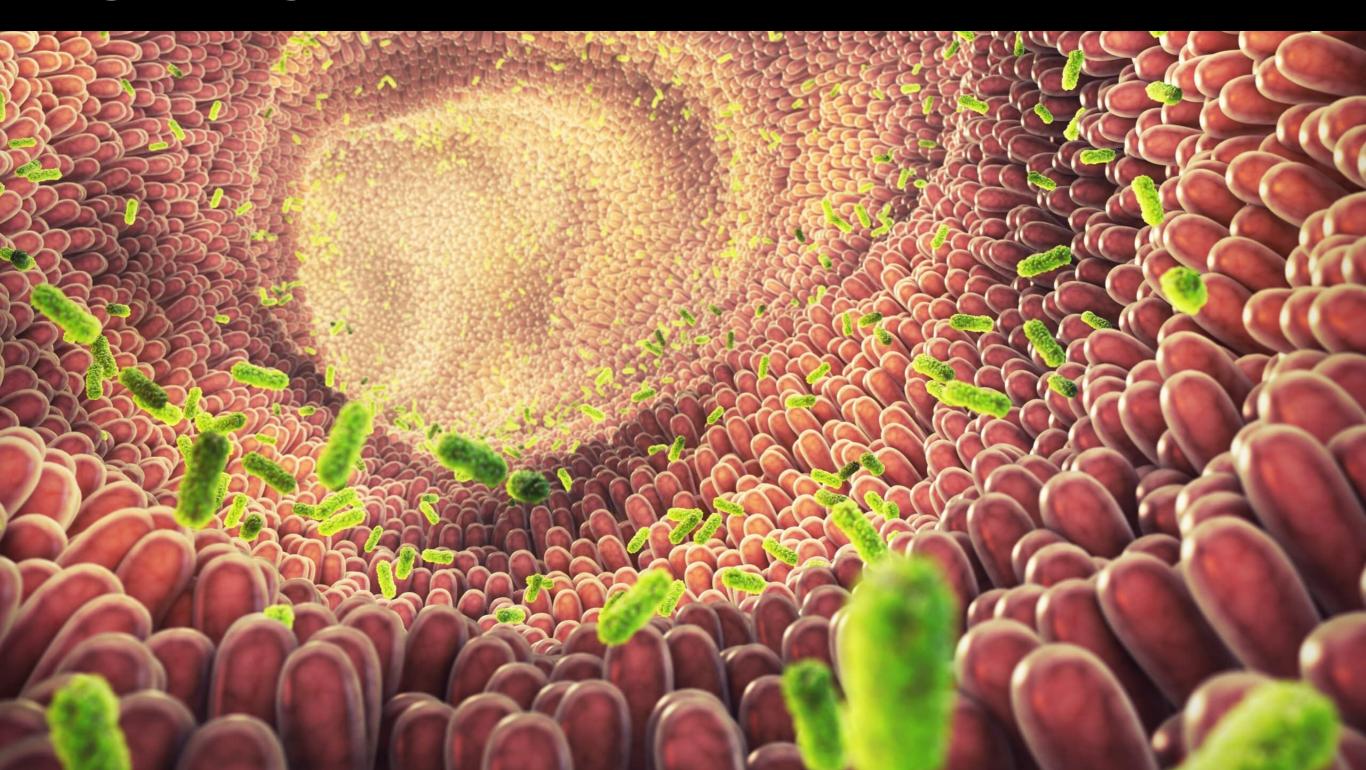
# UNSER DARM



#### "Der Darm ist das Tor zum Leben"

asiatisches Sprichwort

"Der Tod sitzt im Darm"

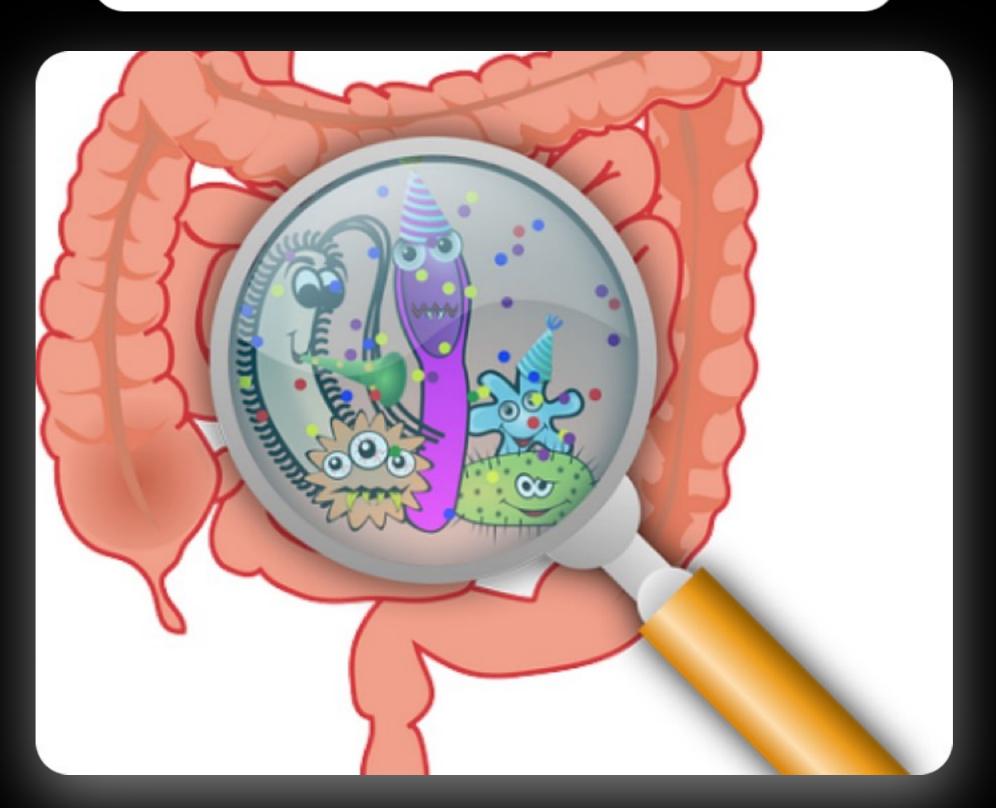
Hippokrates

# Darm gut, alles gut Pharma

«ein kranker Darm ist die Wurzel allen Übels, weil dieses Zentrum den gesamten Körper bis in die kleinste Zelle versorgt!»

Google KI

## "Neues Organ": Das Mircobiom



# WIE KOMMTS...



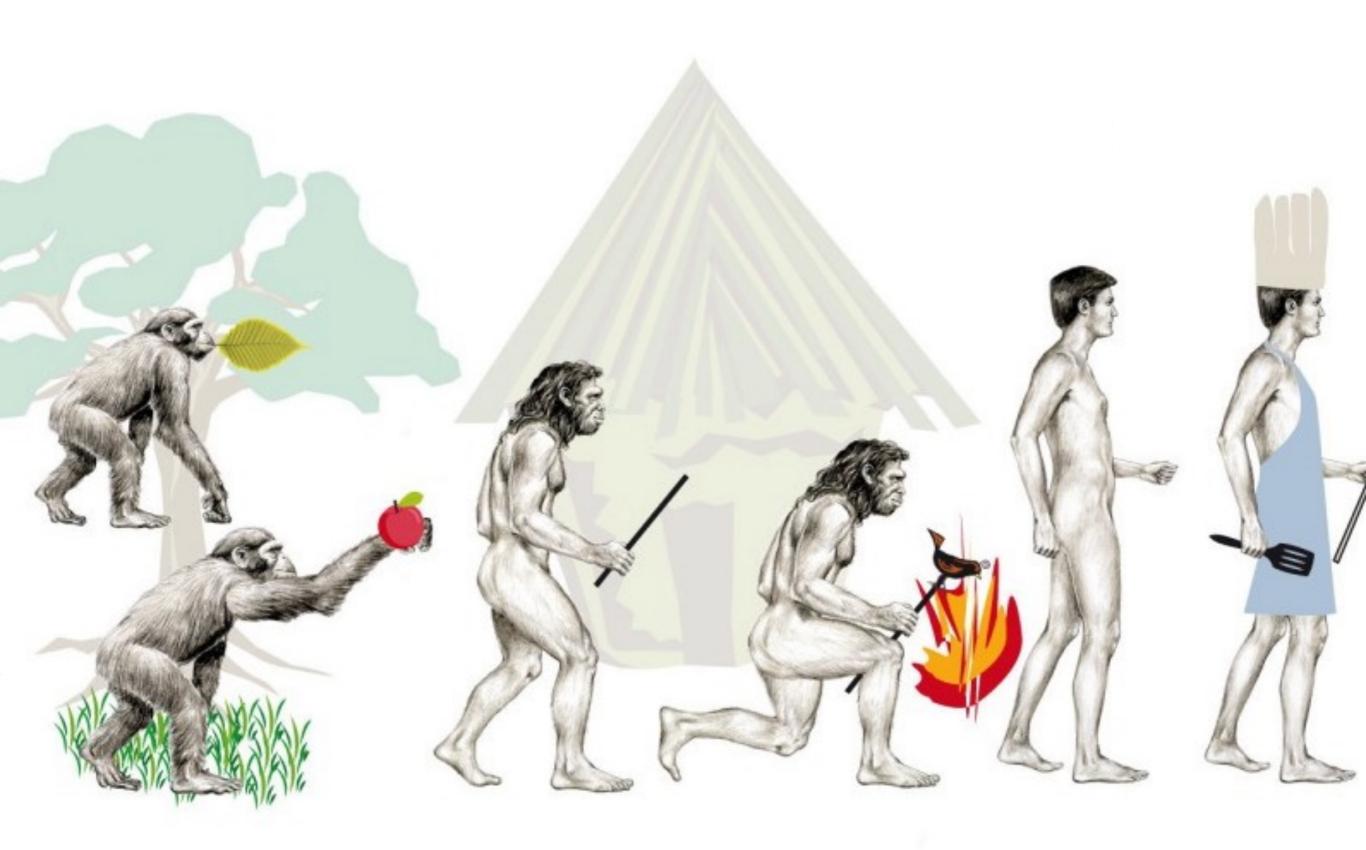
### Grundgedanken

DIE EVOLUTION DES MAGEN- DARMTRAKTES UND DIE ENTWICKLUNG DES NAHRUNGSANGEBOTES SIND UNTERSCHIEDLICH!!!

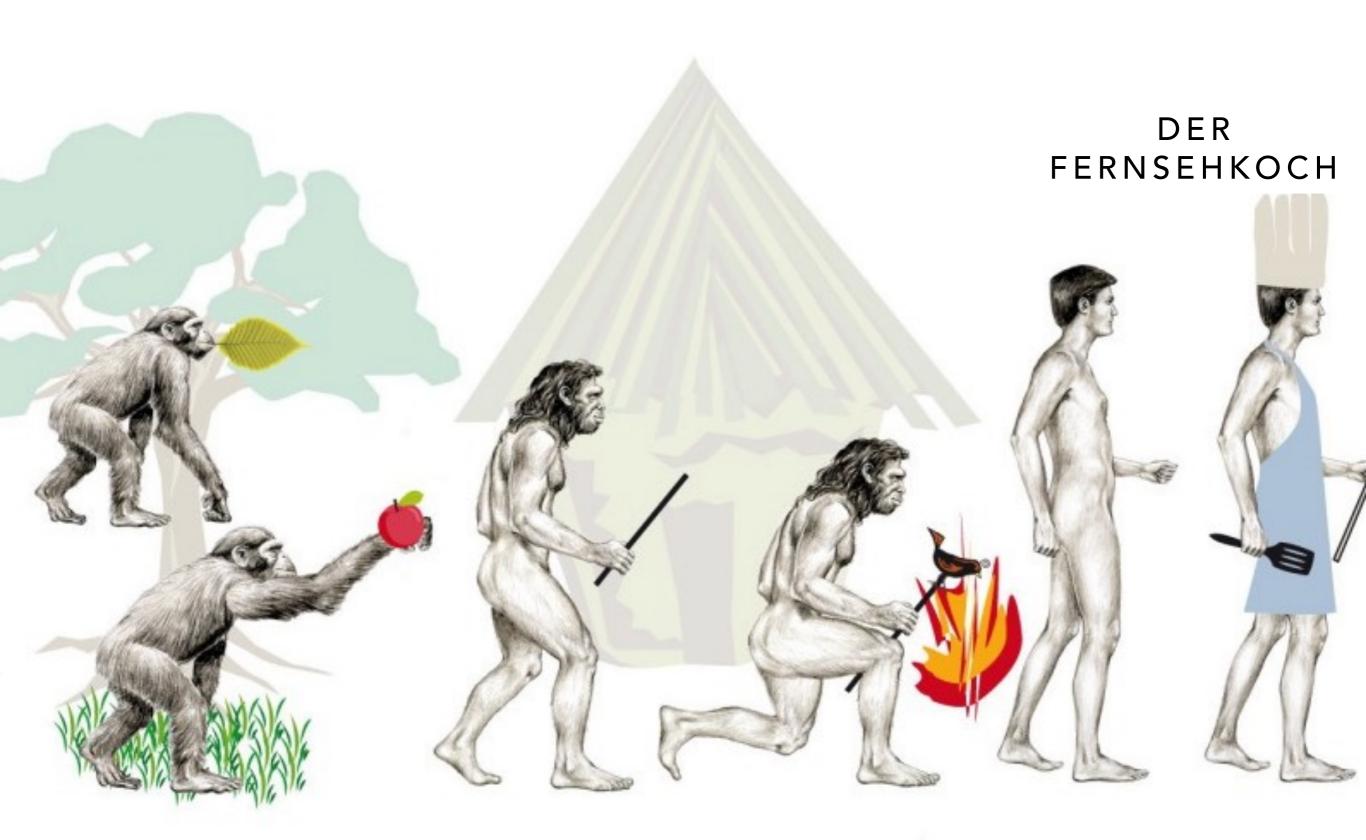
MIKROBIOM NEUES "ORGAN"

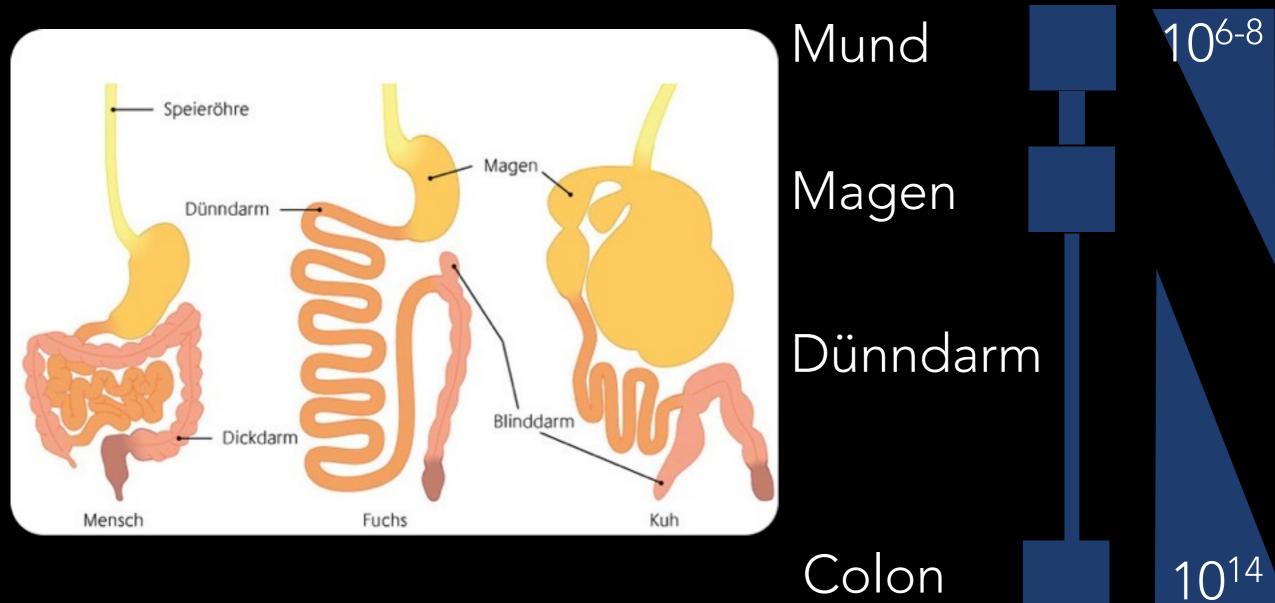
HANDEL UND PRODUKTION: STRESS ODER CHANCE

### Am Anfang war der Rohköstler



### Am Anfang war der Rohköstler

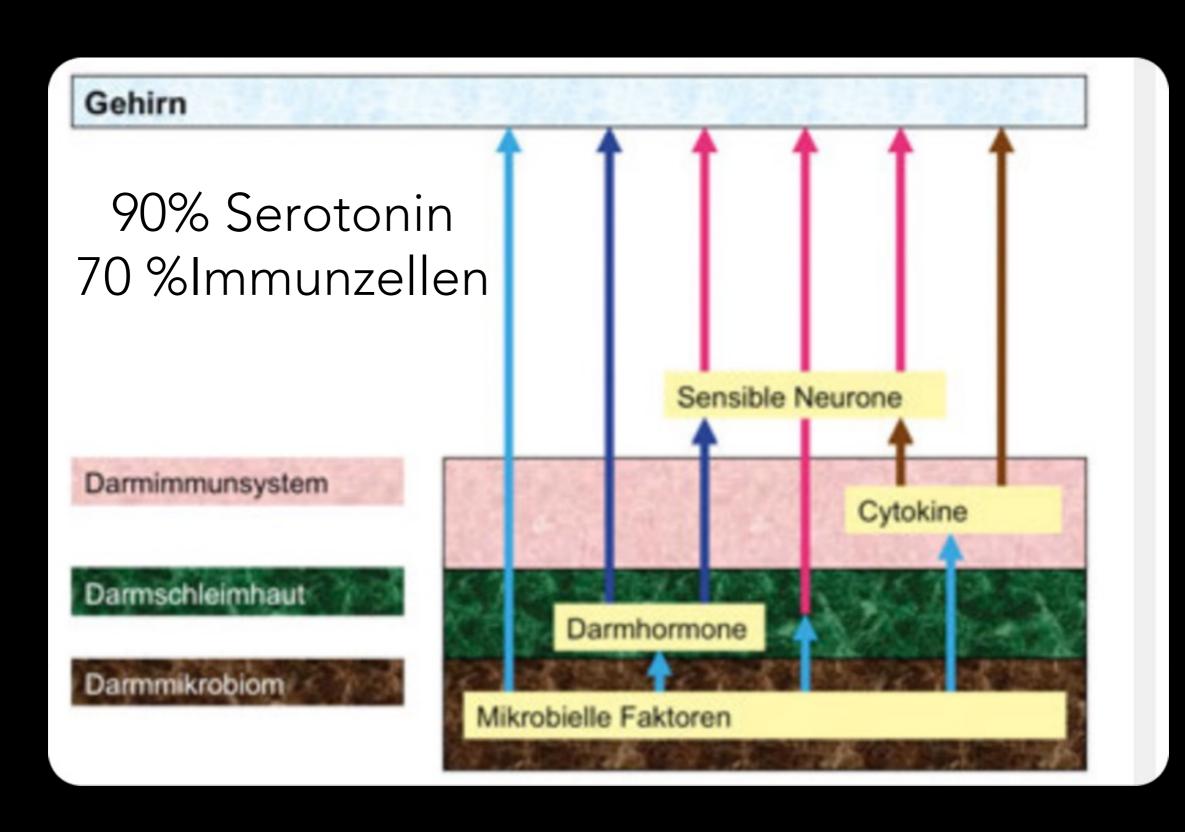




# Warum überhaupt haben wir eine bakterielle Kolonisation?

- "Flora" synthetisiert Vitamine über Eigenbedarf (Vitamin K, Vitamin B12, andere B-vitamine).
- Microbiota verhindert Kolonisation durch Kompetition um Bindungsstellen und Substrat (wohl der wichtigste benefizielle Effekt → Salmonellen Studien)
- Produktion von verschiedenen Substanzen, z.B. Peroxide oder spezifische Bacteriozine, die andere Bakterien hemmen oder sogar abtöten.



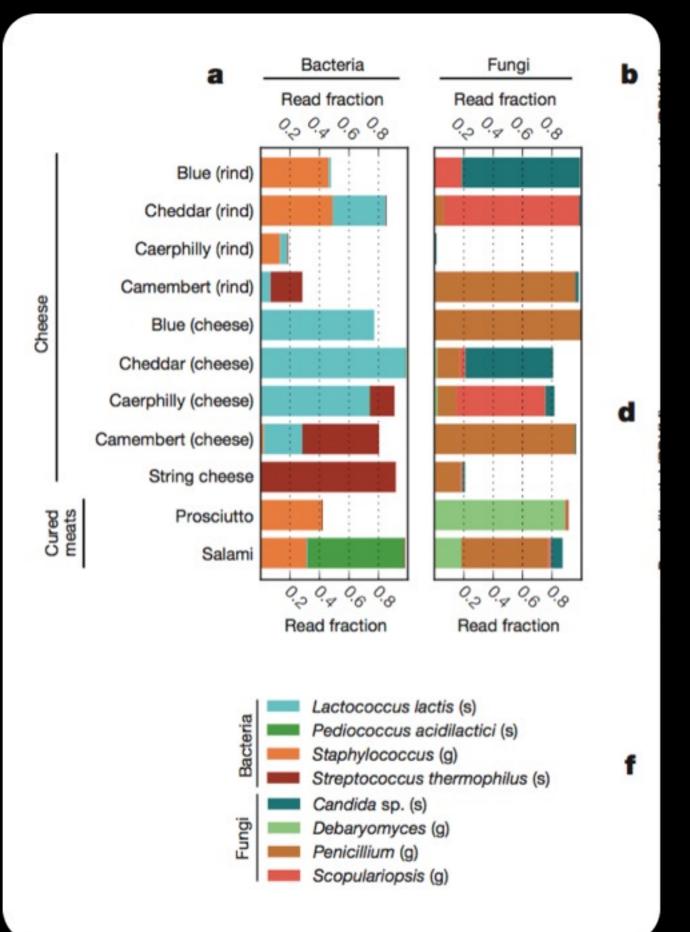




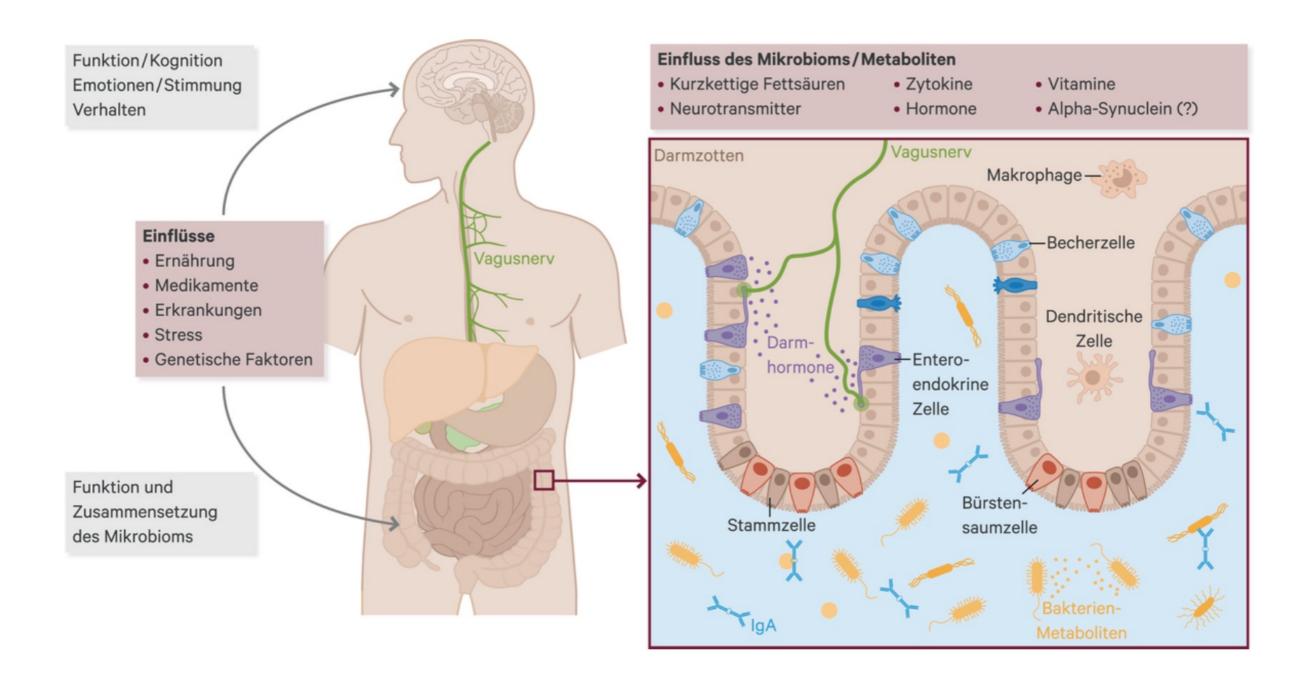
### LETTER

# Diet rapidly and reproducibly alters the human gut microbiome

Lawrence A. David<sup>1,2</sup>†, Corinne F. Maurice<sup>1</sup>, Rachel N. Carmody<sup>1</sup>, David B. Gootenberg<sup>1</sup>, Julie E. Button<sup>1</sup>, Benjamin E. Wolfe<sup>1</sup>, Alisha V. Ling<sup>3</sup>, A. Sloan Devlin<sup>4</sup>, Yug Varma<sup>4</sup>, Michael A. Fischbach<sup>4</sup>, Sudha B. Biddinger<sup>3</sup>, Rachel J. Dutton<sup>1</sup> & Peter J. Turnbaugh<sup>1</sup>







**Tab. 1.** Arzneistoffe mit direkter Wirkung auf das Mikrobiom

Arzneistoffklasse	Arzneistoffe	Betroffene Bakterienstämme	Literatur
Klassische Antipsychotika	Chlorprothixen, Flupentixol, Levomepromazin, Perazin, Promethazin, Zuclopenthixol	Acinetobacter baumannii, Escherichia coli, Klebsiella pneumoniae, Pseudomonas aeruginosa, Staphylococcus aureus	[24]
Atypische Antipsychotika	Aripiprazol, Asenapin, Clozapin, Lurasidon, Olanzapin, Paliperidon, Quetiapin, Risperidon, Ziprasidon	Akkermansia, Lachnospiraceae, Sutterella	[10]
Trizyklische Antidepressiva	Amitriptylin, Clomipramin, Doxepin, Nortriptylin, Opipramol, Trimipramin	Alle: Clostridium leptum  Amitriptylin: Bacillus spp.,  Salmonella typhimurium,  Staphylococcus spp., Vibrio cholerae	[8, 38]
Selektive Serotonin- Wiederaufnahmehemmer	Escitalopram, Fluoxetin, Paroxetin, Sertralin	Alle: Brucellae, Escherichia coli, Eubacterium ramulus, Streptococcus salivarius Escitalopram: Adlercreutzia, Alphaproteobacteria, Coriobacteriaceae, Ruminococcus flavefaciens Fluoxetin: Lactobacillus rhamnosus, Prevotella, Succinivibrio Sertralin: Pseudomonas	[8, 9, 19, 38]



Akute Anpassung- chronische Anpassung

J Clin Invest. 2015 Mar 2;125(3):926-38. doi: 10.1172/JCI76304. Epub 2015 Feb 17.

#### Gut/brain axis and the microbiota.

Mayer EA, Tillisch K, Gupta A.

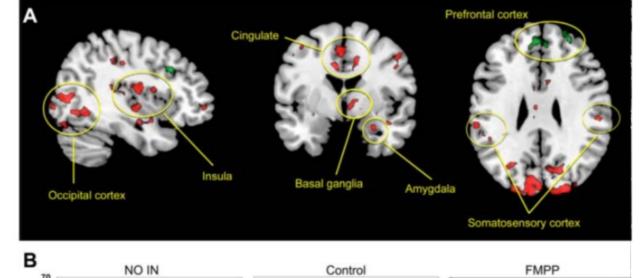
#### **Abstract**

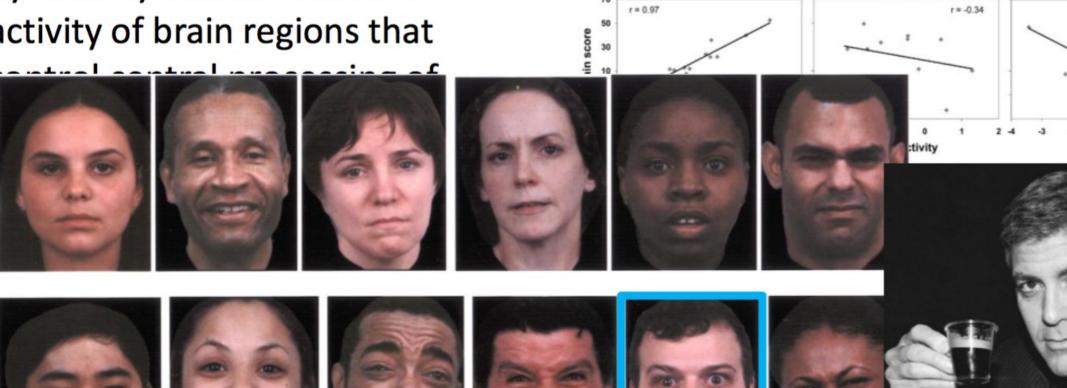
Tremendous progress has been made in characterizing the bidirectional interactions between the central nervous system, the enteric nervous system, and the gastrointestinal tract. A series of provocative preclinical studies have suggested a prominent role for the gut microbiota in these gut brain interactions. Based on studies using rodents raised in a germ-free environment, the gut microbiota appears to influence the development of emotional behavior, stress- and painmodulation systems, and brain neurotransmitter systems. Additionally, microbiota perturbations by probiotics and antibiotics exert modulatory effects on some of these measures in adult animals. Current evidence suggests that a moonanisms, more adocrine and neurocrine pathways, may be invested in gut microbiota-to-brain signaling and that the brain can in turn alter microbial composition and behavior via the autonomic nervous system. Limited information is available on how these in. The may translate to healthy humans or it has ease states involving the brain or the gut/brain axis. Future research needs to locus on confirming that the rodent findings are translatable to human physiology and to diseases such as irritable bowel syndrome, autism, anxiety, depression, and Parkinson's disease.

#### iosquil - sicidenoilli Isniiteeinl

# Beeinflussung des Verhalten auch beim Menschen?

- fermented milk product with probiotic (FMPP) for 4 weeks
- "four-week intake of an FMPP by healthy women affected activity of brain regions that





#### Background unserer Studie I

- Durchschnittliche Gewichtszunahme nach Rauchstopp: 7-8kg<sup>1</sup>
  - Wird einfach nur mehr gefuttert…?
  - Multiple Risk Factor Intervention Trial: quitters nahmen Gewicht zu, obwohl sie weniger Kalorieren einnahmen, generell eher gesündere Diät als Teilnehmer mit Rückfall oder weiter rauchende<sup>2</sup>



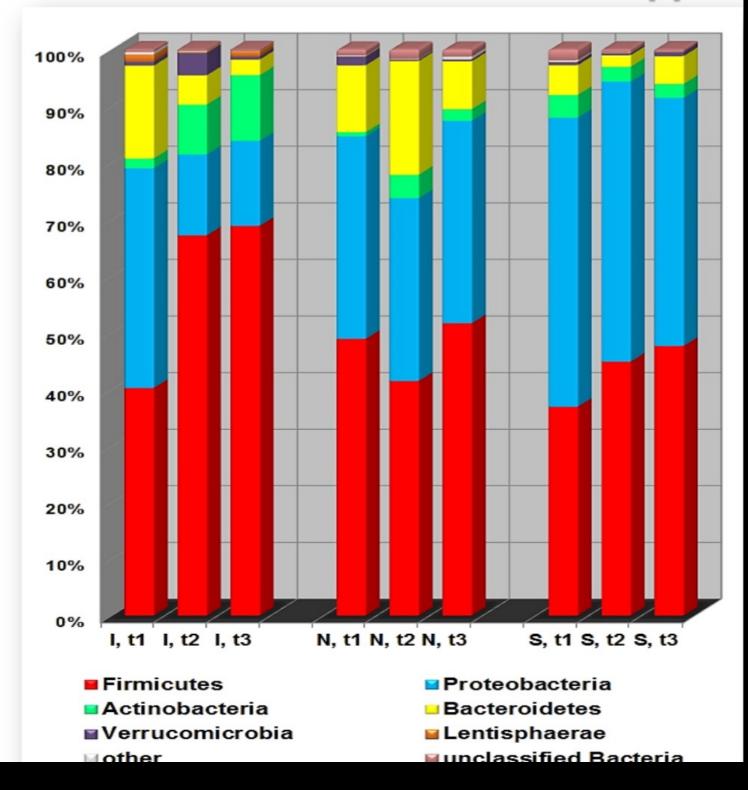
Unerschiedlicher Effekt Rauchen auf CD vs. UC

# Beeinflusst der Raucherstatus die intestinale Mikrobiota???

### qqoizhoush meireillorinox bnerhïsw sioidorxilh elsniizeinl

#### Shift der Phyla: Pyrosequencing

- die 4 grossen Phyla: 97.3% aller
   Sequenzen
- Intervention Group nach Rauchstopp:
  - Zunahme
     Firmicuten und
     Actinobacterien
  - Abnahme der Proteobacterien und Bacteroidetes
- stabile Mikrobiota in beiden Kontroll-Gruppen



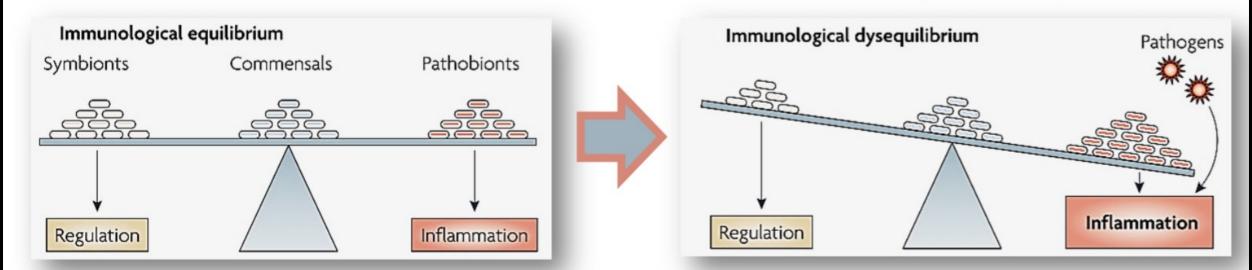






#### Dysbiose, Reduzierte mikrobielle Diversität bei IBD

- Keine Colitis in germ-free mouse models<sup>1</sup>
- Dysbiose bei IBD:
  - Verschiedene Alterationen auf Ebene Phyla, Groups Und Species<sup>2</sup>



- Verminderte Microbielle Diversität<sup>3,4</sup>
- Zunahme von Epithel-adhärenten Bakterien<sup>5,6</sup>
- Zuname von Intrazellulären Bakterien<sup>5,6</sup>
- Erhöhte temporale Instabilität<sup>3</sup>

# Impact of diet in shaping gut microbiota revealed by a comparative study in children from Europe and rural Africa

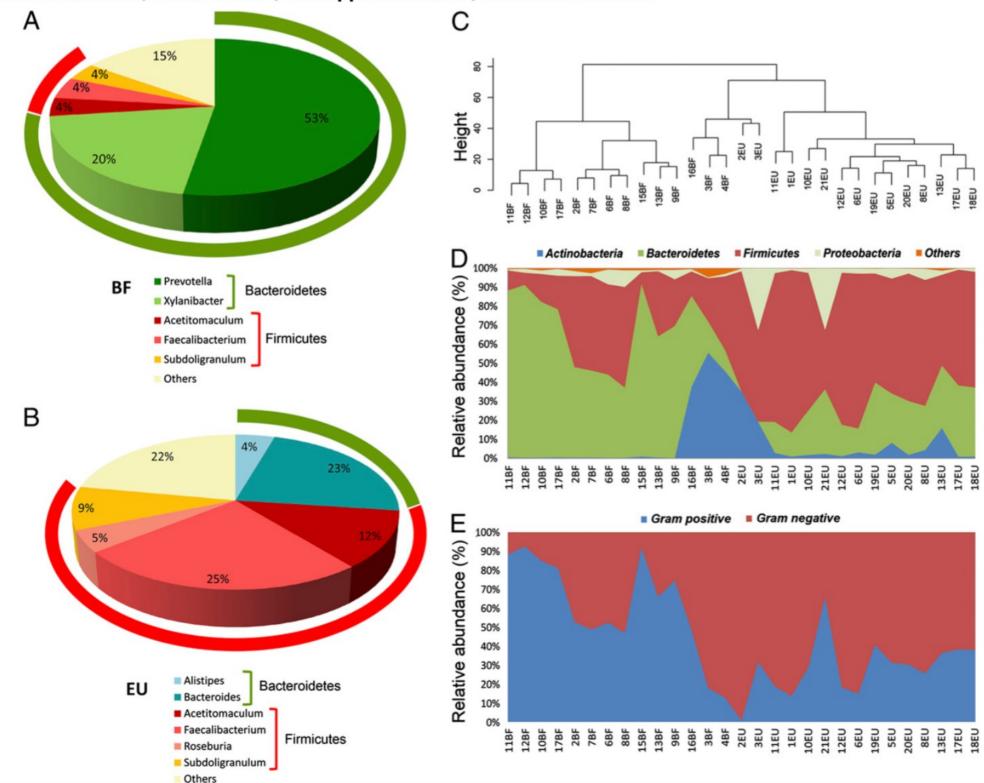
Carlotta De Filippo<sup>a</sup>, Duccio Cavalieri<sup>a</sup>, Monica Di Paola<sup>b</sup>, Matteo Ramazzotti<sup>c</sup>, Jean Baptiste Poullet<sup>d</sup>, Sebastien Massart<sup>d</sup>, Silvia Collini<sup>b</sup>, Giuseppe Pieraccini<sup>e</sup>, and Paolo Lionetti<sup>b,1</sup>



Fig. 1. Life in a rural village of Burkina Faso. (A) Village of Boulpon. (B) Traditional Mossi dwelling. (C) Map of Burkina Faso (modified from the United States ClA's World Factbook, 34). (D) Millet and sorghum (basic components of Mossi diet) grain and flour in typical bowls. (E) Millet and sorghum is ground into flour on a grinding stone to produce a thick porridge called Tô.

# Impact of diet in shaping gut microbiota revealed by a comparative study in children from Europe and rural Africa

Carlotta De Filippo<sup>a</sup>, Duccio Cavalieri<sup>a</sup>, Monica Di Paola<sup>b</sup>, Matteo Ramazzotti<sup>c</sup>, Jean Baptiste Poullet<sup>d</sup>, Sebastien Massart<sup>d</sup>, Silvia Collini<sup>b</sup>, Giuseppe Pieraccini<sup>e</sup>, and Paolo Lionetti<sup>b,1</sup>



# Disentangling type 2 diabetes and metformin treatment signatures in the human gut microbiota

Kristoffer Forslund, Falk Hildebrand, Trine Nielsen, Gwen Falony, Emmanuelle Le Chatelier, Shinichi Sunagawa, Edi Prifti, Sara Vieira-Silva, Valborg Gudmundsdottir, Helle Krogh Pedersen, Manimozhiyan Arumugam, Karsten Kristiansen, Anita Yvonne Voigt, Henrik Vestergaard, Rajna Hercog, Paul Igor Costea, Jens Roat Kultima, Junhua Li, Torben Jørgensen, Florence Levenez, Joël Dore, MetaHIT consortium, H. Bjørn Nielsen, Søren Brunak, Jeroen Raes 

♣ et al.

Affiliations | Contributions | Corresponding authors

Nature **528**, 262–266 (10 December 2015) | doi:10.1038/nature15766

Received 04 March 2015 | Accepted 05 October 2015 | Published online 02 December 2015

#### Diabetes-Medikament ändert Mikroben-Population des Darms



Über die Wirkungsweise von Metformin berichten Wissenschaftler des Europäischen Laboratoriums für Molekularbiologie (EMBL) in Heidelberg und des MetaHIT-Konsortiums in "Nature". Das am häufigsten verschriebene Medikament zur Behandlung von Diabetes Typ 2 hat einen größeren Einfluss auf die Zusammensetzung der Mikroben im Darm als die Krankheit selbst, berichtet Forschungsleiter Peer Bork.

Konkret wurden die Stuhlproben von mehr als 700 Personen verglichen. Unter ihnen befanden sich Patienten mit Diabetes Typ 2 und gesunde Probanden. Dabei fand das Team heraus, dass anhand der Zusammensetzung der Mikroben im Stuhl nicht bestimmt werden konnte, ob die Person an

Diabetes erkrankt war - es sei denn, sie verwendete Metformi

Es zeigte sich, dass Patienten, die das Medikament einnahren, wesentlich mehr E.coli und weniger I. bartletti Bartien als Gesunde und andere Patienten aufwiesen, die kein Metformin nahmen, festgestellt haben, könnten einige der Nebenwirkungen von Metformin erk "" so Erstautor Kristoffer Forslund. Daher wäre es durchaus denkbar, dass Patienten, die Metformin verabreicht bekommen, et an Joghurt oder ein Nahrungsergänzungsmittel erhalten, um ihre Darmflora im Gleichgewicht zu halten.

Quellen

Nature (abstract)

### Erklärung der Nebenwirkungen

In Brief | Published: 09 January 2019

INFECTION

# Probiotics fail to improve preschool gastroenteritis



Nature Reviews Gastroenterology & Hepatology 16, 76 (2019) | Download Citation ±

J Clin Biochem Nutr. 2018 Jul; 63(1): 33-35.

Published online 2018 Jun 20. doi: 10.3164/jcbn.18-57

#### PMCID: PMC6064808

PMID: 30087541

#### A next-generation beneficial microbe: Akkermansia muciniphila

Yuji Naito, 1,2,\* Kazuhiko Uchiyama, 1 and Tomohisa Takagi 1

▶ Author information ▶ Article notes ▶ Copyright and License information <u>Disclaimer</u>

This article has been cited by other articles in PMC.

#### Abstract

Go to: ☑

There have been many reports on the roles of intestinal flora and intestinal environment in health promotion and disease prevention. Beneficial bacteria such as *Bifidobacterium* and lactic acid-producing bacteria have been shown to improve the intestinal environment, and yield a good effect on metabolism, immunity and nerve response. In this review, in addition to these beneficial bacteria, we introduced *Akkermansia muciniphila* as a next-generation beneficial microbe. Several reports indicate that *Akkermansia muciniphila* affects glucose metabolism, lipid metabolism, and intestinal immunity, and that certain food ingredients such as polyphenols may increase the abundance of *Akkermansia muciniphila* in the gut.

### AUSLAGERN

Kochen / Braten

Homo sapiens

Rohkost

Nahrung und Haltbarkeit ausgelagert

Homo high tech

Urmensch 2 Mill

### AUSLAGERN

Microbiom

Hirnmasse

Nahrung und Haltbarkeit ausgelagert

Rohkost

Kochen / Braten

Urmensch 2 Mill

Homo

Homo high tech



"Nature" Artikel Emulgatoren rats and cheese-cake

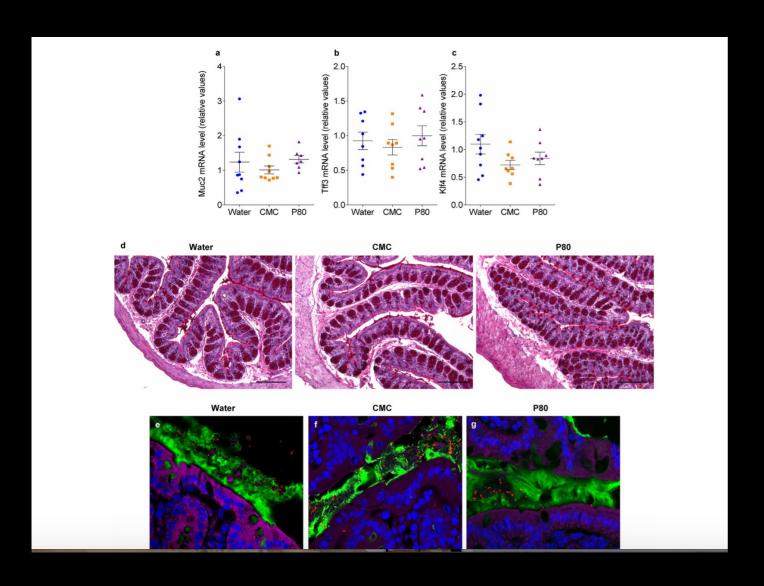
#### MAUSEFUTTER UND EMULGATOREN

Dietary emulsifiers impact the mouse gut microbiota promoting colitis and metabolic syndrome Benoit Chassaing<sup>1</sup>, Omry Koren<sup>2</sup>, Julia K. Goodrich<sup>3</sup>, Angela C. Poole<sup>3</sup>, Shanthi Srinivasan<sup>4</sup>, Ruth E. Ley<sup>3</sup> & Andrew T. Gewirtz<sup>1</sup>





Carboxymethylcellulose Sojalezithin 1%



Doppelt so hohe Inzidienz für entzündliche Veränderungen im MDT

These results support the emerging concept that perturbed host-microbiota interactions resulting in low-grade inflammation can promote adiposity and its associated metabolic effects.

Moreover, they suggest that the broad use of emulsifying agents might be contributing to an increased societal incidence of obesity/metabolic syndrome and other chronic inflammatory diseases.





### Cheesecake-eating rats and the question of food addiction

David H Epstein & & Yavin Shaham

Affiliations | Corresponding author

Nature Neuroscience 13, 529-531 (2010) | doi:10.1038/nn0510-529





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**Article metrics** 

Rats given extended access to high-fat high-sugar food show behavioral and physiological changes that are similar to those caused by drugs of abuse. However, parallels between drug and food "addiction" should be drawn with caution.









#### Cheesecake-eating rats and the question of food addiction

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Article metrics

Rats given extended access to high-fat high-sugar food show behavioral and physiological changes that are similar to those caused by drugs of abuse. However, parallels between drug and food "addiction" should be drawn with caution.





Resultate: massive Gewichtszunahme in der Kombination Trägheit!!!

## WAS MACHEN WIR



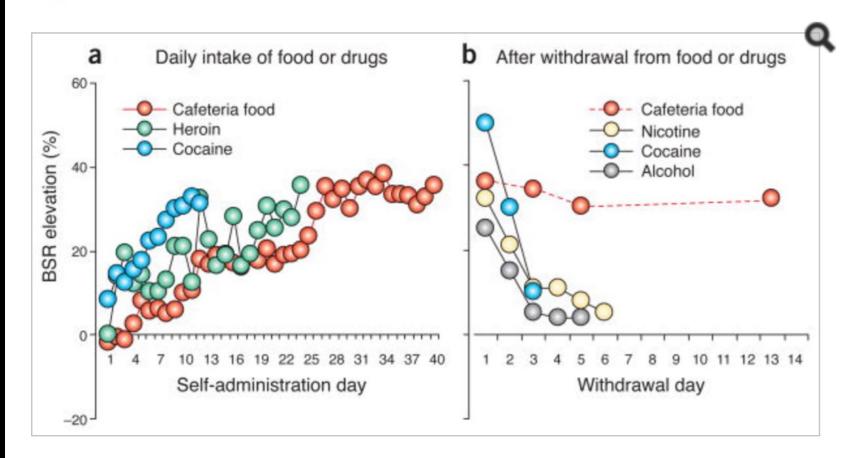






### Wie leben wir ???

Figure 1



Prolonged access to cafeteria food causes persistent elevations in threshold for BSR: comparison with drugs of abuse. (a) BSR threshold during daily intake of cafeteria food or drugs. (b) BSR threshold after loss of access to cafeteria food or drugs. Data were redrawn from Johnson and Kenny<sup>2</sup> and refs. <sup>3-7</sup>. In these studies, rats performed an operant response to obtain rewarding electrical brain stimulation into the median forebrain bundle at the level of the lateral hypothalamus. BSR threshold is defined as the minimum intensity of electrical stimulation that maintains operant responding. Increased BSR threshold is hypothesized to reflect decreased sensitivity of the brain reward system.



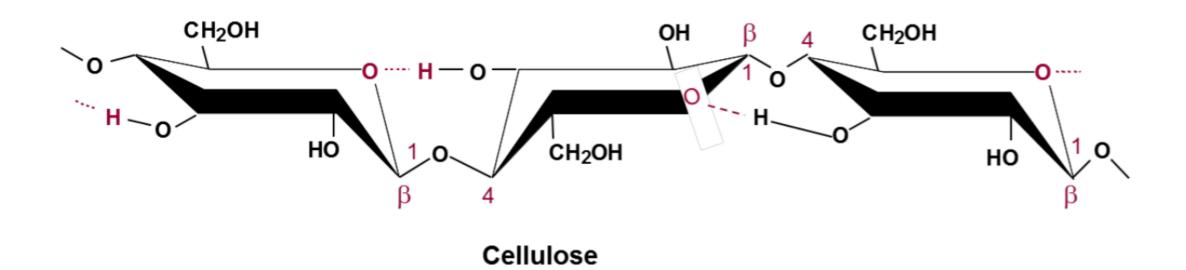
Pflegen sie ihr Microbiom!!!

UND WIE.....

# Polysaccharide (Ballaststoffe) sind die wichtigsten Substrate der Darmbakterien



- Die meisten mit der Nahrung aufgenommenen Polysaccharide sind pflanzlichen Ursprungs.
- Polysaccharide von Pflanzenzellwänden sind die sogenannte Ballaststoffe. Größtenteils werden sie nicht ausgeschieden sondern durch die intestinale Mikrobiota verwertet.
- Im Gegensatz zur Pansenmikrobiota der Wiederkäuer, die Cellulose [β (1 → 4) Glucan] in großem Umfang verwertet, werden im menschlichen Darm hauptsächlich Hemicellulosen verwertet.



(β-1,4-linkages)

## Wie sieht ein gutes Probiotikum aus

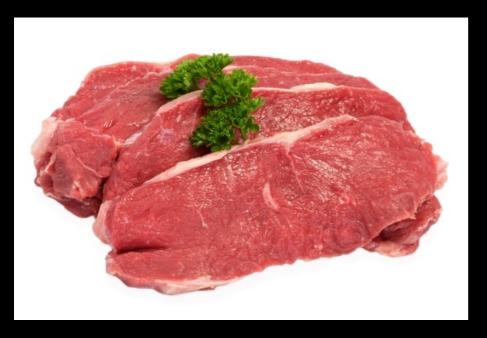
- 1. Viele Bakterienstämme (ideal >20)
- 2. Hohe Keimzahl (ideal >20 Milliarde/kbE)
- 3.Kombination von Vitaminen z.B. B7
- 4. Verpackung im Glas
- 5.Arzneimittelstnadard (GMP)
- 6. Bei Krankheiten oder Mangel

## Natürliche Probiotika

- Joghurt
- Kefir
- Käse (Gruyere, Mozarrela....)
- Sauerkraut
- Saure Gurken
- Apfelessig
- Kombucha, Kimchi













WO SIND PRÄBIOTIKA DRIN FERMENTIERBARE BALLASTSTOFFE (INULIN, GUAR, HAFERFASER- LÖSLICH -UNLÖSLICH)

Avocadoanteile Auberginen Chicorée Zichorienwurzel Topinambur Artischocken Pastinaken Löwenzahnwurzel Porree Zwiebeln Schwarzwurzeln. U.v.a.





# WO SIND PRÄBIOTIKA DRIN



Für 2016 haben die Vereinten Nationen das Internationale Jahr der Hülsenfrüchte ausgerufen. Also: Greifen Sie wieder verstärkt zu gesunden Eiweissbomben wie Bohnen, Kichererbsen oder Linsen!



#### BRITISH MEDICAL JOURNAL

LONDON SATURDAY DECEMBER 20 1958

#### CORONARY HEART DISEASE AND PHYSICAL ACTIVITY OF WORK

EVIDENCE OF A NATIONAL NECROPSY SURVEY\*

.

J. N. MORRIS, F.R.C.P., D.P.H.

MARGARET D. CRAWFORD, M.D.

Social Medicine Research Unit of the Medical Research Council, London Hospital

It has previously been shown that the drivers of London's double-decker buses are more likely to die suddenly from "coronary thrombosis" than the conductors, and that Government clerks suffer more often from rapidly fatal cardiac infarction than do postmen.<sup>13</sup> On the basis of these and similar observations a hypothesis has been stated that men in physically active jobs have a lower incidence of coronary (ischaemic) heart disease in middle-age than men in physically inactive jobs. More important, the disease is not so severe in physically active workers, tending to present in them in relatively benign forms.

It is a principle of epidemiological research of this type to seek evidence from as many, as various, and as independent sources as possible.40 The present report deals with the frequency, in relation to occupation, of ischaemic myocardial fibrosis in men dying from causes other than coronary heart disease itself. These myocardial scars are often evidence of early coronary heart disease and of less severe, or at any rate non-lethal, ischaemia. Thus the present study provides a picture of coronary heart disease in one of its simpler forms, and this may help in elucidating social connexions. At the same time another test of the hypothesis is made, using quite different data from the previous inquiries and upon quite another aspect of the condition. The inquiry it was hoped would provide some information on the pathological mechanisms of any differences with physical activity of work that were found; and in particular we sought to learn something about the relationships of occupation to coronary artery disease. brief, Can the hearts of men be seen to vary with the kind of work they have done? At what phases in the complex of coronary-myocardial disease as a whole can this particular "cause"—physical activity/inactivity be seen to operate? More generally the inquiry was intended to provide a statistical account of the coronary arteries of the middle-aged British male population, 1954-6.42

#### Method

Through the universities and regional hospital boards we appealed personally to all pathologists in the National Health Service who might regularly be in

\*Part of this report was included in the Ernestine Henry Lecture delivered by one of us (J. N. M.) before the Royal College of Physicians of London on June 10, 1958.

charge of post-mortem examinations to co-operate in the inquiry, and the pathologists of 206 hospitals or hospital groups—between 85% and 90% of the "possibles"—very kindly agreed. We cannot be more precise about this fraction, because it is not known whether some of those who did not reply at all to our letters were in fact regularly responsible for appreciable numbers of necronsies.

Each pathologist was invited to provide particulars on a standard form of 25 consecutive unselected necropsies on men aged 45-70 years, no matter what the cause of death or how the case came to necropsy. 5,000 reports were sent to us: trial runs with records from 1944 to 1951 in the Pathological Institute of the London Hos pital39 60 had encouraged the hope that with such numbers stable group patterns would emerge. In all cases macroscopic details were requested of disease in the coronary arteries, its nature, amount, and distribution, with descriptions of any stenosis; and of fibrosis of the myocardium of the left ventricle and interventricular septum, using lesions of 1 cm. in any dimension as markers; and the heart weight. Particulars were also asked about the main causes of death and other important clinical and pathological findings such as valvular heart disease, including especially aortic stenosis. By appealing for detailed descriptions of lesions seen, discouraging evaluations such as "slight" or "much," and giving guidance on terminology if not on technique, we hoped to reduce the variability of such procedures. It was postulated, moreover, that the variability in observation and recording over so many departments of pathology would be "random" in relation to the main factor with which pathology was to be correlated—physical activity of work.

The pathological data on each record relating to disease in the coronary arteries and fibrosis of the myocardium were graded, agreed, and coded by at least two and often three physicians in the unit. It took many months to define appropriate scales; but finally scales emerged that made pathological sense, were simple in use, having no more than five points, and the gradings on which proved highly reproducible by ourselves and by others.

Pathologists were also invited to give details of the last occupation of the deceased, but in many cases this information had to be obtained by the unit from the 5111

**Results:** In models adjusted for a range of covariates including socioeconomic status, smoking, and obesity, high resting heart rate was associated with a modestly elevated rate of mortality from all causes (hazard ratio; 95% confidence interval: tertile 3 vs. tertile 1: 1.17; 0.99, 1.37 p[trend]: 0.07) and respiratory disease (1.69; 1.04, 2.76 p[trend]: 0.03). Of the two markers of physical activity, walking pace was inversely related to mortality ascribed to all causes (slow vs. high walking pace 1.71; 1.53, 1.91 p[trend]: <0.001]), coronary heart disease (2.03; 1.68, 2.47 p[trend]: <0.001), and total cancers (1.25; 0.98, 1.59 p[trend]: 0.04). The corresponding associations for leisure time

CORONARY HEART DISEASE AND PHYSICAL ACTIVITY

RITISH AL JOURNAL

ISH

cial class and, incidentally, no indication excess seen in the national mortality data s I. There is again no class trend in the s (middle column of Table X(A)) and n the heavy. But reading Table X(A) teavy trend previously reported is pretty social class, the light occupations showing han the active in class II, and the light in s IV a higher prevalence than the active ic classes. That is to say, among men of 1-economic circumstances, those in light d a clear excess of ischaemic myocardial ould be expected—Tables X(B) and (C)—the frequency of fibrosis that occurs with of occupation within the social classes is the large healed infarcts; though the X(B) are impossibly small.

ies in these tables are also interesting, inds out on its own, both in showing low workers and in showing little trend with just in these men that the classification of

occupations by "activity" is at its weakest. On the one hand, the light occupations—for example, lift attendant, night watchman—are reserved mostly for disabled persons; therefore the last and "light" job, more than in other classes, is apt to be an inaccurate indicator of the main or usual job during working life. On the other hand, all "general labourers" were perforce graded as "heavy" workers, though, as is well known, many an elderly "labourer," officially or unofficially disabled, does little more than odd jobs on a building site and the like, and may have been so occupied for years before his death. There is an irregularity in social class III because some very heavy workers—for example, stonemasons and boilermakers—have a high prevalence of scars, high for heavy workers; a hint here, perhaps, of the dangers of too much physical activity." The rate for heavy occupations in class IV is particularly low because agricultural workers show little schaemic fibrosis—little even for heavy workers. These special features of classes III and IV persist when the prevalence of scarring in these particular occupations is compared also with that of other heavy workers in the same

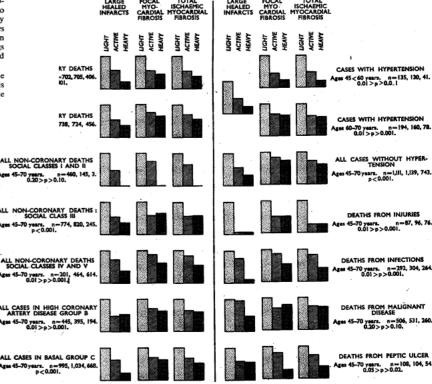


Fig. 2.—Frequency of ischaemic myccardial fibrosis, by physical activity of occupation. 3,800 non-coronary deaths. Males aged 45-70 years. In each block of three columns the first column is the rate among light occupations regarded as 100%. In the second column (active occupations) and the third column (heavy) the rates have been recalculated as proportions of the rate among light.

In each line the first block represents the large fibrous patches; the second block represents the smaller/multiple scars; and the third is the total of ischaemic myocardial fibrosis.

n gives the numbers of cases in light, active, and heavy occupations in that order. The test of significance applies to the total of ischaemic myocardial fibrosis. In deaths from injuries active and heavy occupations have been amalgamated.



#### BRITISH MEDICAL JOURNAL

LONDON SATURDAY DECEMBER 20 1958

#### CORONARY HEART DISEASE AND PHYSICAL ACTIVITY

EVIDENCE OF A NATIONAL NECROPSY SURVEY\*

J. N. MORRIS, F.R.C.P., D.P.H.

MARGARET D. CRAWFORD, M.D.

Social Medicine Research Unit of the Medical Research Council, London Hospita

It has previously been shown that the drivers of London's double-decker buses are more likely to die suddenly from "coronary thrombosis" than the conductors, and that Government clerks suffer more often from rapidly fatal cardiac infarction than do nostmen.43 On the basis of these and similar observations a hypothesis has been stated that men in physically active jobs have a lower incidence of coronary (ischaemic) heart disease in middle-age than men in physically inactive More important, the disease is not so severe in physically active workers, tending to present in them in relatively benign forms.

It is a principle of epidemiological research of this type to seek evidence from as many, as various, and as independent sources as possible.40 The present report deals with the frequency, in relation to occupation, of ischaemic myocardial fibrosis in men dying from causes other than coronary heart disease itself. These myocardial scars are often evidence of early coronary heart disease and of less severe, or at any rate non-lethal, ischaemia. Thus the present study provides a picture of coronary heart disease in one of its simpler forms, and this may help in elucidating social connexions. At the same time another test of the hypothesis is made, using quite different data from the previous inquiries and upon quite another aspect of the condition. The inquiry it was hoped would provide some information on the pathological mechanisms of any differences with physical activity of work that were found; and in particular we sought to learn something about the relationships of occupation to coronary artery disease. In brief, Can the hearts of men be seen to vary with the kind of work they have done? At what phases in the complex of coronary-myocardial disease as a whole can this particular "cause"-physical activity/inactivitybe seen to operate? More generally the inquiry was intended to provide a statistical account of the coronary arteries of the middle-aged British male population 1954-6.42

Through the universities and regional we appealed personally to all pat National Health Service who

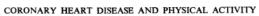
charge of post-mortem examinations to co-operate in the inquiry, and the pathologists of 206 hospitals or hospital groups-between 85% and 90% of the "possibles "-very kindly agreed. We cannot be more precise about this fraction, because it is not known whether some of those who did not reply at all to our letters were in fact regularly responsible for appreciable numbers of

Each pathologist was invited to provide particulars on a standard form of 25 consecutive unselected necropsies on men aged 45-70 years, no matter what the cause of death or how the case came to necropsy. 5,000 reports were sent to us: trial runs with records from 1944 to 1951 in the Pathological Institute of the London Hos pital39 60 had encouraged the hope that with such numbers stable group patterns would emerge. In all cases macroscopic details were requested of disease in the coronary arteries, its nature, amount, and distribution, with descriptions of any stenosis; and of fibrosis of the myocardium of the left ventricle and interventricular septum, using lesions of 1 cm, in any dimension as markers; and the heart weight. Particulars were also asked about the main causes of death and other impoant clinical and pathological findings such as heart disease, including especially aortic stenoappealing for detailed descriptions of lesions couraging evaluations such as "slight" or giving guidance on terminology hoped to reduce the variability was postulated, moreover, th tion and recording of logy would be "rando with which pathology

cord relating to and fibrosis of the myoid coded by at least two iate scales: but finally scales de pathological sense, were simple in re than five points, and the gradings ighly reproducible by ourselves and

also invited to give details of the of the deceased, but in many cases this

Results: In models adjusted for a range of covariates ocioeconomic status, smoking, soch ted ji ra modestly elevated rate of mortality from and obesity, high resting heart rate was a all causes (hazard ratio; 95% confidence) van. erthe 5 vs. tertile 1: 1.17; 0.99, 1.37 p[trend]: 0.07) and respiratory disease (1.69; 1.04, 2.3) p[t] and]: 0.03). Of the two markers of physical activity, walking pace was inversely related to nortality ascribed to all causes (slow vs. high walking pace 1.71; 1.53, 1.91 p[trend]: <0.001]), coronary heart disease (2.03; 1.68, 2.47 p[trend]: <0.001), and total cancers (1.25; 0.98, 1.59 p[trend]: 0.04). The corresponding associations for leisure time



occupations by "activity" is at its weakest. On the one

cial class and, incidentally, no indication excess seen in the national mortality data s I. There is again no class trend in the s (middle column of Table X(A)) and the heavy. But reading Table X(A) neavy trend previously reported is pretty social class, the light occupations showing han the active in class II, and the light in s IV a higher prevalence than the active se classes. That is to say, among men of l-economic circumstances, those in light ed a clear excess of ischaemic myocardial ould be expected—Tables X(B) and (C)—the frequency of fibrosis that occurs with f occupation within the social classes is the large healed infarcts; though the X(B) are impossibly small.

ies in these tables are also interesting inds out on its own, both in showing low workers and in showing little trend with

hand, the light occupations-for example, lift attendant night watchman-are reserved mostly for disabled persons therefore the last and "light" job, more than classes, is apt to be an inaccurate indicator of the main or usual job during working life. On the other hand, all "general labourers" were perforce graded as "heavy" werkers, though, as is well known, many an elderly "labourer," officially or unofficially disabled, does little more than odd jobs on a building site and the like, and may have been so occupied for years before his death. There is an irregularity in social class III because some very heavy example, stonemasons and boilermakershave a high prevalence of scars, high for heavy workers a hint here, perhaps, of the dangers of too much physical activity. The rate for heavy occupations in class IV is particularly low because agricultural workers show little ischaemic fibrosis—little even for heavy workers. These special features of classes III and IV persist when the prevalence of scarring in these particular occupations is compared also with that of other heavy workers in the same

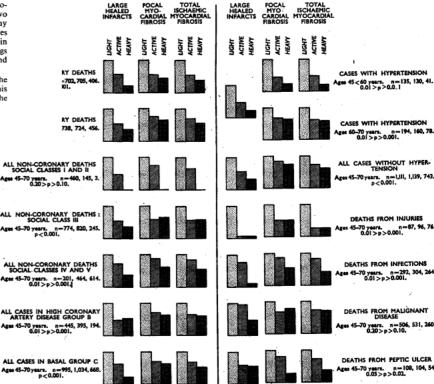


Fig. 2.—Frequency of ischaemic myccardial fibrosis, by physical activity of occupation. 3,800 non-coronary deaths. Males aged 45-70 years. In each block of three columns the first column is the rate among light occupations regarded as 100%. In the second column (active occupations) and the third column (heavy) the rates have been recalculated as proportions of the rate among light

In each line the first block represents the large fibrous patches; the second block represents the smaller/multiple scars; and the

In each mile the inst block represents the large intotos parties, the stock of the total of ischaemic myocardial fibrosis.

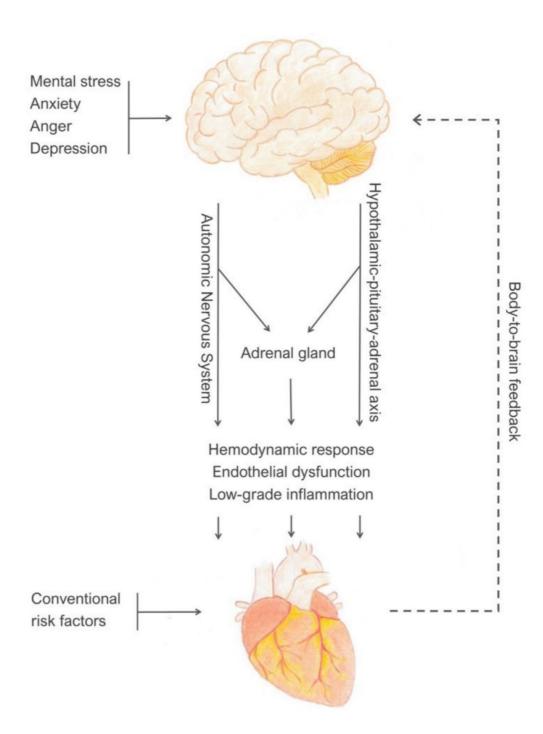
n gives the numbers of cases in light, active, and heavy occupations in that order. The test of significance applies to the total of ischaemic myocardial fibrosis. In deaths from injuries active and heavy occupations have been amalgamated.

Review

#### Mental Stress and Cardiovascular Health—Part I

Federico Vancheri <sup>1,\*</sup>, Giovanni Longo <sup>2</sup>, Edoardo Vancheri <sup>3</sup> and Michael Y. Henein <sup>4,5,6</sup>

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### CV Risk like smoking

### WO SIND PROBIOTIKA DRIN

- Lactobacillus
- Typ: Streptococcus thermophilus, Lactobacillus delbrueckii subsp. bulgaricus)

Oberflächen von Früchten, Beeren (Waschen ?!?!?!)
 Kontamination (PPI)

• Fermentierte Lebensmittel (Kimchi / Sauerkraut)

### Merke:

Essen sie nichts wofür Reklame gemacht wird!

Kochen sie mit bekannten natürlichen Zutaten selbst

Essen sie weniger und vornehmlich pflanzlich (Kochen, Braten ist Verdauung ausserhalb des Körpers!)

Bewegung und "Denken" ist auch essentiell

Nehmen sie sich Zeit für sich! Viel Zeit, und beobachten sie, was wie wirkt.

### Facts

Emulgatoren

Diversität des Darmes schützt !!!

Grösstes Immunorgan und intensivster Austausch mit der Umwelt, Production von Serotonin!!!

Nahrung immens wichtig, Foodmap, Zucker,

In Balance befindliche Beziehung des Immunsystems und des Microbioms

Microbiom wandelt sich schnell (dramatische Reaktionen Oder nur wenig Symptome.)

Alternative Facten
Abnehmen durch Transplantation,
einfache Therapieansätze, komplexes System

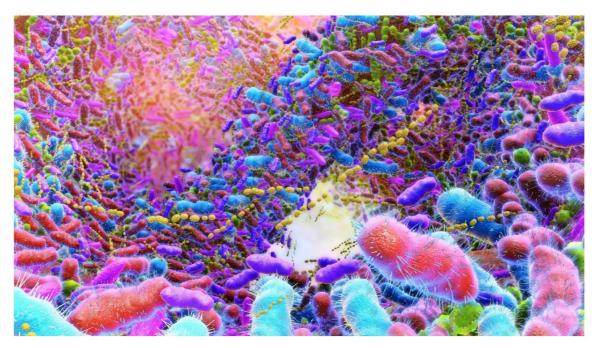
## Das Mikrobiom im Darm ist wichtig für die Gesundheit. Lohnt sich ein Test?

Untersuchungen der Darmflora sind gross in Mode. Doch selbst Fachleute verstehen den Zusammenhang zwischen den Bakterien im Darm und unserer Gesundheit nicht genau.

Martin Lindner

Os.01.2025, 05.30 Uhr ○ 3 min

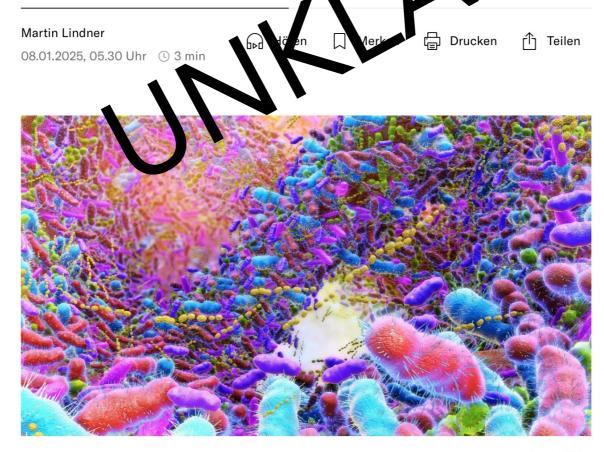
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Vielfältige Mitbewohner: Im Dickdarm leben in jedem Gramm Darminhalt etwa 100 Milliarden Mikroorganismen. Nopparit/Getty

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Review > Probiotics Antimicrob Proteins. 2024 Dec;16(6):2316-2328.

doi: 10.1007/s12602-024-10363-8. Epub 2024 Sep 25.

## Efficacy of Probiotics in Overweight and Obesity Control: An Umbrella Review and Subgroup Meta-Analysis

A Sadeghi <sup>1</sup>, R Daroudi <sup>1</sup>, M Davari <sup>2</sup>, Z Gharib-Naseri <sup>3</sup>, J Jafarzadeh <sup>1</sup>, M Tajvar <sup>4</sup>

Affiliations + expand

PMID: 39320636 DOI: 10.1007/s12602-024-10363-8

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#### **Abstract**

Numerous primary and secondary studies have consistently demonstrated that probiotics, including lactobacillus and Bifidobacterium, possess a potential anti-obesity effect. However, it is worth noting that some studies have yielded contrasting results. Considering this, our study aims to present a comprehensive overview of published systematic reviews and meta-analyses, focusing on the efficacy and safety of probiotics in managing obesity. To achieve this objective, we conducted an umbrella review following the PRISMA protocol and Cochrane guidelines. We searched databases such as Embase, PubMed, Cochrane Library, and Google Scholar for relevant systematic reviews and meta-analyses published in English, without imposing any date restrictions. Our inclusion criteria encompassed studies evaluating the anti-obesity impact of probiotics, with a specific focus on changes in body mass index (BMI), fat mass percentage (FMP), body weight (BW), and body fat mass (BFM). These studies were meticulously reviewed by two independent reviewers. Our analysis included five systematic reviews and 18 meta-analyses that met the predefined inclusion and exclusion criteria. The meta-analyses revealed statistically significant reductions in the following parameters: BMI, a decrease of 0.30 kg/m<sup>2</sup> (p < 0.00001, 95% CI - 0.36 to - 0.25); BFM, a reduction of 0.86 kg (p < 0.00001, 95% CI - 1.02 to - 0.71); BW, a decrease of 0.59 kg (p < 0.00001, 95% CI - 0.74 to - 0.44); and FMP, a substantial decline of 78% (p < 0.00001, 95% CI - 1.02 to - 0.54). In summary, our umbrella review suggests that existing evidence supports the potential benefits of probiotics in managing obesity and overweight. However, it is essential to acknowledge that the credibility of this evidence is somewhat limited due to the inclusion of studies with poor-quality designs and relatively small participant numbers. To establish the true efficacy of probiotics in obesity management, we recommend conducting robust studies involving larger participant cohorts.

Keywords: Bifidobacterium; Lactobacillus; Obesity; Overweight; Probiotics; Umbrella review.

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