

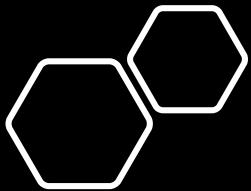
# Persoonallisuuden piirteet bakteereissani

Anna Aatsinki, LT,

tutkijatohtori

FinnBrain-tutkimus,

Turun Yliopisto

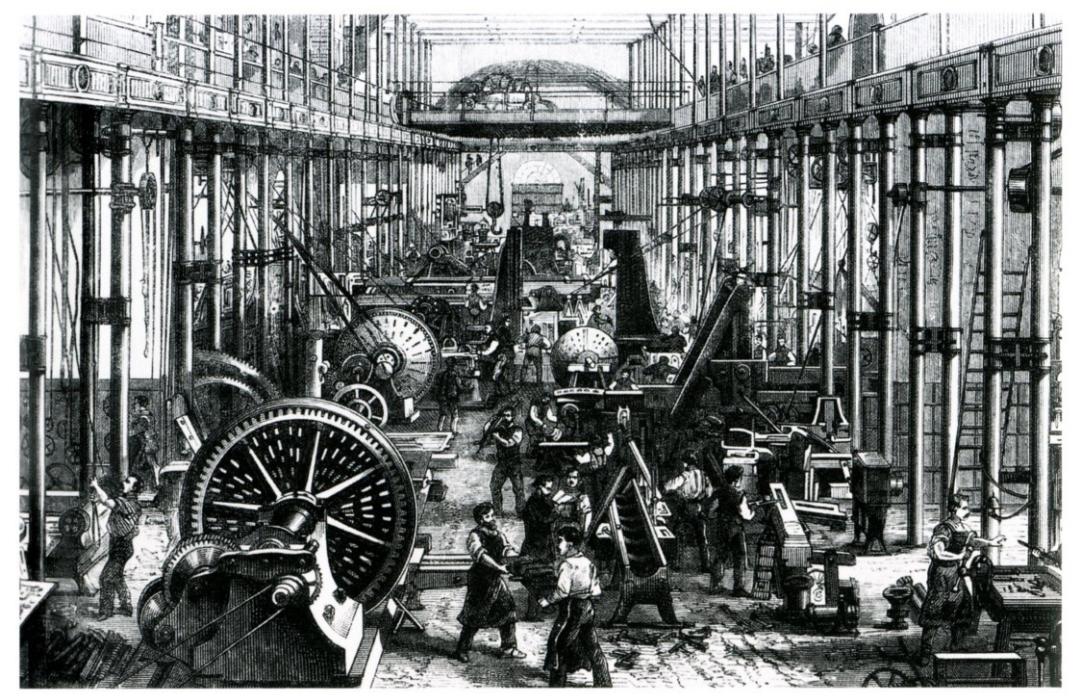


# Esityksen sisältö

- Suoli-aivoakselin historiaa
- Suoli-aivo –akseli eläimillä ja sen mekanismit
- Havaintoja ihmisillä
- Esimerkkejä “interventioista”



# Historian havinaa



March 12, 1898

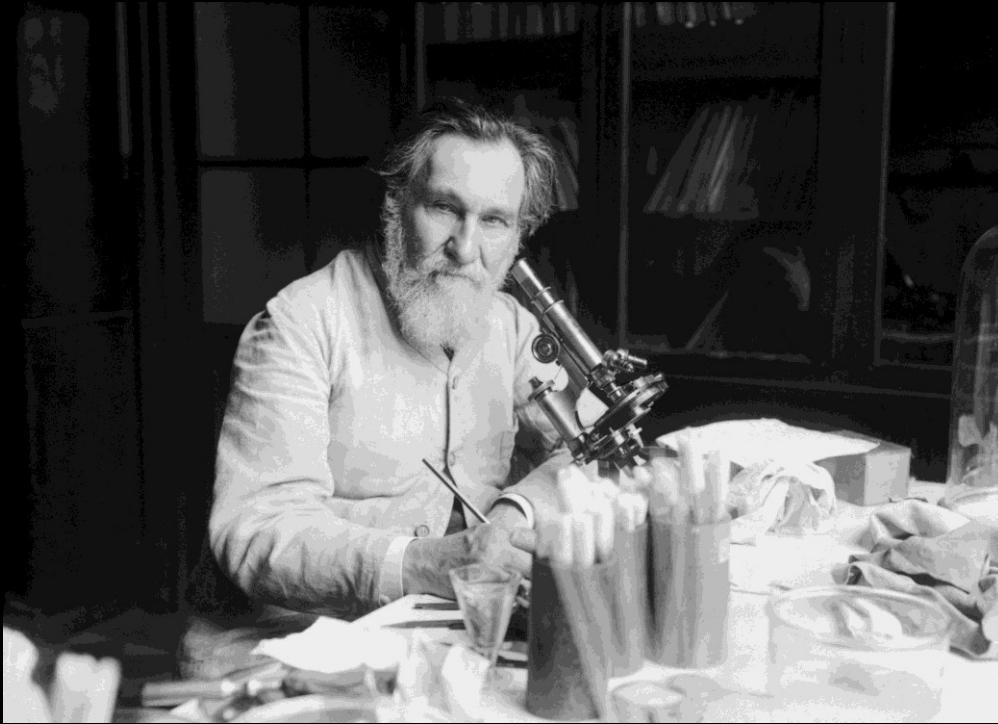
## AUTO-INTOXICATION IN ITS RELATIONS TO THE DISEASES OF THE NERVOUS SYSTEM.

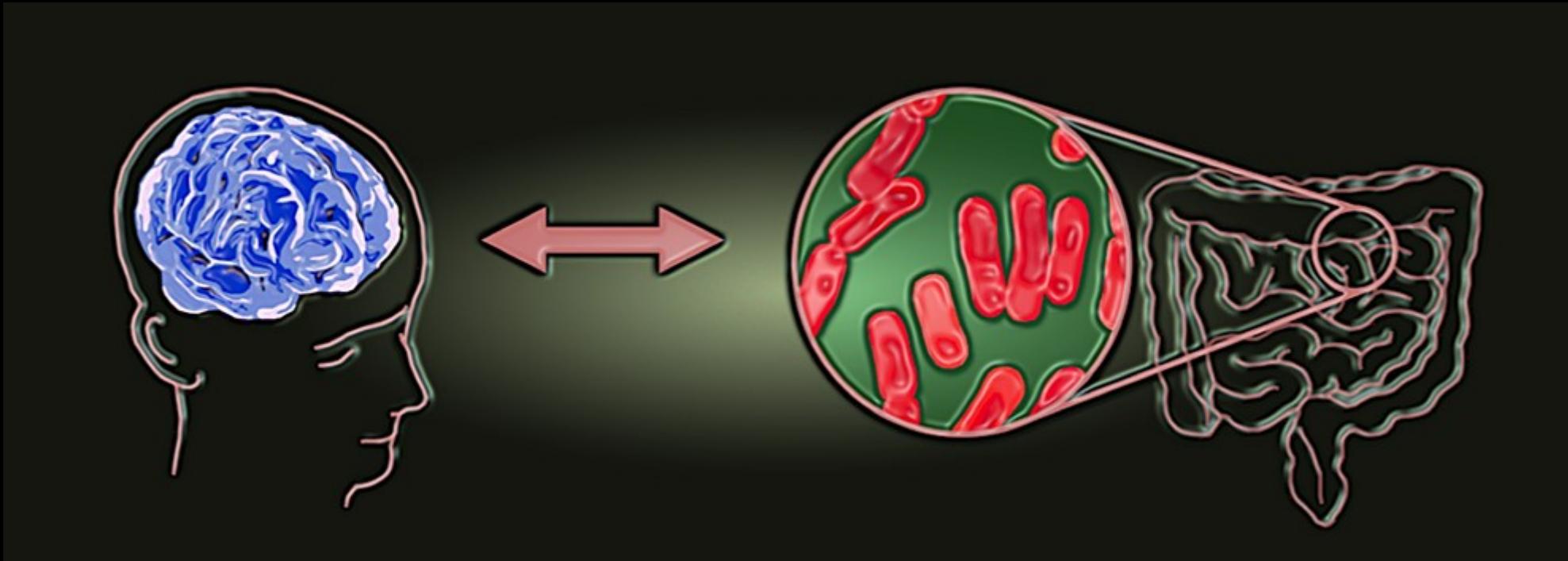
DANIEL R. BROWER, M.A., M.D.

» [Author Affiliations](#)

JAMA. 1898;XXX(11):575-577. doi:10.1001/jama.1898.72440630001001

# Historian havinaa





Original Articles | Published: February 1986

## Variations of brain histamine levels in germ-free and nephrectomized rats

Linda R. Hegstrand & R. Jean Hine

*Neurochemical Research* 11, 185–191 (1986) | Cite this article

85 Accesses | 21 Citations | 3 Altmetric | Metrics

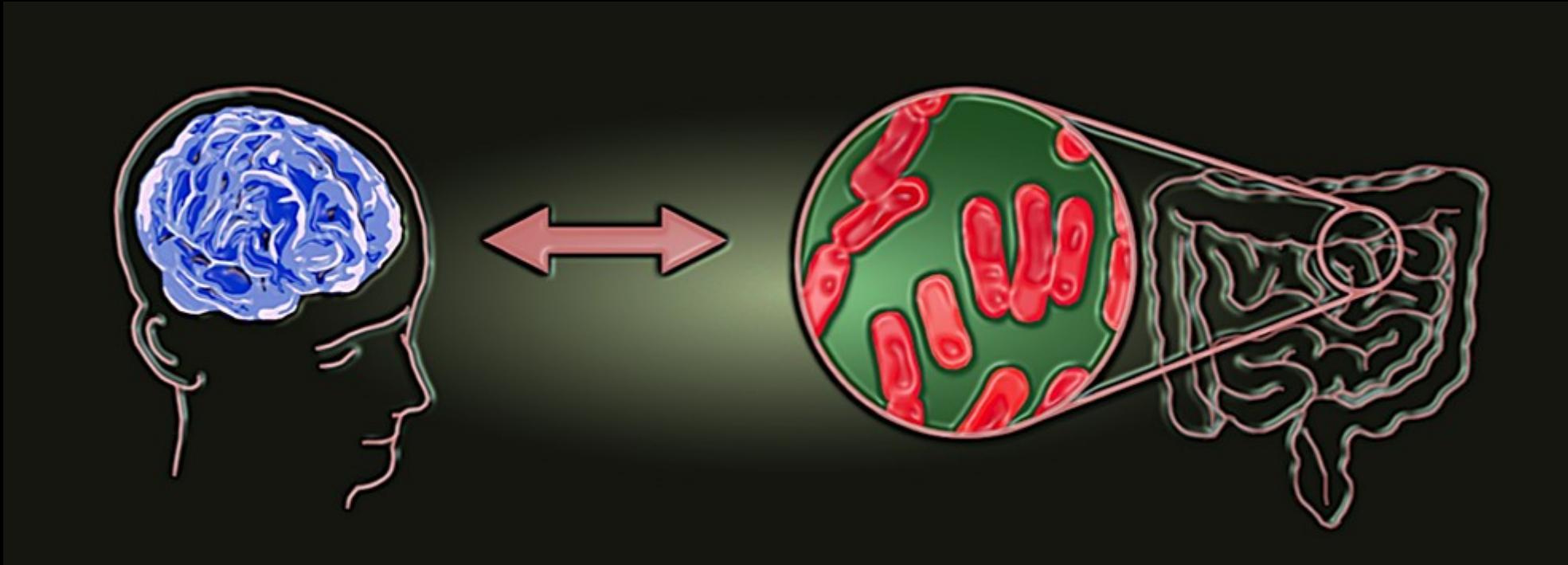
## ARTICLES

### BIOLOGICAL FREUDIANISM Lasting Effects of Early Environmental Influences

René Dubos, Ph.D. (M.D. Hon.), Dwayne Savage, Ph.D., and Russell Schaedler, M.D.

The Rockefeller University, New York

PEDIATRICS, Vol. 38, No. 5, November 1966



## Microbiota-Gut-Brain axis

=

Bidirectional communication between gut microbiota and central nervous system



Havaintoja  
eläinmalleissa

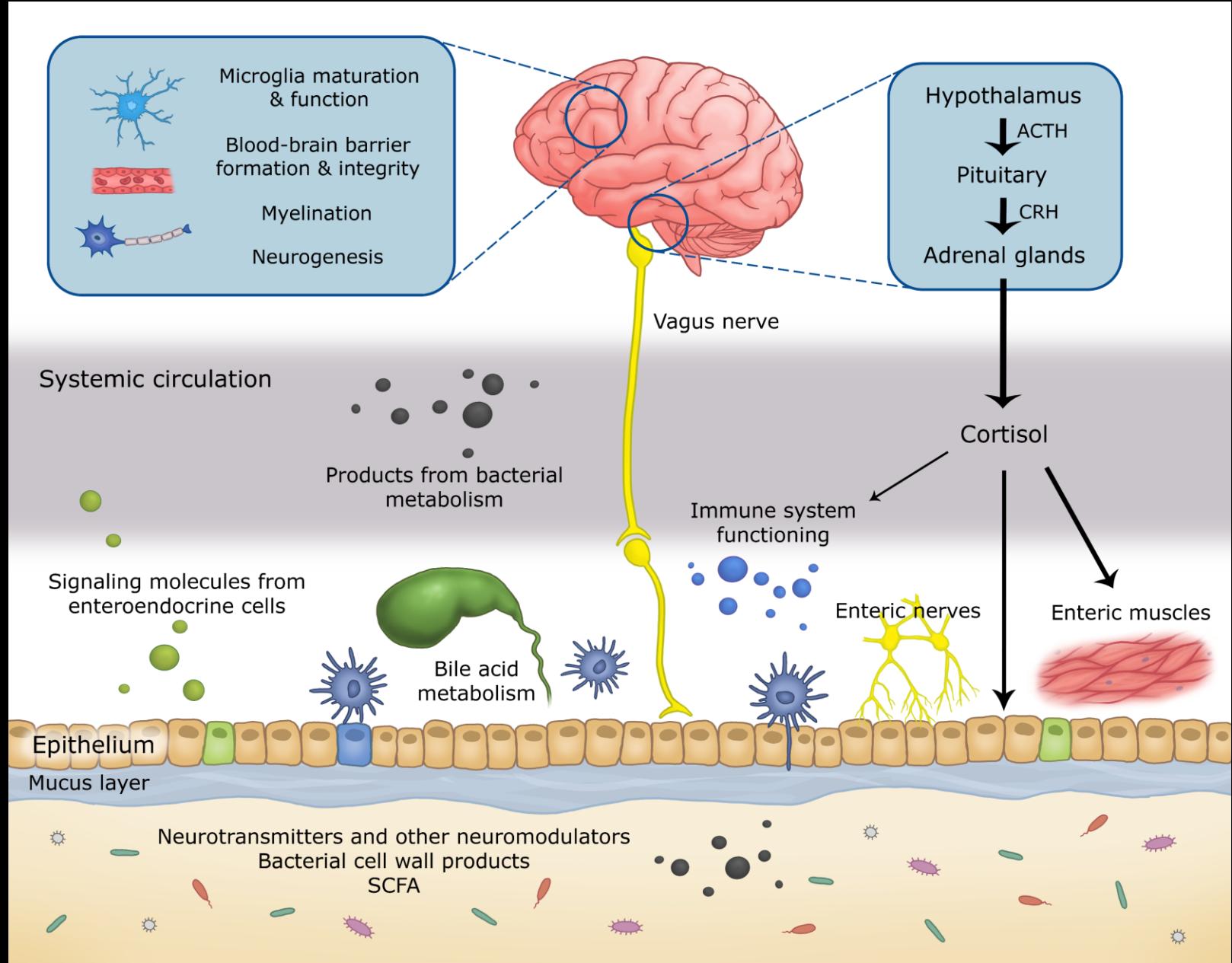
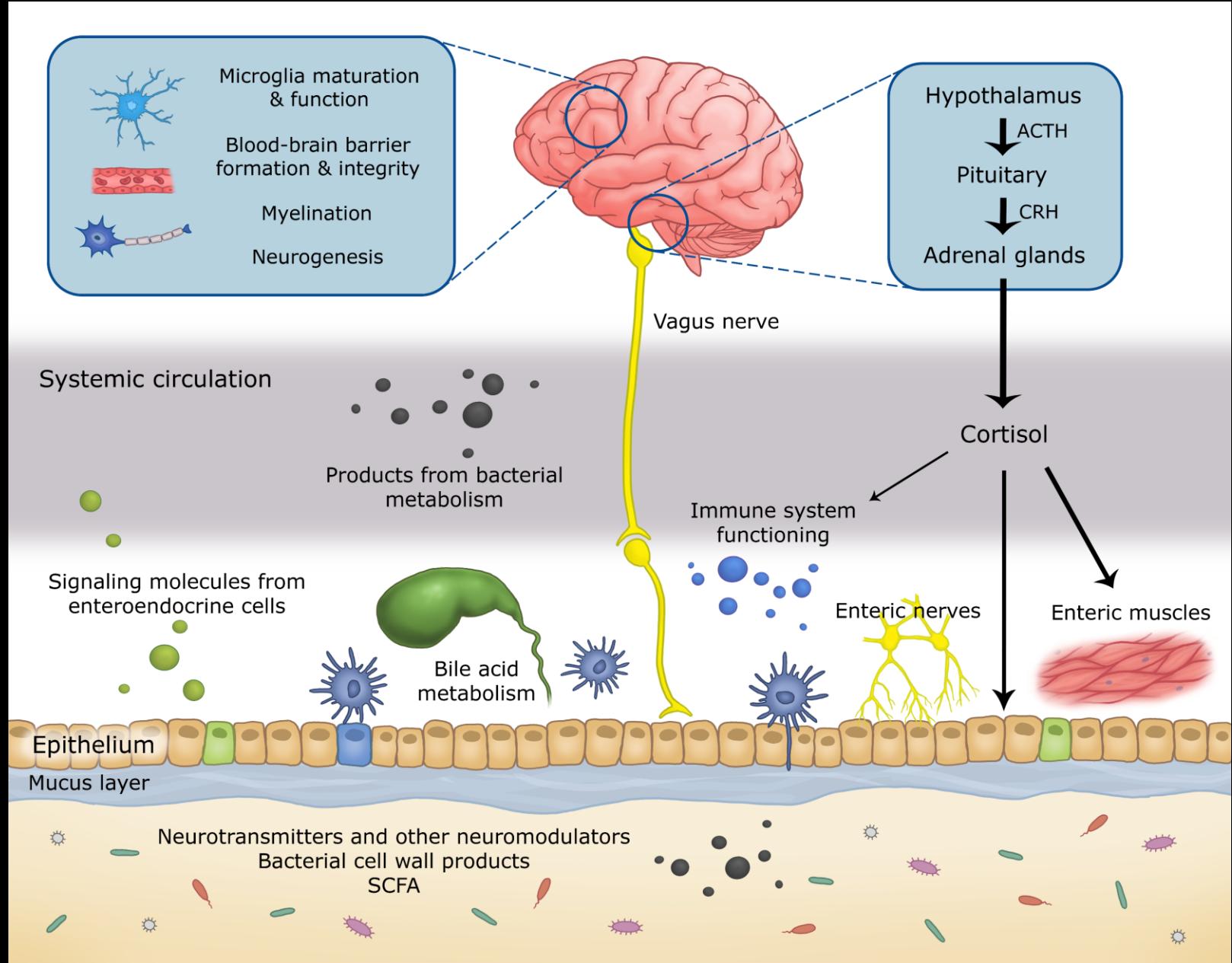


Image: Heidi Huhtanen

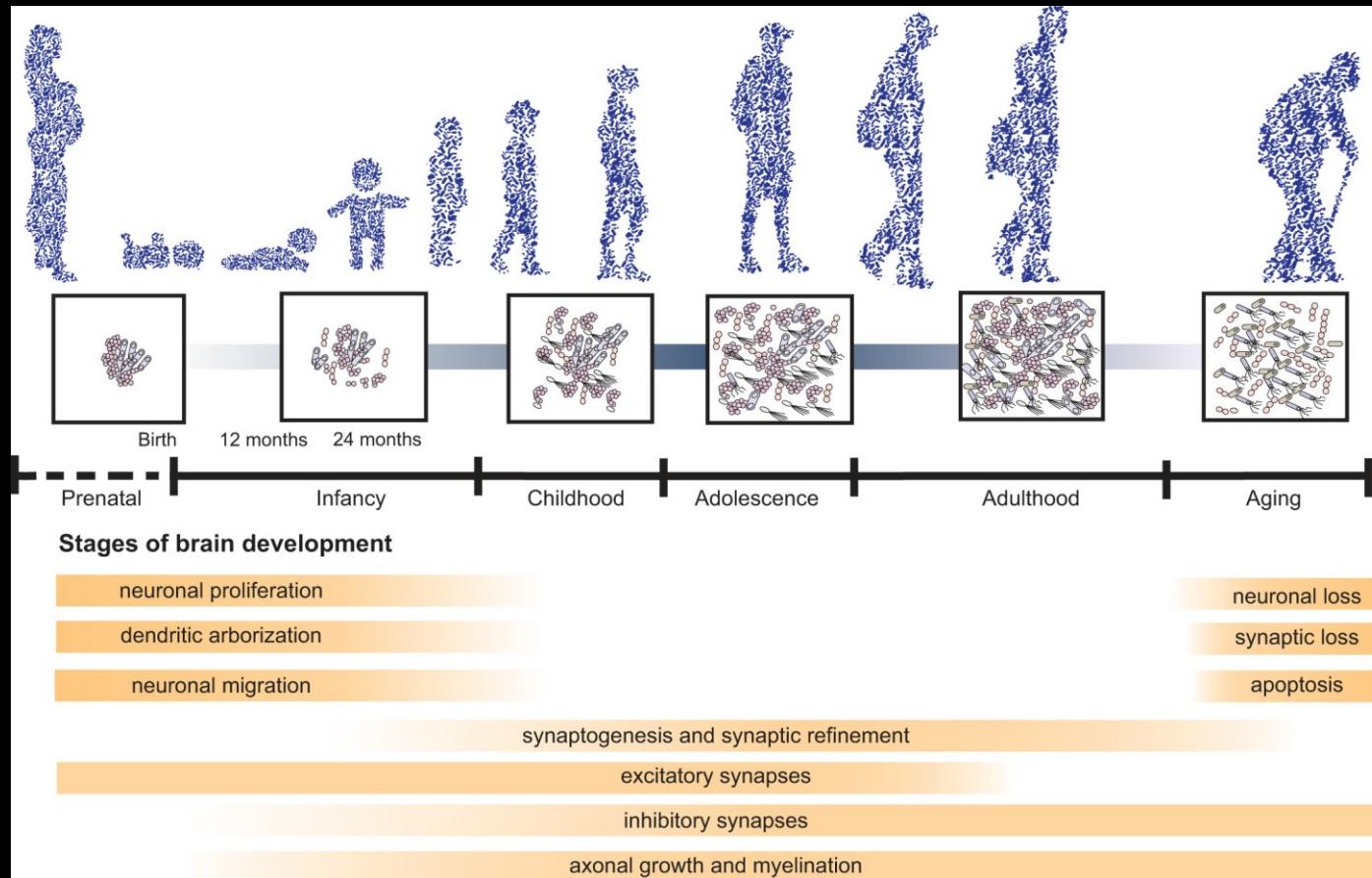


## Phenotypes affected:

- Social behavior
- Cognition
- Fear regulation
- Stress-responsivity
- Food intake and appetite

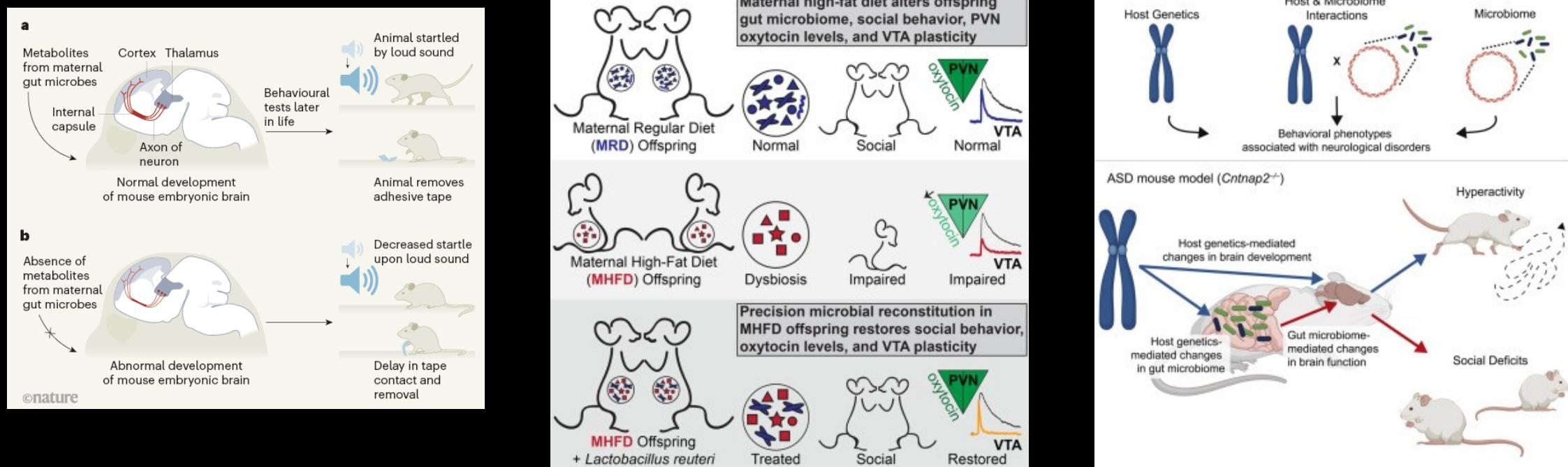
Cryan et al. 2019  
doi:10.1152/physrev.00018.  
2018

# Mikrobisto elämänkaaren aikana



DOI: (10.1152/physrev.00018.2018)

# Varhainen mikrobisto ja sosiaaliset aivot



→ Varhainen kontakti mikrobeihin tai niiden aineenvaihduntatuotteisiin ohjelmoi käyttäytymisen kehitystä yhdessä perimän kanssa

Ref: Vuong, 2020, Nature; Meckel, Kiraly, 2020, Nature; Buffington et al. 2016, Cell; Buffington et al., Cell, 2021; Garcias, 2016 eLife

# Varhainen mikrobisto ja kehitys

ARTICLE IN PRESS



## Archival Report

### Infant Gut Microbiome Associated With Cognitive Development

Alexander L. Carlson, Kai Xia, M. Andrea Azcarate-Peril, Barbara D. Goldman, Miye Ahn, Martin A. Styner, Amanda L. Thompson, Xiujuan Geng, John H. Gilmore, and Rebecca C. Knickmeyer



Original Investigation | Pediatrics

### Association of the Infant Gut Microbiome With Early Childhood Neurodevelopmental Outcomes An Ancillary Study to the VDAART Randomized Clinical Trial

Joanne E. Sordillo, ScD; Susan Korrick, MD; Nancy Laranjo, BA; Vincent Carey, PhD; George M. Weinstock, PhD; Diane R. Gold, MD; George O'Connor, MD; Megan Sandel, MD; Leonard B. Bacharier, MD; Avraham Beigelman, MD; Robert Zeiger, MD, PhD; Augusto A. Litonjua, MD; Scott T. Weiss, MD

### Gut microbiome and brain functional connectivity in infants-a preliminary study focusing on the amygdala

Wei Gao<sup>1,2</sup> · Andrew P. Salzwedel<sup>1</sup> · Alexander L. Carlson<sup>3</sup> · Kai Xia<sup>4</sup> · M. Andrea Azcarate-Peril<sup>5,6</sup> · Martin A. Styner<sup>4,7</sup> · Amanda L. Thompson<sup>8,9</sup> · Xiujuan Geng<sup>10</sup> · Barbara D. Goldman<sup>11,12</sup> · John H. Gilmore<sup>4</sup> · Rebecca C. Knickmeyer<sup>4,13,14</sup>

Received: 22 September 2018 / Accepted: 21 December 2018

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Research paper

### Maternal prenatal gut microbiota composition predicts child behaviour

Samantha L. Dawson<sup>a, b</sup>, Martin O'Hely<sup>a, b</sup>, Felice N. Jacka<sup>a, b, h</sup>, Anne-Louise Ponsonby<sup>c, d</sup>, Christos Symeonides<sup>b, d</sup>, Amy Loughman<sup>a</sup>, Fiona Collier<sup>a, b, f</sup>, Margarita Moreno-Betancur<sup>b, d</sup>, Peter Sly<sup>b, g</sup>, David Burgner<sup>b, d, e</sup>, Mimi L.K. Tang<sup>b, d, e</sup>, Richard Saffery<sup>b, d</sup>, Sarah Ranganathan<sup>b, d, e</sup>, Michael A. Conlon<sup>i</sup>, Leonard C Harrison<sup>d, j</sup>, Susanne Brix<sup>k</sup>, Karsten Kristiansen<sup>l</sup>, Peter Vuillermin<sup>a, b, f</sup> , the BIS Investigator Group

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ISSN: 1528-3542

<https://doi.org/10.1037/emo00009>

Research paper

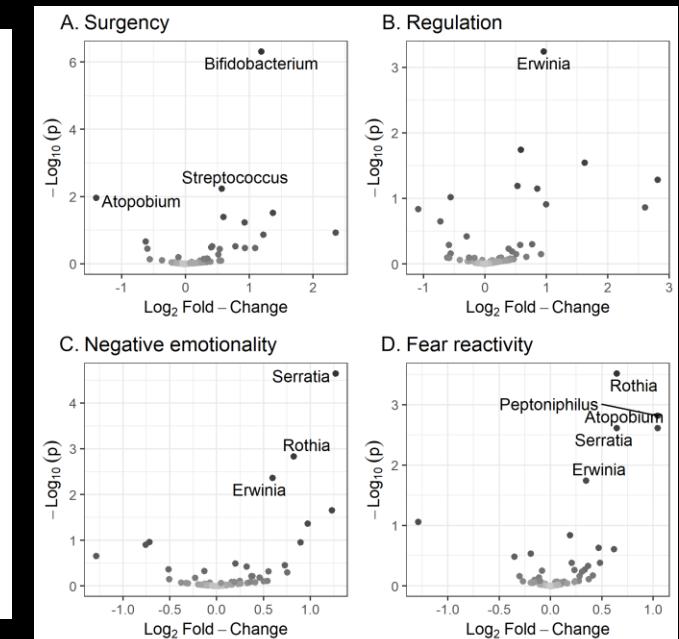
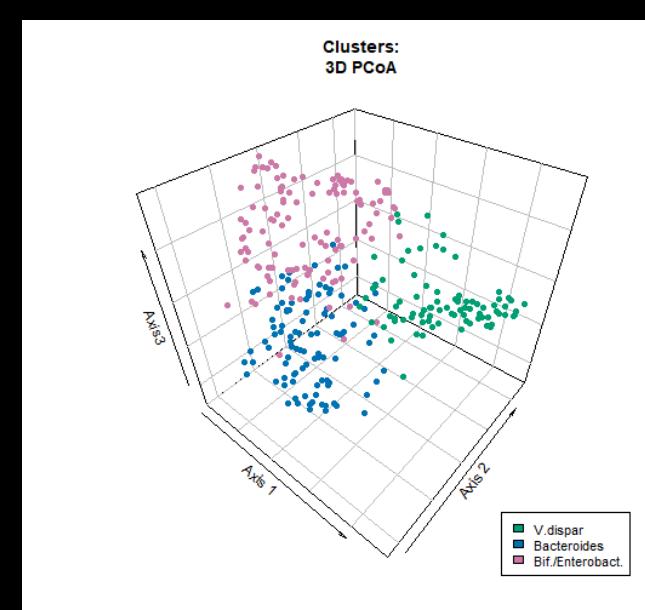
### Gut microbiota composition during infancy and subsequent behavioural outcomes

Amy Loughman<sup>a</sup>, Anne-Louise Ponsonby<sup>b, h</sup>, Martin O'Hely<sup>a, b</sup>, Christos Symeonides<sup>b, d, e</sup>, Fiona Collier<sup>a, b, c</sup>, Mimi L.K. Tang<sup>b, d, e</sup>, John Carlin<sup>b, d</sup>, Sarah Ranganathan<sup>b, d, e</sup>, Katrina Allen<sup>b, d, e</sup>, Angela Pezic<sup>b</sup>, Richard Saffery<sup>b, d</sup>, Felice Jacka<sup>a, b</sup>, Leonard C. Harrison<sup>d, f</sup>, Peter D. Sly<sup>b, g</sup>, Peter Vuillermin<sup>a, b, c, \*</sup>, the BIS Investigator Group

### Infant Fecal Microbiota Composition and Attention to Emotional Faces

Anna-Katarina Aatsinki<sup>1</sup>, Eeva-Leena Kataja<sup>1, 2</sup>, Eveliina Munukka<sup>3</sup>, Leo Lahti<sup>4</sup>, Anniina Keskitalo<sup>1, 5</sup>, Riikka Korja<sup>2, 1</sup>, Saara Nolvi<sup>6, 1, 7</sup>, Tuomo Häkiö<sup>2</sup>, Saija Tarro<sup>1</sup>, Hasse Karlsson<sup>1, 8, 9, 10</sup>, and Linnea Karlsson<sup>1, 8, 9, 11</sup>

# Varhainen mikrobisto ja temperamentti



Infant behavioral questionnaire revised (Rothart et al. 1981) at 6 mo, n=301

Aatsinki et al., Brain, Behavior, and Immunity, 2019

Gut microbiota composition is associated with newborn functional brain connectivity and behavioral temperament

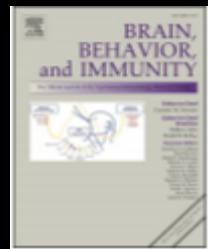
Caroline M. Kelsey <sup>a,b,c,\*</sup>, Stephanie Prescott <sup>d,e</sup>, John A. McCulloch <sup>d</sup>, Giorgio Trinchieri <sup>d</sup>, Tara L. Valladares <sup>a</sup>, Caitlin Dreisbach <sup>f,g</sup>, Jeanne Alhusen <sup>f</sup>, Tobias Grossmann <sup>a,b,\*</sup>

Gut microbiome composition is associated with temperament during early childhood

Lisa M. Christian <sup>a, b, c, d, e</sup>, Jeffrey D. Galley <sup>b, e</sup>, Erin M. Hade <sup>f</sup>, Sarah Schoppe-Sullivan <sup>e</sup>, Claire Kamp Dush <sup>g</sup>, Michael T. Bailey <sup>b, e</sup>

Gut microbiota composition is associated with temperament traits in infants

Anna-Katriina Aatsinki <sup>a</sup>, Leo Lahti <sup>b</sup>, Henna-Maria Uusitupa <sup>a</sup>, Eveliina Munukka <sup>c, d</sup>, Anniina Keskkitalo <sup>d, e</sup>, Saara Nolvi <sup>a</sup>, Siobhain O'Mahony <sup>f</sup>, Sami Pietilä <sup>a</sup>, Laura L. Elo <sup>f</sup>, Erkki Eerola <sup>a</sup>, Hasse Karlsson <sup>a</sup>, Linnea Karlsson <sup>a, i</sup>



## Association between Gut Microbiota and Infant's Temperament in the First Year of Life in a Chinese Birth Cohort



Ying Wang <sup>1,†</sup>, Xiaoli Chen <sup>1,†</sup>, Yun Yu <sup>1</sup>, Yanqun Liu <sup>1,\*</sup>, Qing Zhang <sup>1,\*</sup> and Jinbing Bai <sup>2</sup>

Development of the infant gut microbiome predicts temperament across the first year of life

Molly Fox<sup>1,2</sup>, S. Melanie Lee<sup>2,3</sup>, Kyle S. Wiley<sup>1,2</sup>, Venu Lagishetty<sup>3,4,5</sup>, Curt A. Sandman<sup>6</sup>, Jonathan P. Jacobs<sup>3,4,5</sup> and Laura M. Glynn<sup>7</sup>

OPEN ACCESS | Research Article | 21 January 2020



Gut Feelings Begin in Childhood: the Gut Metagenome Correlates with Early Environment, Caregiving, and Behavior

Authors: Jessica E. Flannery, Keaton Stagaman <sup>1,2</sup>, Adam P. Burns <sup>1,2</sup>, Roxana J. Hickey, Leslie E. Roos, Ryan J. Giuliano, Philip A. Fisher, and Thomas J. Sharpton | AUTHORS INFO & AFFILIATIONS



# Temperament and gut microbiota

## Positive emotionality

- Bifidobacterium + positive emotionality (Fox 2021)
- Klebsiella – positive emotionality (Fox 2021)
- Lachnospiraceae + positive emotionality (Fox 2021)
- Bacteroides fragilis – impulsivity (Flannery 2020)

## Self-regulation

- Bifidobacterium + Soothability (Wang 2020)
- Hungatella + Cuddliness (Wang 2020)
- Bifidobacterium adolescentis – inhibitory control (Flannery 2020)
- Bacteroides fragilis + inhibitory control (Flannery 2020)
- Eubacterium siraeum + impulsivity (Flannery 2020)

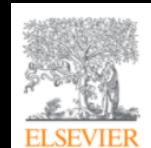
## Negative emotionality

- Ruminococcus + negative emotionality (Fox 2021)
- Lactobacillus – negative emotionality (Fox 2021)
- Parabacteroides distasonis + fear (Flannery 2020)
- Bilophilia + fear (Flannery 2020)
- Bacteroides fragilis – sadness (Flannery 2020)

- Impulsivity + tryptophan metabolism (Flannery 2020)
- Fear + melatonin biosynthesis (Flannery 2020)



# Aikuisilla



Brain, Behavior, and Immunity

Volume 69, March 2018, Pages 374-385

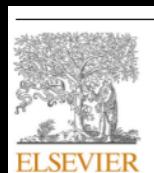


Full-length Article

Correlation between gut microbiota and personality in adults: A cross-sectional study

Han-Na Kim <sup>a</sup>, Yeojun Yun <sup>a</sup>, Seungho Ryu <sup>b, c</sup>, Yoosoo Chang <sup>b, c</sup>, Min-Jung Kwon <sup>b, d</sup>, Juhee Cho <sup>b, e, g</sup>, Hocheol Shin <sup>b, f</sup>, Hyung-Lae Kim <sup>a</sup>✉

- Neuroottisuus pos. yhteydessä Gammaproteobakteerien pitoisuuteen



Contents lists available at ScienceDirect

Human Microbiome Journal

journal homepage: [www.sciencedirect.com/journal/human-microbiome-journal](http://www.sciencedirect.com/journal/human-microbiome-journal)



Original Article

Gut microbiome composition and diversity are related to human personality traits

Katerina V.-A. Johnson

University of Oxford, Department of Experimental Psychology, New Radcliffe Building, Radcliffe Observatory Quarter, Oxford OX2 6GG, United Kingdom



# Intervention paikka: probiootit

# Intervention paikka: probiootit

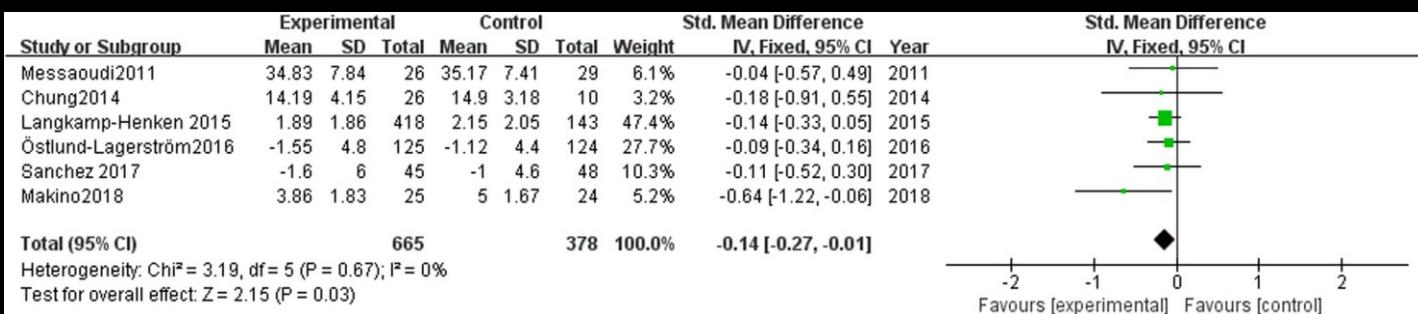
Brain and Behavior

Open Access

REVIEW | Open Access | CC BY

## Efficacy of probiotics on stress in healthy volunteers: A systematic review and meta-analysis based on randomized controlled trials

Ning Zhang, Yanan Zhang, Menglin Li, Weiguang Wang, Zhenzhu Liu, Chongcheng Xi, Xunying Huang, Jintao Liu, Junwei Huang, Dong Tian, Jie Mu, Xing Liao, Shuangqing Zhai



Review article

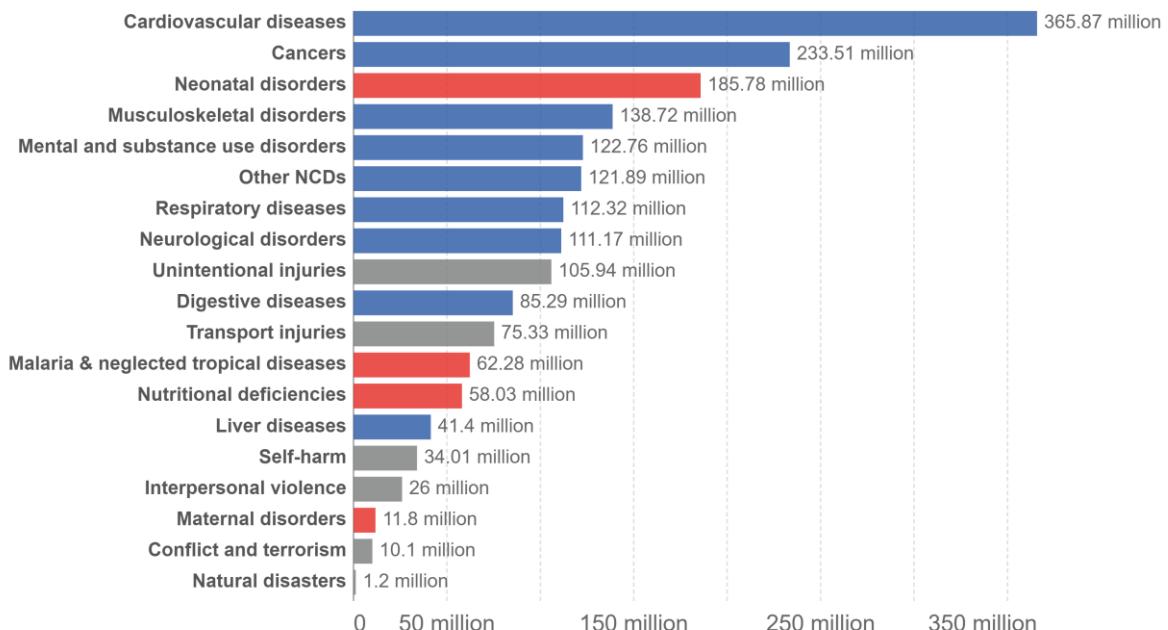
## The effect of probiotics on cognitive function across the human lifespan: A systematic review

Jessica Eastwood <sup>a</sup>, Gemma Walton <sup>b</sup>, Saskia Van Hemert <sup>c</sup>, Claire Williams <sup>a</sup>, Daniel Lampert <sup>a</sup>

# Muuttuva tautitaakka

## Burden of disease by cause, World, 2017

Total disease burden, measured in Disability-Adjusted Life Years (DALYs) by sub-category of disease or injury. DALYs measure the total burden of disease – both from years of life lost due to premature death and years lived with a disability. One DALY equals one lost year of healthy life.



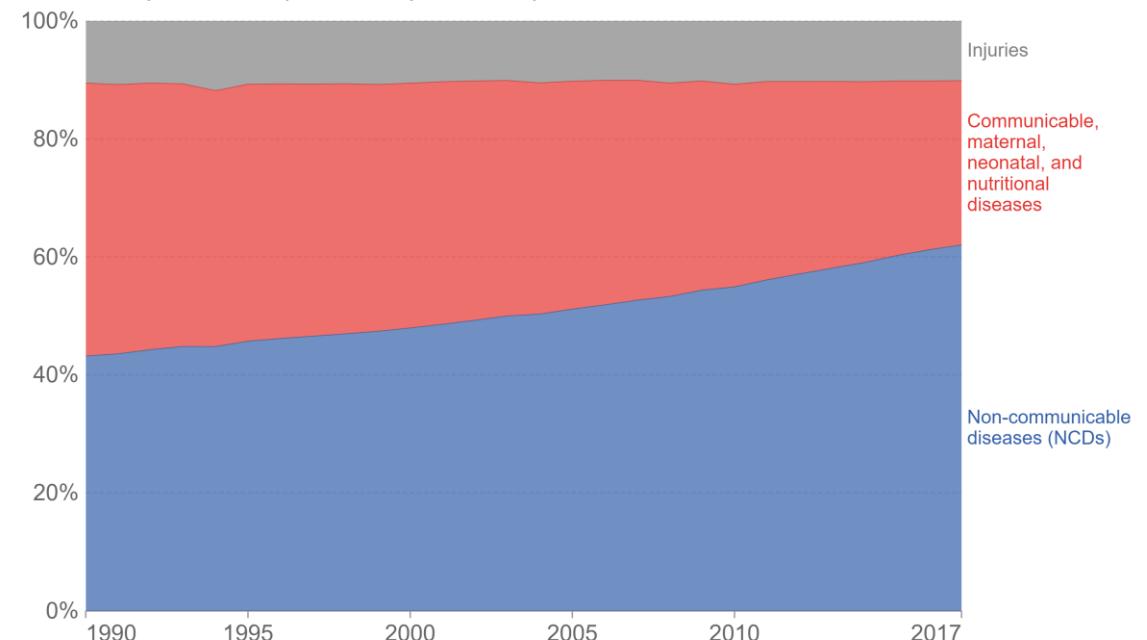
Source: IHME, Global Burden of Disease

[OurWorldInData.org/burden-of-disease](http://OurWorldInData.org/burden-of-disease) • CC BY

Our World  
in Data

## Total disease burden by cause, World, 1990 to 2017

Total disease burden measured as Disability-Adjusted Life Years (DALYs) per year. DALYs measure the total burden of disease – both from years of life lost due to premature death and years lived with a disability. One DALY equals one lost year of healthy life.

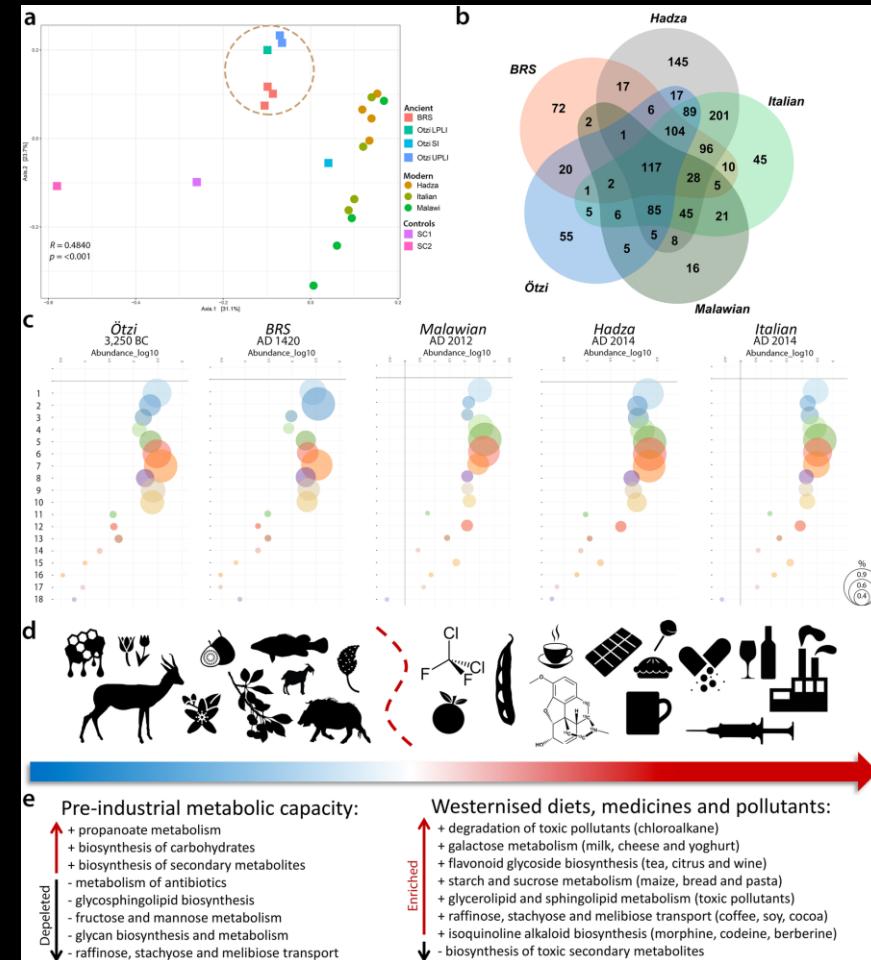
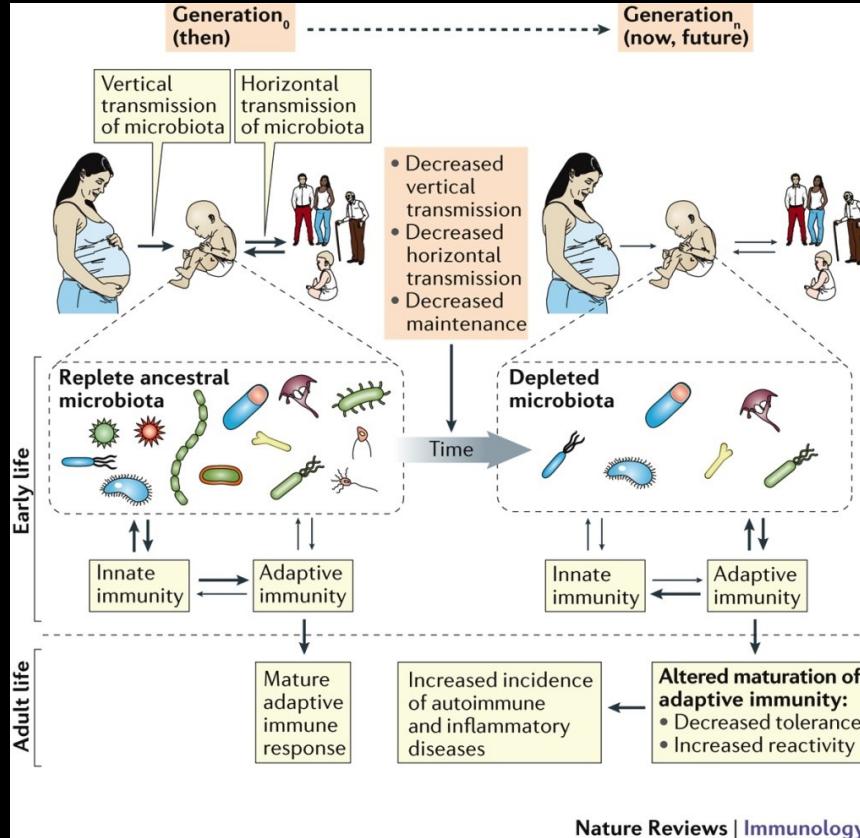


Source: IHME, Global Burden of Disease

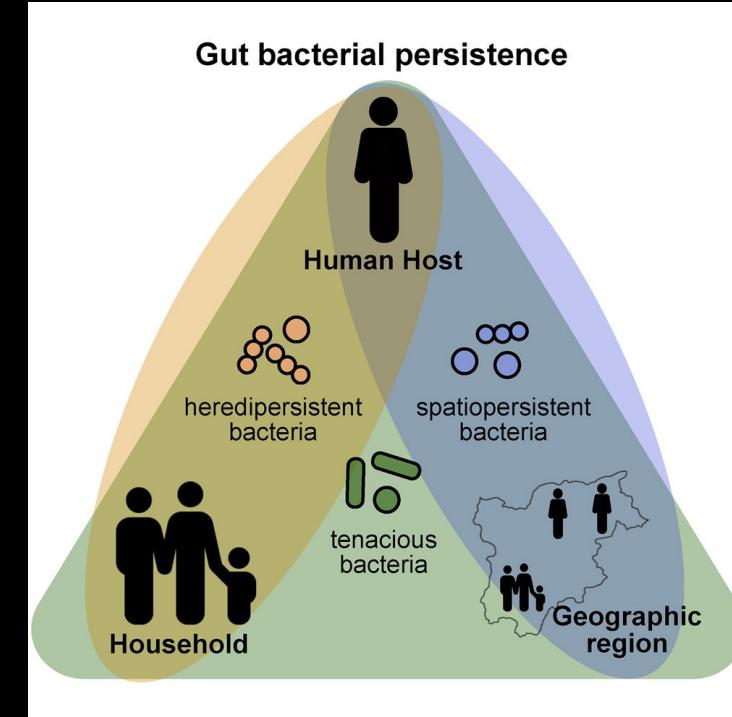
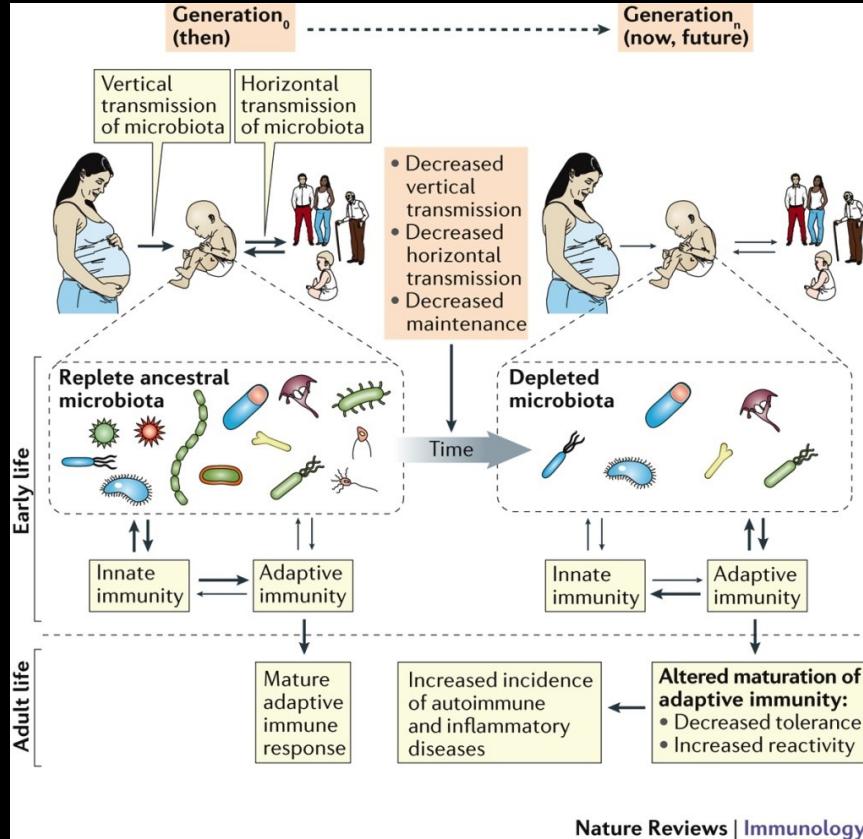
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# Muuttuva mikrobisto



# Muuttuva mikrobisto

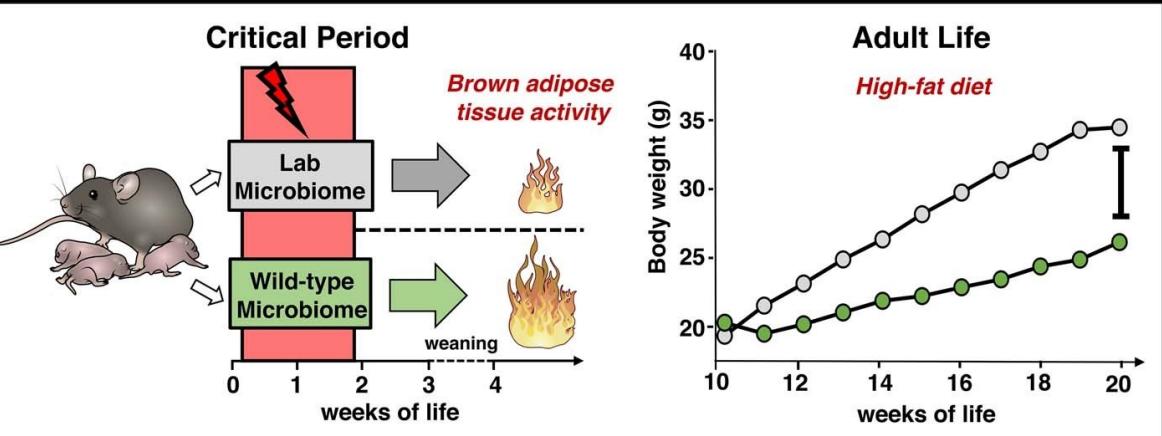
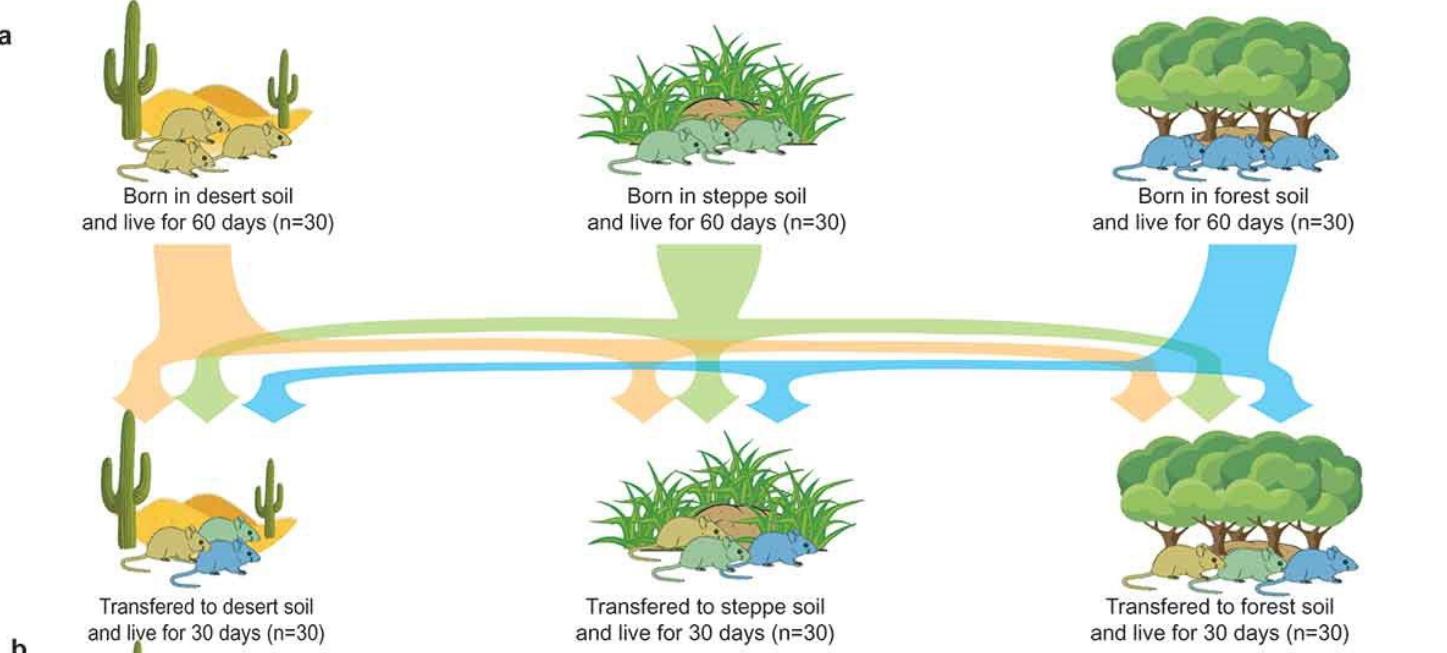
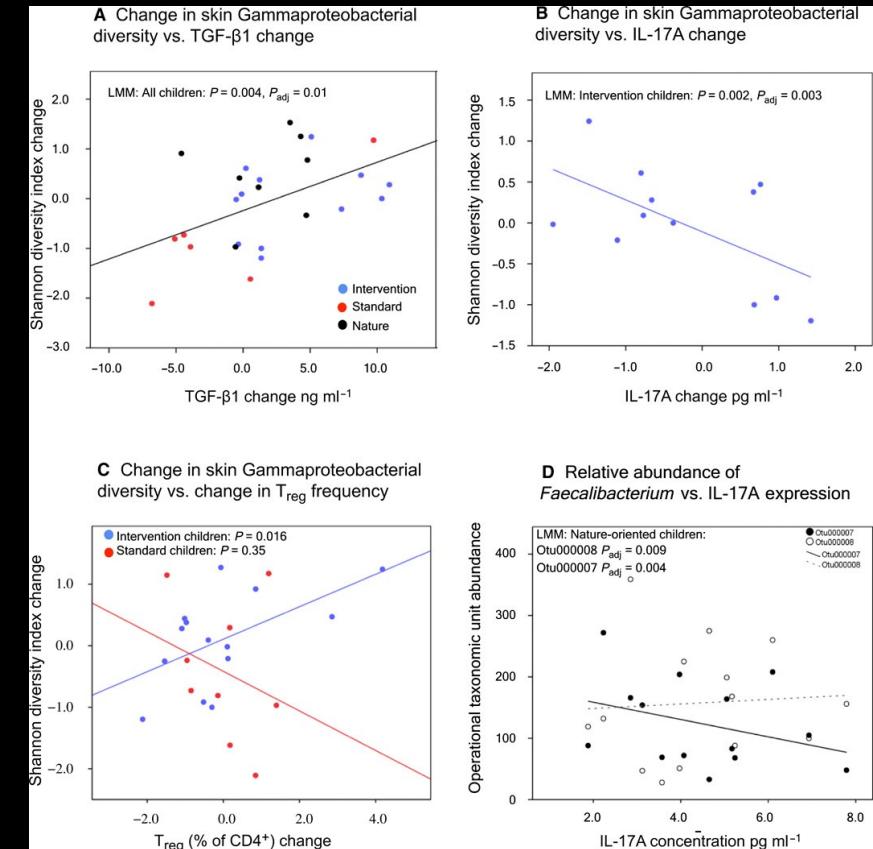


# Impact of outdoor nature-related activities on gut microbiota, fecal serotonin, and perceived stress in preschool children: the Play&Grow randomized controlled trial

Tanja Sobko, Suisha Liang, Will H. G. Cheng & Hein M. Tun 

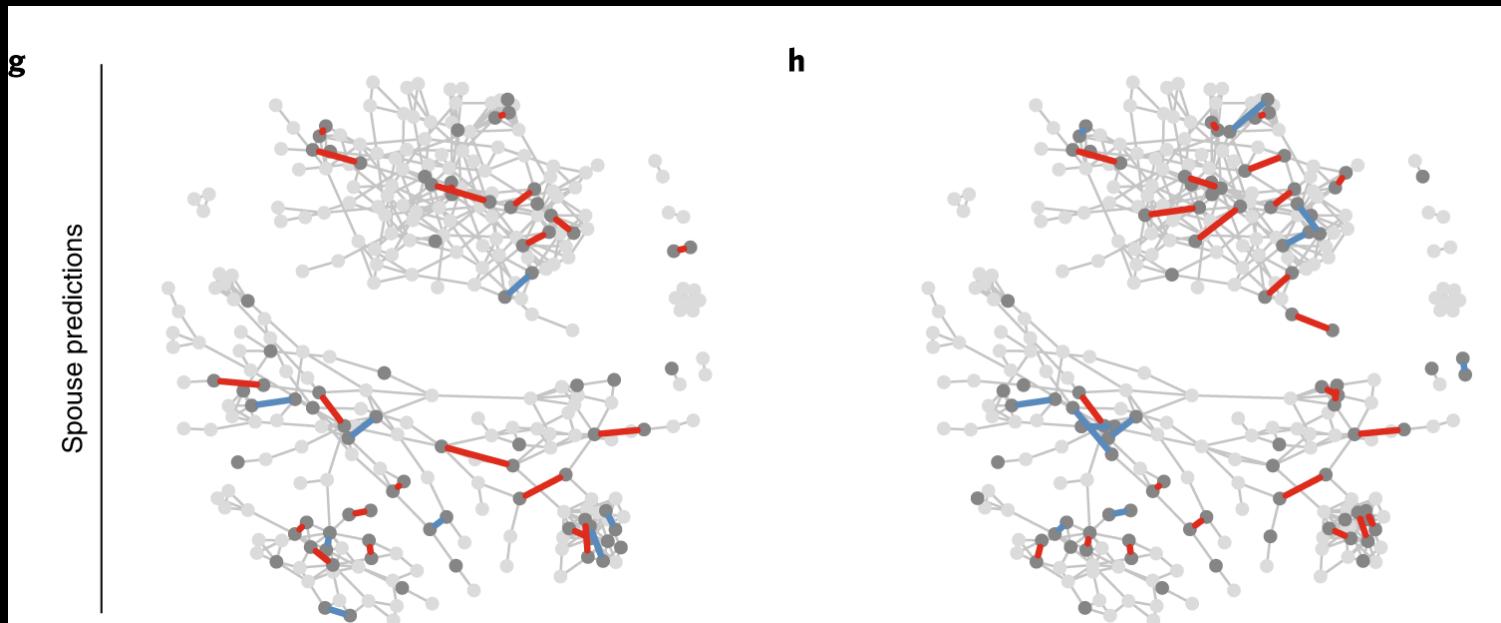
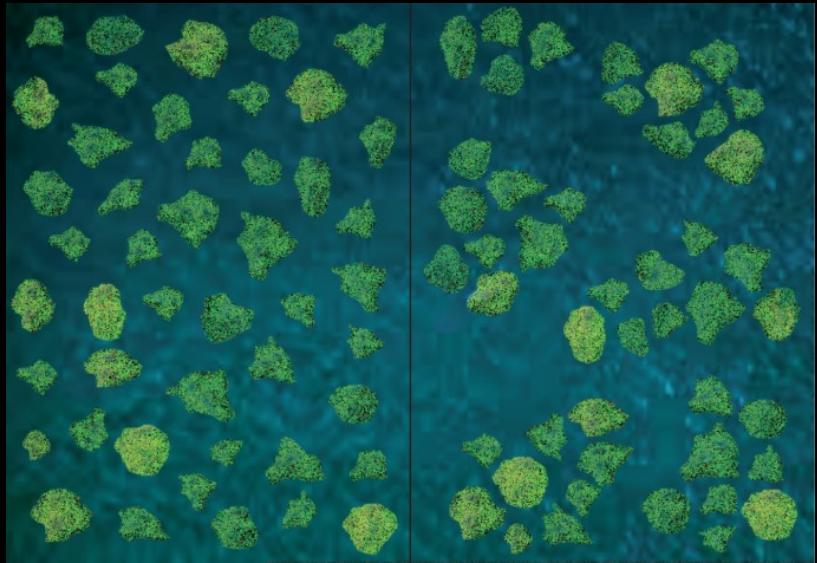
*Scientific Reports* 10, Article number: 21993 (2020) | [Cite this article](#)

3111 Accesses | 16 Altmetric | [Metrics](#)



Liu et al. 2021 Gut Microbes; ROSLUND et al. 2020, Sci Adv; Hild et al. 2021 Nat Metabolism.

# Sosiaaliset verkostot ja mikrobisto



Perspective | Published: 22 June 2020

## Microbial transmission in animal social networks and the social microbiome

[Amar Sarkar](#) [Siobhán Harty](#), [Katerina V.-A. Johnson](#), [Andrew H. Moeller](#), [Elizabeth A. Archie](#), [Laura D. Schell](#), [Rachel N. Carmody](#), [Timothy H. Clutton-Brock](#), [Robin I. M. Dunbar](#) & [Philip W. J. Burnet](#)

[Nature Ecology & Evolution](#) 4, 1020–1035 (2020) | [Cite this article](#)

5070 Accesses | 22 Citations | 65 Altmetric | [Metrics](#)

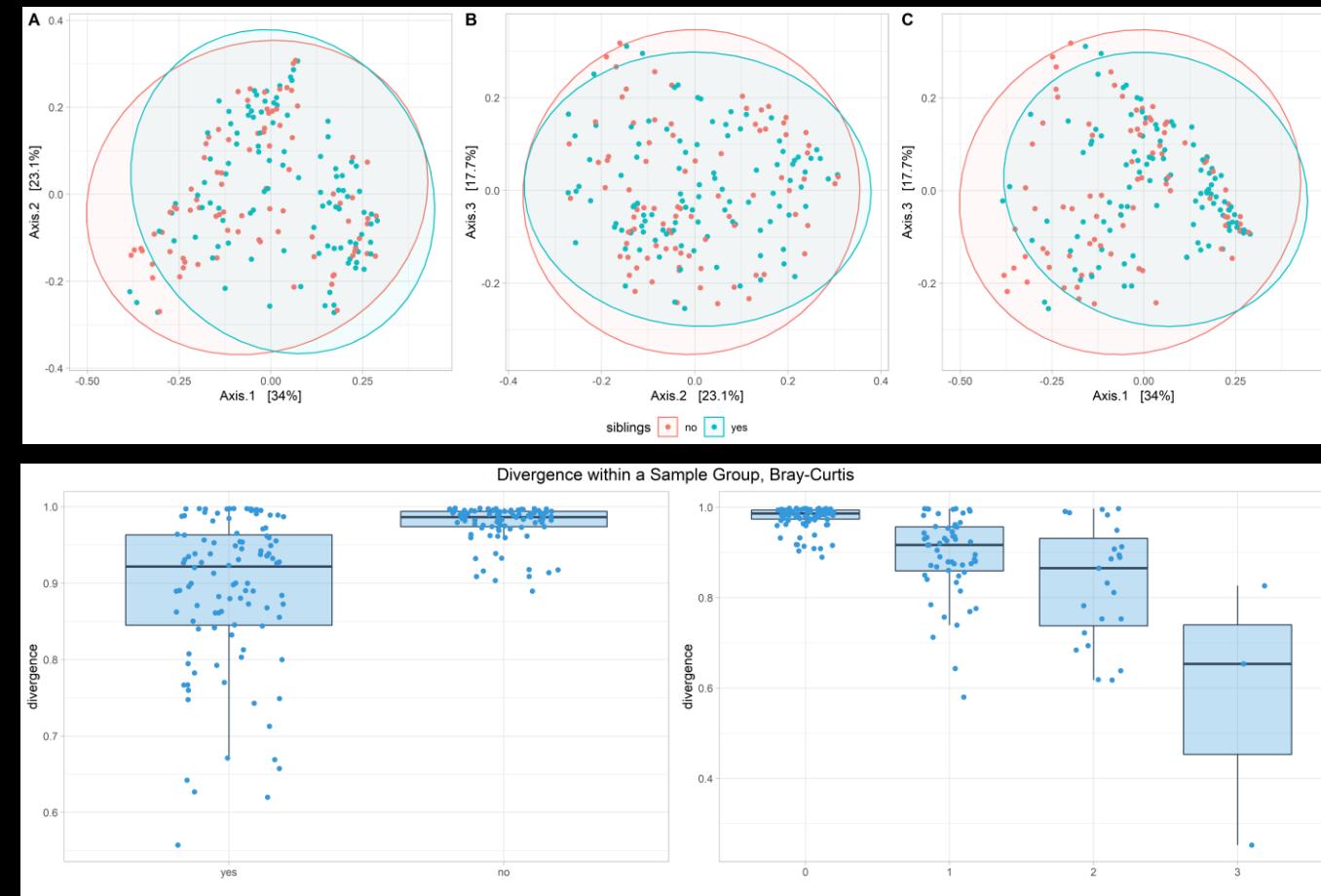
Letter | Published: 25 March 2019

## Transmission of human-associated microbiota along family and social networks

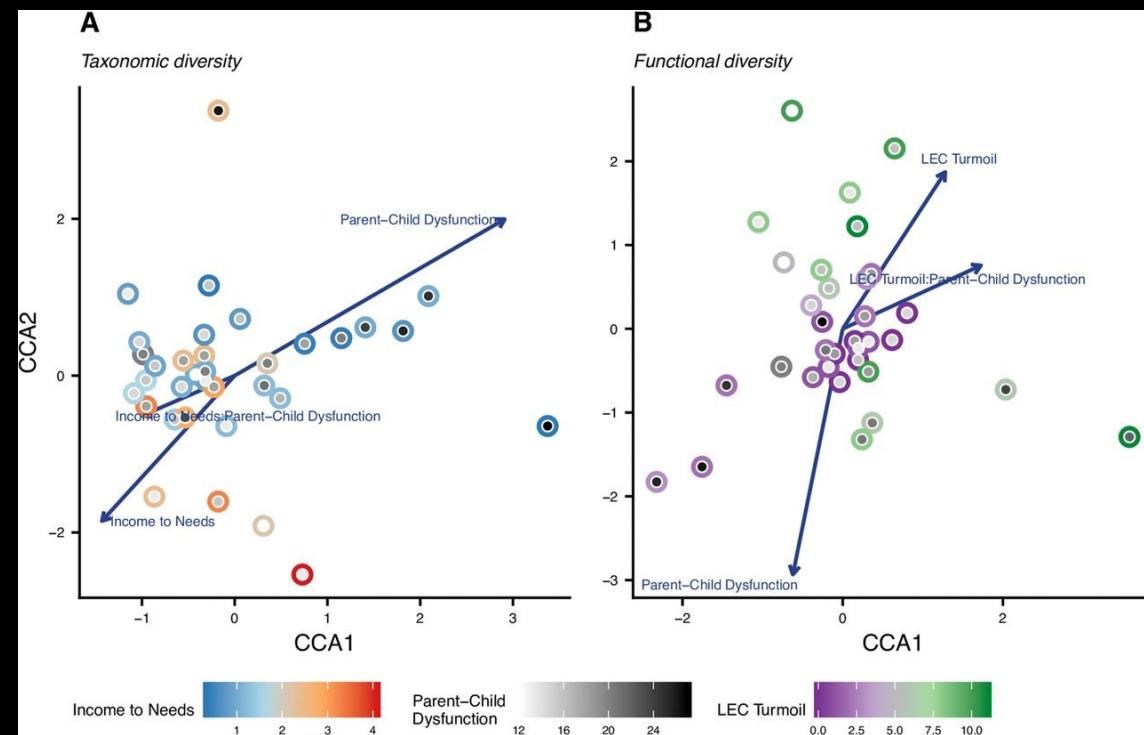
[Ilana L. Brito](#) [Thomas Gurry](#), [Shijie Zhao](#), [Katherine Huang](#), [Sarah K. Young](#), [Terrence P. Shea](#), [Waisea Naisilisili](#), [Aaron P. Jenkins](#), [Stacy D. Jupiter](#), [Dirk Gevers](#) & [Eric J. Alm](#)

[Nature Microbiology](#) 4, 964–971 (2019) | [Cite this article](#)

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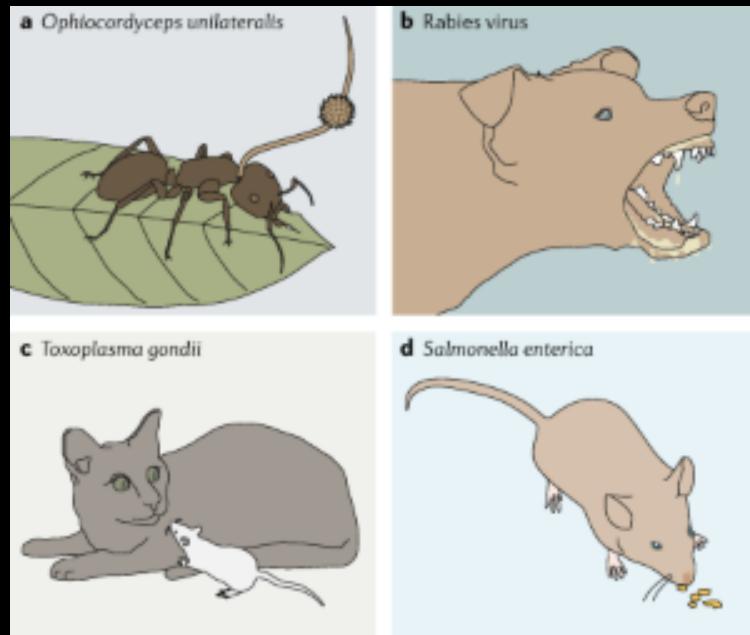
Infant saliva cortisol stress reactivity at 2.5 mo age, n=191,  
Keskitalo, Aatsinki et al. 2021 Stress



## Gut Feelings Begin in Childhood: the Gut Metagenome Correlates with Early Environment, Caregiving, and Behavior

Jessica E. Flannery,<sup>a</sup> Keaton Stagaman,<sup>b</sup> Adam R. Burns,<sup>c</sup> Roxana J. Hickey,<sup>d,e</sup> Leslie E. Roos,<sup>f</sup> Ryan J. Giuliano,<sup>f</sup> Philip A. Fisher,<sup>a</sup> Thomas J. Sharpton<sup>b,g</sup>

# Mikrobisto ja käyttäytyminen



Perspective | [Published: 24 April 2018](#)

OPINION

## Why does the microbiome affect behaviour?

[Katerina V.-A. Johnson](#) & [Kevin R. Foster](#)

[Nature Reviews Microbiology](#) **16**, 647–655 (2018) | [Cite this article](#)

**27k** Accesses | **96** Citations | **709** Altmetric | [Metrics](#)

# Take-home

- Mikrobisto voi olla yhteydessä käyttäytymiseen myös ihmisillä
  - välittyy mikrobiston aineenvaihduntatuotteiden, vagushermon toiminnan, immuunijärjestelmän kautta
  - Syntyy mikrobiston evoluution ja kiplailun sivutuotteena
- Varhainen mikrobisto voi olla yhteydessä sosioemotionaalisiin kehityskaariin
- Epäselvää minkä verran yksittäisillä bakteerikannoilla voi muuttaa käyttäytymistä aikuisilla
- Myös muista lähteistä tulevilla mikrobeilla voi olla vaikutusta suoliston ekosysteemiin ja sen isäntään kohdistuviin vaikuttuksiin



Adj. Prof. Linnea Karlsson

Prof. Hasse Karlsson

Adj. Prof. Leo Lahti

Henna-Maria Kailanto, Eveliina  
Munukka, alumni

Saara Nolvi, Eeva-Leena Kataja,  
Anniina Keskitalo, post docs

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Teemu Kallonen, Turku  
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Alex Dickens, Santosh  
Lamichhane, Turku Bioscience  
Siobhain O'Mahony, University  
College Cork

Ana Joao Rodrigues, University of  
Minho

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The Academy of Finland, Yrjö  
Jahnsson Foundation, Signe and  
Ane Gyllenberg Foundation,  
Finnish State Grants for Clinical  
Research and Waterloo Foundatio,  
Jalmari and Rauha Ahokas  
Foundation and Finnish State  
Grants for Clinical Research

Personal Funding:  
Finnish Medical Foundation,  
Psychiatry Research Foundation,  
Maire Tapnonen foundation, Turku  
University Foundation, Signe and  
Ane Gyllenberg Foundation,  
Aivosäätiö, Instrumentarium  
Science Foundation, Finnish State  
Grants for Clinical Research, Emil  
Aaltonen Foundation