

Disclosure Slide



College of Psychiatrists
of Ireland
Wisdom • Learning • Compassion

WINTER CONFERENCE 2018

FROM THE BIOME TO BIPOLAR

15th & 16th NOVEMBER 2018
The BREHON HOTEL / INEC Killarney

**** External CPD credits (Thursday: 7 credits / Friday: 6 credits)*

- **Speakers Bureau, Janssen**
- **APC Microbiome Ireland research funded in part by Dupont Nutrition Biosciences APS, Cremo SA, Alkermes Inc., 4D Pharma PLC, Alimentary Health, Mead Johnson Nutrition, Nutricia Danone, Suntory Wellness**
- **This support has neither influenced nor constrained the content of this presentation**



Interfacing Food & Medicine



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WINTER CONFERENCE 2018

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13th & 16th NOVEMBER 2018
The BREHON HOTEL / INEC Killarney

*** External CPD credits (Thursday: 7 credits / Friday: 6 credits)

Interfacing Food and Medicine

The Little Things that Matter Most in Psychiatry: An Update on Gut Microbes and Their Effects on Brain Function and Host Behaviour

Dr Gerard Clarke

Department of Psychiatry and Neurobehavioural Science
and APC Microbiome Ireland

University College Cork

Killarney November 16th 2018



UCC
University College Cork, Ireland
Coláiste na hOllscoile Corcaigh

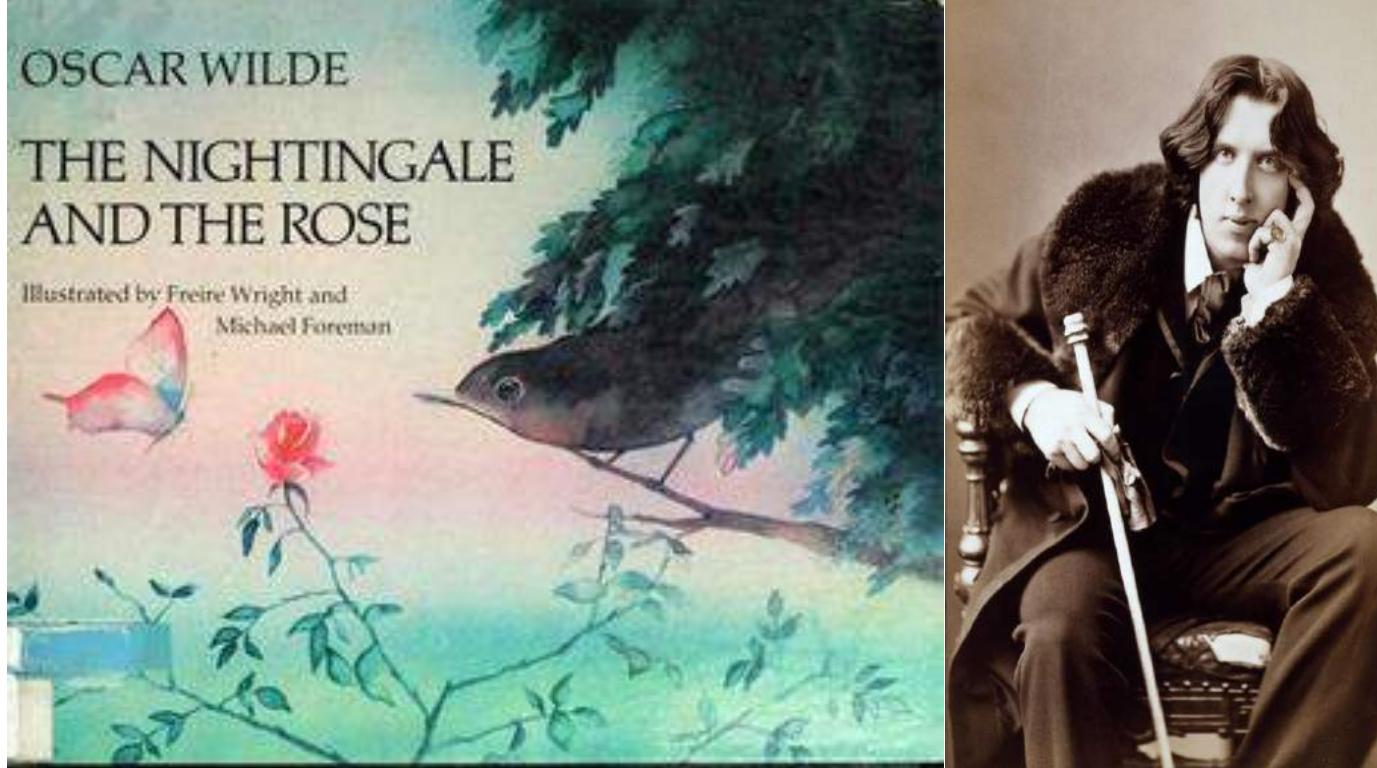
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eagasc
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Gut Feelings

- Brain-Gut-Microbiome Axis

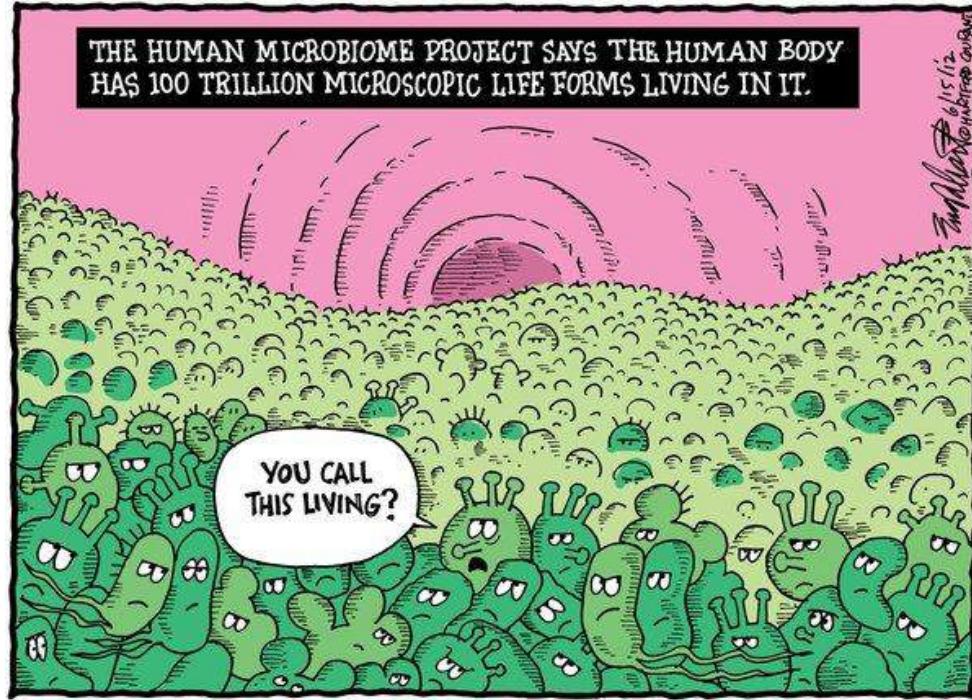
- Health and Disease

- 'Mind altering microbes'

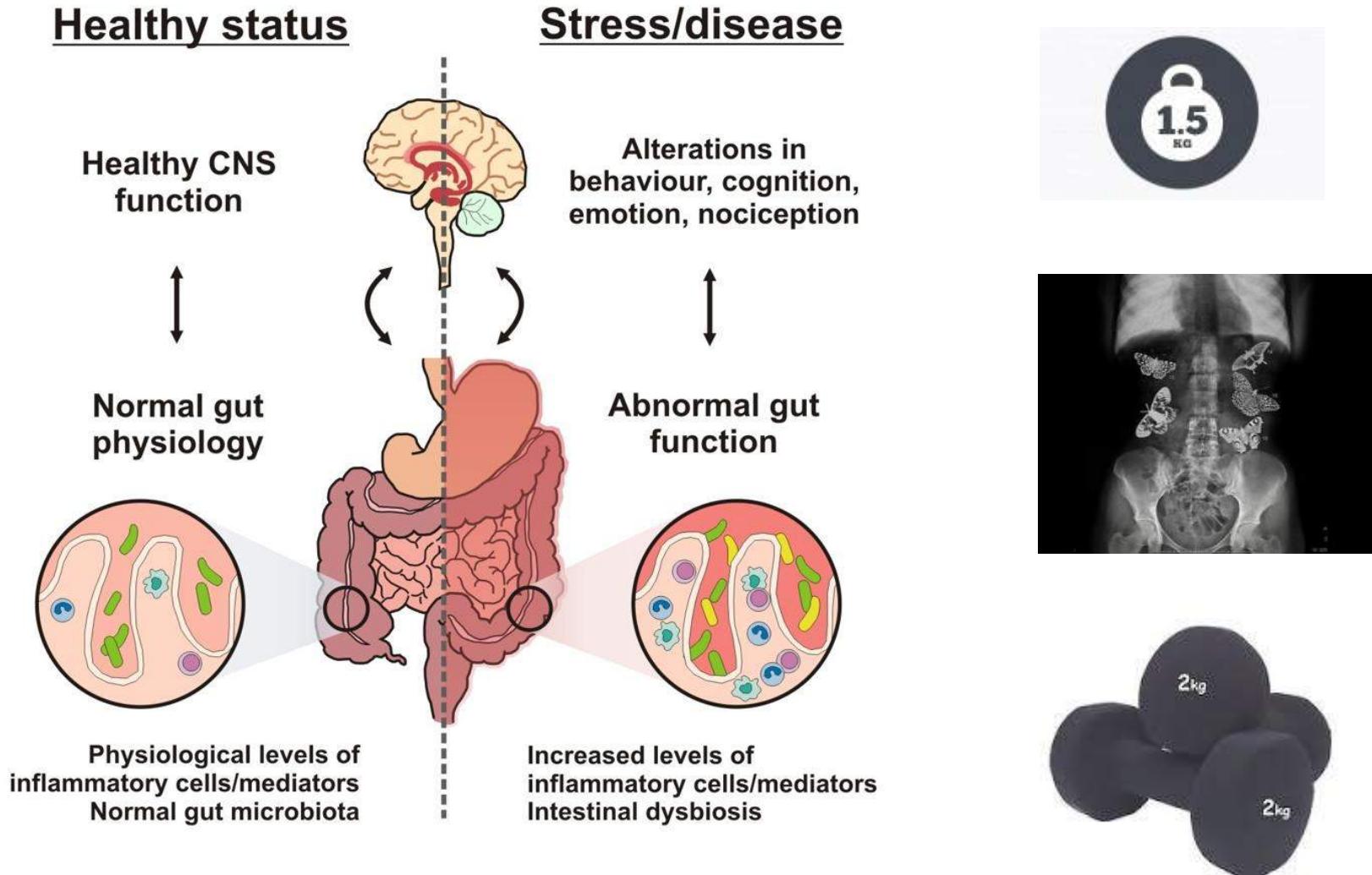
- Anxiety, Depression,
Stress and Cognition

- Tryptophan availability and metabolism

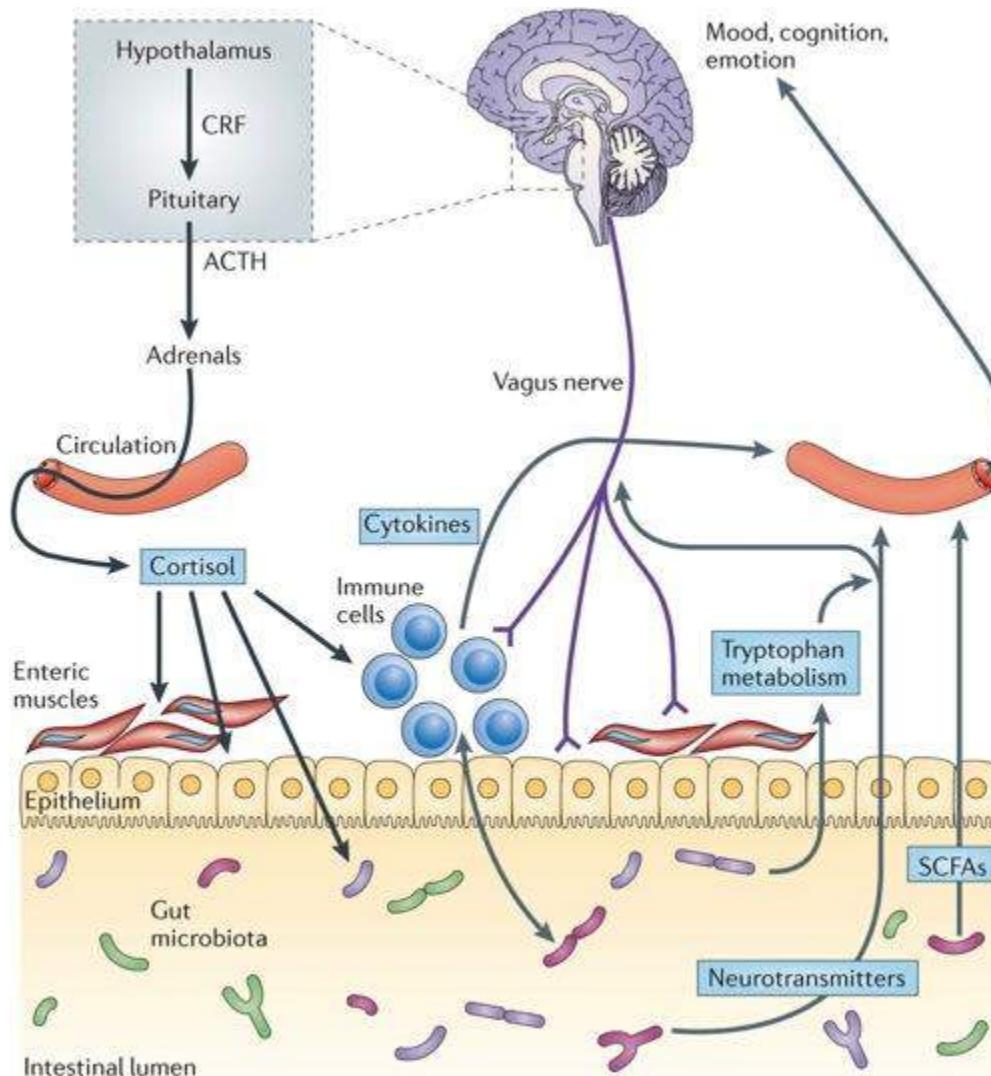
- Translational implications and opportunities for intervention?



Brain-Gut-Microbiome Axis



Signalling Along the Brain-Gut-Microbiota axis

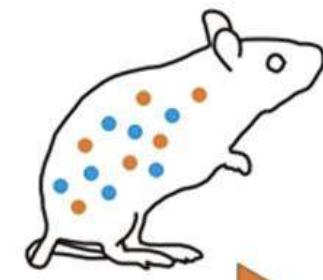
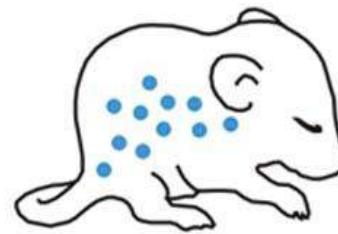


Where do we get our microbiota from?



Interfacing Food & Medicine

<http://apc.ucc.ie>



Prenatal
(sterile)

Birth

Postnatal
(colonized)



Grenham et al., 2011

Article

Cell

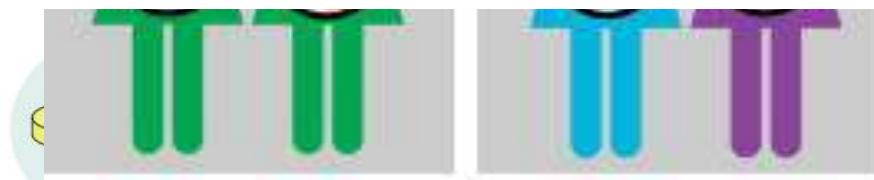
210 | NATURE | VOL 555 | 8 MARCH 2018

doi:10.1038/nature25973

ARTICLE

Environment dominates over host genetics in shaping human gut microbiota

Daphna Rothschild^{1,2*}, Omer Weissbrod^{1,2*}, Elad Barkan^{1,2*}, Alexander Kurilshikov³, Tal Korem^{1,2}, David Zeevi^{1,2}, Paul I. Costea^{1,2}, Anastasia Godneva^{1,2}, Iris N. Kalka^{1,2}, Noam Bar^{1,2}, Smadar Shilo^{1,2}, Dar Lador^{1,2}, Arnau Vich Vila^{3,4}, Niv Zmora^{5,6,7}, Meirav Pevsner-Fischer⁵, David Israeli⁸, Noa Kosower^{1,2}, Gal Malka^{1,2}, Bat Chen Wolf^{1,2}, Tali Avnit-Sagi^{1,2}, Maya Lotan-Pompan^{1,2}, Adina Weinberger^{1,2}, Zamir Halpern^{7,9}, Shai Carmi¹⁰, Jingyuan Fu^{3,11}, Cisca Wijmenga^{3,12}, Alexandra Zhernakova³, Eran Elinav^{5,§} & Eran Segal^{1,2§}

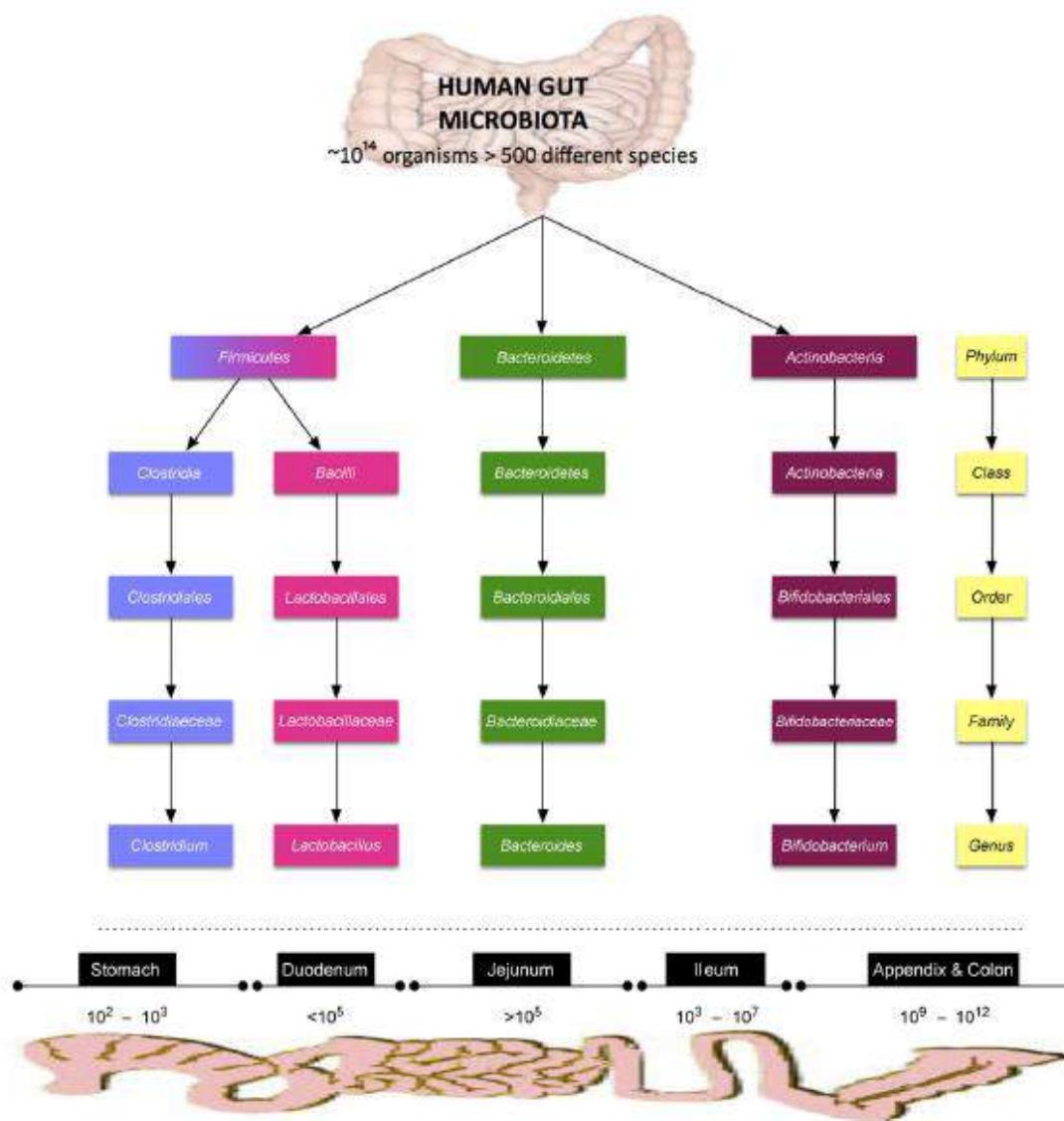
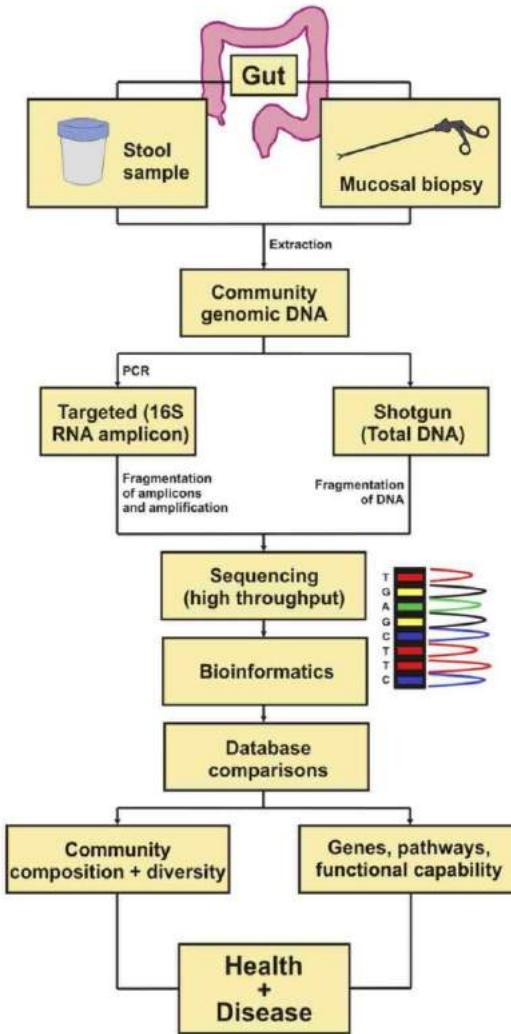


MZ twins have a more similar
microbiota than DZ twins

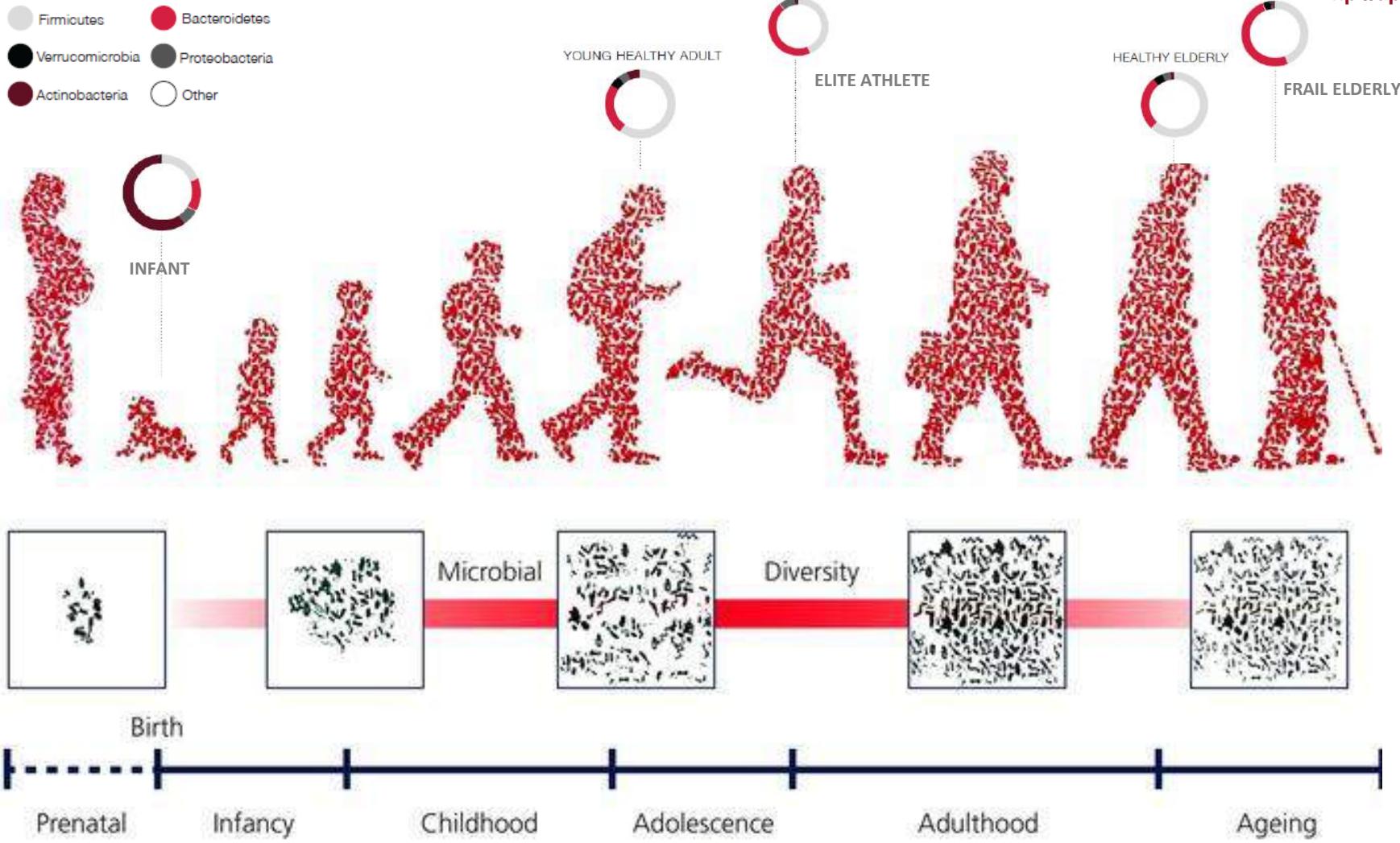
Figure 1 | Factors that can influence the composition and function of the human gut microbiota.

It's a gut feeling: How the gut microbiota affects the state of mind

Adam D. Farmer, Holly A. Randall and Qasim Aziz

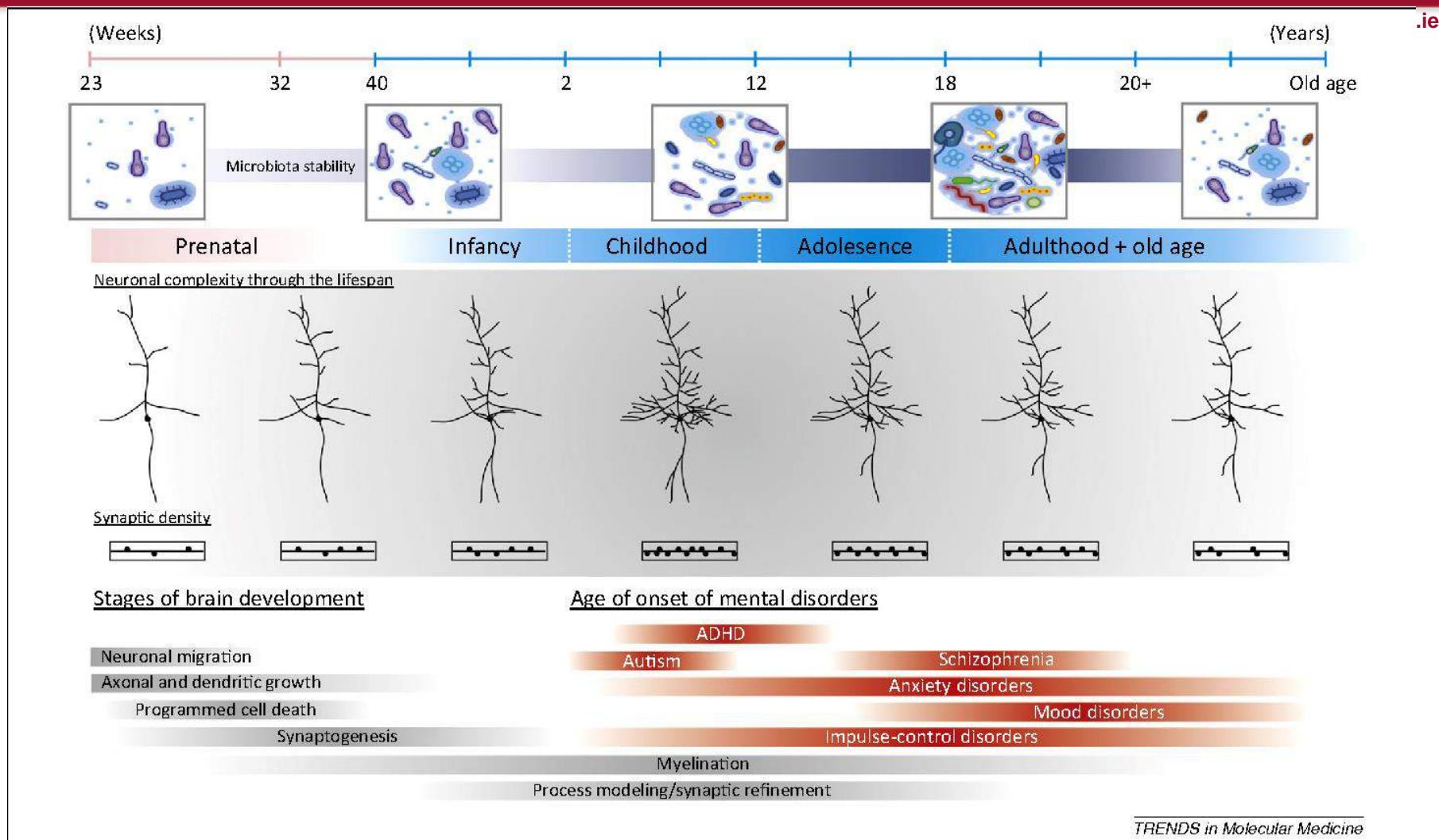


GI microbiota over lifetime

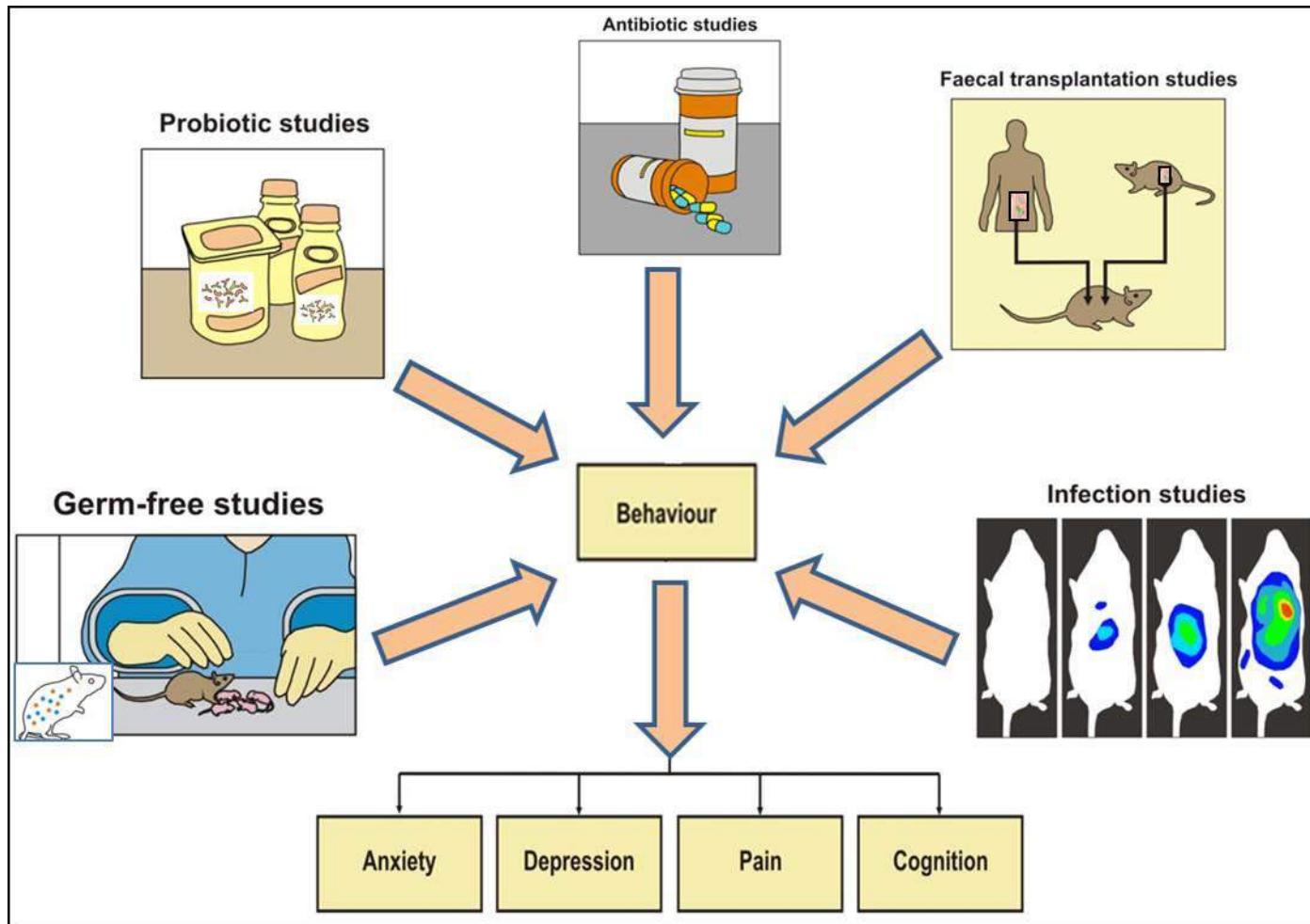


Cryan and Dinan, J Physiology 2017

Microbiota and Neurodevelopment

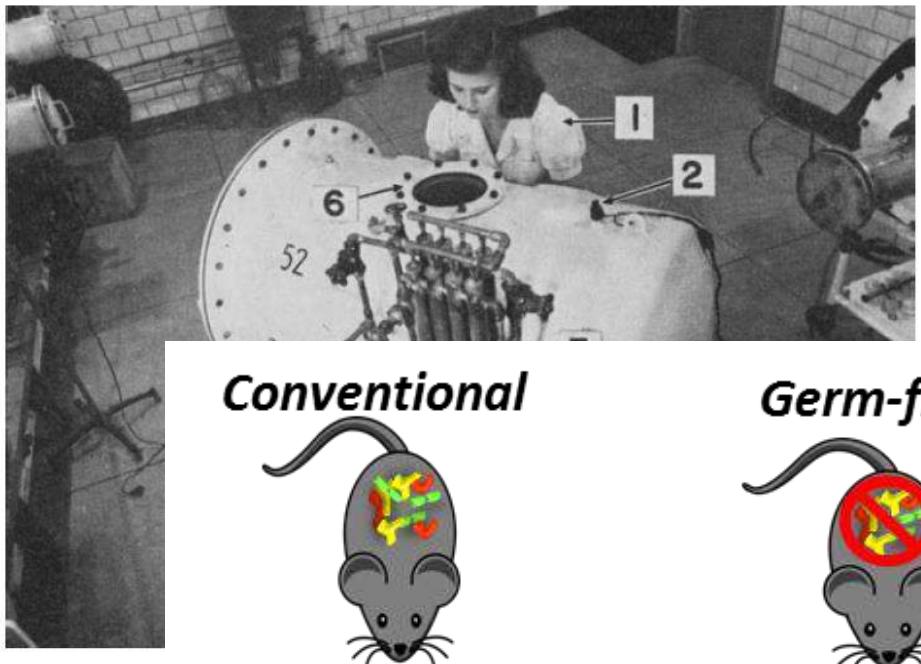


Microbiota, Brain and Behaviour

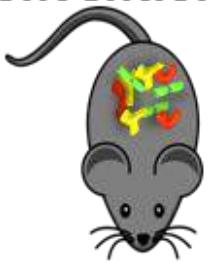


Clarke et al., Encyclopedia Metagenomics 2013

Germ-Free Living?



Conventional



Germ-free



Germ-free colonised

Figure 2. Reyniers's isolator; (1) technician, (2) electrical outlet, (3) air outlet, (4) mobile truck, (5) entrance/exit autoclave, (6) viewing port.
Source: J. A. Reyniers, P. C. Trexler, and R. F. Ervin, "Rearing Germ-Free Albino Rats," *LOBUND Rep.* 1 (1946): 1–84, 5. © University of Notre Dame. Reprinted with permission.

Kirk, R, *Bulletin of the History of Medicine*, 2012

Mr. William Shearer, said today at a news conference.

"He said that we had all these tubes and wires that taste sour. 'They're getting tired. Why don't we just pull out all of these tubes and let me go home.' Shearer said.

The end came just over two weeks after the joyous moment when David stepped out of his mobile for the first time, kissed his mother and felt the loving warmth of a human touch.

With David gone, everybody in the hospital felt it. There were tears all around. "It was like a funeral," said Dr. Shearer. "The nurses cried and even some tough police officers cried," said Houston police Lt. Jim Mills.

His family, whose last name has never been released to protect their privacy, had been delivered to the hospital by plane from Canada.

"They seemed limp and exhausted," Mills said.

David left the two-room enclosure Feb. 7 because it was the only way doctors could find flu-like symptoms attributed to an experimental bone marrow transplant he received in October from a donor in Canada.

David, who had talked of getting out of his bubble since the age of 3 and once

■ weeks after the joyous moment when David stepped out of his mobile for the first time, kissed his mother and felt the loving warmth of a human touch.

He was delivered by Caesarean section under extremely sterile conditions on Sept. 21, 1971, and put into a sterile incubator inside a fine mesh enclosure home that grew with as he did.

Everything that touched him clothes, food and books, was sterilized and passed through an airlock into the bubble.

David initially spent most of his time at the hospital, then shared time at home with his parents and his brother along with one for the family's station wagon.

In 1981, he was spending all but two weeks a year at home. A sixth-grader at the time of his death, he attended school by telephone and consistently got high grades and tests showed he was brighter than average.

"It was necessary to take the calcified bones from the hospital spokesman said.

But in January, David became ill for the first time in his life, developing diarrhea and vomiting.

After leaving the bubble, he developed open blisters on his hands and began receiving blood transfusions. Other internal bleeding occurred and could not be stopped.

It was necessary to take the calcified bones from the hospital spokesman said.

McNutt said. But in January, David became ill for the first time in his life, developing diarrhea and vomiting.

After leaving the bubble, he developed open blisters on his hands and began receiving blood transfusions. Other internal bleeding occurred and could not be stopped.

Doctors said Feb. 13 that tests showed David had graft vs.-host disease, a condition in which the transplanted immune system attacks the body.

The boy's death was his most important contribution to medicine, Shearer said.

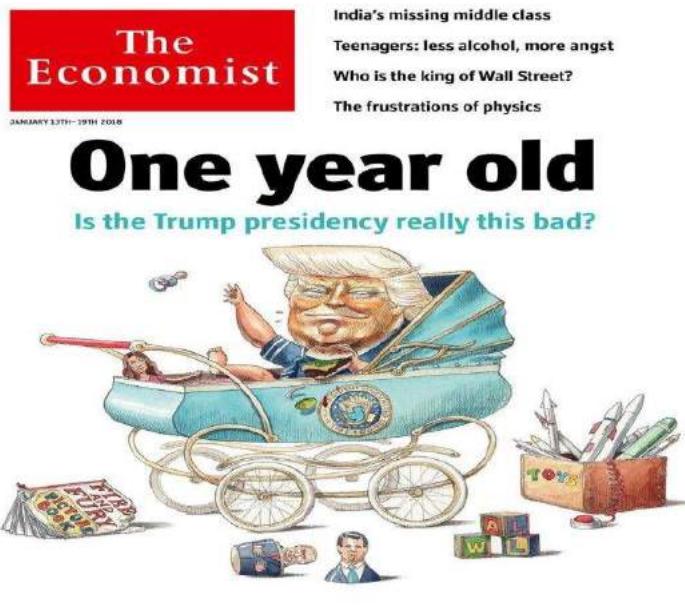
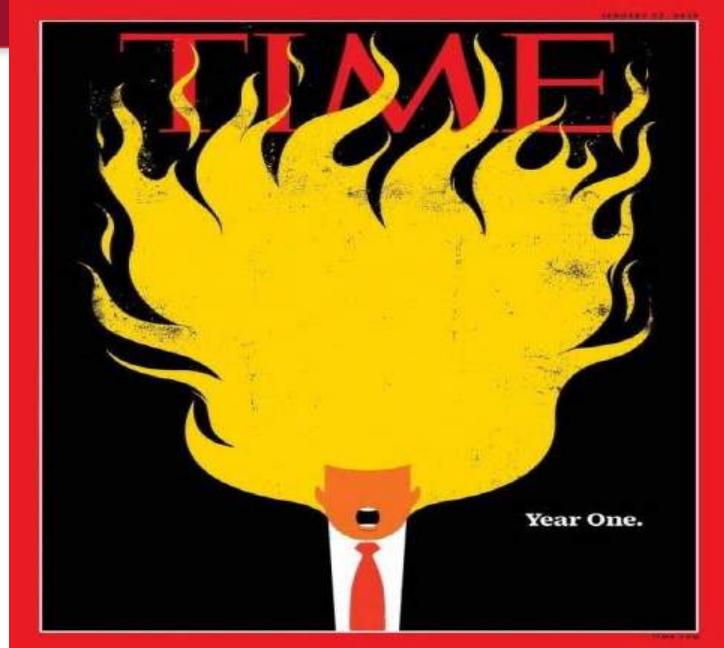
Shearer said apparently as a result of a proliferation of a type of lymphocyte — an abnormal growth of B-cells — not from the transplant as had been believed.

Shearer said as had been believed.

That discovery, made after Shearer performed an autopsy, was a major finding and of great medical significance, he said.

David's funeral was scheduled for Saturday morning. David's family requested that it be private, the hospital said.

Stressors



Microbiota Controls Stress Response

J Physiol 558.1 (2004) pp 263–275

263

250

200

150

100

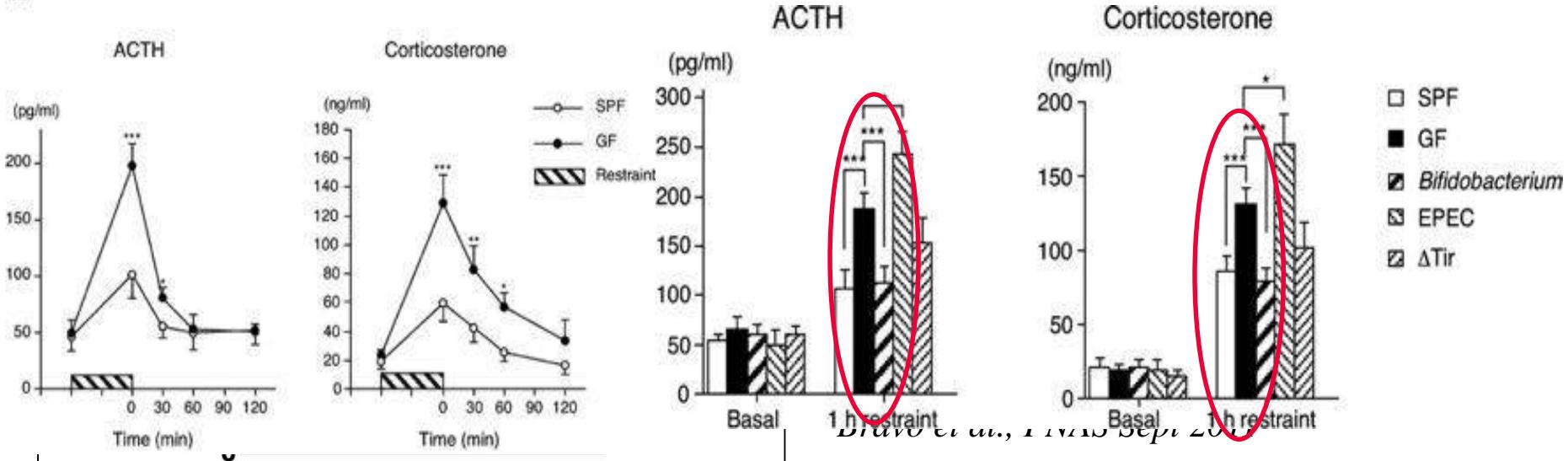
50

Postnatal microbial colonization programs the hypothalamic–pituitary–adrenal system for stress response in mice

Nobuyuki Sudo^{1,2}, Yoichi Chida¹, Yuji Aiba^{3,4}, Junko Sonoda¹, Naomi Oyama¹, Xiao-Nian Yu¹, Chiharu Kubo¹ and Yasuhiro Koga³

¹Department of Psychosomatic Medicine and ²Department of Health Care Administration & Management, Graduate School of Medical Sciences, Kyushu University, Fukuoka, Japan, ³Department of Infectious Diseases, Tokai University School of Medicine, Isehara, Kanagawa, Japan and ⁴Wakamoto Pharmaceutical Co. Ltd, Ohi-machi, Kanagawa, Japan

A



Microbiota Determines Amygdala Volume & Dendritic Morphology

Research Report

Adult microbiota-deficient mice have distinct dendritic morphological changes: differential effects in the amygdala and hippocampus

Pauline Luczynski¹, Seán O' Whelan³,
Colette O'Sullivan³, Gerard Clarke^{1,2},
Fergus Shanahan¹, Timothy G. Dinan^{1,2}
and John F. Cryan^{1,3,*}

DOI: 10.1111/ejn.13291

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European Journal of
Neuroscience

Accepted Article (Accepted,
unedited articles published
online and citable. The final
edited and typeset version of
record will appear in future.)

SEARCH

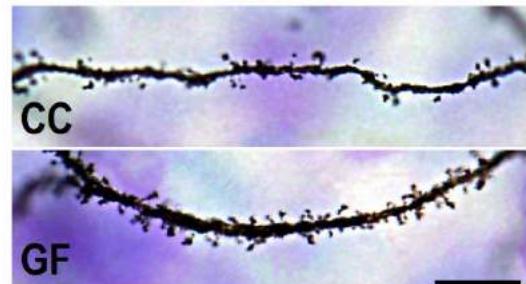
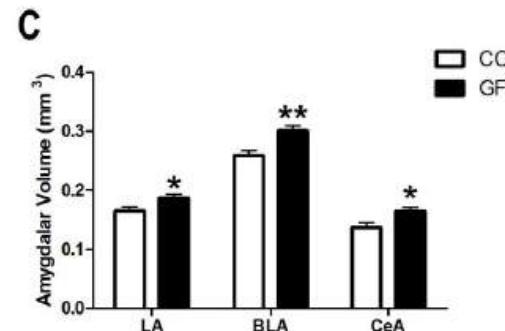
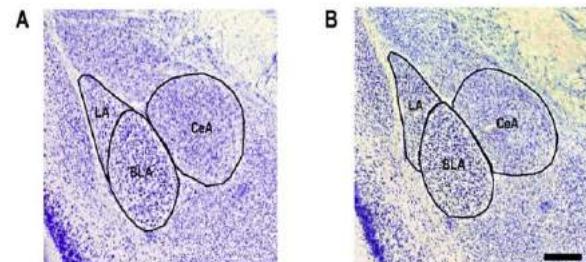
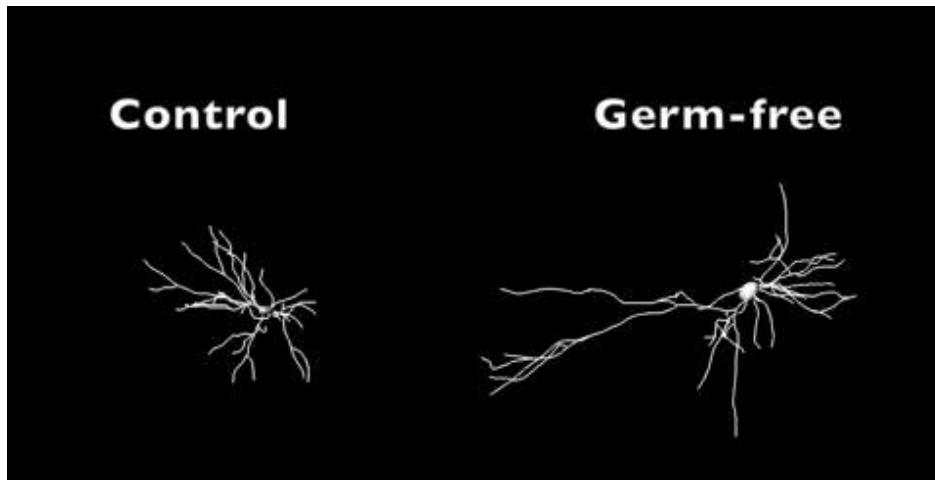
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Dendritic Hypertrophy of Basolateral Amygdala Neurons



CC = Conventionally Colonised
GF = Germ Free

ORIGINAL ARTICLE

The microbion

AE Hoban^{1,2}, RM Stilling^{1,2}, G N

Volume 40 No. 1 January 2018 · ISSN 0265-9247

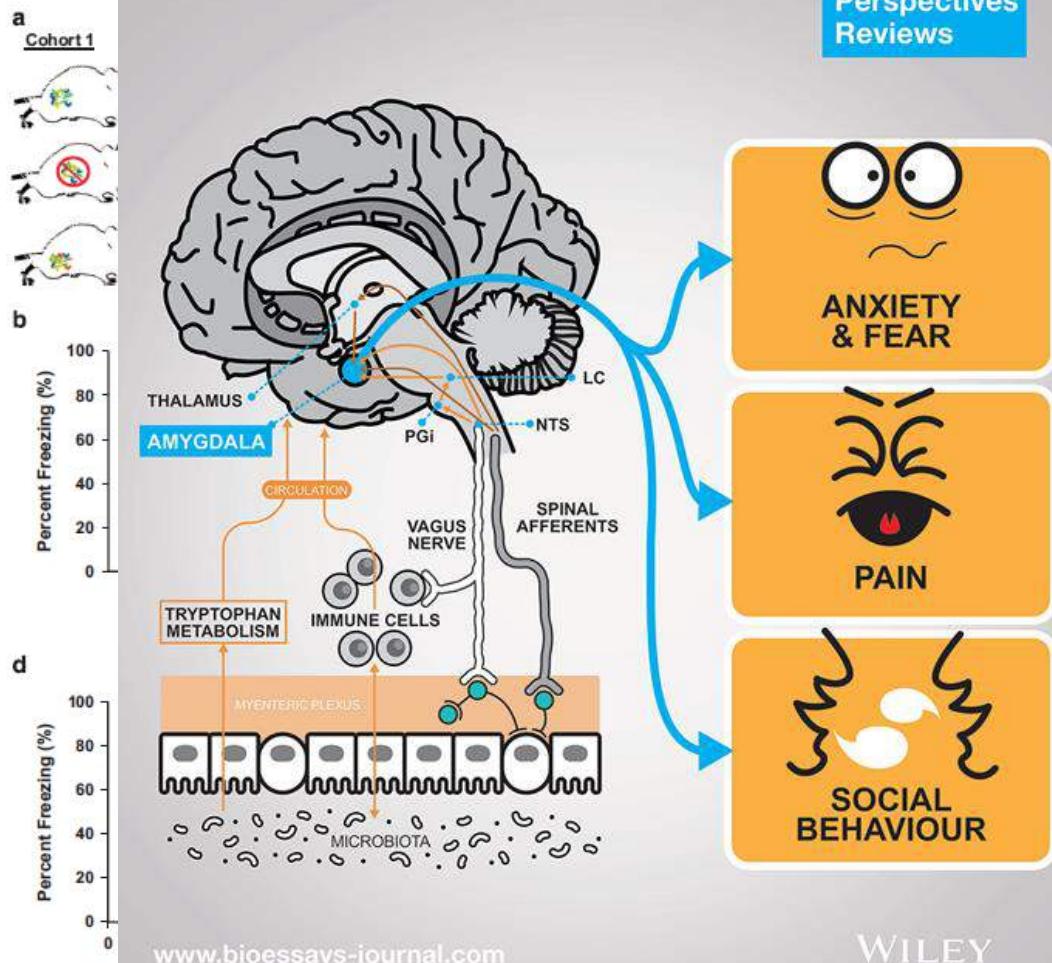
BioEssays

Ideas that Push the Boundaries

1/18

 Hypotheses
 Perspectives
 Reviews

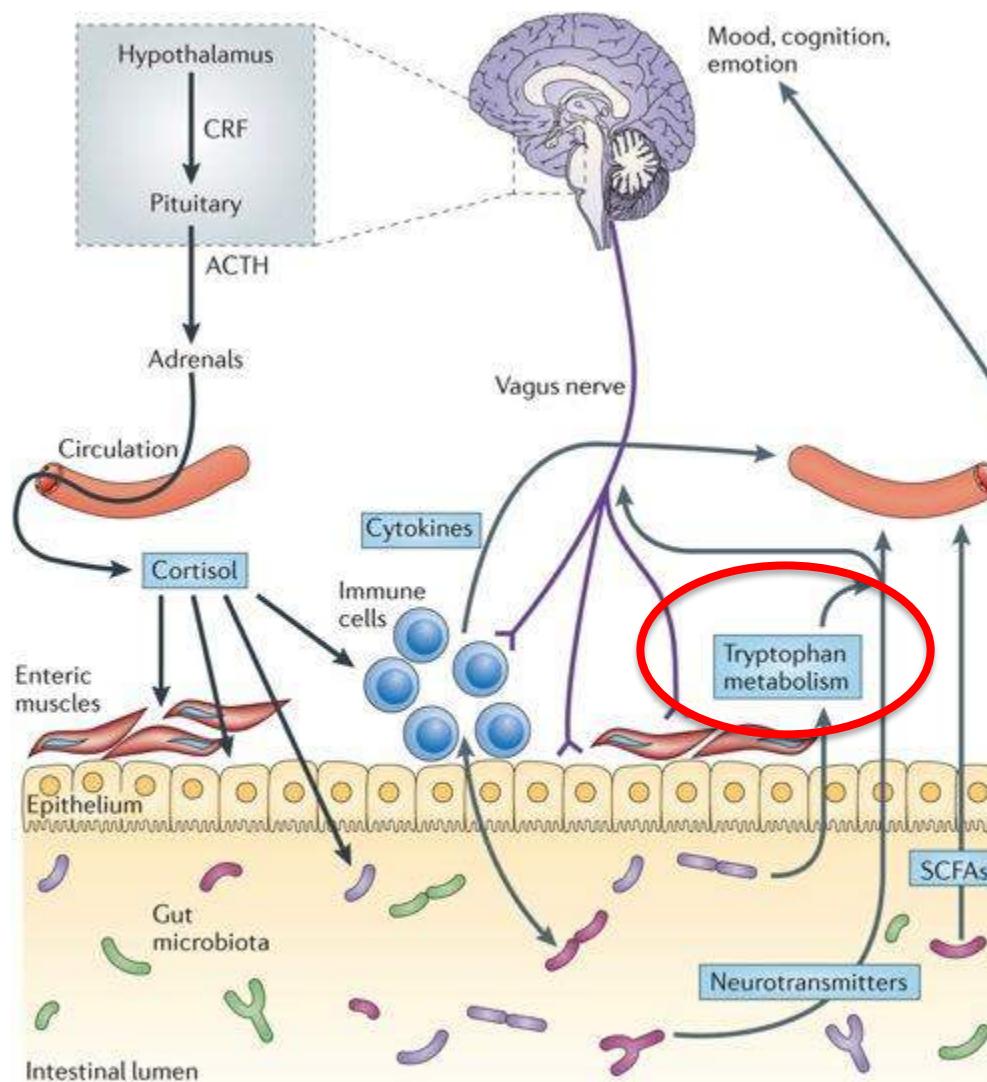
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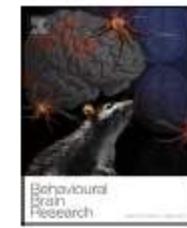


Anatomical correlates of abnormal fear and anxiety can be localised to the amygdala in germ-free animals using cued fear conditioning

WILEY

Signalling Along the Brain-Gut-Microbiota axis





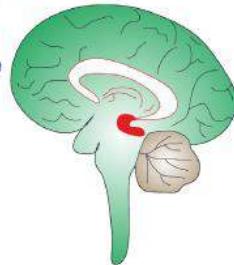
Review

Serotonin, tryptophan metabolism and the brain-gut-microbiome axis

S.M. O'Mahony ^{a,b,1}, G. Clarke ^{a,c,*,1}, Y.E. Borre ^a, T.G. Dinan ^{a,c}, J.F. Cryan ^{a,b}

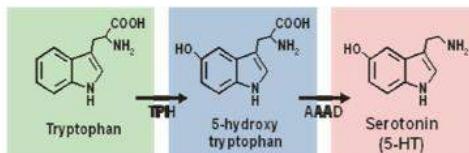
Behavioural Effects

- Visceral pain
- Emotion
- Stress response
- Appetite
- Addiction
- Sexuality



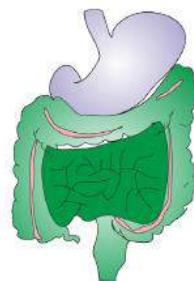
CNS Effects

- Motor control
- Circadian rhythm
- Cerebellar regulation
- Body temperature
- CNS vascular tone

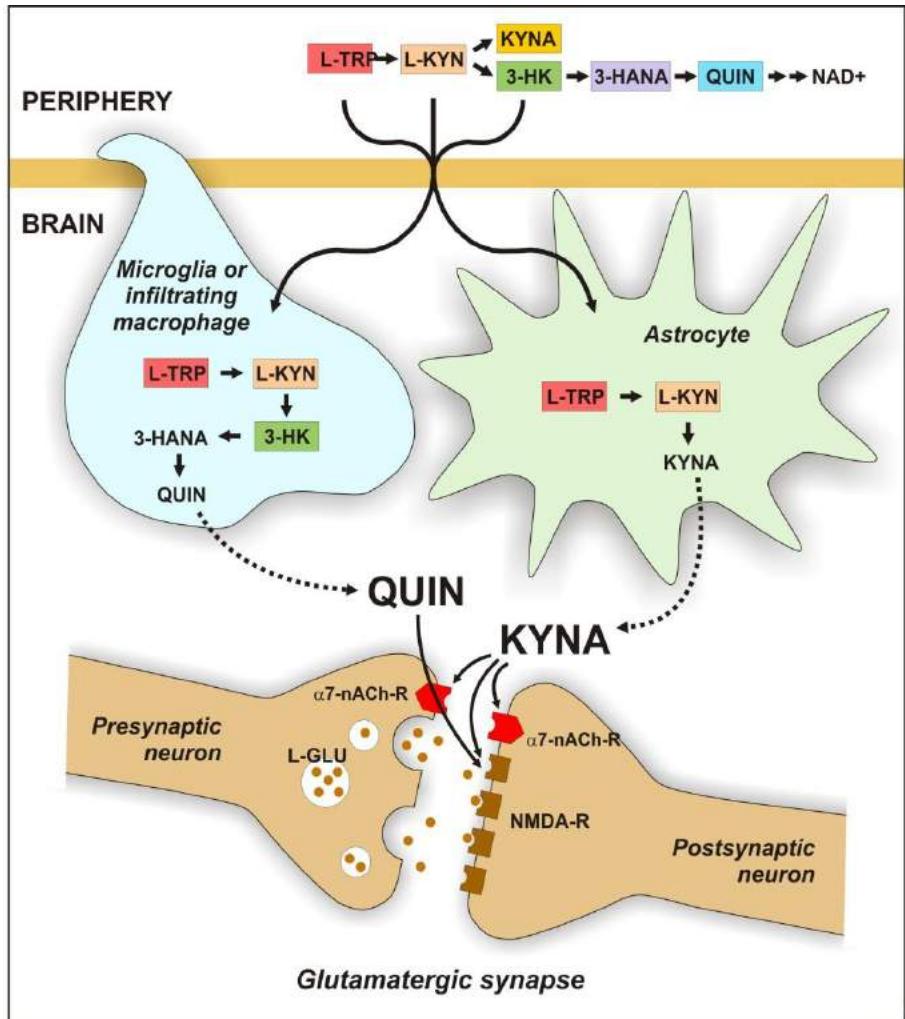


GI Effects

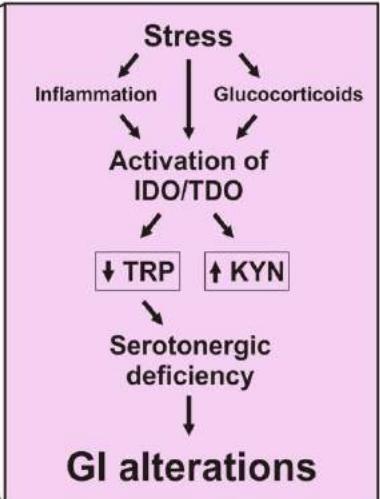
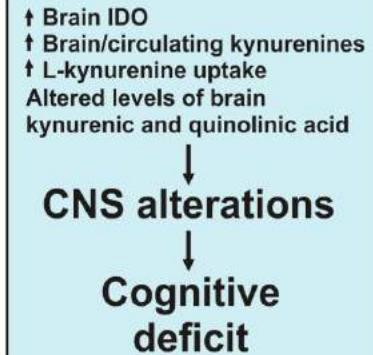
- Gastric secretion
- Gastrointestinal motility
- Intestinal secretions
- Colonic tone
- Pancreatic secretion



The Kynurenine Pathway



Altered TRP metabolism:



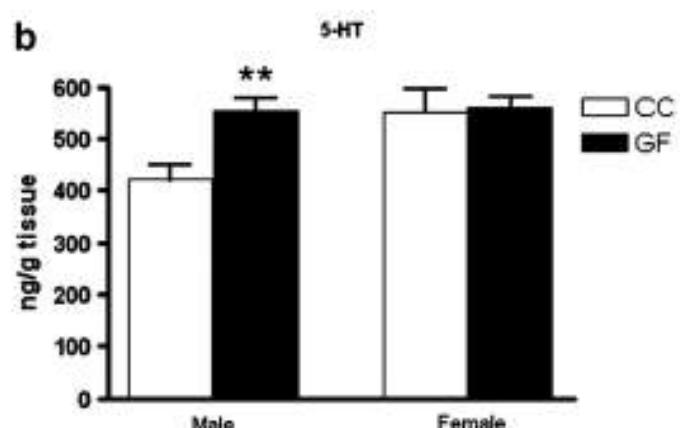
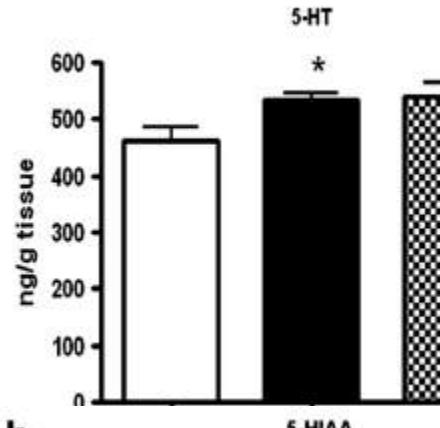
Microbiota Regulates Tryptophan Metabolism

ORIGINAL ARTICLE

The microbiome-gut-brain axis during early life regulates the hippocampal serotonergic system in a sex-dependent manner

G Clarke^{1,2}, S Grenham¹, P Scully¹, P Fitzgerald¹, RD Moloney¹, F Shanahan^{1,3}, TG Dinan^{1,2} and JF Cryan^{1,4}

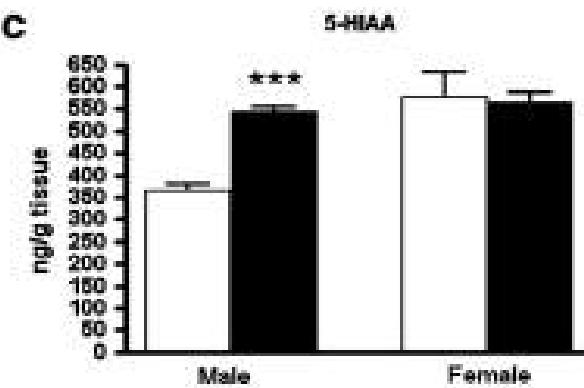
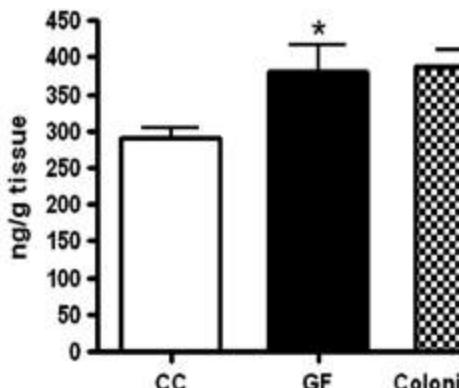
a



CC
GF
Colonised GF

CC
GF
Colonised GF

b



CC
GF
Colonised GF

OPEN

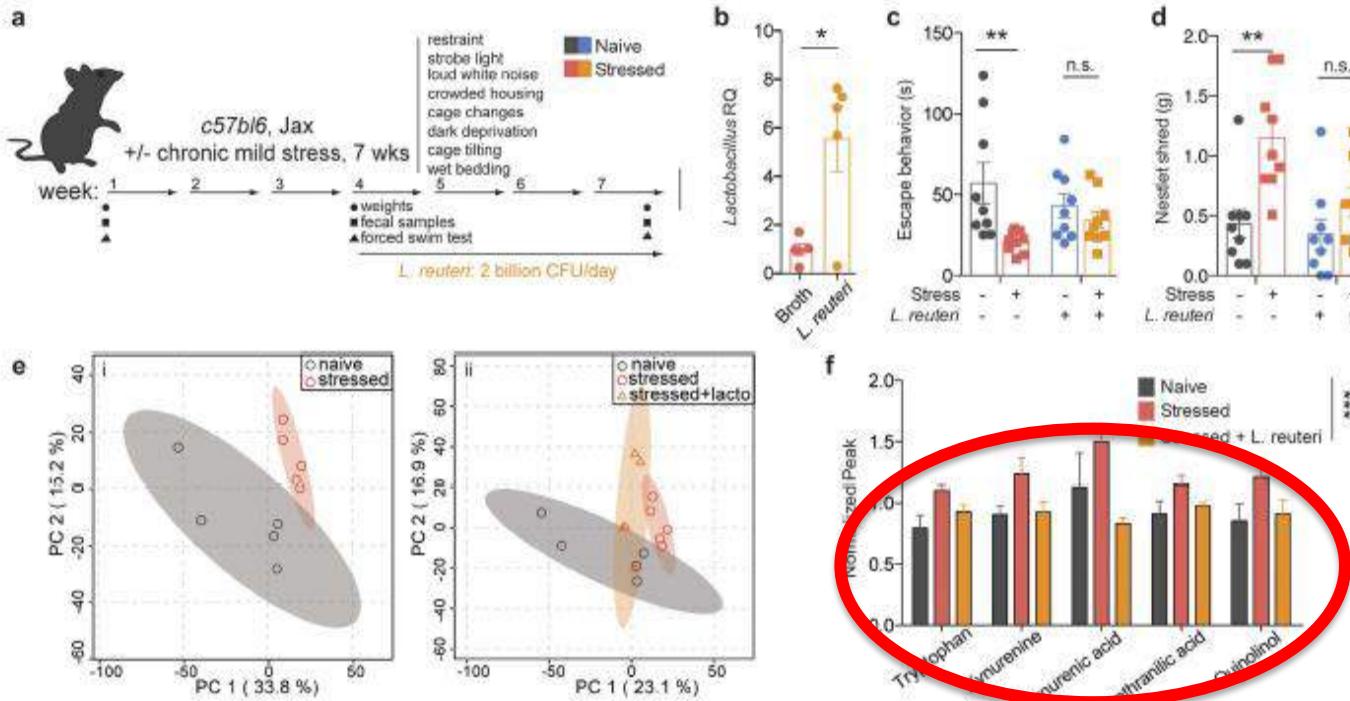
Microbiota alteration is associated with the development of stress-induced despair behavior

Received: 11 October 2016

Accepted: 31 January 2017

Published: 07 March 2017

Ioana A. Marin^{1,2,3}, Jennifer E. Goertz^{1,2}, Tiantian Ren⁴, Stephen S. Rich⁵,
Suna Onengut-Gumuscu⁵, Emily Farber⁵, Martin Wu⁴, Christopher C. Overall^{1,2},
Jonathan Kipnis^{1,2,3,*} & Alban Gaultier^{1,2,3,*}



Restoring intestinal *Lactobacillus* levels normalized stress-induced behavior and suppressed kynurenic acid production

Figure 3. Treatment with probiotic *L. reuteri* ameliorates the escape behavior induced by chronic stress.

Indigenous Bacteria from the Gut Microbiota Regulate Host Serotonin Biosynthesis

Jessica M. Yano,¹ Kristie Yu,¹ Gregory P. Donaldson,¹ Gauri G. Shastri,¹ Phoebe Ann,¹ Liang Ma,² Cathryn R. Nagler,³ Rustem F. Ismagilov,² Sarkis K. Mazmanian,¹ and Elaine Y. Hsiao^{1,*}

¹Division of Biology and Biological Engineering, California Institute of Technology, Pasadena, CA 91125, USA

²Division of Chemistry and Chemical Engineering, California Institute of Technology, Pasadena, CA 91125, USA

³Department of Pathology and Department of Medicine, I

*Correspondence: ehsiao@caltech.edu

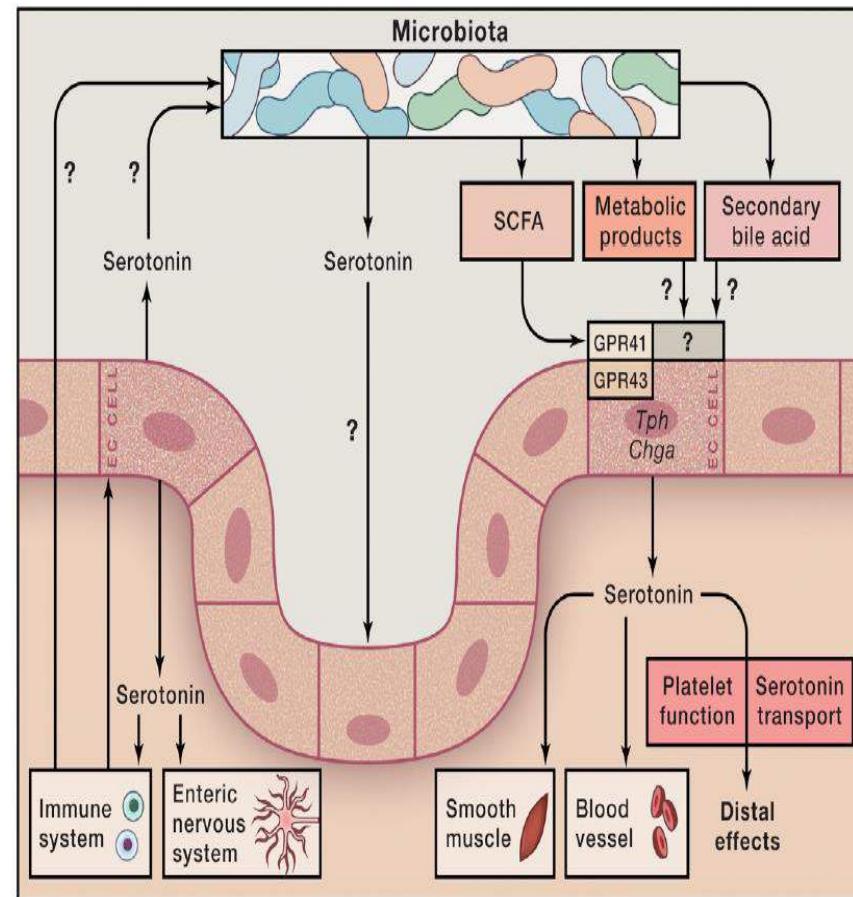
<http://dx.doi.org/10.1016/j.cell.2015.02.047>

Leading Edge

Previews

Gut Microbiota: The Link to Your Second Brain

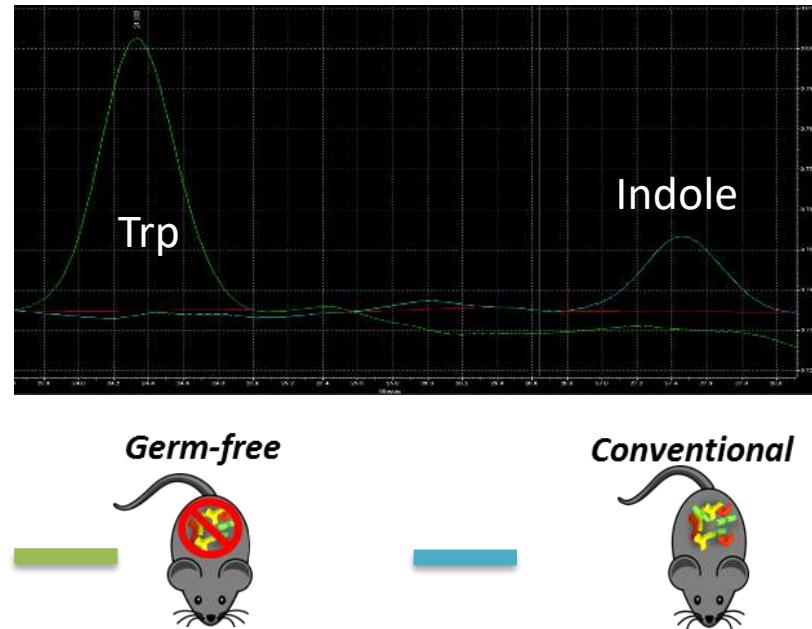
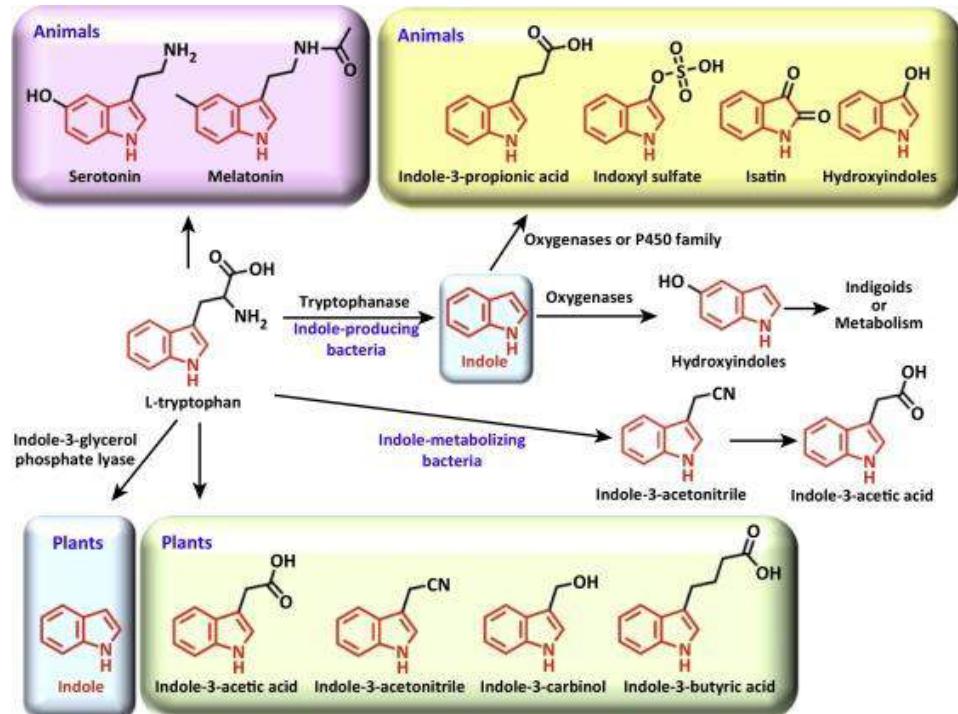
Vanessa Ridaura^{1,2} and Yasmine Belkaid^{1,2,*}



Review

Roles of Indole as an Interspecies and Interkingdom Signaling Molecule

Jin-Hyung Lee,¹ Thomas K. Wood,² and Jintae Lee^{1,*}



Lyte et al., Unpublished data

Microglial control of astrocytes in response to microbial metabolites

Veit Rothhammer¹, Davis M. Borucki¹, Emily C. Tjon¹, Maisa C. Takenaka¹, Chun-Cheih Chao¹, Alberto Ardura-Fabregat², Kalil Alves de Lima¹, Cristina Gutiérrez-Vázquez¹, Patrick Hewson¹, Ori Staszewski², Manon Blain³, Luke Healy³, Tradite Neziraj¹, Matilde Borio¹, Michael Wheeler¹, Loic Lionel Dragin⁴, David A. Laplaud⁵, Jack Antel³, Jorge Ivan Alvarez⁴, Marco Prinz^{2,6} & Francisco J. Quintana^{1,7*}

NEWS & VIEWS

| NATURE | 1

<https://doi.org/10.1038/d41586-018-05113-0>

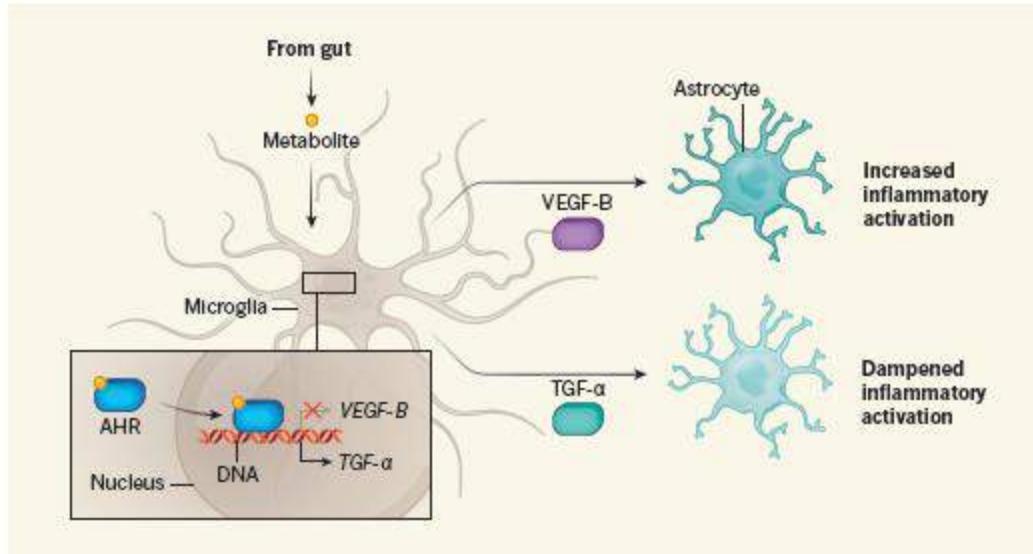
IMMUNOLOGY

Gut molecules control brain inflammation

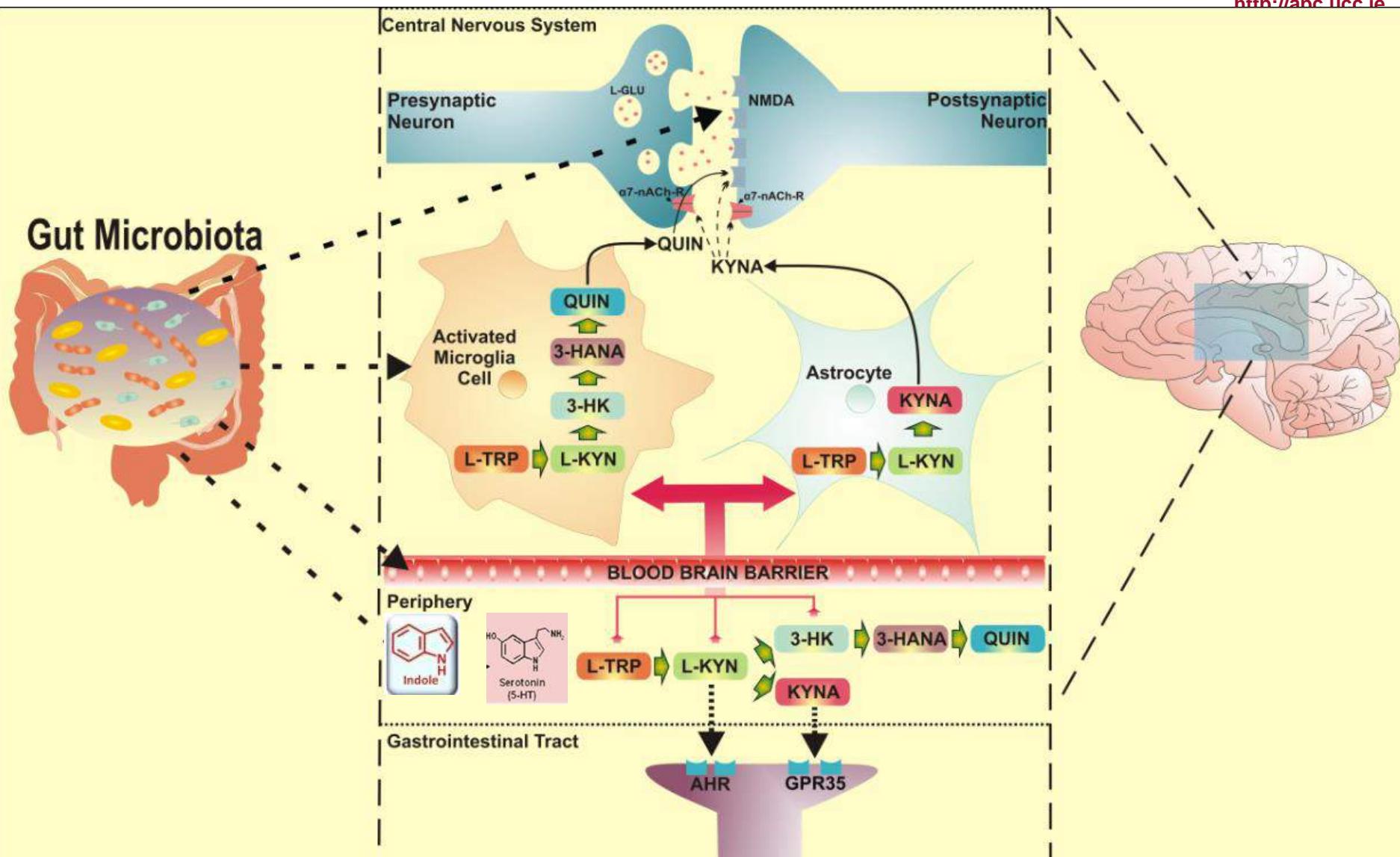
Metabolite molecules produced by the gut's microbes activate immune cells in the brain called microglia, which signal to astrocyte cells to mediate responses to inflammation in the central nervous system.

HARTMUT WEKERLE

microglia inhibits inflammation in the CNS.



Summary



Translational Research

Larson

ARTICLE IN PRESS

Annals of Epidemiology xxx (2016) 1–7

Contents lists available at ScienceDirect

Annals of Epidemiology

journal homepage: www.annalsofepidemiology.org

 ELSEVIER



Review article

Brain-gut-microbiota axis: challenges for translation in psychiatry

John R. Kelly MD^{a,b}, Gerard Clarke PhD^{a,b}, John F. Cryan PhD^{a,c}, Timothy G. Dinan MD, PhD^{a,b,*}

^aAlimentary Pharmabiotic Centre, APC Microbiome Institute, University College Cork, Cork, Ireland

^bDepartment of Psychiatry and Neurobehavioural Science, University College Cork, Cork, Ireland

^cDepartment of Anatomy and Neuroscience, University College Cork, Cork, Ireland

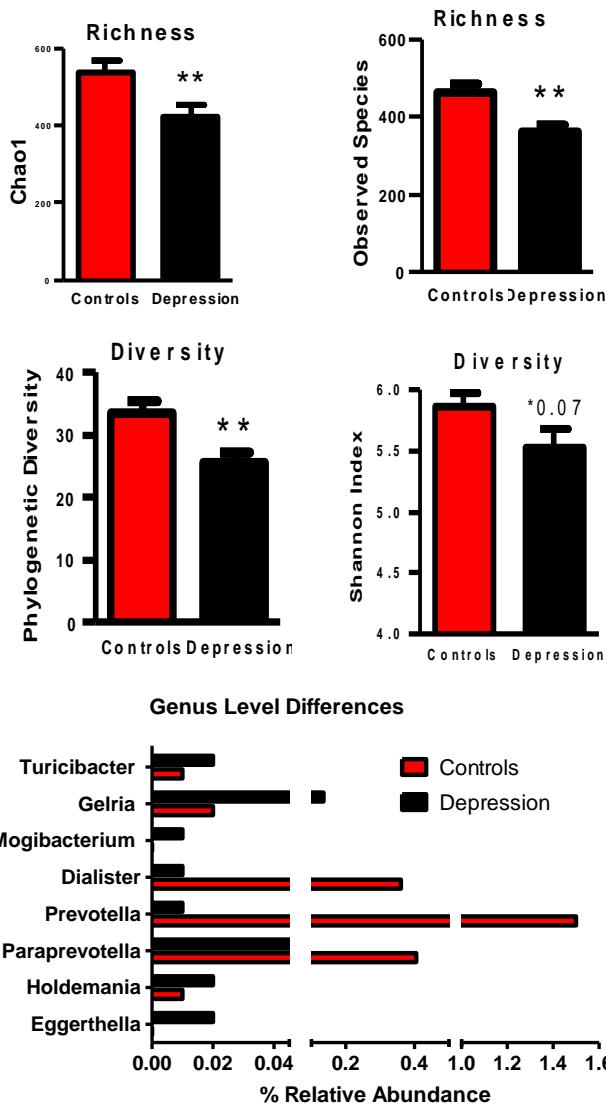
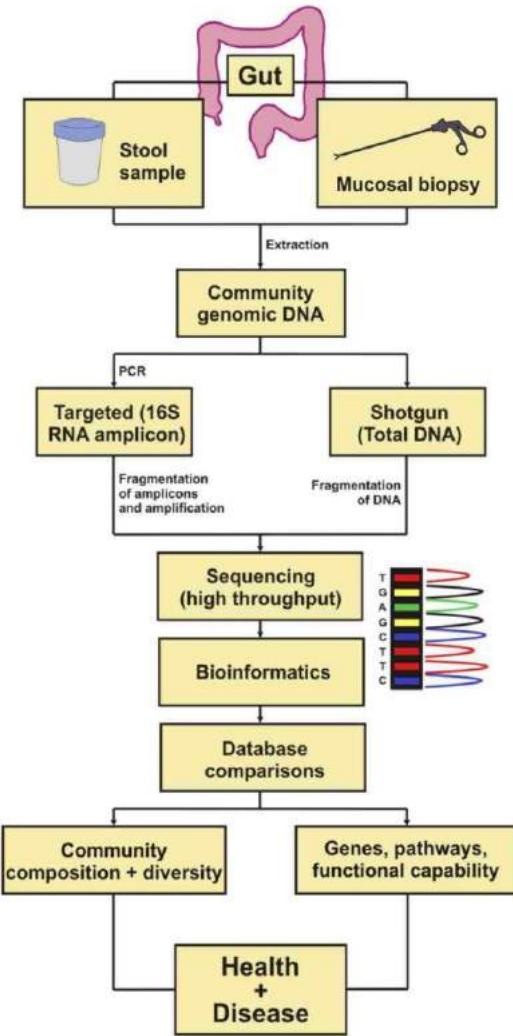
How to recognize the moods of an Irish setter

Cryan et al., Trends in Pharmacol. Sci. 2002

Gary Larson



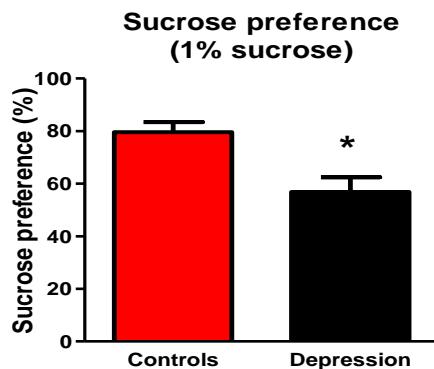
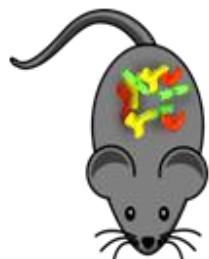
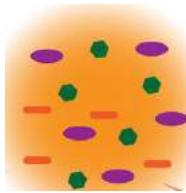
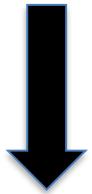
Altered Microbiota in Depression



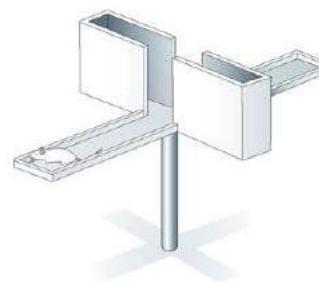
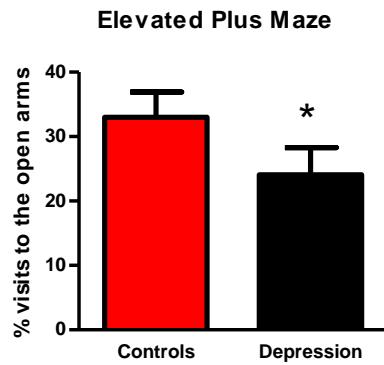
Reduced microbial diversity in depression

Prevotella, a genus of Gram-negative bacteria, is reduced in depression

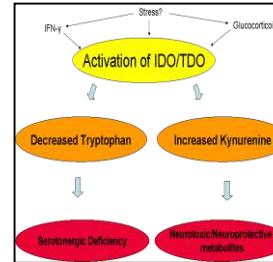
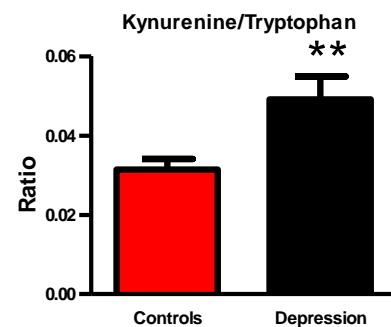
Transfer of Depressive Phenotype



Anhedonia-like behaviours transferred via gut microbiota



Anxiety-like behaviours transferred via gut microbiota

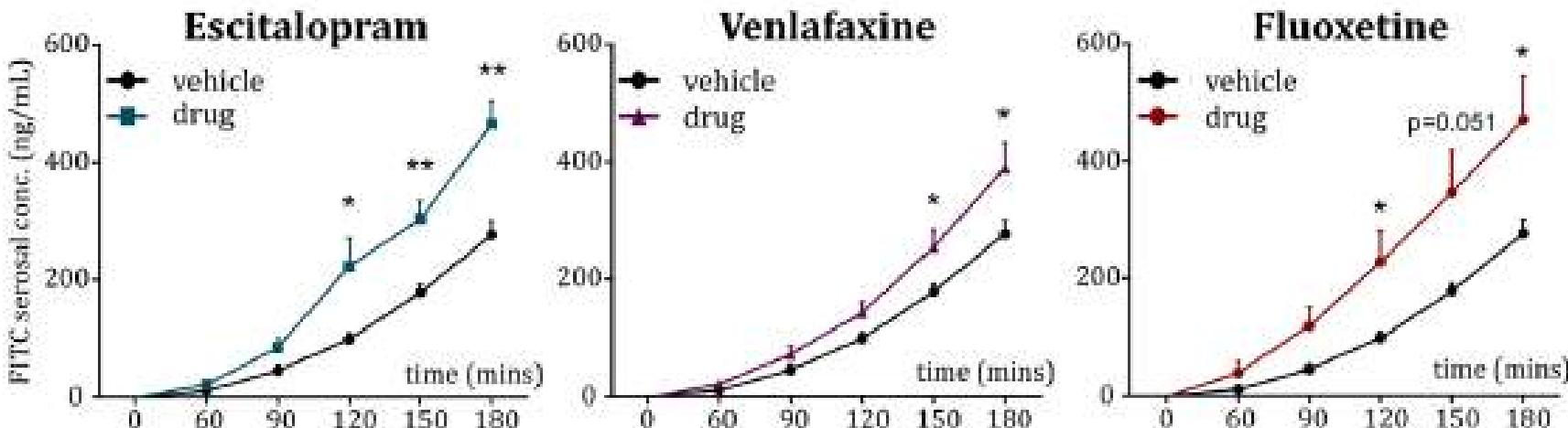


Tryptophan metabolism Profile transferred via gut microbiota

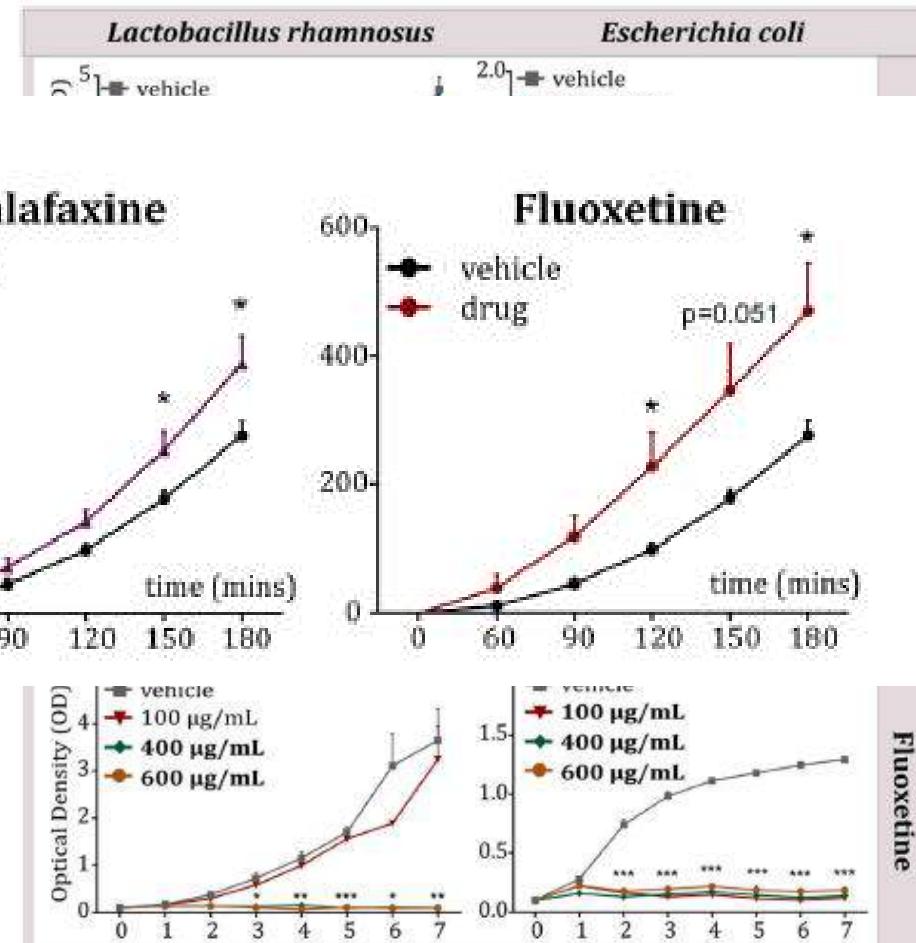
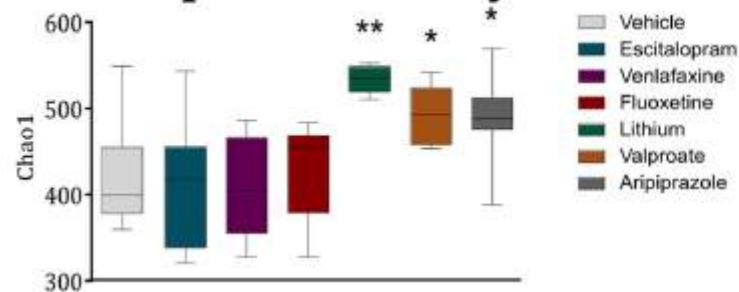
Differential effects of psychotropic drugs on microbiome composition and gastrointestinal function

Sofia Cussotto^{1,2} · Conall R. Strain^{1,3} · Fiona Fouhy^{1,3} · Ronan G. Strain^{1,3} · Veronica L. Peterson^{1,2} · Gerard Clarke^{1,4} · Catherine Stanton^{1,3,4} · Timothy G. Dinan^{1,4} · John F. Cryan^{1,2}

a Intestinal Permeability - Ileum



a Alpha Diversity



OPEN

Citation: *Transl Psychiatry* (2013) 3, e309; doi:10.1038/tp.2013.83

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www.nature.com/tp

ORIGINAL ARTICLE

Antipsychotics and the gut microbiome: olanzapine-induced metabolic dysfunction is attenuated by antibiotic administration in the rat

KJ Davey^{1,2}, PD Cotter^{1,3}, O O'Sullivan^{1,3}, F Crispie³, TG Dinan^{1,4}, JF Cryan^{1,5} and SM O'Mahony^{1,5}

Psychopharmacology (2012) 221:155–169

DOI 10.1007/s00213-011-2555-2

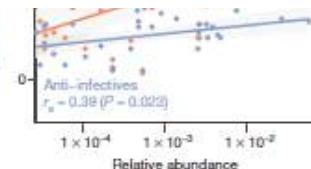
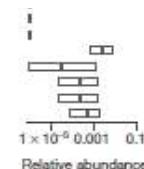
ORIGINAL INVESTIGATION

Gender-dependent consequences of chronic olanzapine in the rat: effects on body weight, inflammatory, metabolic and microbiota parameters

Kieran J. Davey · Siobhain M. O'Mahony · Harriet Schellekens · Orla O'Sullivan · John Bienenstock · Paul D. Cotter · Timothy G. Dinan · John F. Cryan



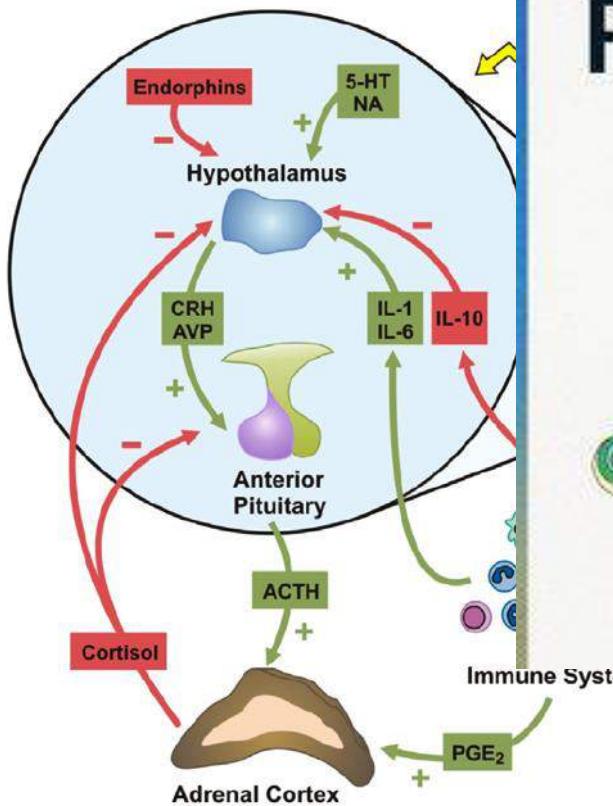
Clostridium difficile
Fusobacterium nucleatum
Bacteroides xylosoxolvens
Akkermansia muciniphila
Escherichia coli [A]1
Escherichia coli ED1a
Bilophila wadsworthia



REVIEW**Psychobiotics:**

Timothy G. Dinan, Catherine Stanton

Here, we define a psychobiotic as a live microorganism that confers a health benefit on the host beyond its nutritional value. As such as gamma-aminobutyric acid and serotonin, psychobiotics possess antidepressant properties. So far, psychobiotics have been shown to alleviate depression syndrome, where positive benefits are emerging of benefits in alleviating symptoms of anxiety and depression. The inflammatory actions of certain psychobiotics in the gut may explain the scale placebo-controlled studies are needed.

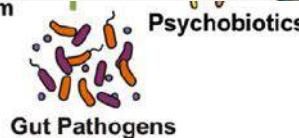


"Up-to-the-minute research and practical advice on the gut-brain axis—perhaps the most exciting area of science today!"
—ROB KNIGHT, author of *Follow Your Gut*

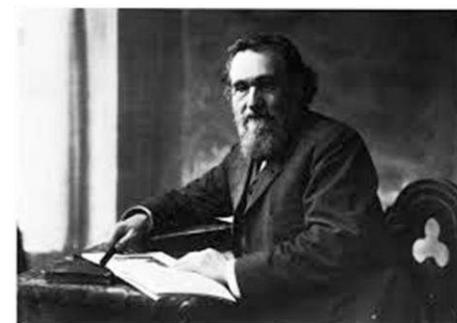
THE PSYCHOBIOYTIC REVOLUTION

Mood, Food, and the New Science of the Gut-Brain Connection

SCOTT C. ANDERSON

with
JOHN F. CRYAN, PH.D. &
TED DINAN, M.D., PH.D.**opic**

duces a health benefit in patients delivering neuroactive substances in rodents suggests that certain active spinal cord, or neuroendocrine changes in patients with irritable bowel syndrome. Evidence is emerging that these benefits may be related to the anti-inflammatory activity. Results from large

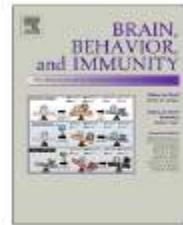


Elie Metchnikoff (1845-1916)
Nobel Prize 1908



Contents lists available at ScienceDirect

Brain, Behavior, and Immunity

journal homepage: www.elsevier.com/locate/ybrbi

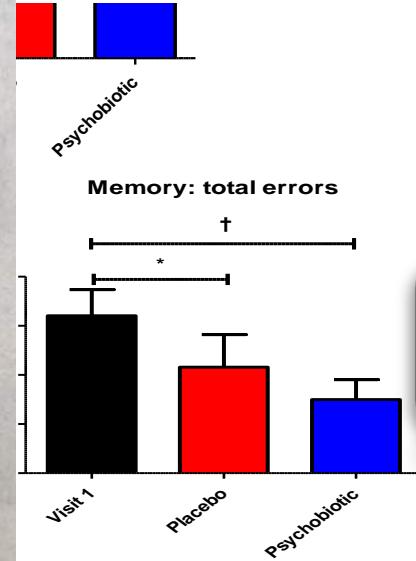
Full-length Article

Lost in translation? The potential psychobiotic *Lactobacillus rhamnosus* (JB-1) fails to modulate stress or cognitive performance in healthy male subjects

John R. Kelly^{a,b}, Andrew P. Allen^{a,b}, Andriy Temko^c, William Hutch^d, Paul J. Kennedy^a, Niloufar Farid^b, Eileen Murphy^e, Geraldine Boylan^d, John Bienenstock^f, John F. Cryan^{a,g}, Gerard Clarke^{a,b}, Timothy G. Dinan^{a,b,*}



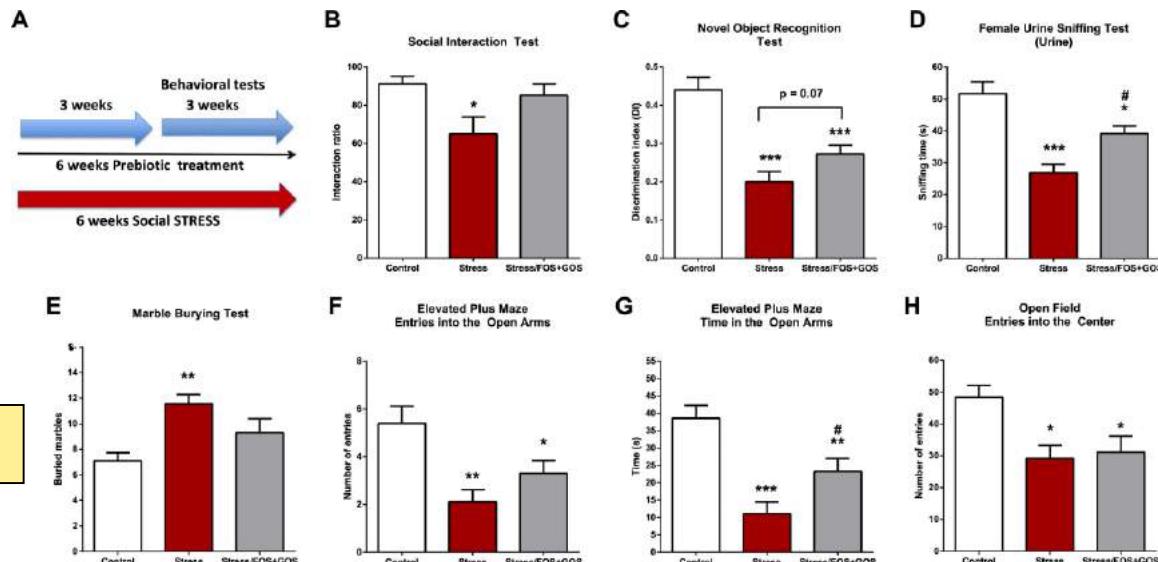
Fz Mobility (Hz)



Targeting the Microbiota-Gut-Brain Axis: Prebiotics Have Anxiolytic and Antidepressant-like Effects and Reverse the Impact of Chronic Stress in Mice

Aurelijus Burokas, Silvia Arboleya, Rachel D. Moloney, Veronica L. Peterson, Kiera Murphy, Gerard Clarke, Catherine Stanton, Timothy G. Dinan, and John F. Cryan

Prebiotic reverses consequences of chronic stress

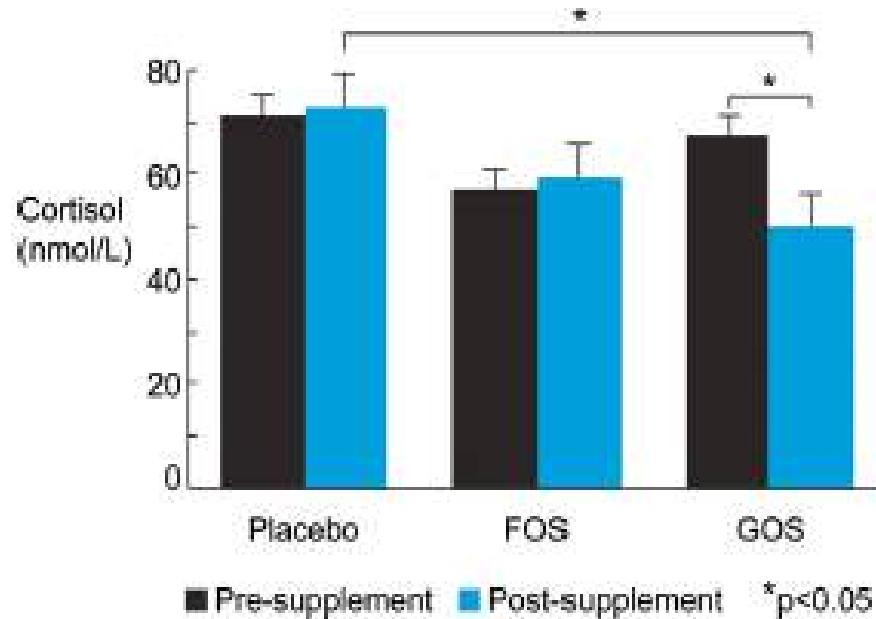


Anxiety

ORIGINAL INVESTIGATION

Prebiotic intake reduces the waking cortisol response and alters emotional bias in healthy volunteers

Kristin Schmidt · Philip J. Cowen · Catherine J. Harmer ·
George Tzortzis · Steven Errington · Philip W. J. Burnet

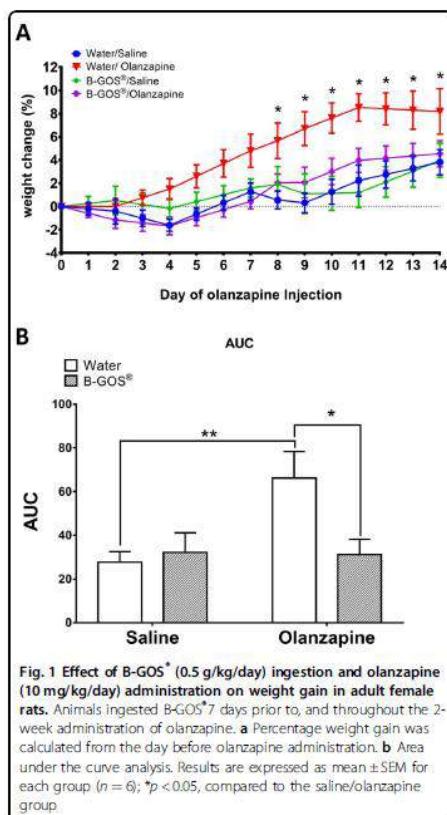


ARTICLE

Open Access

Prebiotic attenuation of olanzapine-induced weight gain in rats: analysis of central and peripheral biomarkers and gut microbiota

Amy Chia-Ching Kao¹, Sonia Spitzer¹, Daniel C. Anthony², Belinda Lennox¹ and Philip W. J. Burnet¹



Let food be thy medicine

Feeding the microbiota-gut-brain axis: diet, microbiome, and neuropsychiatry



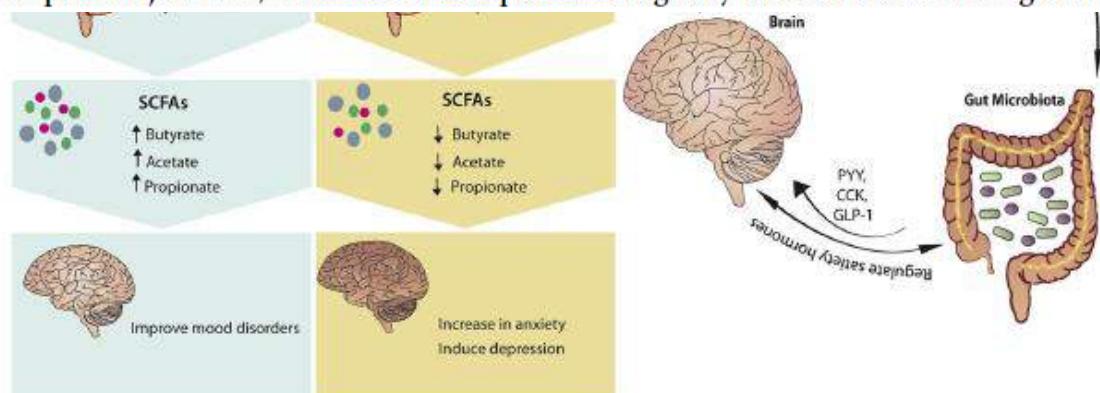
Nutritional medicine as mainstream in psychiatry

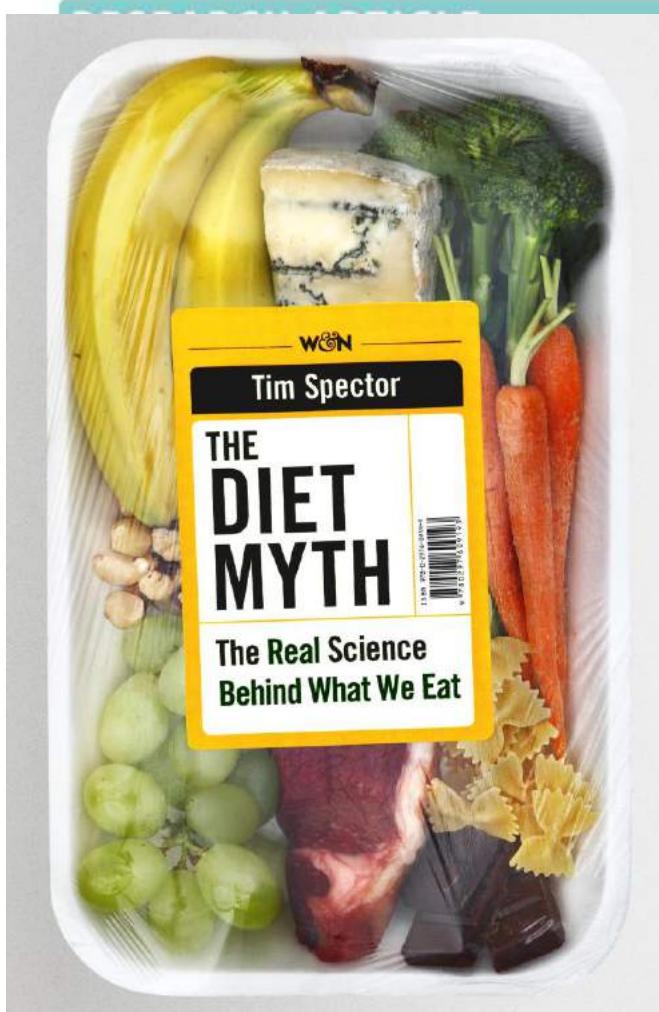


CrossMark

Jerome Sarris, Alan C Logan, Tasnime N Akbaraly, G Paul Amminger, Vicent Balanzá-Martínez, Marlene P Freeman, Joseph Hibbeln, Yutaka Matsuoka, David Mischoulon, Tetsuya Mizoue, Akiko Nanri, Daisuke Nishi, Drew Ramsey, Julia J Rucklidge, Almudena Sanchez-Villegas, Andrew Scholey, Kuan-Pin Su, Felice N Jacka, on behalf of The International Society for Nutritional Psychiatry Research

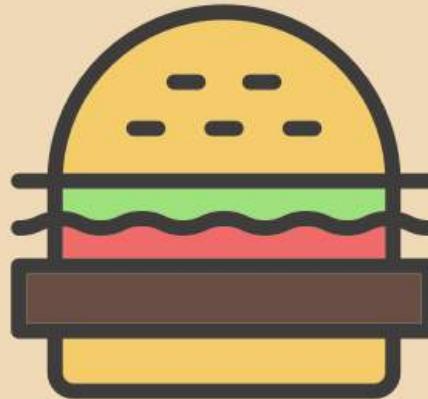
Psychiatry is at an important juncture, with the current pharmacologically focused model having achieved modest Lancet Psychiatry 2015





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10 Days



Professor Spector's son Tom spent 10 days eating only McDonald's



Fig. 2 MADRS scores for dietary support and social support control groups at baseline and endpoint. Effect size: Cohen's $d = -1.16$ (95% CI $-1.73, -0.59$). Baseline data $n = 67$; 12 week data $n = 56$

REVIEW

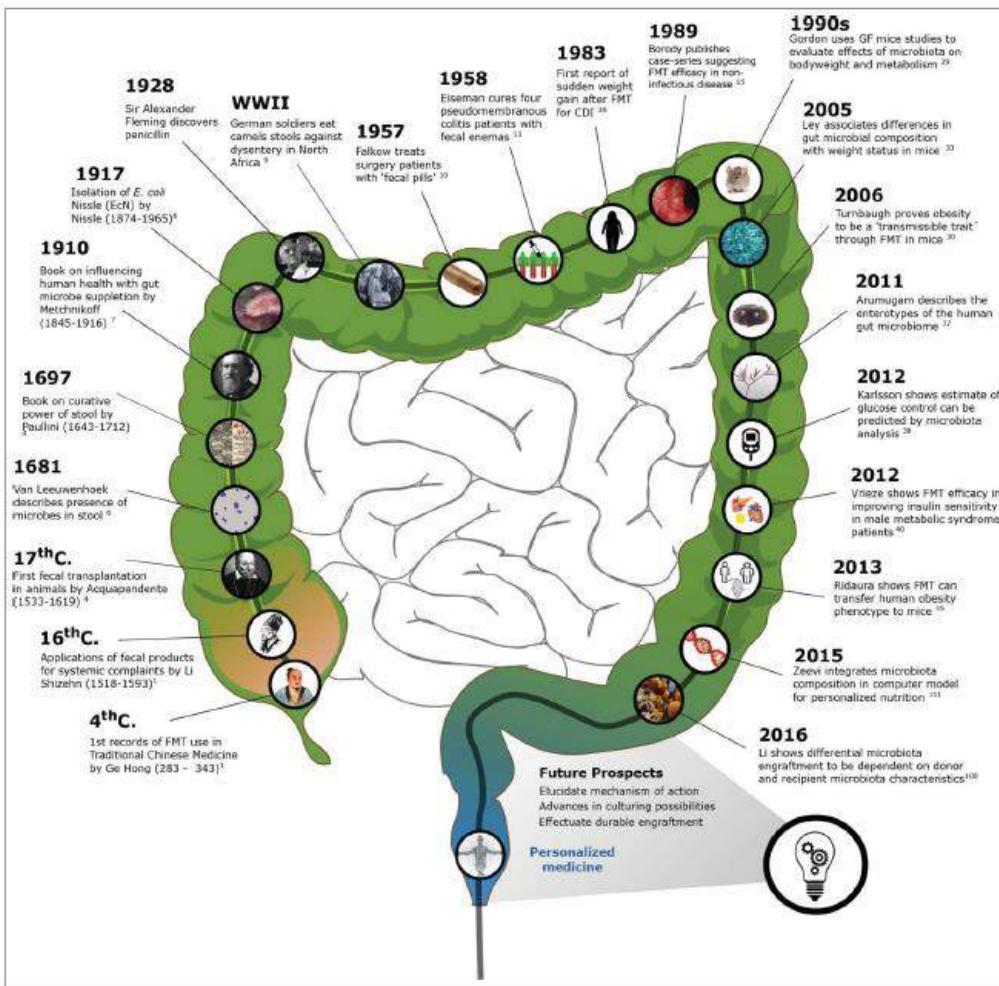
OPEN ACCESS

Fecal microbiota transplantation in metabolic syndrome: History, present and future

P. F. de Groot^a, M. N. Frissen  ^a, N. C. de Clercq^a, and M. Nieuwdorp^{a,b,c,d}



Andrea Levy, *The Plain*



The New York Times
HEALTH
A Promising

By PAM BELLUCK OCT. 11, 2014



therapy

Thursday, October 25, 2012
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fecal transplants?



European consensus conference on faecal microbiota transplantation in clinical practice

Giovanni Cammarota,¹ Gianluca Ianiro,¹ Herbert Tilg,² Mirjana Rajilić-Stojanović,³ Patrizia Kump,⁴ Reetta Satokari,⁵ Harry Sokol,⁶ Perttu Arkkila,⁷ Cristina Pintus,⁸ Ailsa Hart,⁹ Jonathan Segal,⁹ Marina Alois,¹⁰ Luca Masucci,¹¹ Antonio Molinaro,¹² Franco Scaldaferri,¹ Giovanni Gasbarrini,¹ Antonio Lopez-Sanroman,¹³ Alexander Link,¹⁴ Pieter de Groot,¹⁵ Willem M de Vos,^{5,16} Christoph Högenauer,⁴ Peter Malfertheiner,¹⁴ Eero Mattila,¹⁷ Tomica Milosavljević,¹⁸ Max Nieuwdorp,^{12,15,19} Maurizio Sanguinetti,¹¹ Magnus Simren,²⁰ Antonio Gasbarrini,¹ The European FMT Working Group

Box 1 Key issues to select potential donors at the preliminary interview

INFECTIOUS DISEASES

- History of, or known exposure to, HIV, HBV or HCV, syphilis, human T-lymphotropic virus I and II, malaria, trypanosomiasis, tuberculosis
- Known systemic infection not controlled at the time of donation
- Use of illegal drugs
- Risky sexual behaviour (anonymous sexual contacts; sexual contacts with prostitutes, drug addicts, individuals with HIV, viral hepatitis, syphilis; work as prostitute; history of sexually transmittable disease)
- Previous reception of tissue/organ transplant
- Previous (<12 months) reception of blood products
- Recent (<6 months) needle stick accident
- Recent (<6 months) body tattoo, piercing, earring, acupuncture
- Recent medical treatment in poorly hygienic conditions
- Risk of transmission of diseases caused by prions
- Recent parasitosis or infection from rotavirus, *Giardia lamblia* and other microbes with GI involvement
- Recent (<6 months) travel in tropical countries, countries at high risk of communicable diseases or traveller's diarrhoea
- Recent (<6 months) history of vaccination with a live attenuated virus, if there is a possible risk of transmission
- Healthcare workers (to exclude the risk of transmission of multidrug-resistant organisms)
- Individual working with animals (to exclude the risk of transmission of zoonotic infections)

GI, METABOLIC AND NEUROLOGICAL DISORDERS

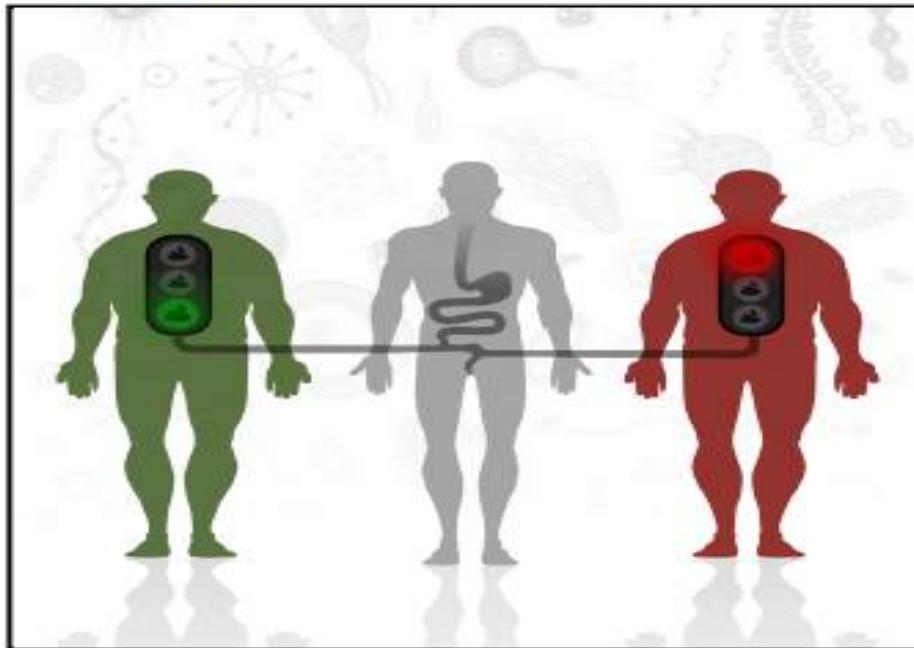
- History of IBS, IBD, functional chronic constipation, coeliac disease, other chronic GI disorders
- History of chronic, systemic autoimmune disorders with GI involvement
- History of, or high risk for, GI cancer or polypsis
- Recent appearance of diarrhoea, haematochezia
- History of neurological/neurodegenerative disorders
- History of psychiatric conditions
- Overweight and obesity (body mass index >25)

DRUGS THAT CAN IMPAIR GUT MICROBIOTA COMPOSITION

- Recent (<3 months) exposure to antibiotics, immunosuppressants, chemotherapy
- Chronic therapy with proton pump inhibitors

Improvement of Insulin Sensitivity after Lean Donor Feces in Metabolic Syndrome Is Driven by Baseline Intestinal Microbiota Composition

Graphical Abstract



Highlights

- Lean donor FMT in obese metabolic syndrome patients improves insulin sensitivity
- Beneficial effects of lean donor FMT are transient
- Improvement in insulin sensitivity is linked to changes in plasma metabolites
- Response to lean donor FMT is driven by baseline fecal microbiota composition

Authors

Ruud S. Kootte, Evgeni Levin,
Jarkko Salojärvi, ..., Erik S.G. Stroes,
Albert K. Groen, Max Nieuwdorp

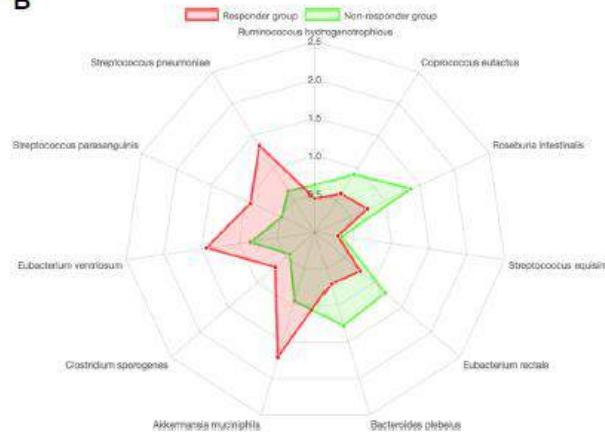
Correspondence

m.nieuwdorp@amc.uva.nl

In Brief

Kootte et al. show that fecal microbiota transplantation from lean donors to obese patients with metabolic syndrome improves insulin sensitivity, a transient effect associated with changes in microbiota composition and fasting plasma metabolites. Baseline fecal microbiota composition in recipients predicts the response to lean donor fecal microbiota transplantation.

B



Early Life Factors

ACTA PÆDIATRICA
NURTURING THE CHILD

Acta Paediatrica ISSN 0803-5253

REVIEW ARTICLE

Priming for health: gut microbiota acquired in early life regulates physiology, brain and behaviour

G Clarke (g.clarke@ucc.ie)^{1,2*}, SM O'Mahony^{1,3*}, TG Dinan^{1,2}, JF Cryan^{1,3}

1.Alimentary Pharmabiotic Centre, University College Cork, Cork, Ireland

2.Department of Psychiatry, University College Cork, Cork, Ireland

3.Department of Anatomy and Neuroscience, University College Cork, Cork, Ireland

Keywords

Behaviour, Brain Development, Breastfeeding, Early Life, Microbiota

Correspondence

G Clarke, Department of Psychiatry/Alimentary Pharmabiotic Centre, 1.15 Biosciences Institute, University College Cork, Cork, Ireland.
 Tel: +353 214 901 408 |

ABSTRACT

The infant gut microbiome is dynamic, and radical shifts in composition occur during the first 3 years of life. Disruption of these developmental patterns, and the impact of the microbial composition of our gut on brain and behaviour, has attracted much recent attention. Integrating these observations is an important new research frontier.

Conclusion: Early-life perturbations of the developing gut microbiota can impact on the central nervous system and potentially lead to adverse mental health outcomes.



Drug Discovery Today • Volume 17, Numbers 9/10 • May 2012



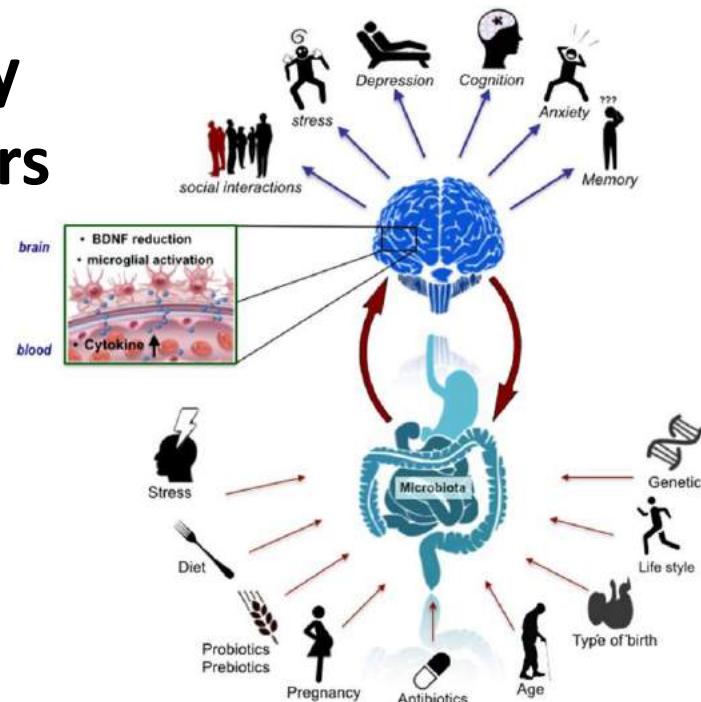
REVIEWS

Can we vaccinate against depression?

Graham A.W. Rook¹, Charles L. Raison² and Christopher A. Lowry³

Summary & Conclusions

- Gut microbiota is both stress-susceptable and can regulate stress response
- Regulates behaviours and physiology relevant to neuropsychiatric disorders
- Tryptophan availability/Kynurenone metabolism
- Translation to clinic?
- Microbial-based strategies for the treatment of stress-related disorders?



Kelly et al., Frontiers Neuroscience 2017

The allium

Science news you won't read nowhere else



Pope Francis To Award Sainthood To All Microbiome Researchers

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Vatican City – Pope Francis announced today that he was going to award automatic sainthood to all microbiome researchers worldwide for “Doing God’s Work”.

Acknowledgements



Laboratory of NeuroGastroenterology



**BRAIN &
BEHAVIOR**
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EUROPEAN COOPERATION
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Genes in irritable bowel
syndrome

CRF-C
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Interfacing Food & Medicine



The Daily Mail, London Sept 2011
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