





LMKaplan0@gmail.com

## **The Microbiome and Obesity Management**

## Lee M. Kaplan, MD, PhD

Professor of Medicine and Chief, Section of Obesity Medicine Director, Dartmouth Weight and Wellness Center Geisel School of Medicine at Dartmouth

LMKaplan0@gmail.com

4 September 2024

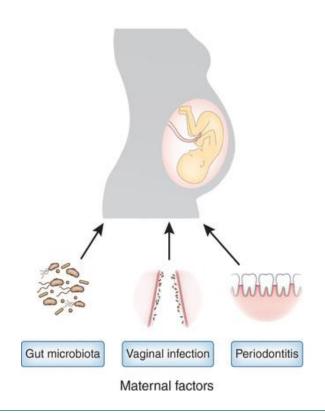


IFSO 2024 • Melbourne, Victoria, Australia

- Gut microbiota is composed of bacteria, viruses, archaea, and fungi
- Can be considered an organ that resides in the lumen of the gut
- Contains 150-fold more genetic information than the human genome

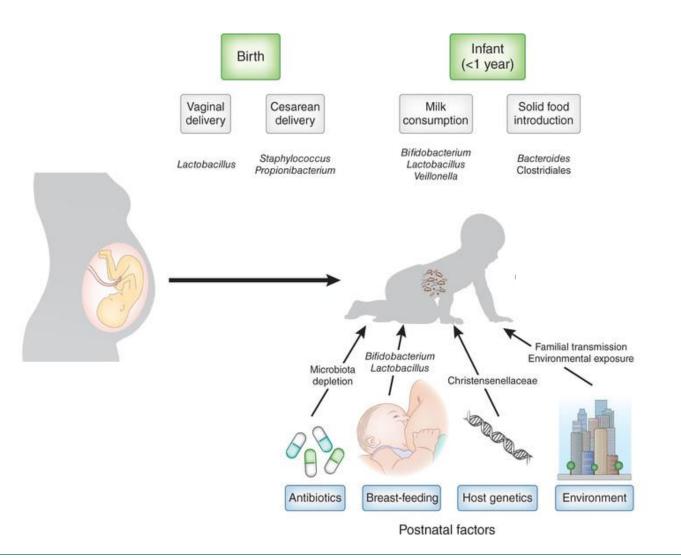






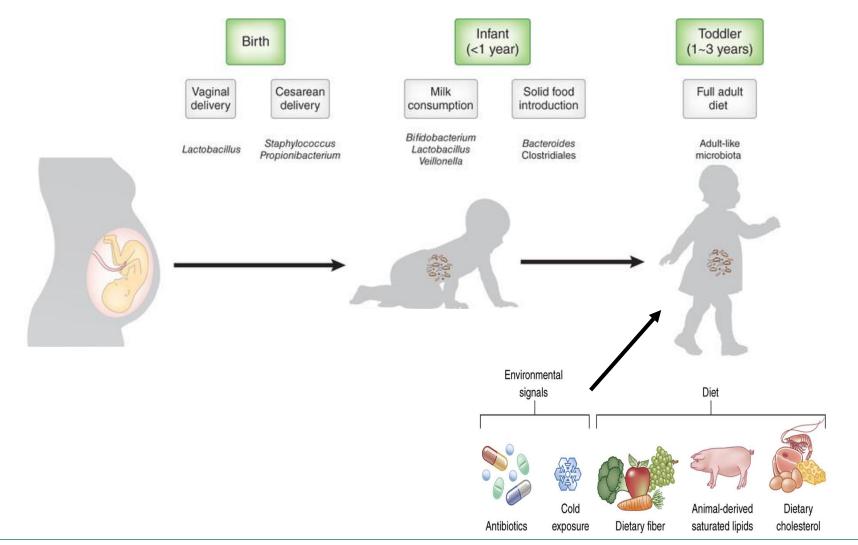
Shoeder and Backhed. (2016) Nature Medicine, Tamburini et al. (2016) Nature Med





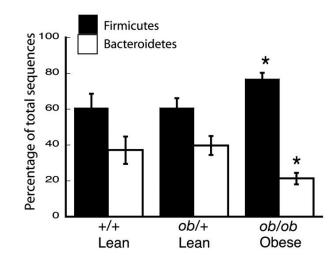
Shoeder and Backhed. (2016) Nature Medicine, Tamburini et al. (2016) Nature Med





Shoeder and Backhed. (2016) Nature Medicine, Tamburini et al. (2016) Nature Med

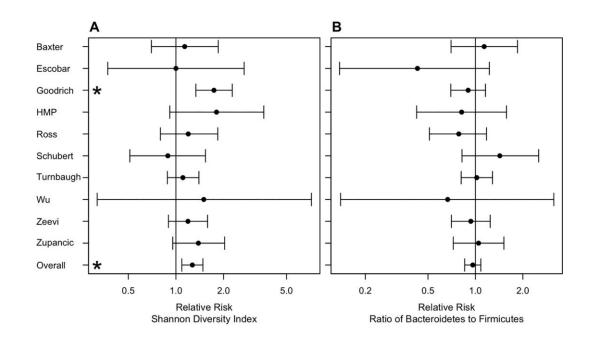




- The first studies showed a prominent bacterial signature of obesity in mice
- Some human studies recapitulated this signature

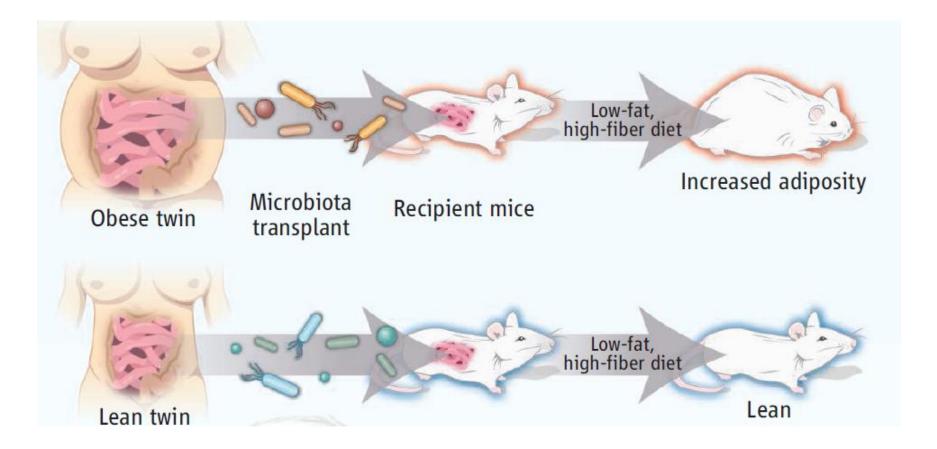






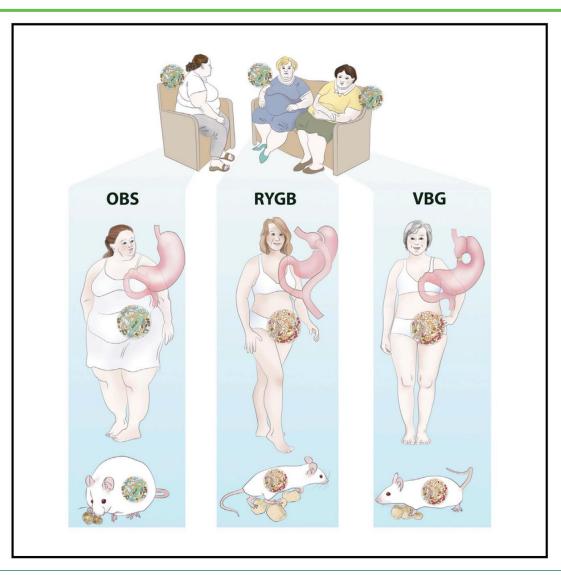
- The first studies showed a prominent bacterial signature of obesity in mice
- Some human studies recapitulated this signature
- Large meta-analysis showed there was no clear taxonomic obesity signature





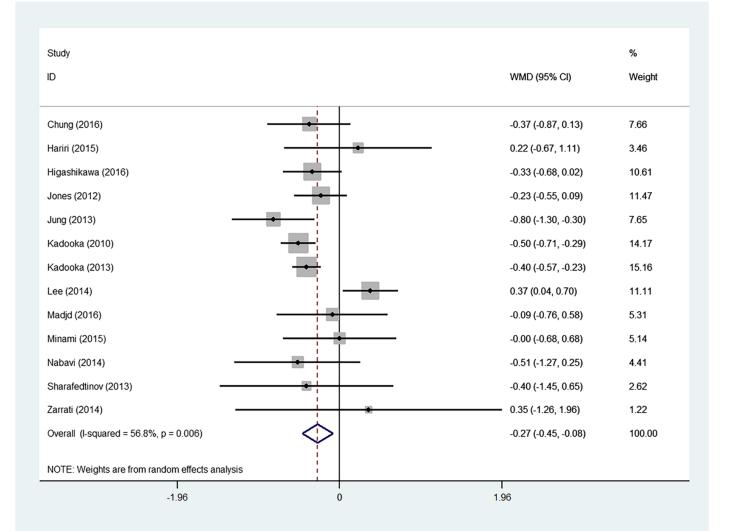
Ridaura VK et al., Science 2013, Walker AW et al., 2013, Science 2013

Bartmouth GEISEL SCHOOL OF MEDICINE



Tremaroli, V et al., Cell Metabolism 2015





Borgeraas et al. (2017) Obesity Reviews, Wiciński et al. (2020) Microorganisms





- Very few safety problems that have been identified with pro- or prebiotics
- May be some weight independent beneficial effects, but more work needs to be done
- Weight effects have very small effect size (~1.3 lbs weight difference)



- Many different strains and compounds are used
- Most probiotic species are not native to the human intestine (aerobic)
- We do not know how well they "engraft" into or change the ecosystem
- Very few metabolic clinical benefits have been seen



GASTROENTEROLOGY 2012;143:913-916

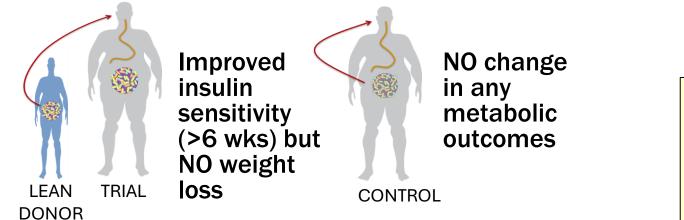
**BRIEF REPORT** 

### **BRIEF REPORT**

#### Transfer of Intestinal Microbiota From Lean Donors Increases Insulin Sensitivity in Individuals With Metabolic Syndrome

ANNE VRIEZE,\* ELS VAN NOOD,\* FRITS HOLLEMAN,\* JARKKO SALOJÄRVI,<sup>‡</sup> RUUD S. KOOTTE,<sup>§</sup> JOEP F. W. M. BARTELSMAN,<sup>∥</sup> GEESJE M. DALLINGA-THIE,<sup>§</sup> MARIETTE T. ACKERMANS,<sup>¶</sup> MIREILLE J. SERLIE,<sup>#</sup> RAISH OOZEER,\*\* MURIEL DERRIEN,\*\* ANNE DRUESNE,\*\* JOHAN E. T. VAN HYLCKAMA VLIEG,\*\* VINCENT W. BLOKS,<sup>‡‡</sup> ALBERT K. GROEN,<sup>‡‡</sup> HANS G. H. J. HEILIG,<sup>§§</sup> ERWIN G. ZOETENDAL,<sup>§§</sup> ERIK S. STROES,<sup>§</sup> WILLEM M. DE VOS,<sup>‡,§§</sup> JOOST B. L. HOEKSTRA,\* and MAX NIEUWDORP\*.<sup>§</sup>





#### **Exciting initial observation**

... but multiple follow-up human studies have NOT been able to reproduce this result



#### Vrieze A et al., *Gastroenterology*, 2012 Copyright © 2024 The Obesity and Metabolism Institute. All rights reserved.



#### RESEARCH ARTICLE

Fecal microbiota transplantation for the improvement of metabolism in obesity: The FMT-TRIM double-blind placebo-controlled pilot trial

Elaine W. Yu<sup>1,2</sup>\*, Liu Gao<sup>1</sup>, Petr Stastka<sup>1</sup>, Michael C. Cheney<sup>1</sup>, Jasmin Mahabamunuge<sup>3</sup>, Mariam Torres Soto<sup>3</sup>, Christopher B. Ford<sup>4</sup>, Jessica A. Bryant<sup>4</sup>, Matthew R. Henn<sup>4</sup>, Elizabeth L. Hohmann<sup>2,3</sup>

**Clinical Gastroenterology and Hepatology 2019** 

# **Effects of Fecal Microbiota Transplantation With Oral Capsules in Obese Patients**

Jessica R. Allegretti,\* Zain Kassam,<sup>‡</sup> Benjamin H. Mullish,<sup>§</sup> Austin Chiang,<sup>∥</sup> Madeline Carrellas,\* Jonathan Hurtado,\* Julian R. Marchesi,<sup>§</sup> Julie A. K. McDonald,<sup>§</sup> Alexandros Pechlivanis,<sup>§</sup> Grace F. Barker,<sup>§</sup> Jesus Miguens Blanco,<sup>§</sup> Isabel Garcia-Perez,<sup>§</sup> Wing Fei Wong,<sup>¶</sup> Ylaine Gerardin,<sup>‡</sup> Michael Silverstein,<sup>‡</sup> Kevin Kennedy,<sup>#</sup> and Christopher Thompson\*

#### **Conclusion:**

NO significant effect of microbiota Transplantation on body weight



## Gut microbiota in the clinic tomorrow

Diseases of the Microbiota

Diseases Directly Affected by the Microbiota

Diseases Affected by Pathways Regulated by Microbiota

Diseases Apparently Influenced by the Microbiota





Irritable Bowel Syndrome Cognitive Disorders Depression









LMKaplan0@gmail.com

## **The Microbiome and Obesity Management**

## Lee M. Kaplan, MD, PhD

Professor of Medicine and Chief, Section of Obesity Medicine Director, Dartmouth Weight and Wellness Center Geisel School of Medicine at Dartmouth

LMKaplan0@gmail.com

4 September 2024



IFSO 2024 • Melbourne, Victoria, Australia