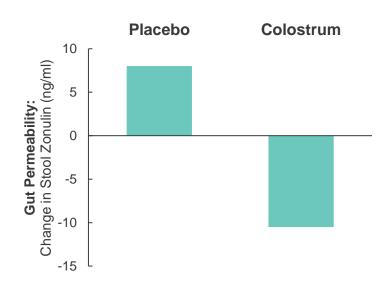
Suwarba et al (2006). Paediatrica Indonesiana, 46(5-6):127-133



Colostrum Gut Health Clinical Evidence Snapshot

Clinical Study:

- Healthy adult athletes
- 1g/d Colostrum for 20 days
- Reduced intestinal permeability reduced "leaky gut"



Halasa et al (2017). Nutrients, 9(4):370

Clinical Study:

- Healthy adult athletes
- 1g/d Colostrum for 20 days
- Reduced intestinal permeability only with first day milking Resulted in reduced "leaky gut"

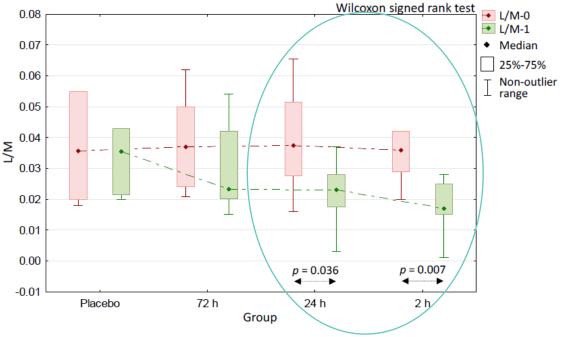


Figure 2. Comparison of lactulose/mannitol differential sugar absorption test results obtained before (L/M-0) and after (L/M-1) 20 days of supplementation with whey (placebo) and colostrum milked at 72 h, 24 h, and 2 h time points after delivery.

Halasa et al (2020). Medicina, 56, 495

Developmental Growth





Clinical Evidence: Developmental Growth

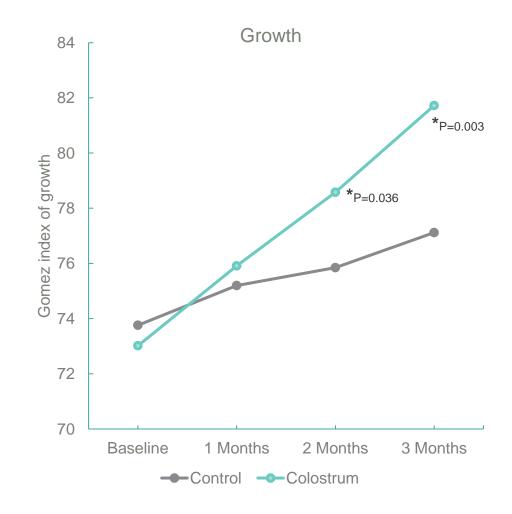
Study

- · Children 1 year to 10 years old
- 120 participants
- Mild or moderate unexplained failure to thrive
- 85-95% of 50th percentile for height
- 60-90% of 50th percentile for weight
- · Participant blinded, randomized trial
- Half received 40 mg bovine colostrum per kg body weight daily for 3 months

Results

Compared to control:

- After 2 months consuming bovine colostrum
 - Faster growth rate
- After 3 months consuming bovine colostrum
 - Faster growth rate



Panahi et al (2010). Journal of Pediatric Gastroenterology and Nutrition, 50(5):551-554

Sports Performance





Clinical Evidence: Sports Performance

First Author	Year	Population	Sample Size	Intervention	Duration	Main Outcomes
Kotsis	2018	Soccer players (4-5 sessions per week + one match)	Total: 22 Control: 11 Colostrum: 11	Control: 3.2g of WPC Tx: 3.2g BC Specific training program plan 3-days preand 4-days post-LIST	6 weeks	Squat jump height: BC > CON Countermovement jump height: BC > CON CRP: BC <con, bc="" ii-6:=""> CON</con,>
Antonio	2001	18-35 yo males and females Already on resistance training min 3x per week	Total: 20 Control: 11 Colostrum: 9	Control: 20g of WPC Tx: 20g BC	8 weeks	Lean body mass: BC > BC baseline (2.3%, 1.5kg)
Buckley	2002	18-35 yo males Already participating in regular physical activity	Total: 30 Control: 13 Colostrum: 17	Control: 60g of WPC Tx: 60g BC 8 weeks of endurance running Nutrition plan	8 weeks	Second bout running performance: BC > CON (+0.3 mph)
Buckley	2003	18-35 yo males Already doing regular physical activity	Total: 51 Control: 26 Colostrum: 25	Control: 60g of WPC Tx: 60g BC 8 weeks resistance and plyometric training	8 weeks	Peak anerobic cycling power: BC > CON Max vertical jump: BC > CON
Duff	2014	Males (59.1 +/- 5.4y) and Females (59.0 +/-6.7y)	Total: 40 Control: 21 Colostrum: 19	Control: 60g of WPC Tx: 60g BC 8 week resistance training program	8 weeks	Leg press 1-RM strength: BC > CON (21% vs 5% improvement) Bone resorption marker (N-telopeptides): BC < CON
Jones	2016	Meta-analysis and systematic review Clinical studies on atheletes with UTRI endpoints	5 studies met inclusion criteria	10g to 20g of BC	8, 10, or 12 weeks	URS days: BC < CON URS episodes: BC < CON

