Christopher E. Mason

**Associate Professor** Department of Physiology and Biophysics &

The Institute for Computational Biomedicine at the Weill Cornell Medical College and the

Tri-Institutional Program on Computational Biology and Medicin

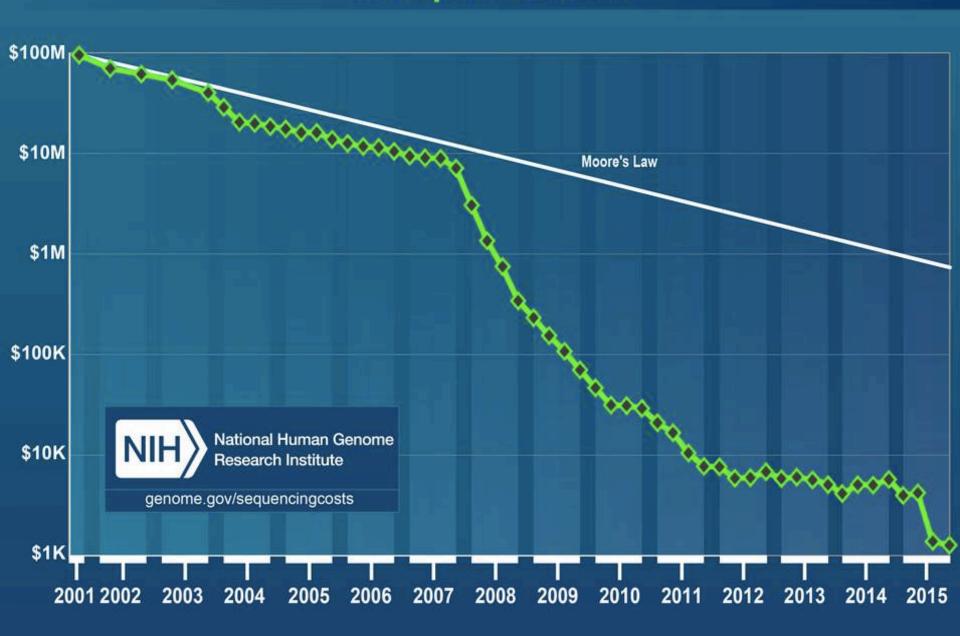
Fellow of the Information Society Project, Yale Law School

March 14<sup>th</sup> , 2016

(0)

## Revolution

#### Cost per Genome





NATURE | NEWS FEATURE





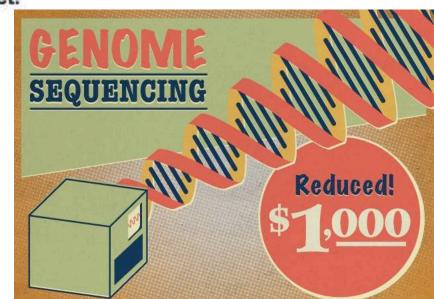


### Technology: The \$1,000 genome

With a unique programme, the US government has managed to drive the cost of genome sequencing down towards a much-anticipated target.

Erika Check Hayden

19 March 2014



## But! There is more than one genome:

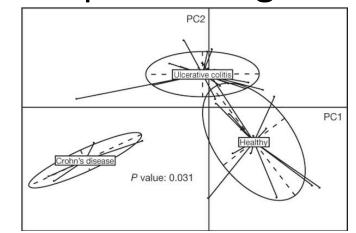
In your body's cellular democracy,

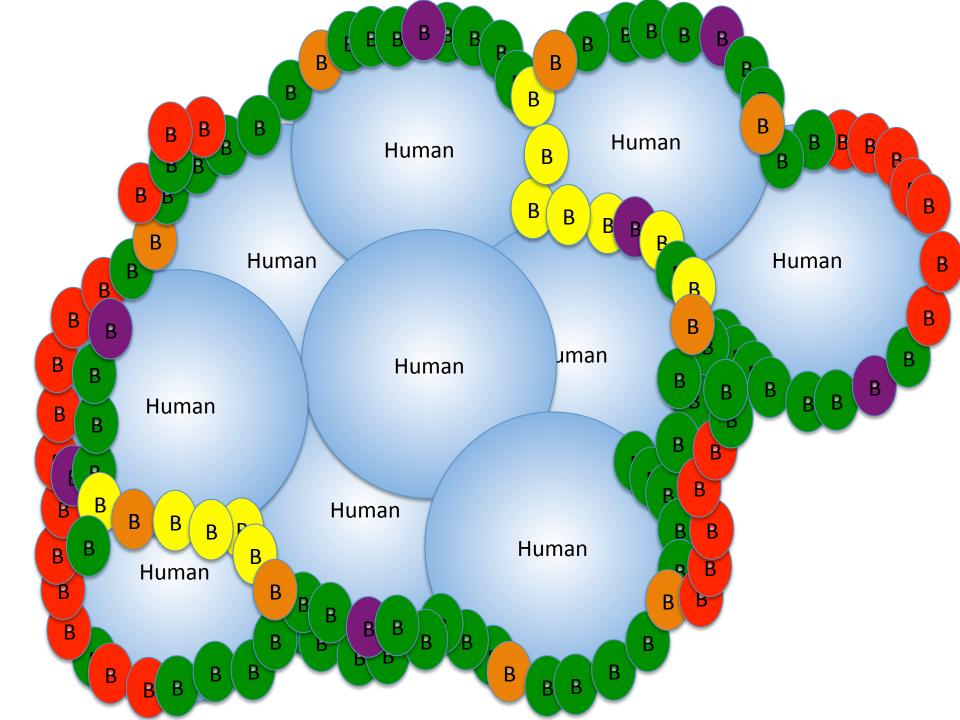
## YOU are a minority party:

1.3-10X bacterial:human cells (Zhu et al., 2010, Sender et al, 2016)

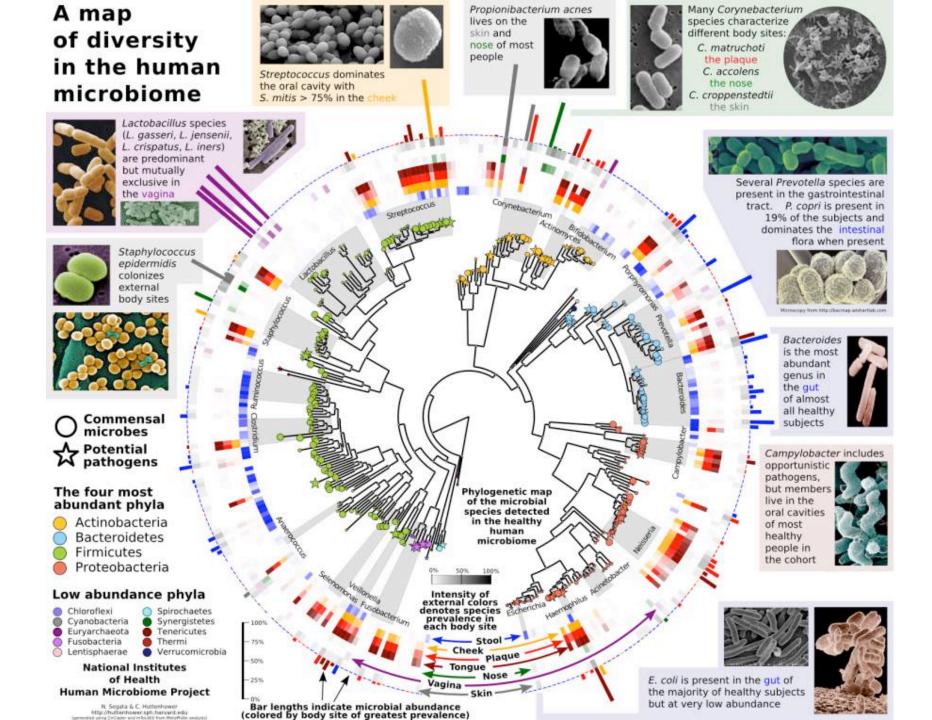
150:1 bacterial:human active transcripts in the gut

microbiome (Qin et al, 2010)









(1)

## Microbiome

Hope

#### nature

Proceedings of the National Academy of Sciences of the United States of A

CURRENT ISSUE // ARCHIVE // NEWS & MULTIMEDIA // FOR AUTHORS // ABOUT PNAS

vol. 109 no. 2 >



Performing your original search "microbiome disease links science" in PNAS retrieves 80

Metagenomic systems biology of the human g reveals topological shifts associated with obes inflammatory bowel disease

Sharon Greenblum, Peter J. Turnbaugh, and Elhanan Borenstein a,c,d,1

Author Affiliations



ARTICLE PREVIEW

view full access options

NATURE | LETTER

< ⊠ ⊜

Sharon Greenblum, 594-599, doi: 10.1073/pnas

Diet rapidly and reproducibly alters the human gut microbiome

Lawrence A. David, Corinne F. Maurice, Rachel N. Carmody, David B. Gootenberg, Julie E. Button, Benjamin E. Wolfe, Alisha V. Ling, A. Sloan Devlin, Yug Varma, Michael A. Fischbach, Sudha B. Biddinger, Rachel J. Dutton & Peter J. Turnbaugh

Affiliations | Contributions | Corresponding author

Nature (2013) | doi:10.1038/nature12820

Received 18 April 2013 | Accepted 29 October 2013 | Published online 11 December 2013

- Acne
- Antibiotic-associated diarrhea
- Asthma/allergies
- Autism
- Autoimmune diseases
- Cancer
- Dental cavities
- Depression and anxiety
- Diabetes
- Eczema
- Gastric ulcers
- Hardening of the arteries
- Inflammatory bowel diseases
- Malnutrition
- Obesity

learn.genetics.utah.edu/content/microbiome/disease/





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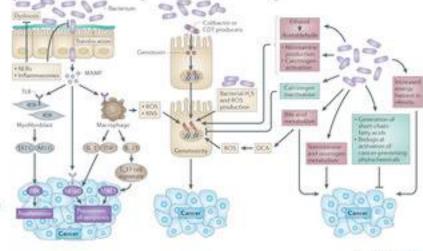
NATURE REVIEWS CANCER | PERSPECTIVES | OPINION

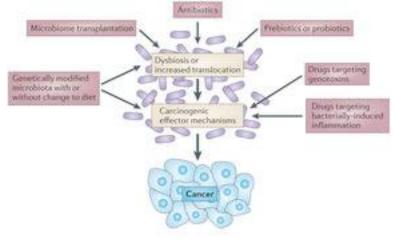
#### The microbiome and cancer

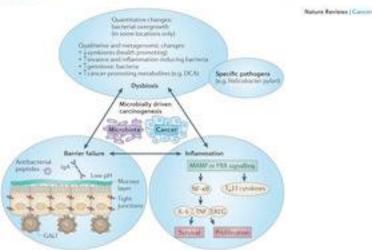
Robert F. Schwabe & Christian Jobin

Affiliations | Corresponding authors

Nature Reviews Cancer 13, 800–812 (2013) | doi:10.1038/nrc3610 | Published online 17 October 2013









HEALTH

## You Are Your Bacteria: How the Gut Microbiome Influences Health

The bacteria in our gut already plays an important role in digestion. But new studies indicate that our bacteria could play a major role in whether or not we become obese



Home > Science Magazine > 8 June 2012 > Hood, 336 (6086): 1209



stay thin.

Published Online June 6 2012 Science 8 June 2012: Vol. 336 no. 6086 p. 1209 DOI: 10.1126/science.1225475

EDITORIAL

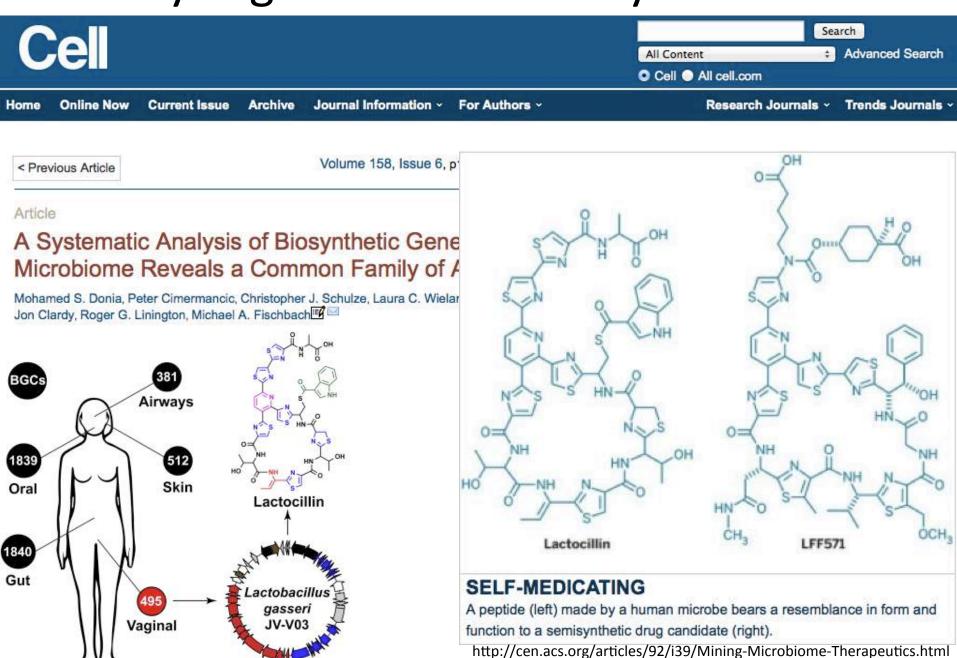
Tackling the Microbion

Leroy Hood

36%

of the small molecules in your blood are from the microbiome

## Many organisms work on your behalf!





Microbiologist Satoshi Omura holds a news conference Monday evening at Kitasato University in Tokyo after winning the 2015 Nobel Prize in medicine for his work on debilitating diseases caused by parasitic worms. | KYODO

NATIONAL / SCIENCE & HEALTH

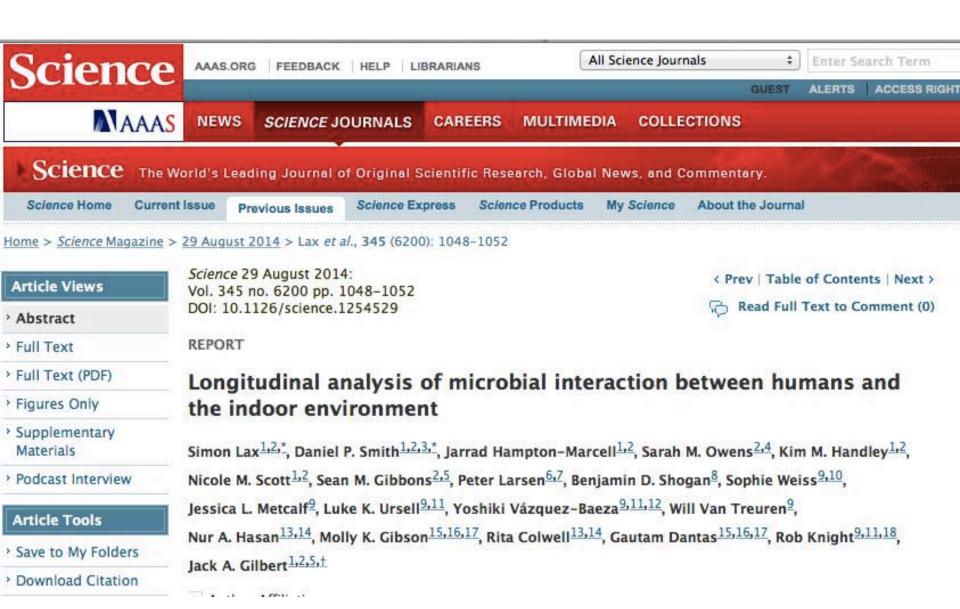
## Japanese microbiologist Satoshi Omura shares Nobel Prize for medicine

## Omura thanks his microbial helpers

"I was helped by microorganisms. I wonder if I deserve the prize," he said.

# Where do they come from?

## We all carry a "microbiome cloud"





### Mothers Give Us a Lot (Beyond Love)

**OPEN**  ACCESS Freely available online



Essay

## Mom Knows Best: The Universality of Maternal Microbial Transmission

Lisa J. Funkhouser<sup>1\*</sup>, Seth R. Bordenstein<sup>1,2\*</sup>

1 Department of Biological Sciences, Vanderbilt University, Nashville, Tennessee, United States of America, 2 Department of Pathology, Microbiology, and Immunology, Vanderbilt University, Nashville, Tennessee, United States of America

#### Summary

The sterile womb paradigm is an enduring premise in biology that human infants are born sterile. Recent studies suggest that infants incorporate an initial microbiome before birth and receive copious supplementation of maternal microbes through birth and breastfeeding. Moreover, evidence for microbial maternal transmission is increasingly widespread across animals. This collective knowledge

While maternal transmission of microbes in humans has attracted considerable attention in the last few years, nearly a century's worth of research is available for vertical transmission of symbionts in invertebrates [22]. Similar to gut bacteria in humans that assist nutrient intake, many insect-associated bacteria function as nutritional symbionts that supplement the nutrient-poor diet of their host with essential vitamins or amino acids [23,24]. Since these

that human infants develop within a sterile environment and acquire their initial bacterial inoculum while traveling through the maternal birth canal [28]. More than a century later, the sterile womb hypothesis remains dogma, as any bacterial presence in the uterus is assumed to be dangerous for the infant. Indeed, studies of preterm deliveries have found a strong correlation between intrauterine infections and preterm labor, especially when birth occurs less than 30 weeks into the pregnancy

#### Mouth

- Amniotic fluid bacteria with an oral origin include Fusobacterium nucleatum, Streptococcus spp., Bergeyella spp., Porphyromonas gingivalis, Rothia dentocariosa, and Filfactor alocis.
- Oral bacteria are likely transmitted to the uterus via the blood stream, especially if gingiva inflammation is present.

#### Internal Breast

- Asceptically-collected breast milk contains 100-600 OTUs. Abundant genera include Streptoccocus, Staphylococcus, Serratia, Corynebacteria, Lactococcus, Weisella, and Leuconostoc.
- An entero-mammary pathway brings gut bacteria to the mammary gland via lymph and blood circulation. External sources of milk microbes include the maternal skin and infant oral cavity.



#### **External Breast**

 The sebaceous skin of the breast is an external source of microbes for breast milk (Staphylococcus and Corynebacteria) but may also provide skin bacteria not common in milk (such as Propionibacteria) to the suckling infant.

#### Uterus

- Contrary to the "sterile womb" paradigm, bacteria are found in umbilical cord blood, amniotic fluid, fetal membranes, and meconium of healthy, term infants.
- Meconium microbiomes are dominated by Enterobacteriaceae (Escherichia and Shigella) and lactic acid bacteria (Leuconostoc, Enterococcus, and Lactococcus).
- Microbes likely gain access to the womb through ascension from the vagina and/or through the blood stream for bacteria of intestinal or oral origin.

#### Vagina

- Vaginal microbial communities vary significantly among women of different ethnicities and could influence which microbes
  are transferred to an infant.
- The vaginal microbiota becomes less diverse during pregnancy while certain Lactobacillus species become enriched.
- The initial microbiota of vaginally-born infants resembles that of their mother's vagina, while that of C-section infants is dominated by skin microbes not related to those of their mother.

# Sharing

Caring

## Fecal Microbiota Transplants (FMT)



Gastroenterol Hepatol (N Y). 2012 March; 8(3): 191-194

PMCID: PMC3365524

#### Fecal Transplantation for the Treatment of Clostridium difficile Infection

Lawrence J. Brandt, MD, MACG, AGAF, FASGE\*

Author information ► Copyright and License information ►



enable diet-induced obesity

Vaibhav Upadhyay, Valeriy Poroyko, Tae-jin Kim, Suzanne Devkota, Sherry Fu, Donald Liu, Alexei V Tumanov, Ekaterina P Koroleva, Liufu Deng, Cathryn Nagler, Eugene B Chang, Hong Tang & Yang-Xin Fu

Lymphotoxin regulates commensal responses to



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#### **FDA Comes to Grips with Fecal Transplants**

Efforts are under way to standardize fecal transplants, which helps re-establish healthy microbe populations in the guts of patients

By Beth Mole and Nature magazine

### The POWER of POOP

promoting safe, accessible fecal microbiota transplant for all who need it

http://thepowerofpoop.com/

Affiliations | Contributions | Corresponding author



Q Search

## 95% success rate for *C. diff* 70-80% success rate for ulcerative colitus

- David Shepard, MD

thepowerofpoop.com



#### Welcome to OpenBiome.

#### What we do

We work with clinicians to make FMT easier, cheaper, safer and more widely available. We do so by providing hospitals with screened, filtered, and frozen material ready for clinical use. This service eliminates the time, staff, protocols, and facilities needed to screen and prepare material from new donors for each treatment. With OpenBiome, all that's needed to deliver FMT is a doctor and an endoscope.

#### Why we're here

We founded OpenBiome, a nonprofit 501(c)(3) organization, after watching a friend and family member suffer through 18 months of C. difficile and 7 rounds of vancomycin before finally receiving a successful, life-changing Fecal Microbiota Transplantation (FMT). The remarkable efficacy of this treatment and the great lengths required to receive it convinced us that we needed to help expand access. After many discussions with local clinicians and the FDA, we launched OpenBiome in 2012 to make FMT faster and easier for patients and











EXPLORE

**HOW IT WORKS** 

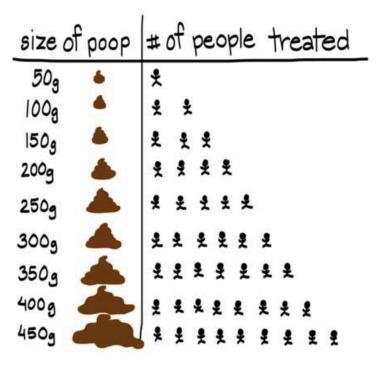
START YOUR CAMPAIGN



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#### OpenBiome's Give a Sh!t Campaign

Medford, Massachusetts, United States



THE MOST IMPORTANT THING YOU'LL DO ALL DAY!



# Sharing

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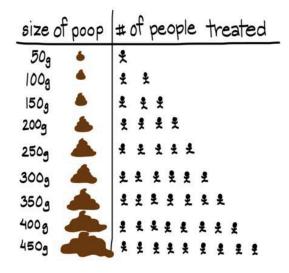












THE MOST IMPORTANT THING YOU'LL DO ALL DAY!



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Volume 506

News & Comment Research Issue 7488

Comment

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NATURE | COMMENT

Archive

#### Policy: How to regulate faecal transplants

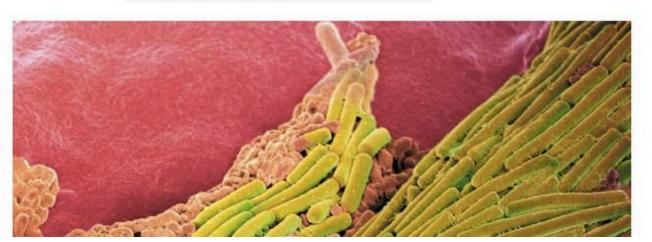
Mark B. Smith, Colleen Kelly & Eric J. Alm

19 February 2014

For medical use, human stool should be considered a tissue, not a drug, argue Mark B. Smith, Colleen Kelly and Eric J. Alm.



Policy · Therapeutics · Infection · Microbiology Subject terms:







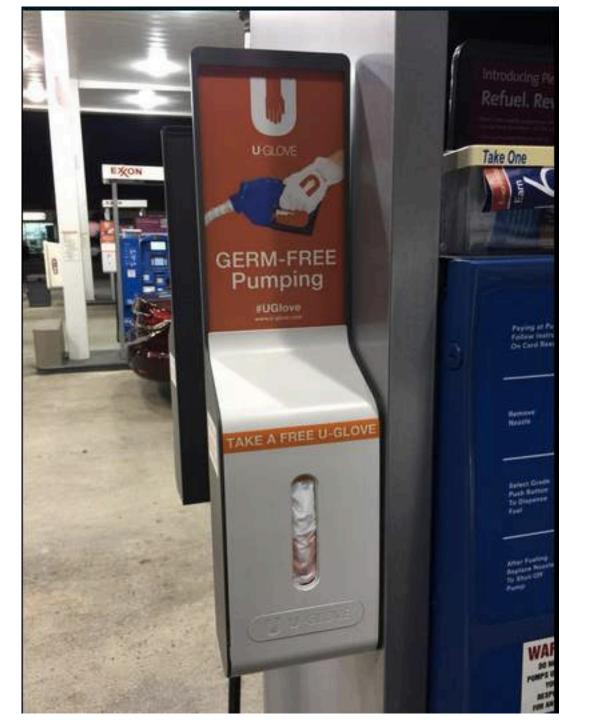


(2)

## Microbiome











# PLEASE.....do not swim when you have diarrhea

Swimmers who are sick with diarrhea—or who have been sick in the last two weeks—risk contaminating pool water with germs. Certain germs that cause diarrhea can live from minutes to days in pools, even if the pool is well-maintained.

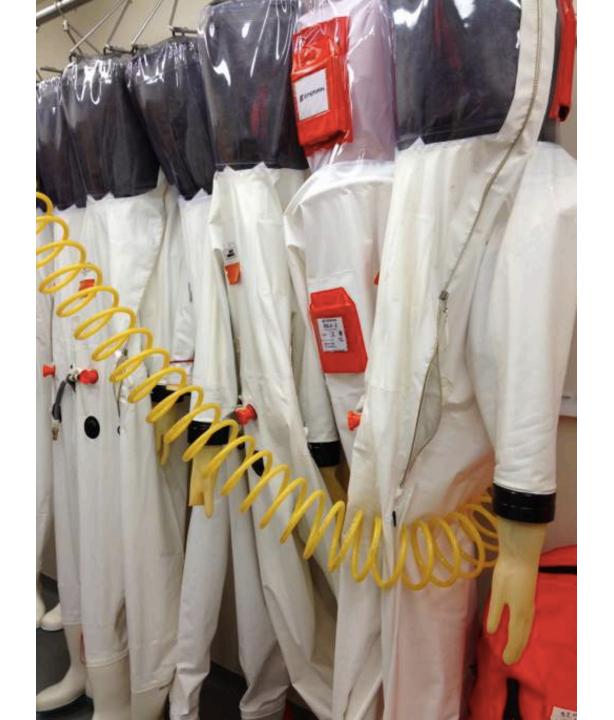
Microscopic amounts of infected fecal matter can contaminate an entire pool or hot tub and make others sick if they swallow the water

# Sharing

Caring













"FEAR is the main source of SUPERSTITION,

and one of the main sources of cruesty.



To conquer fear is the beginning of wisdom."

Bertrand Russell

## FEAR

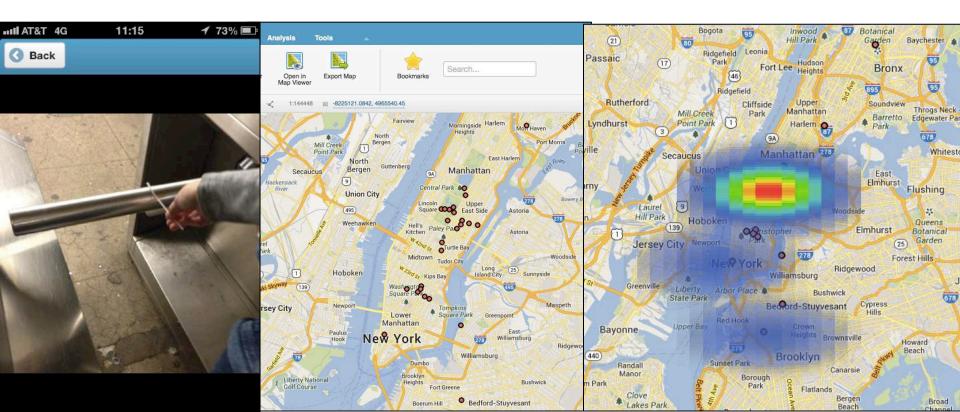
## KNOWLEDGE





## (3) City-scale metagenomics

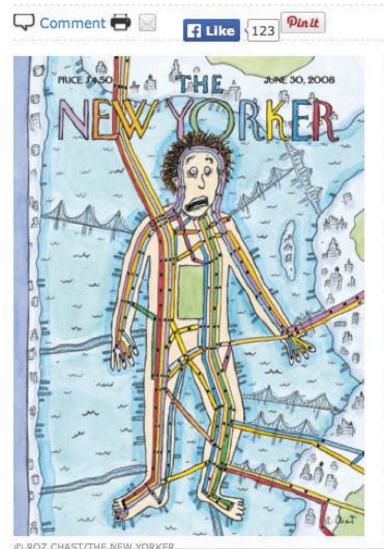
hww.pathomap.org



### Metropolome

Researchers take advantage of rapid and cheap DNA sequencing technologies to map the bacterial microbiome of New York City.

By Jef Akst | December 1, 2013





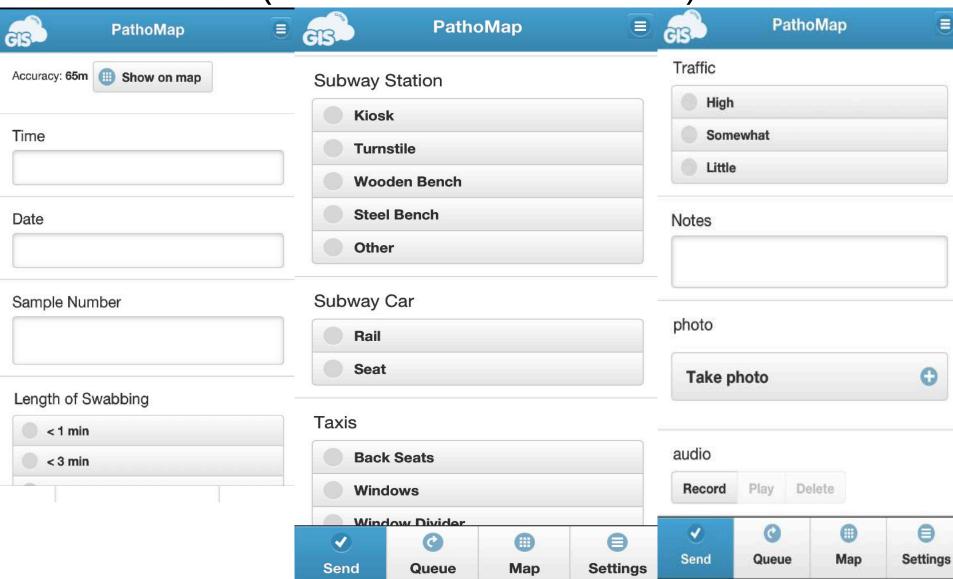


hen Weill Cornell Medical College genomicist Chris Mason would drop his 6-month-old daughter off at day care in 2011, he'd encounter more than a room filled with active toddlers. He'd step into a microbial nightmare. "Seeing how they all shared toys and put them in their mouths, I wanted to swab everything before and after," he recalls.

Mason learned that there was an ongoing day-care microbiome project, led by researchers at Drexel University and Brooklyn College. But he didn't necessarily want to study just the day care's microbiome. Rather, the experience sparked a bigger idea: Mason wondered if he could explore the bacterial microbiome of New York City.

Last summer, Mason led a team of five NYC undergraduate students in the collection of more than 1,400 samples: three from each of the city's 468 subway stations. "We did swabs of the kiosks, the turnstiles, and then trains at each one of the stations," Mason says. "They pretty much just went and swabbed every day all day for about a month and a half." For each sample collected, the students took a photo of the spot, and their smart phones noted the GPS location.

## Open-Source GIS Cloud App (iOS and Android)

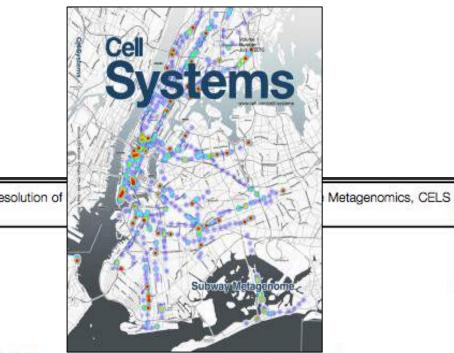


## First city-scale metagenome profile



Cell Systems

**Article** 



OPEN **ACCESS** Cell<sup>2</sup>ress

### Geospatial Resolution of Human and Bacterial **Diversity with City-Scale Metagenomics**

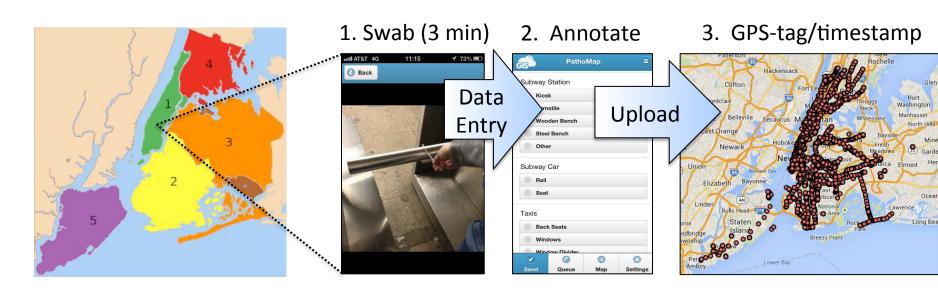
Ebrahim Afshinnekoo, 1,2,3,21 Cem Meydan, 1,2,21 Shanin Chowdhury, 1,2,4 Dyala Jaroudi, 1,2 Collin Boyer, 1,2 Nick Bernstein, 1,2 Julia M. Maritz, 5 Darryl Reeves, 1,2,6 Jorge Gandara, 1,2 Sagar Chhangawala, 1,2 Sofia Ahsanuddin, 1,2,7 Amber Simmons, 1,2 Timothy Nessel, 8 Bharathi Sundaresh, 8 Elizabeth Pereira, 8 Ellen Jorgensen, 9 Sergios-Orestis Kolokotronis, 10 Nell Kirchberger, 1,2 Isaac Garcia, 1,2 David Gandara, 1,2 Sean Dhanraj, 7 Tanzina Nawrin, 7 Yogesh Saletore, 1,2,6 Noah Alexander, 1,2 Priyanka Vijay, 1,2,6 Elizabeth M. Hénaff, 1,2 Paul Zumbo, 1,2 Michael Walsh, 11 Gregory D. O'Mullan,3 Scott Tighe,12 Joel T. Dudley,13 Anya Dunaif,14 Sean Ennis,15,16 Eoghan O'Halloran,15 Tiago R. Magalhaes, 15,16 Braden Boone, 17 Angela L. Jones, 17 Theodore R. Muth, 7 Katie Schneider Paolantonio, 5 Elizabeth Alter, 18 Eric E. Schadt, 13 Jeanne Garbarino, 14 Robert J. Prill, 19 Jane M. Carlton, 5 Shawn Levy, 17 and Christopher E. Mason1,2,20,\*

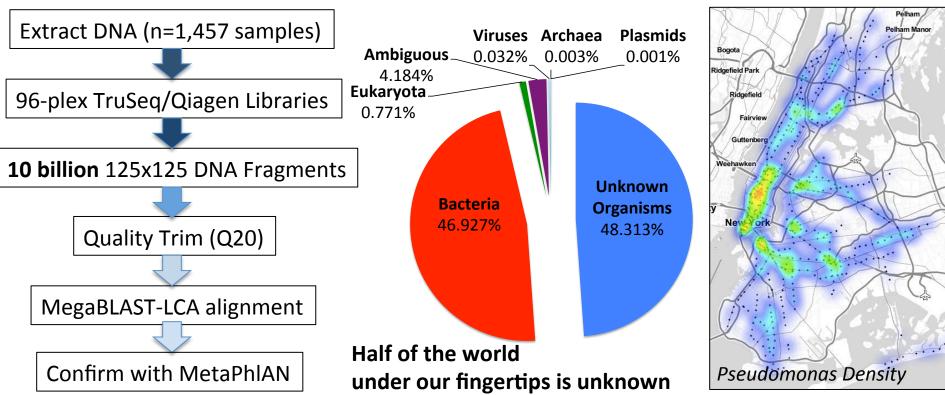
1,427 samples from the NYC Subway:

5.5 million riders/day

656 miles

468 stations











#### Where Certain Types of Bacteria Were Found

Select a category below to see the stations where researchers found the associated types of bacteria.



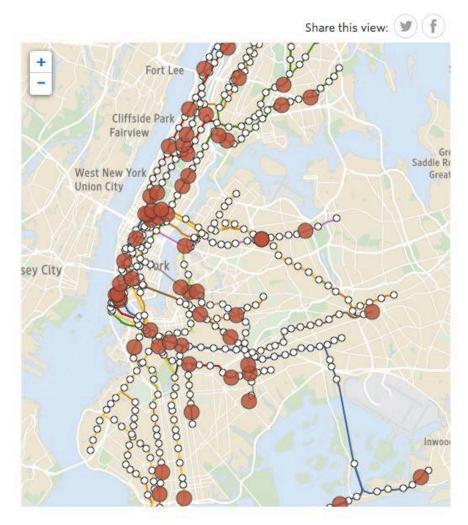
#### CLOSE X

#### Leuconostoc citreum

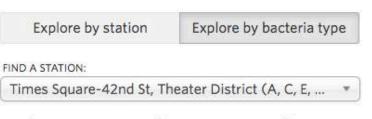
Associated with Kimchi and sauerkraut

A species called Leuconostoc citreum is widely used in the fermentation of the Korean dish kimchi and is also used to ferment cabbage to make sauerkraut.

Medical-device infections			Meningitis		
Mozzarella cheese			Oil cleanup		
Radiatio	n resis	tance	Res	piratory a	ailmen
Sepsis	Staph	Staph infections		Sunscreen	
Swiss cheese		Tetanus		Toxic cleanup	
Urinary-	tract in	fectio	ne		

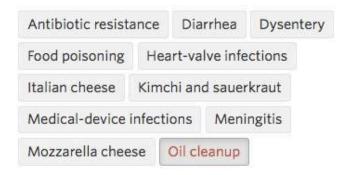


### Pseudomonas putida can help absorb chemicals



#### Where Certain Types of Bacteria Were Found

Select a category below to see the stations where researchers found the associated types of bacteria.

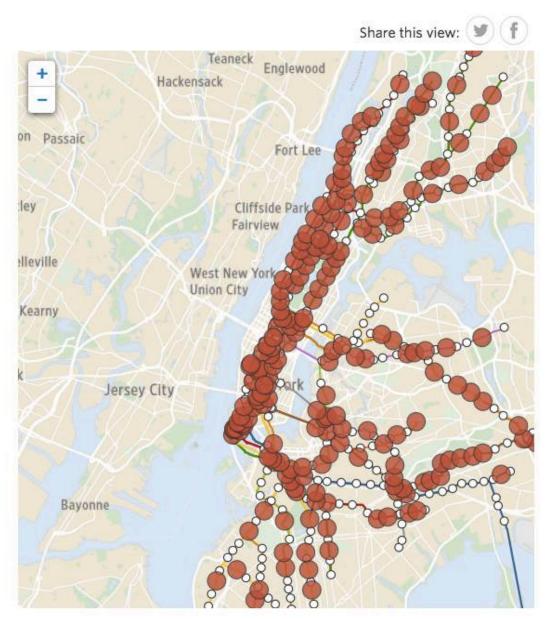


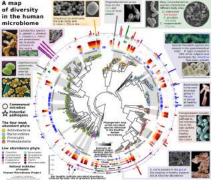
CLOSE X

#### Pseudomonas putida

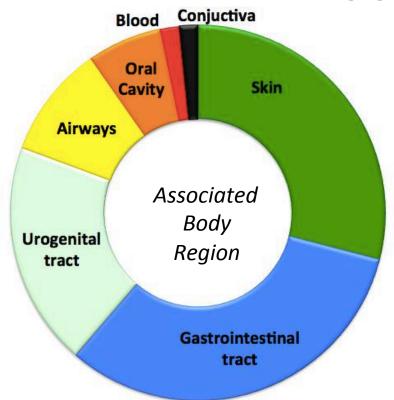
Associated with Oil cleanup

A bacteria species called Pseudomonas putida can break down organic pollutants such as toluene and oil. It is the first patented organism in the world, and, because the bacteria are living creatures, the patent was fought all the way up to the U.S. Supreme Court. Some strains can live on pure caffeine.

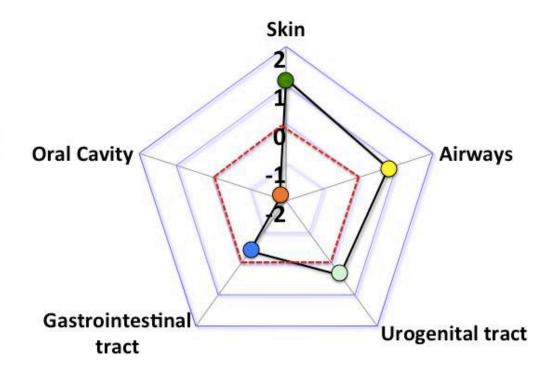




# HMP Comparison Shows That the Subway "Looks Like Skin"

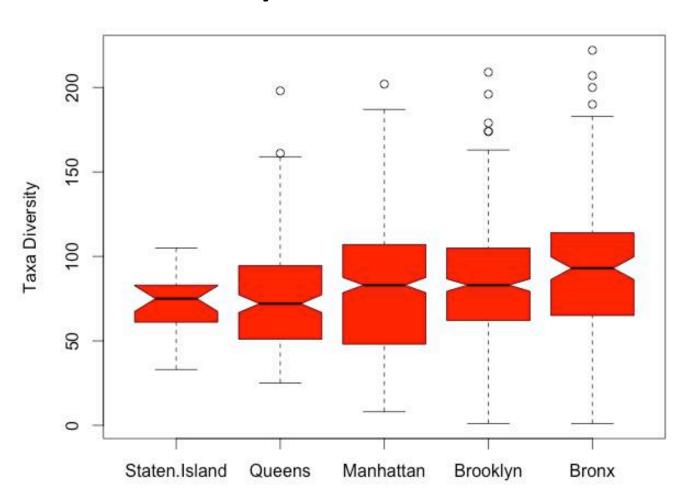


Staphylococcus epidermidis Staphylococcus aureus Acinetobacter radioresistens Propionibacterium acnes

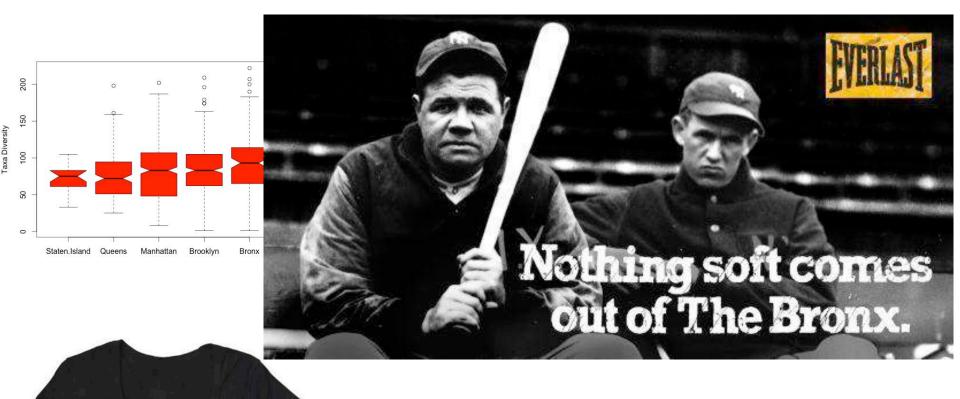


Log<sub>2</sub> Ratio of (Observed/Expected)

## The Bronx is the most diverse! Brooklyn comes in at #2.



### This makes some sense



BROOKLYN IS FOR LOVERS

>600 species ride the subway with you!

Mostly harmless.

Pathogenicity markers absent.

NY Magazine, November 5<sup>th</sup>, 2013



#### Corynebacterium glutamicum

The stairwell at the Eighth Avenue L stop housed this tiny helper in industrial amino-acid production. Researchers also see promise for its use in bioremediation: It removes arsenic from the environment.

#### phingomonas

Turnstiles at the Bedford Avenue L tested positive for this bacterium discovered by scientists two decades ago, in the Elbe River in Germany, and known for its ability to digest toxic compounds sometimes found in chemical spills.



#### radioresistens

The MTA underground (namely turnstiles) harbors a bacterium so robust it can resist exposure to radiation. Scientists found it on the Mars Odyssey orbiter before liftoff and worry it could compromise future life-detection missions.

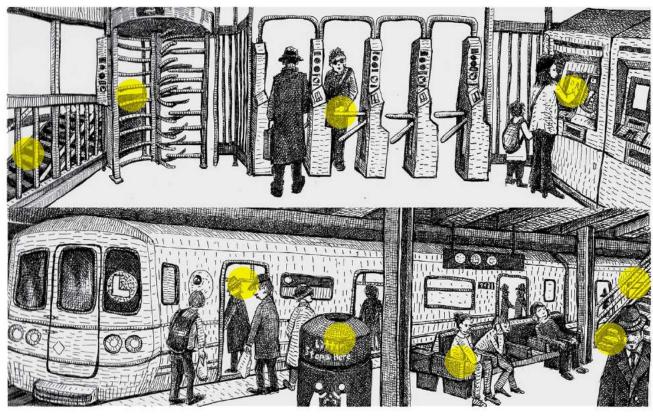


#### actobacillus

A swab from a Union Square turnstile revealed this bacterium, which is related to the kind used to make yogurt; it's even been tested as a probiotic treatment for children with a form of the skin disease eczema.



Mason's team collected this germ, typically found in the Indian subcontinent, from a Bedford Avenue kiosk. It belongs to a genus of bugs responsible for shigellosis, a diarrheal disease that infects up to 800 people a year in New York State.





#### Lysinibacillus sphaericus

Several samples, including one from a 7 train handrail, carried a microbe used to fight insect-borne disease; it produces a toxin that kills mosquito larvae-a useful defense against malaria and West Nile virus.



#### Aerococcus

This bug, found on a Beverley Road garbage can, among other sites, prevs upon weakened immune systems, has been linked to the heart disease endocarditis. and causes gaffkemia, a fatal disease for lobsters.



#### Exiguobacterium sibiricum

#### Found on a Bedford Avenue bench,

Exiguobacterium sibiricum was named for Siberia, where researchers first detected it in a permafrost sample that dates back as far as 3 million years.



#### Enterococcus faecalis

Mason discovered this bacterium, usually found in fecal matter, in several high-traffic areas, like a Times Square handrail. It's behind urinary-tract infections and hospital-

acquired illnesses that now

show antibiotic resistance.



#### Bacillus megaterium

Doctors worldwide have this inhabitant of the Eighth Avenue L stairwell to thank:

It produces key enzymes used to make synthetic penicillin and to test blood-glucose levels in diabetics.

## Anthrax-specific fragments found by Kraken, MetaPhlAn, SURPI, BWA. But scant pXO, no PlcR.

No strong evidence of pathogen. Organism is different from pathogen.

read-lab-confederation.github.io/nyc-subway-anthrax-study/	C A histogenetic	x → ☆ 自 ♣ 1
Searching for anthrax in the New York City subway metagenome.	Study	Number of matched 31-mers with Anthrax 31-mers
April 23, 2015	SRR1749083	0
None 23, 2013	SRR1748708	2
Robert A Petit III Matthew Ezewudo Sandeep J. Joseph Timothy D. Read	SRR1748707	0

#### Introduction

In January 2015 Chris Mason and his team published an in-depth analysis of metagenomic data (environmental shotgun DNA sequence) from samples isolated from public surfaces in the New York City (NYC) subway system. Along with a ton of really interesting findings, the authors claimed to have detected DNA from the bacterial biothreat pathogens *Bacillus anthracis* (which causes anthrax) and *Yersinia pestis* (causes plague) in some of the samples. This predictably led to a huge interest from the press and scientists on social media. The authors followed up with an re-analysis of the data on microbe.net, where they showed some results that suggested the tools that they were using for species identification overcalled anthrax and plague.

B. anthracis is a Gram-positive bacterium that forms tough spores as part of its lifecycle. The 5.2 M basepair (Mb) main chromosome is very similar to those of other bacteria in species informally called the 'Bacillus cereus group' (including B. cereus, B. thuringiensis and B. mycoides). Bacillus cereus group strains in general are commonly found in soil but B. anthracis itself is very rare and generally associated with livestock grazing sites with a past history of anthrax.

What sets *B. anthracis* apart from close relatives is the presence of two plasmids: pXO1 (181kb), which carries the lethal toxin genes and pXO2 (94kb), which includes genes for a protective capsule. Without one of these plasmids, *B. anthracis* is considered attenuated in virulence and unable to cause classic anthrax. Other *B. cereus* group bacteria can have plasmids very similar to pXO1 and pXO2 but missing the important virulence genes. Rarely, other *B. cereus* group carry pXO1 and appear to cause anthrax-like disease. Its a confusing situation, not helped by the current overly-narrow species definitions. This recent review gives more information.

Table 13: Annotated 31-mers mapped to SRR1748708.

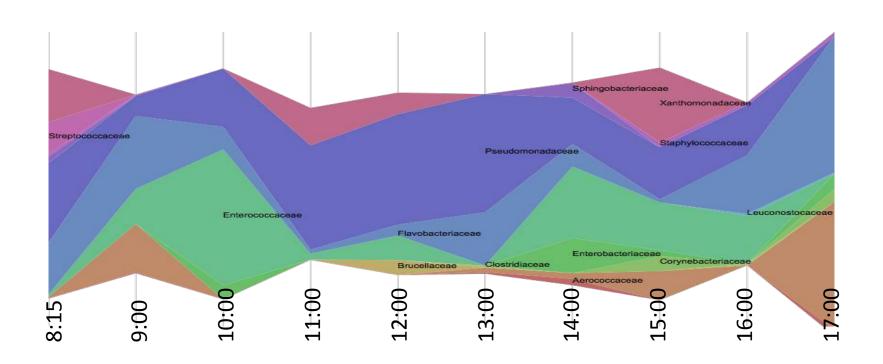
Anthrax-specific K-mers identified from the NYC subway sy SRR1748708 sample.

SNP position on the B. anthracis Ames ancestor genome	31-mer sequence	
4083672	TGTGCCCCATCCTGAGCATACAACTTTATAA	2
4136672	ACATAGAACAAGTGACATTCTATCAAACGGT	2

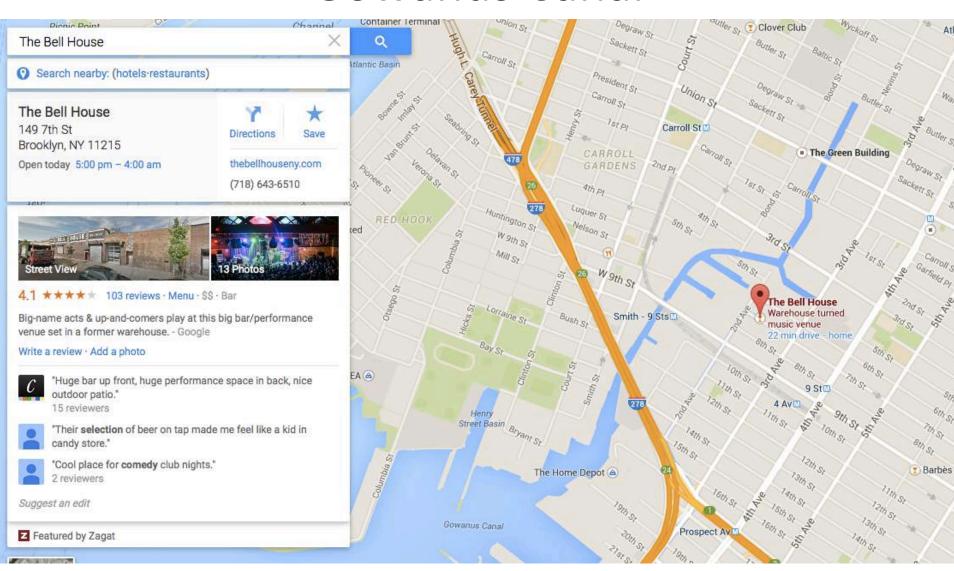
http://read-lab-confederation.github.io/nyc-subway-anthrax-study/

## **(4)** 'molecular echoes'

## Hourly dynamics of a populated kiosk are far more heterogeneous



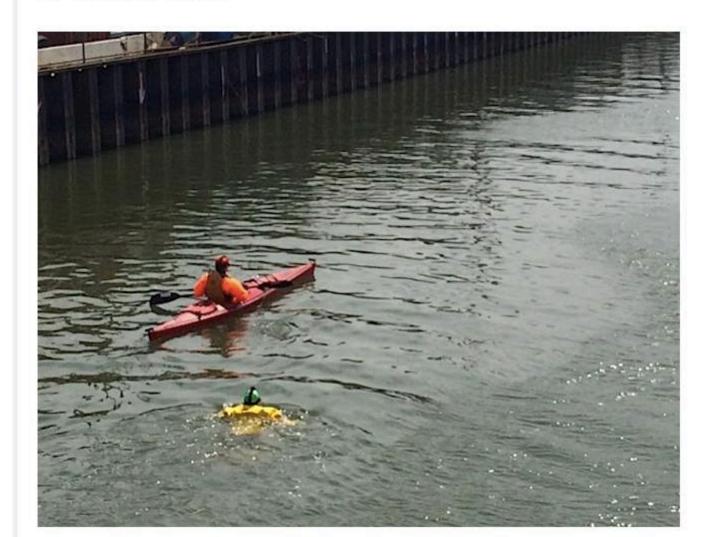
## Some areas are more stable: Gowanus Canal

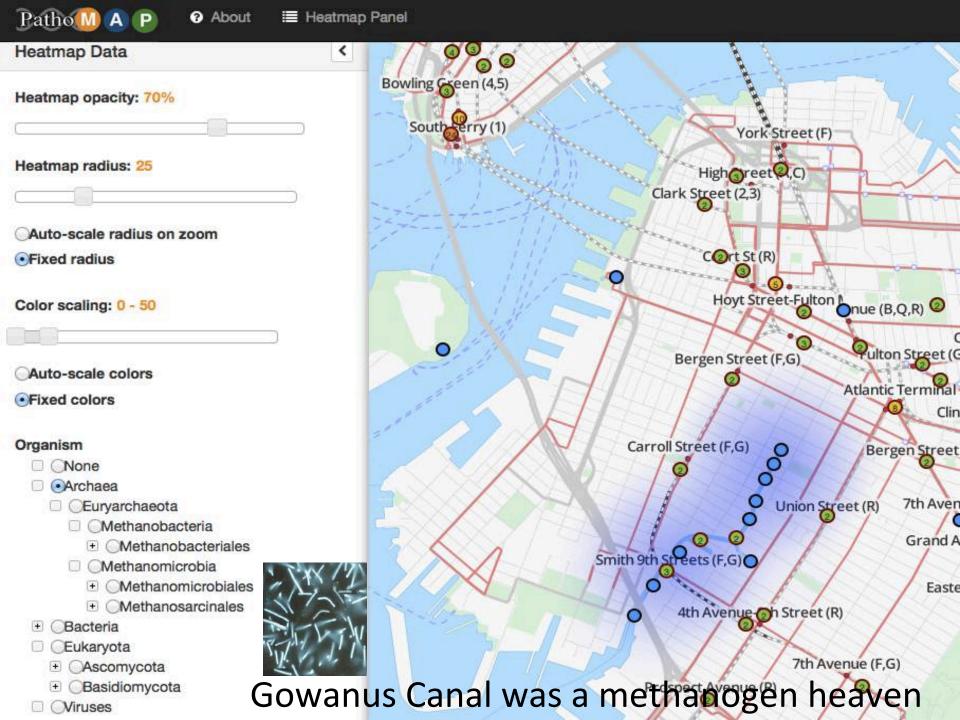




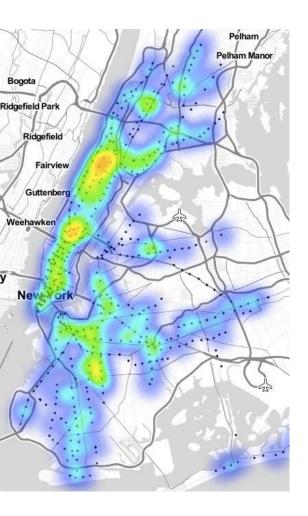


## Man Swims In Gowanus Canal, Concludes "It's Not Safe To Swim In There"

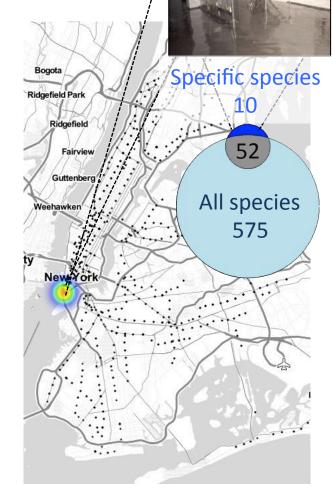




Species diversity varies by area of the city



Bogota Ridgefield Park Ridgefield Guttenberg Weehawken New York



Hurricane-Flooded

Enterococcus faecium

Staphylococcus aureus

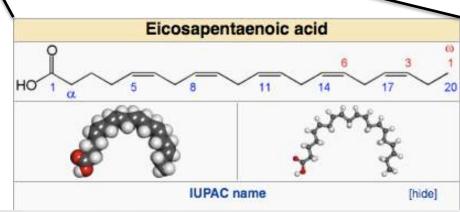
Pseudoalteromonas haloplanktis

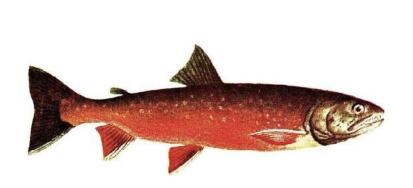


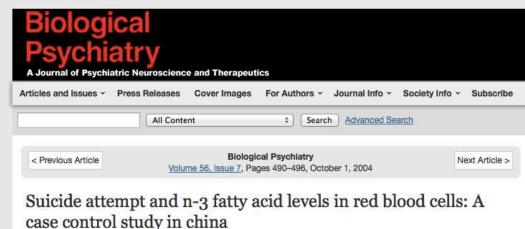
### Shewanella frigidimarina

An Antarctic species with the ability to produce eicosapentaenoic acid. It grows anaerobically by dissimilatory Fe (III) reduction.[1] Its cells are motile and rod shaped

EPA is obtained in the human diet by eating oily fish or fish oil, e.g. cod liver, herring, mackerel, salmon, menhaden and sardines.







Frolova, G. M.; Gumerova, P. A.; Romanenko, L. A.; Mikhailov, V. V. (2011). "Characterization of the lipids of psychrophilic bacteria Shewanella frigidimarina isolated from sea ice of the Sea of Japan". Microbiology 80 (1): 30–36



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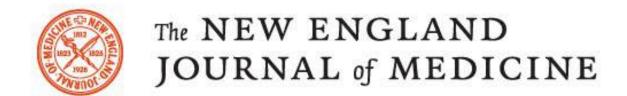
TRENDING:

PORTALS

**NEW YORK BOTANICAL GARDEN** 

ROBERTA'S

### Licking Subway Poles "Probably Fine," Says Expert



HOME

ARTICLES & MULTIMEDIA \*

ISSUES \*

SPECIALTIES & TOPICS \*

FOR AUTHORS \*

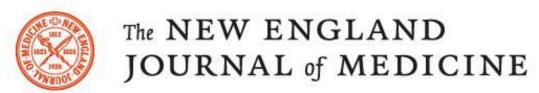
CME >

#### ORIGINAL ARTICLE

#### Randomized Trial of Peanut Consumption in Infants at Risk for Peanut Allergy

George Du Toit, M.B., B.Ch., Graham Roberts, D.M., Peter H. Sayre, M.D., Ph.D., Henry T. Bahnson, M.P.H., Suzana Radulovic, M.D., Alexandra F. Santos, M.D., Helen A. Brough, M.B., B.S., Deborah Phippard, Ph.D., Monica Basting, M.A., Mary Feeney, M.Sc., R.D., Victor Turcanu, M.D., Ph.D., Michelle L. Sever, M.S.P.H., Ph.D., Margarita Gomez Lorenzo, M.D., Marshall Plaut, M.D., and Gideon Lack, M.B., B.Ch. for the LEAP Study Team N Engl J Med 2015; 372:803-813 | February 26, 2015 | DOI: 10.1056/NEJMoa1414850

## Replicated!



HOME

ARTICLES & MULTIMEDIA \*

ISSUES \*

SPECIALTIES & TOPICS \*

FOR AUTHORS \*

CME >

ORIGINAL ARTICLE

#### Effect of Avoidance on Peanut Allergy after Early Peanut Consumption

George Du Toit, M.B., B.Ch., Peter H. Sayre, M.D., Ph.D., Graham Roberts, D.M., Michelle L. Sever, M.S.P.H., Ph.D., Kaitie Lawson, M.S., Henry T. Bahnson, M.P.H., Helen A. Brough, M.B., B.S., Ph.D., Alexandra F. Santos, M.D., Ph.D., Kristina M. Harris, Ph.D., Suzana Radulovic, M.D., Monica Basting, M.A., Victor Turcanu, M.D., Ph.D., Marshall Plaut, M.D., and Gideon Lack, M.B., B.Ch., for the Immune Tolerance Network LEAP-On Study Team March 4, 2016 DOI: 10.1056/NEJMoa1514209

Abstract

Article

References

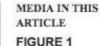
Citing Articles (1)

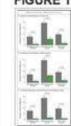
Metrics

Share: 🚮 🐸 🐼 🛅 🔠

#### BACKGROUND

In a randomized trial, the early introduction of peanuts in infants at high risk for allergy was shown to prevent peanut allergy. In this follow-up study, we investigated whether the rate of peanut allergy remained low after 12 months of peanut avoidance among participants who had consumed peanuts during the primary trial (peanut-consumption group), as compared with those who had avoided peanuts (peanut-avoidance group).





Primary Outcome

## The Hygiene Hypothesis

The hygiene hypothesis is a hypothesis that states that a lack of early childhood exposure to infectious agents, symbiotic microorganisms (e.g. gut flora or probiotics), and parasites increases susceptibility to allergic diseases by suppressing the natural development of the immune system.

"Infants born by cesarean delivery are at increased risk of asthma, obesity and type 1 diabetes, whereas breastfeeding is variably protective against these and other disorders."

- Rob Knight





http://blog.ted.com/how-microbes-could-cure-disease-rob-knight-at-ted2014/



## Effects of early-life exposure to allergens and bacteria on recurrent wheeze and atopy in urban children

Susan V. Lynch, PhD,<sup>a\*</sup> Robert A. Wood, MD,<sup>b\*</sup> Homer Boushey, MD,<sup>a</sup> Leonard B. Bacharier, MD,<sup>c</sup>
Gordon R. Bloomberg, MD,<sup>c</sup> Meyer Kattan, MD,<sup>d</sup> George T. O'Connor, MD,<sup>e</sup> Megan T. Sandel, MD,<sup>e</sup>
Agustin Calatroni, MS,<sup>f</sup> Elizabeth Matsui, MD,<sup>b</sup> Christine C. Johnson, PhD,<sup>g</sup> Henry Lynn, PhD,<sup>f</sup> Cynthia M. Visness, PhD,<sup>f</sup>
Katy F. Jaffee, MS,<sup>f</sup> Peter J. Gergen, MD,<sup>h</sup> Diane R. Gold, MD, MPH,<sup>i</sup> Rosalind J. Wright, MD, MPH,<sup>i</sup> Kei Fujimura, PhD,<sup>a</sup>
Marcus Rauch, PhD,<sup>a</sup> William W. Busse, MD,<sup>j</sup> and James E. Gern, MD<sup>j</sup>
San Francisco, Calif, Baltimore, Md, St Louis, Mo,
New York, NY, Boston, Mass, Chapel Hill, NC, Detroit, Mich, and Madison, Wis

Background: Wheezing illnesses cause major morbidity in infants and are frequent precursors to asthma. Objective: We sought to examine environmental factors associated with recurrent wheezing in inner-city environments. Methods: The Urban Environment and Childhood Asthma study examined a birth cohort at high risk for asthma (n = 560)in Baltimore, Boston, New York, and St Louis. Environmental assessments included allergen exposure and, in a nested case-control study of 104 children, the bacterial content of house dust collected in the first year of life. Associations were determined among environmental factors, aeroallergen sensitization, and recurrent wheezing at age 3 years. Results: Cumulative allergen exposure over the first 3 years was associated with allergic sensitization, and sensitization at age 3 years was related to recurrent wheeze. In contrast, first-year exposure to cockroach, mouse, and cat allergens was negatively associated with recurrent wheeze (odds ratio, 0.60, 0.65, and 0.75, respectively;  $P \le .01$ ). Differences in house dust bacterial content in the first year, especially reduced exposure to specific Firmicutes and Bacteriodetes, was associated with atopy and

atopic wheeze. Exposure to high levels of both allergens and this subset of bacteria in the first year of life was most common among children without atopy or wheeze.

Conclusions: In inner-city environments children with the highest exposure to specific allergens and bacteria during their first year were least likely to have recurrent wheeze and allergic sensitization. These findings suggest that concomitant exposure to high levels of certain allergens and bacteria in early life might be beneficial and suggest new preventive strategies for wheezing and allergic diseases. (J Allergy Clin Immunol 2014;134:593-601.)

Key words: Asthma, atopy, allergen exposure, microbial exposure, inner city

Wheezing illnesses affect 35% to 50% of children by the age of 3 years<sup>1,2</sup> and are a leading cause for outpatient visits and hospitalizations.<sup>3,4</sup> Wheezing in nonatopic children is often transient, but recurrent wheezing in children with early allergic sensitization or other signs of atopy during the preschool years

### Cockroaches are the best for young bodies?

**596 LYNCH ET AL** 

### TABLE III. Association between bedroom dust allergen expos

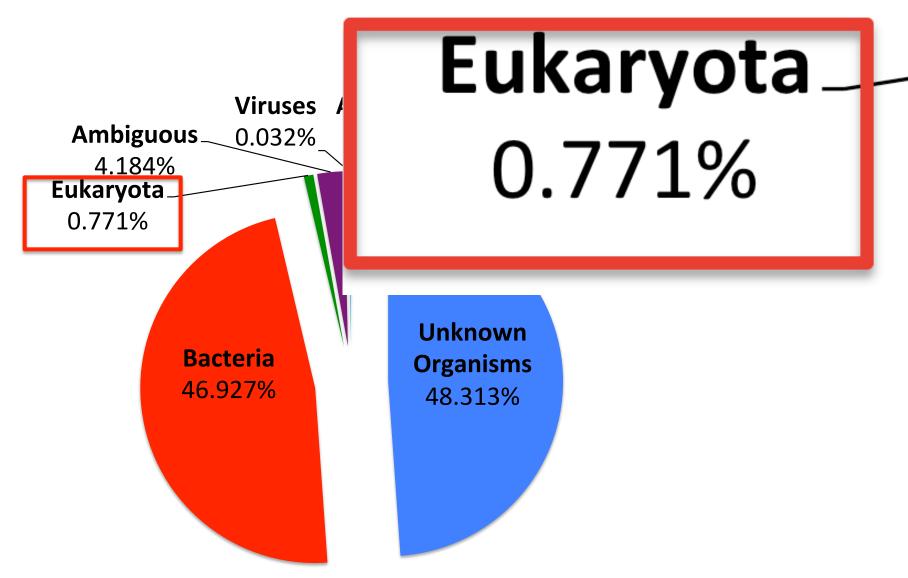
		Unadjusted		
	No.	Odds ratio (95% CI)	P value	
Exposures (year 1)				
Cat	362	0.75 (0.61-0.92)	.005	
Dog	359	1.01 (0.80-1.27)	.97	
Cockroach	359	0.60 (0.45-0.80)	<.001	
Mouse	366	0.65 (0.52-0.82)	<.001	
Dust mite (Dermatophagoides farinae)	364	0.97 (0.78-1.20)	.78	

# (5) Other DNA left behind?

## Metagenomics

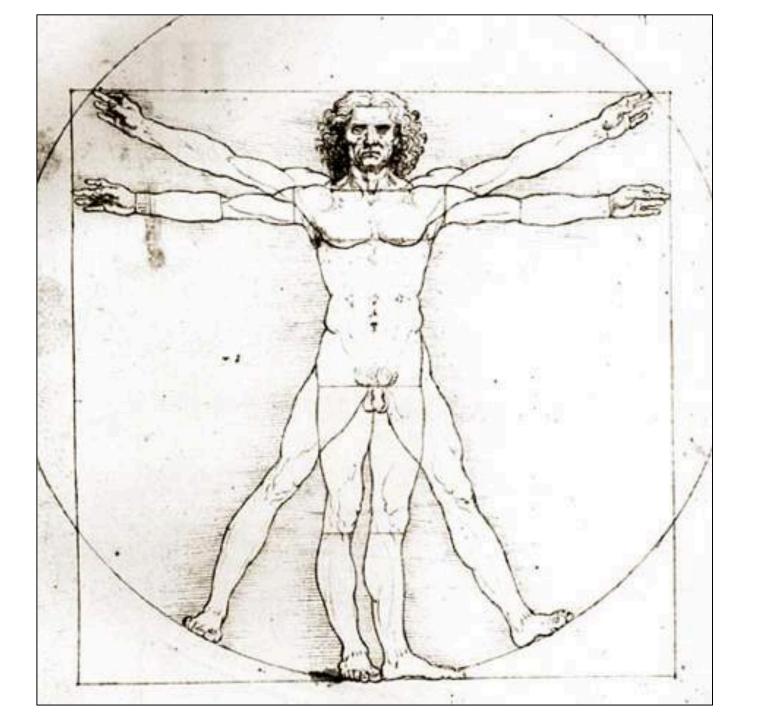
All kingdoms of life and their DNA

### What else is there?

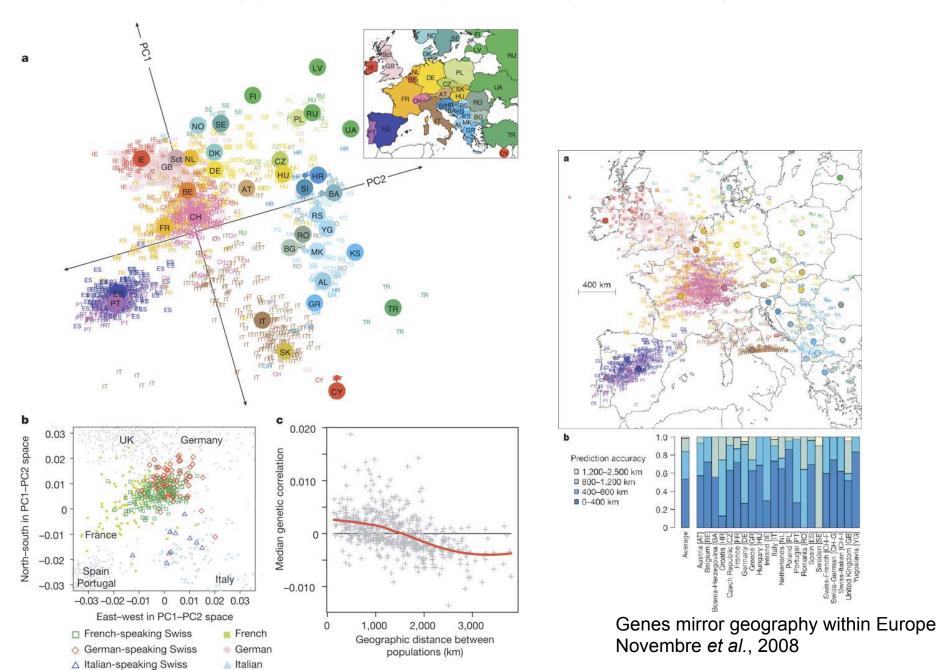


## **BLAST Results**





### Genotype data can predict your birthplace

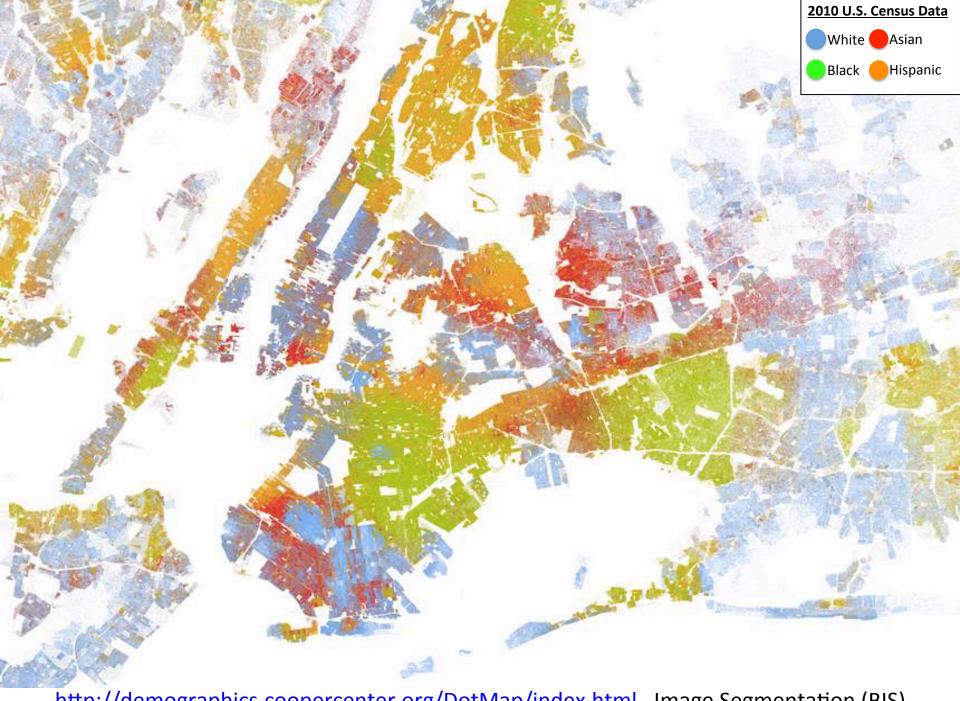


## Artist creates faces from DNA left in public

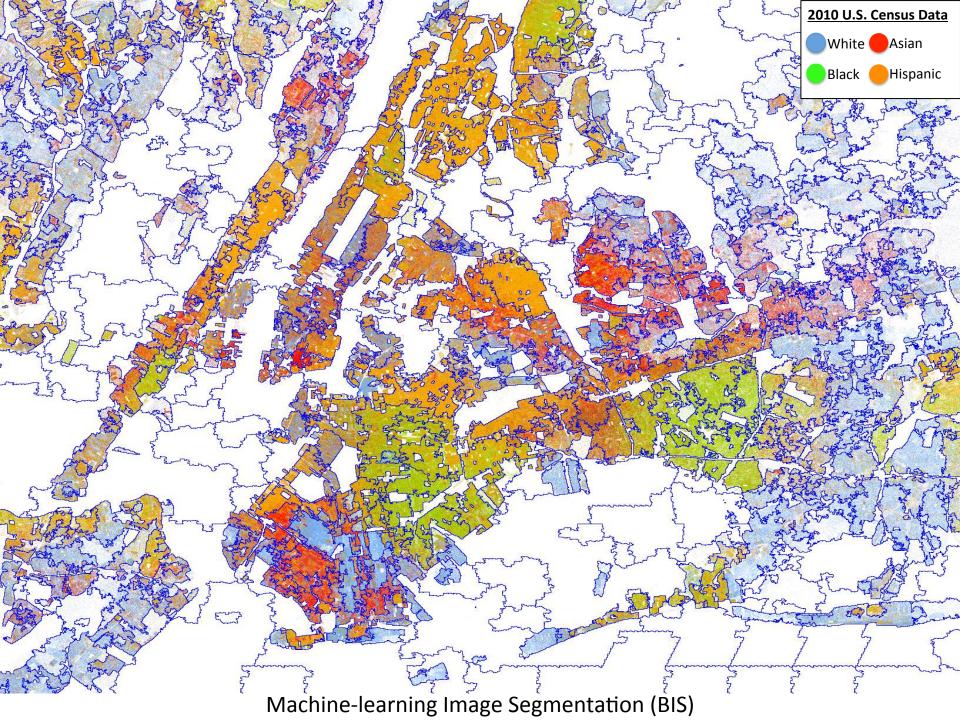
By Natalie Angley, CNN updated 3:10 PM EDT, Wed September 4, 2013 | Filed under: Innovations

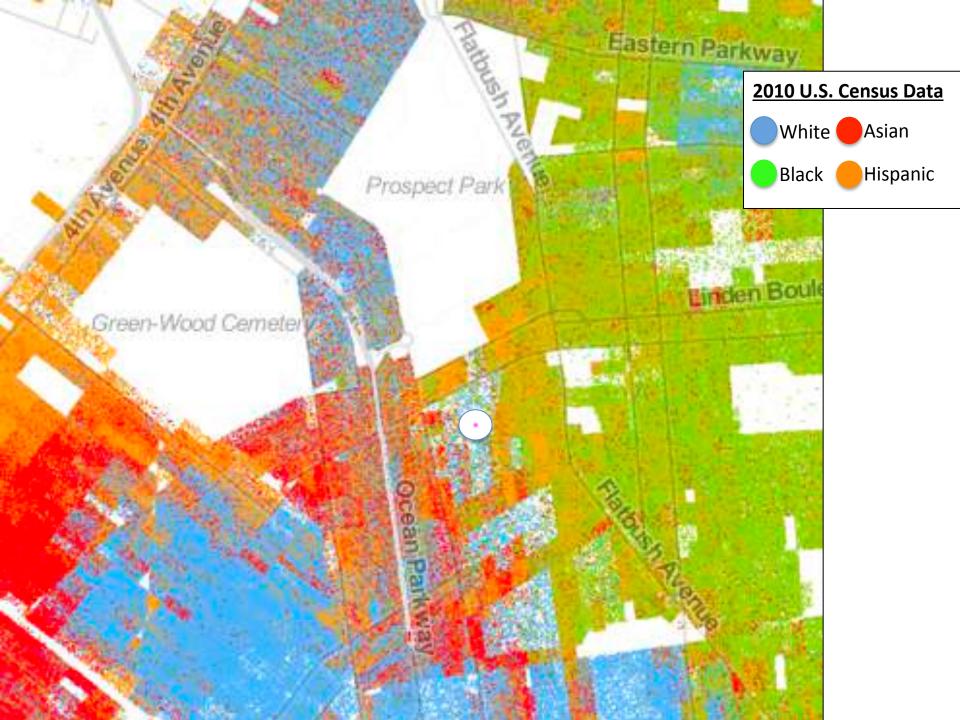


from Heather Dewey-Hagborg's "Stranger Visions" at Genspace (Brooklyn) http://www.cnn.com/2013/09/04/tech/innovation/dna-face-sculptures/

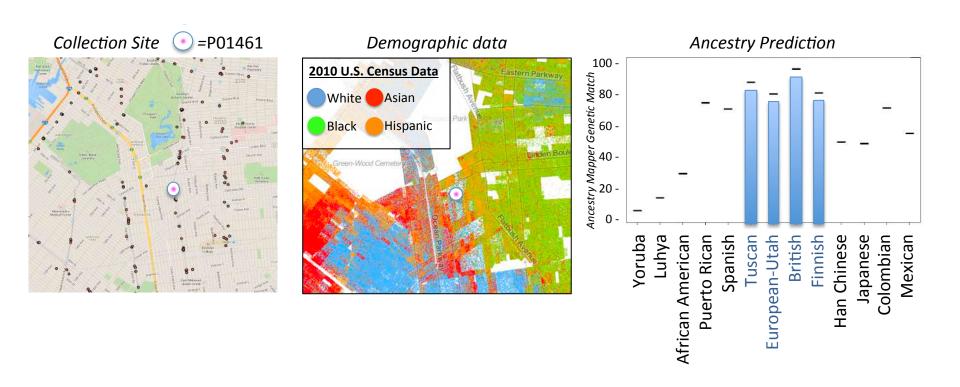


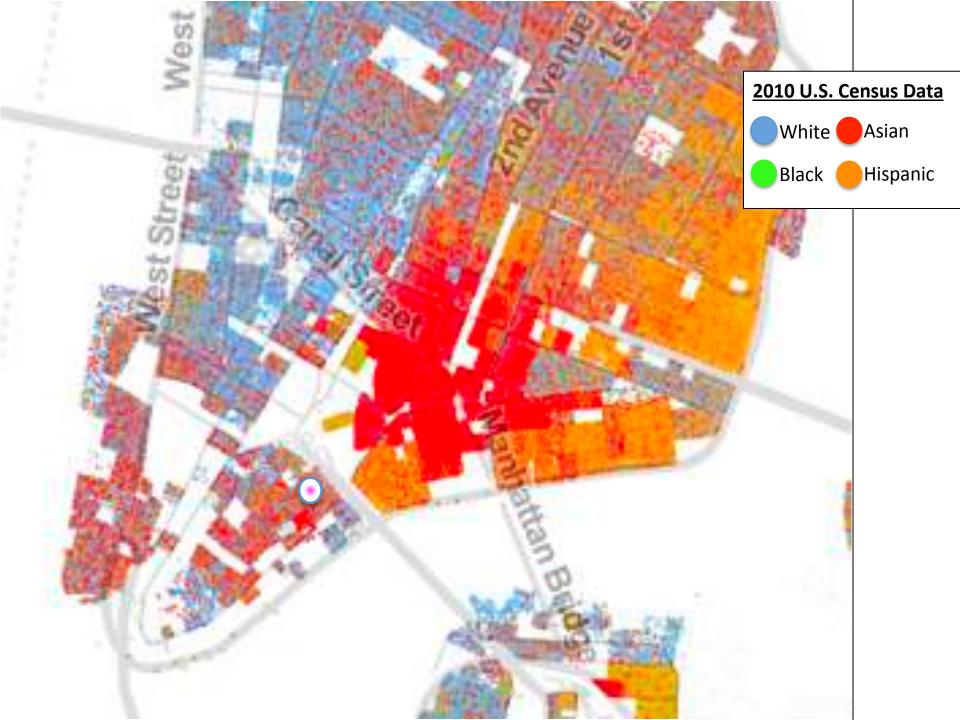
http://demographics.coopercenter.org/DotMap/index.html. Image Segmentation (BIS)



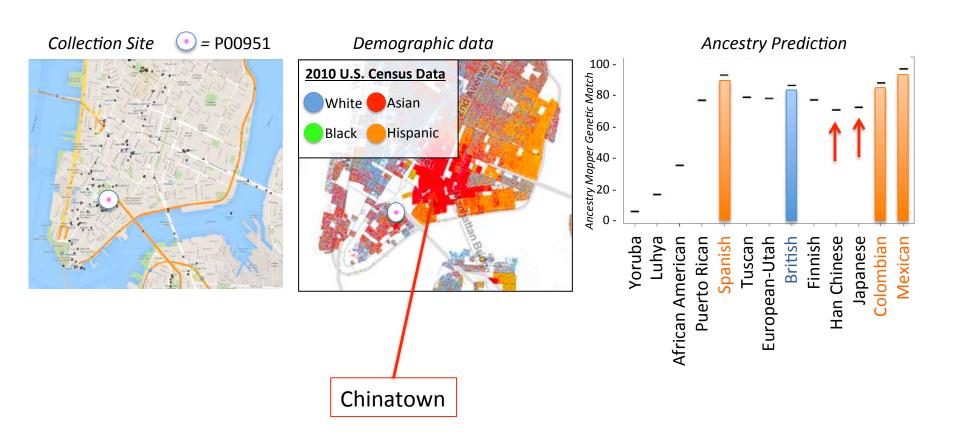


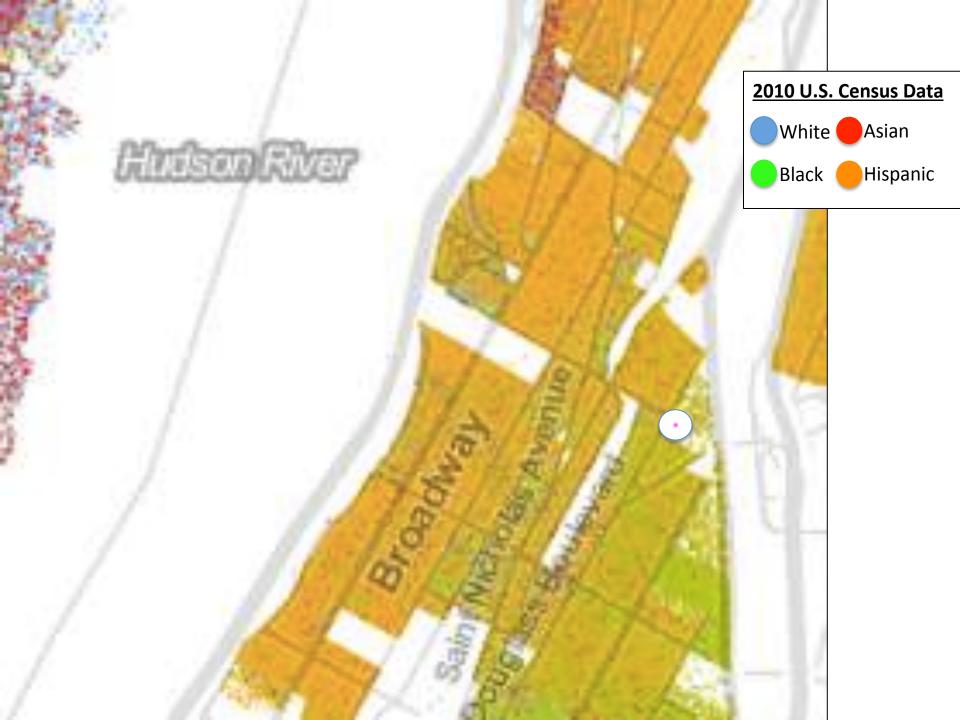
## Predicted Ancestry of DNA left behind can Mirror Census Data in "White" areas



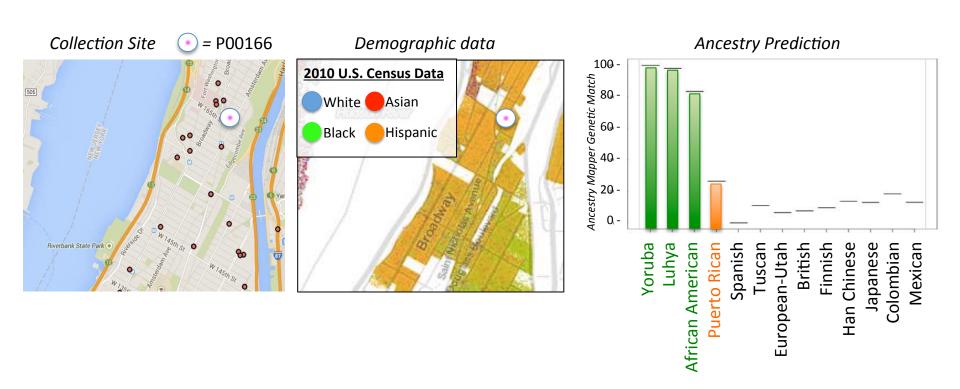


# Alleles appear more Hispanic and more Asian in downtown Manhattan

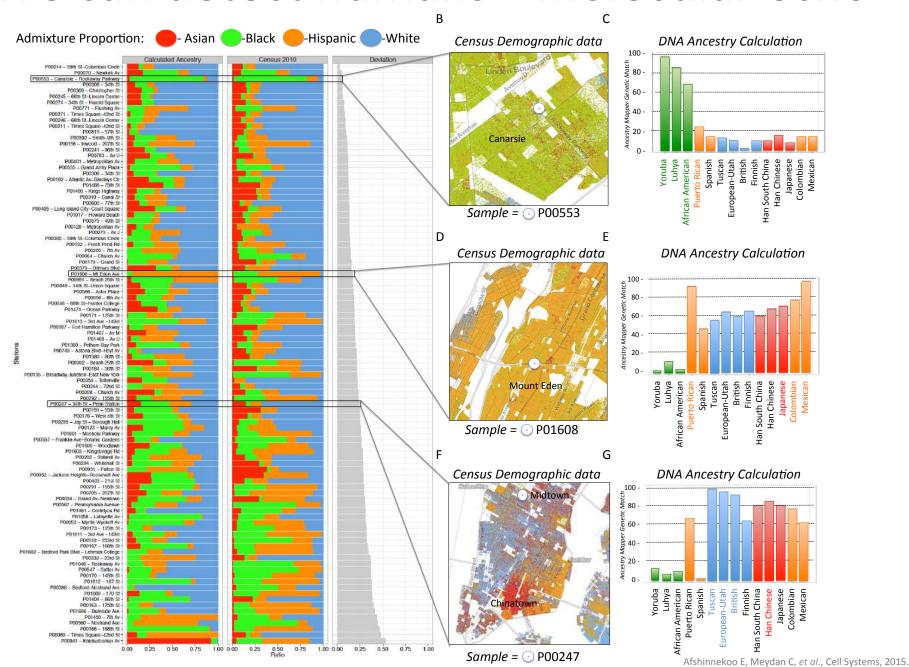




# North Harlem and Washington Heights show more Yoruban alleles and Puerto Rican alleles



## We can detect humans' "molecular echo"



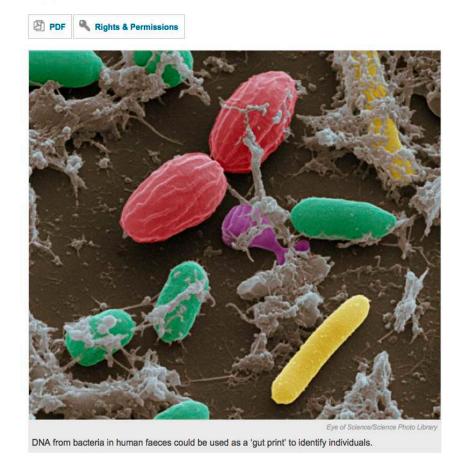


#### Microbiomes raise privacy concerns

DNA from microbes living on the human body can be used to identify individuals.

#### **Ewen Callaway**

11 May 2015



Call it a 'gut print'. The collective DNA of the microbes that colonize a human body can uniquely

### You can choose what DNA to leave behind



A G





Biononymous.me

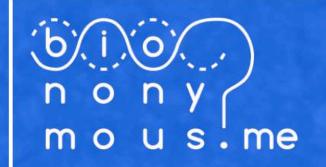
**DIY Guides** 

Legal

Reading

Forum

Resources



A hub for community research into biological privacy.

#### ABOUT

Biological surveillance is the means by which biological science is used to track, monitor, analyze, and turn bodies into data.

It is the extraction of DNA and microbes from our skin, nails, hair and body fluids.

It is the analysis of identifying body parts like faces, fingerprints and irises.

It is the the tracking of life itself by body heat, pulse, perspiration, and involuntary movement.

It is the vulnerability we each face every day by the very situation of being human, by simply having a body.

We believe that privacy should be a choice.

#### CONTRIBUTORS

**Heather Dewey-Hagborg** 

Fei Liu

Aurelia Moser

**Adam Harvey** 

Paul Vanouse

Ben Brucato

Jeremy Gruber

Claire Pentecost

**Brian Holmes** 

Marc R. Dusseiller

Elizabeth Joh

Alondra Nelson

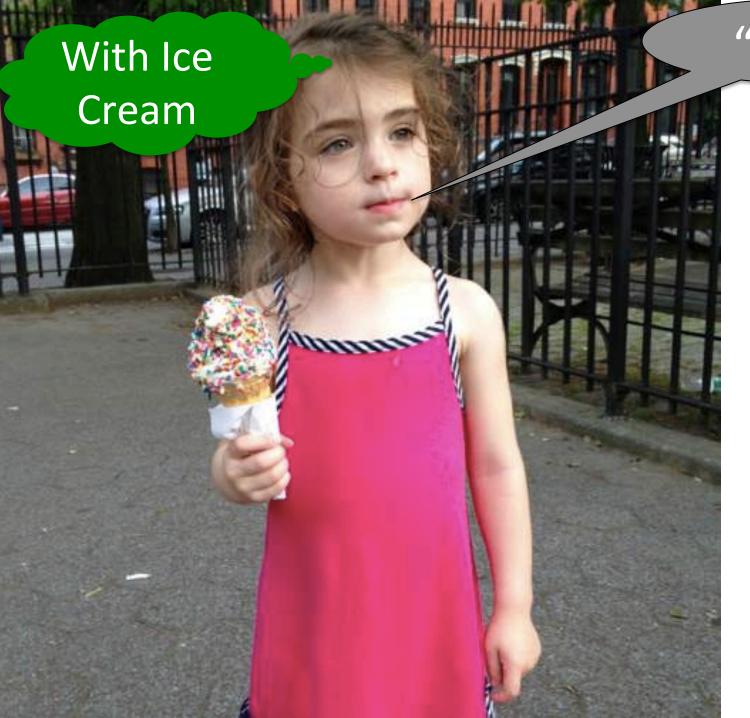
Ignacio Nieto



(6)

## Next Steps

# Should I ride the subway?



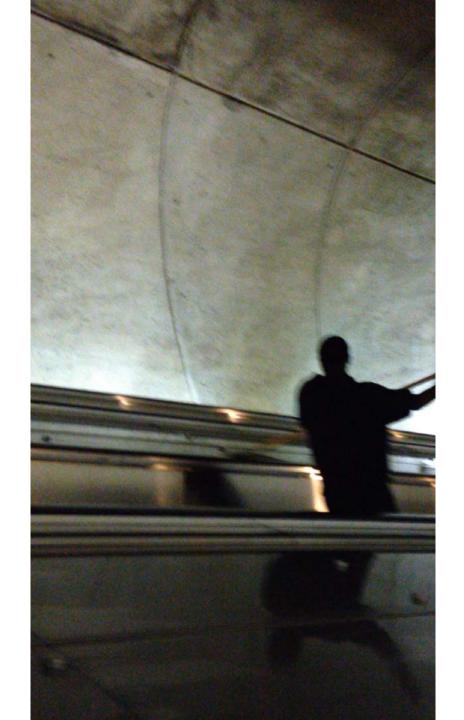
"Yes!"







Washington D.C.



# Global Context

#### MetaSUB: Metagenomics & Metadesign of Subways & Urban Biomes



#### www.metasub.org

		City Details							Site Principal Investigator		
Site	Status	City	Country	Gov't Status	Total # Stations	Transit System Length (km)	Riders / year (billions)	Population (people)	Department	University / Institute	Contact PI
1	Ongoing	Beijing	China	Pending	318	527	3.387	21,150,000	Beijing Children's Hospital	Capital Medical University	Yongli Guo / Yiming
2	Planned	Seoul	South Korea	Approved	377	464	2.563	<b>10</b> , <mark>117,909</mark>	Microbiology	Institut Pasteur Korea	Soojin Jang
3	Ongoing	Shanghai	China	Approved	337	548	2.519	24,150,000	Personalized Genomics	Fudan University	Leming Shi
4	Planned	Moscow	Russia	Started	196	328	2.491	<b>11,5</b> 03,501	Genomics	Institute for Physical)Chemical Medicine,	Dmitry Alexeev
5	Ongoing	Tokyo	Japan	Approved	295	316	2.351	13,185,502	Computational Biology Research Ctr.	National Institute of Advanced Science	Haruo Suzuki
6	Ongoing	Guangzhou	China	Pending	130	240	2.280	<b>12,700</b> ,800	Systems Biology	Sun Yat-sen University	Zhi Xie
7	Ongoing	New York City	USA	Pending	468	436	1.708	8,405,837	Physiology and Biophysics	Weill Cornell Medical College	Christopher Mason
8	Ongoing	Mexico City	Mexico	Pending	195	227	1.685	<mark>8</mark> ,851,080	National Institute of Public Health	IANPHI Mexico Secretariat	Celia M. Alpuche Aranda / Jesus Martinez
9	Ongoing	Hong Kong	China	Pending	87	175	1.618	7,219,700	Computer Science and Engineering	City University	Patrick Lee
10	Ongoing	Paris	France	Pending	303	214	1.527	2,273,305	Analytical Genomics	University Pierre et Marie Curie	Hugues Richard / Ingrid Lafontaine
11	Ongoing	Barcelona	Snain	Approved	131	178	1 036	4 576 394	Genomic and Epigenomic	Centre for Genomic Regulation	Stephan Ossowski / Daniela Bezdan

4,576,394

5,399,343

**11,89**5,893

**11,0**00,000

2,627,000

4,180,000

1,502,604

4,576,394

6,763,000

577,050

1,377,000

530,847

Centre for Genomic Regulation

Pennsylvania State University

University of Sao Paolo

Memorial Sloan Kettering

University of Rome Sapienza

Harvard School of Public

Health

Protection and Societal

Security Division

University of Technology

Corpogen

University of Sheffield

Univeristy of Waikato

IPATIMUP

Variation
Biochemistry and Molecular

Biology

Genetics and Bioinformatics

Computational Biology

Molecular Biology Section

Biostatistics

Protection and Societal

Security Division

Computational

Metagenomics

Metagenomics

rtment of Animal & Plant Sci Environmental Research

Institute

Population Genetics

Stephan Ossowski / Daniela Bezdan

Stephan Schuster

Houtan Noushmehr / Emmanuel Dias-Neto

Sikander Hyat

Romano Florigio Lista / Anna Anselmo

Curtis Huttenhower

Marius Dybwad

Aaron Darling /Catherine Burke

Carlos Ruiz / Maria Zambrano

Eran Elhaik

Christopher Dada

Manuela Oliveira

1.036

0.922

0.921

0.799

0.279

0.166

0.083

0.031

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0.011

0.004

City Details

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Oslo

Sydney

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Sheffield

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Lisbon

Spain

Singapore

Brasil

India

Italy

USA

Norway

Australia

Colombia

UK

New Zealand

Portugal

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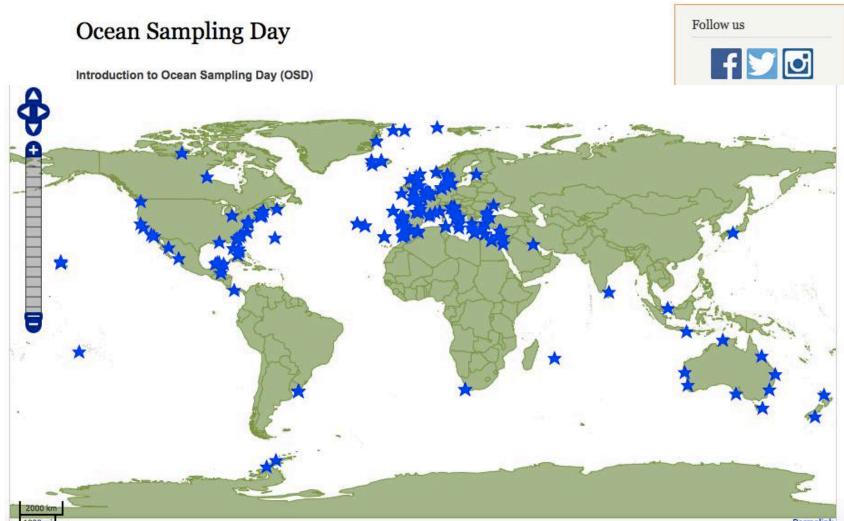
43

## Global Swab Metagenome Festival

http://www.metasub.org/interactive-map.html







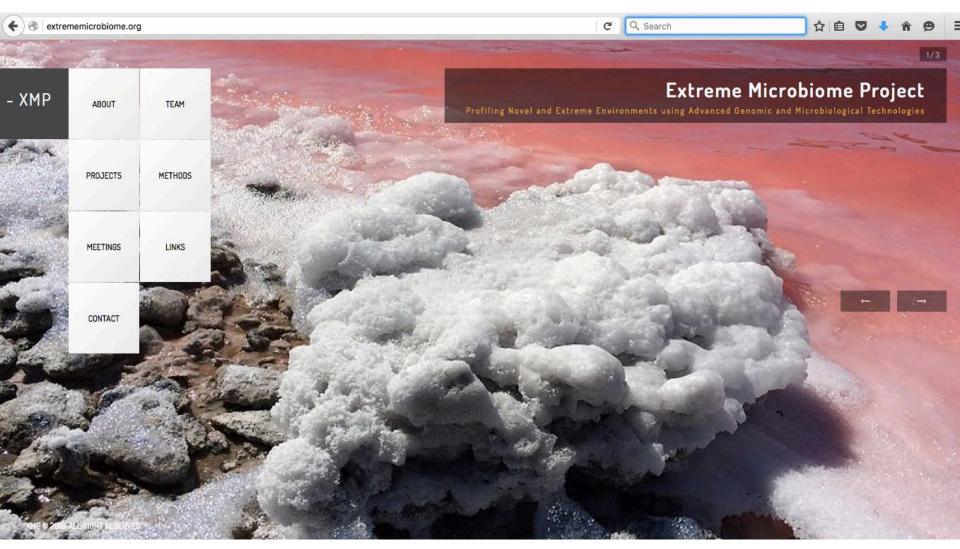
## The Olympiome – Rio 2016 Before, During, and After





# Extreme Context

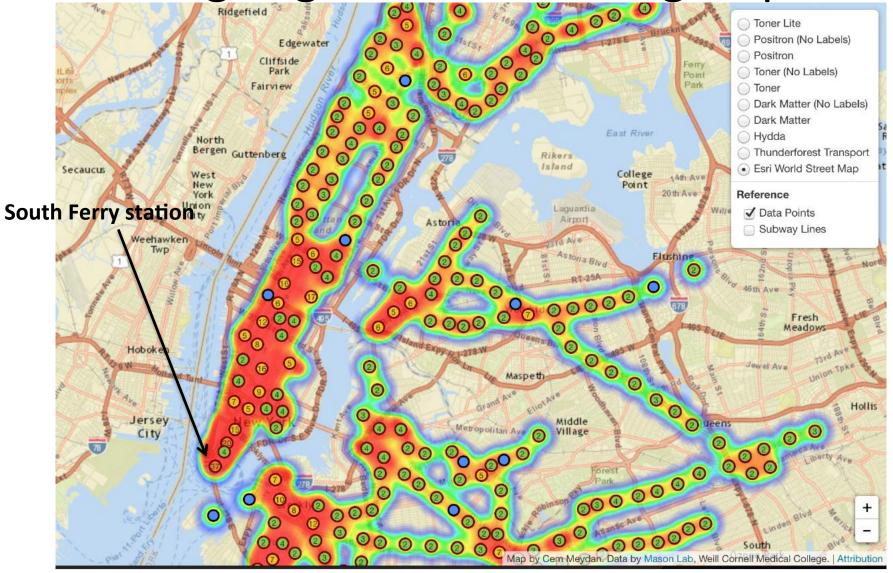
# Extreme Microbiomes for New Biology and Drug Discovery

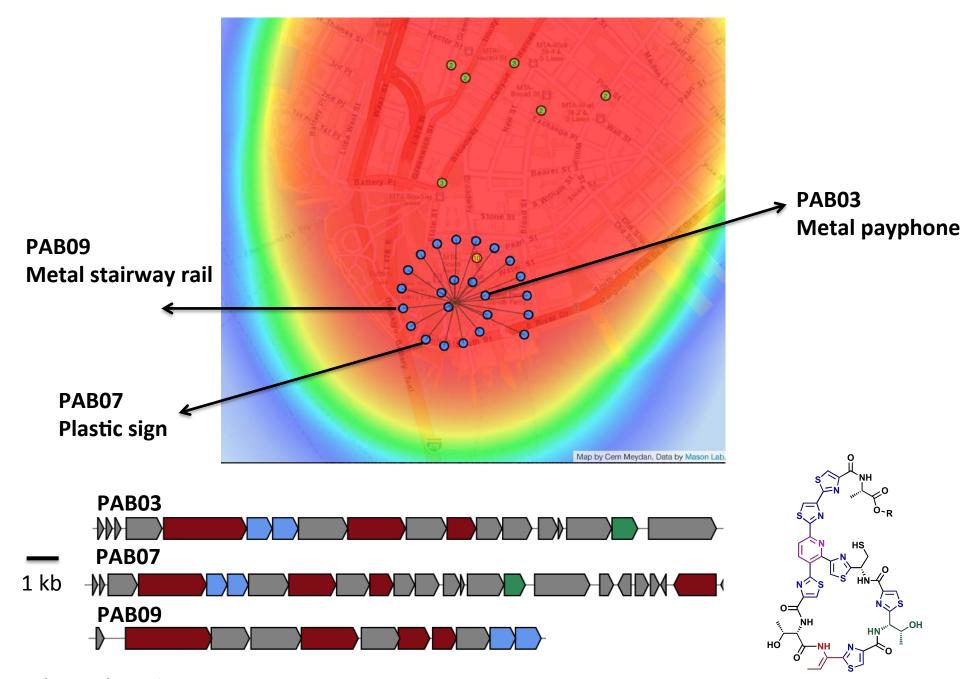


http:/www.extrememicrobiome.org



Biosynthetic Gene Clusters show new drugs right under our fingertips

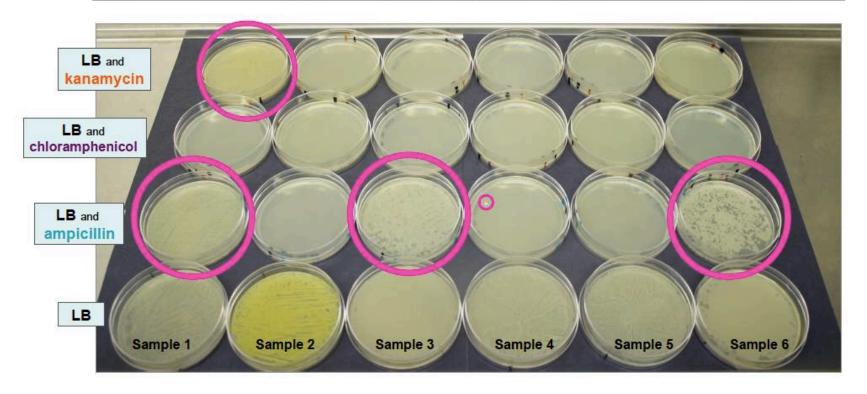




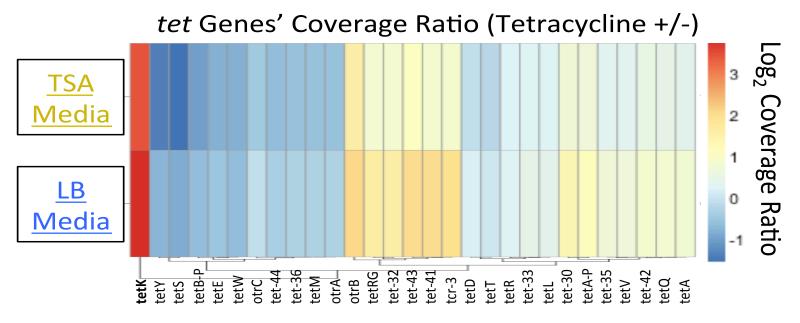
## **(7)** Precision Metagenomics

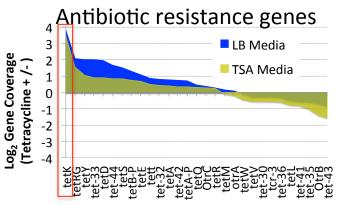
## Evidence of live & antibiotic resistant bacteria

Bacterial Gro	wth from S	Subway Sta	tion Sample	s on Antibi	otic Plates	
	Sample 1	Sample 2 63rd St.	Sample 3 68th St.	Sample 4 96th St.	Sample 5	Sample 6
LB + kanamycin	growth	no growth	no growth	no growth	no growth	no growth
LB + chloramphenicol LB + ampicillin LB	no growth growth growth	no growth no growth growth	no growth growth growth	no growth one colony growth	no growth no growth growth	no growth growth growth



## Metagenomics reveals the likely source of Tetracycline resistance (TetK) on both media





## Examine hospital settings – at Chicago (Jack Gilbert) and now at WCMC

Home Goals

Overview

Design

Timeline

Consortium

**Findings** 

Affiliations



This study aims to collect microbial samples from surfaces, air, staff, and patients from the University of Chicago's new hospital pavilion in order to better understand the factors that influence bacterial population development in healthcare environments.

**Study Design** 



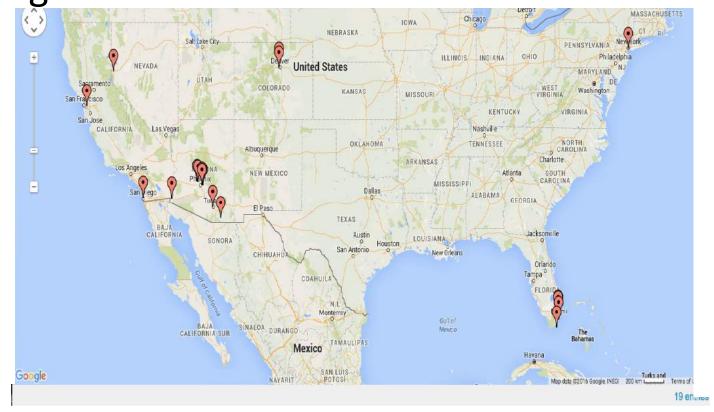
## What if wasn't the hospital that carried the infection?

- Currently conducting pilot study in ambulances across the US.
- Building HAI/AAI genomic database and backend.
- Developing new data analysis pipelines.



### **Current Ambulance Pilot Study**

- 1407 samples collected from ambulances across the US.
- 137 ambulances, in 20 cities, 6 states and 5 regions.

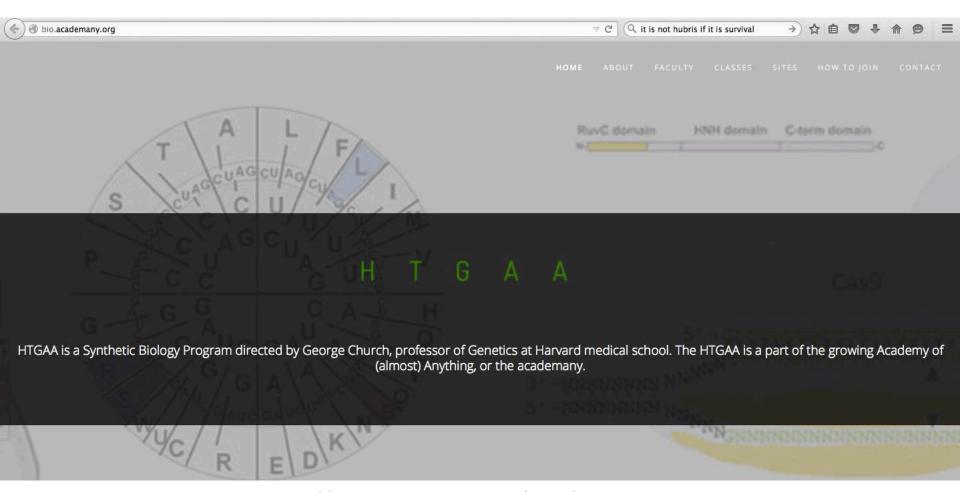


(8)

## Designer and Borrowed Metagenomics

## Understanding of biology is measured by the ability to engineer it and predict the outcome.

# How to Grow Almost Anything, HTGAA, including synthetic genomes and your modified microbiome

















http://science.nasa.gov/science-news/science-at-nasa/2007/11may\_locad3/



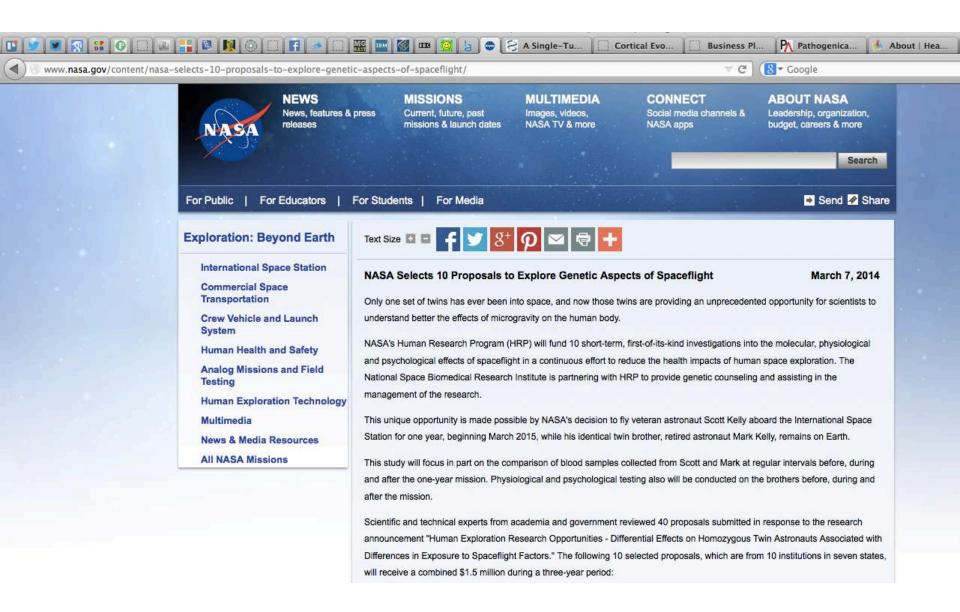


Participatory Medicine with Twin Astronauts





### Longitudinal, Integrative Systems Biology



"All the News That's Fit to Print"

## The New York Times

Late Edition

Today, mostly cloudy, snow showers, windy, colder, high 40. Tonight, evening flurries, clearing, cold, low 26. Tomorrow, mostly sunny, high 44. Weather map is on Page C8.

VOL. CLXIV . . . No. 56,819 +

© 2015 The New York Times

NEW YORK, SATURDAY, MARCH 28, 2015

2 50

#### PILOT CONCEALED MENTAL ILLNESS, AUTHORITIES SAY

#### SCRUTINY AFTER CRASH

In Screening, Onus Is on Employees to Be Forthcoming

This article is by Melissa Eddy, Dan Bilefsky and Nicola Clark.

DÜSSELDORF, Germany — Andreas Lubitz, the pilot at the controls of the Germanwings jetliner that crashed into the French Alps on Tuesday, had a mental illness but kept the diagnosis hidden from his employer, the authorities said Friday.

A psychiatric diagnosis might explain why Mr. Lubitz, a 27-year-old German, did not disclose his full medical record to Germanwings and its parent company, Lufthansa. Lufthansa company policy requires notification of conditions that could affect flying or a pilot's license.

Prosecutors said Friday that among the items found at Mr. Lubitz's home were several doctors' notes stating that he was too ill to work, including on the day of the crash; one of the notes had been torn up. These documents "support the preliminary assessment that the deceased hid his illness from his employer and colleagues," the prosecutors said in a statement.

But there remained considerable confusion about the precise nature and severity of his psychiatric condition. A German hospital said it had evaluated Mr. Lubitz twice in the past two months but added that he had not been there for assessment or treatment of depression.

The crash killed Mr. Lubitz and the other 149 people on board the Airbus A320 jettiner bound from Barcelona, Spain, to Düsseldorf, Germany, on Tuesday morning.

Germany, on Tuesday morning.

Airlines and their regulatory agencies have longstanding processes to screen for pilots whose



BILL INGALLS/ NAS.

#### Getting a Lift to 2016

A NASA astronaut, Scott J. Kelly, and two Russians left Saturday from Kazakhstan in a Soyuz TMA-16M spacecraft, headed to the International Space Station, Mr. Kelly will stay about a year.

#### Venture Capital Firm Prevails in Bias Case Riveting Silicon Valley

#### Sweeping Victory for Kleiner Perkins in Suit on Diversity in Tech Industry

#### By DAVID STREITFELD

SAN FRANCISCO — One of Silicon Valley's most famous venture capital firms prevailed on Friday over a former partner in a closely watched suit claiming gender discrimination, but hardly got away unscathed.

The plaintiff, Ellen Pao, had accused the firm, Kleiner Perkins Caufield & Byers, of discriminating against her in the course of her employment and eventual dismissal.

The decision handed Kleiner a sweeping victory in a case that had mesmerized Silicon Valley with its salacious details while simultaneously amplifying concerns about the lack of diversity in the technology industry.

Even with her loss in the case, Ms. Pao's suit succeeded in prompting debate about women in technology and venture capital, said Deborah Rhode, a law professor at Stanford University.

"This case sends a powerful signal to Silicon Valley in general and the venture capital industry in particular," Ms. Rhode said. "Defendants who win in court sometimes lose in the world outside it."

Kleiner and its lawyers did little to celebrate the win, with the lawyer Lynne C. Hernile saying that it "never occurred to me for a second that a careful and attentive jury like this would find either discrimination or retaliation." Kleiner issued a statement saying it was committed to supporting women.

Ms. Pao waved to the jury as she left the courtroom for the last time, a smile fixed on her face. "If I've helped to level the playing field for women and minorities in venture capital, then the battle was worth it." she said in a brief



HISTORICAL DIAMOSTTY DATE

Ellen Pao brought the suit against her former employer.

news conference.

Her suit, filed in Superior Court here, claimed that Kleiner did not promote her because of her gender, that it retaliated against her for complaining, that it failed to prevent gender discrimination and that it fired her in 2012 for complaining.

The suit asked \$16 million in compensatory damages plus punitive damages. Ms. Pao is now interim chief of the social media news site Reddit.

After the jurors rejected each of her four claims, they were found to be one vote short on a claim about her termination. For two hours, doubt reigned, the media unspooled possible outcomes and the jury went back to work. In the end, the problem seemed more juror confusion than anything else, and the claim went down with the others.

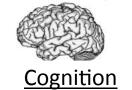
The jurors said in interviews they did not take on the role of "conscience of this community."

Continued on Page A3

#### In Defeat, Some Gains

Ellen Pao got people talking about the status of women, Farhad Manjoo writes. Page Bl.



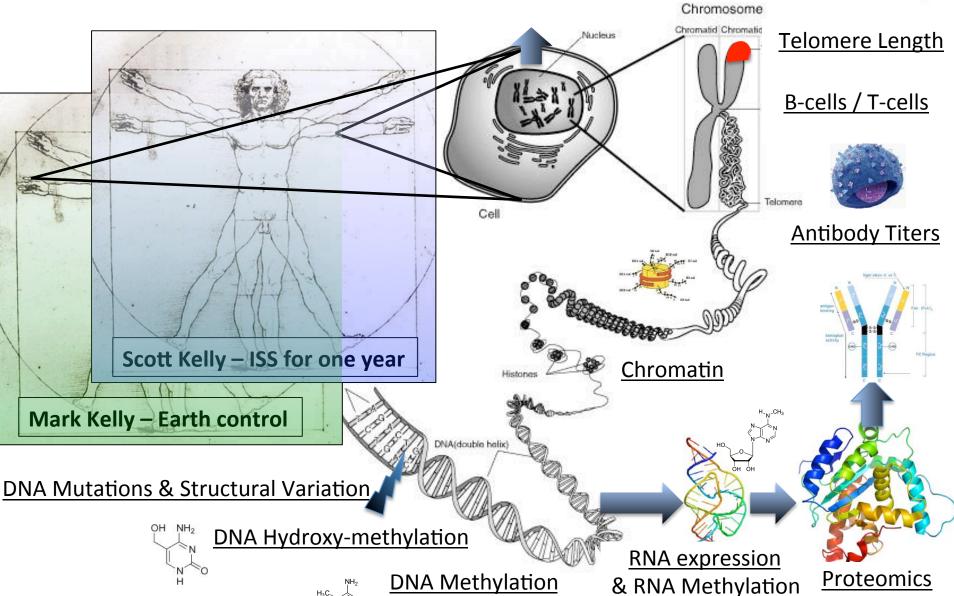




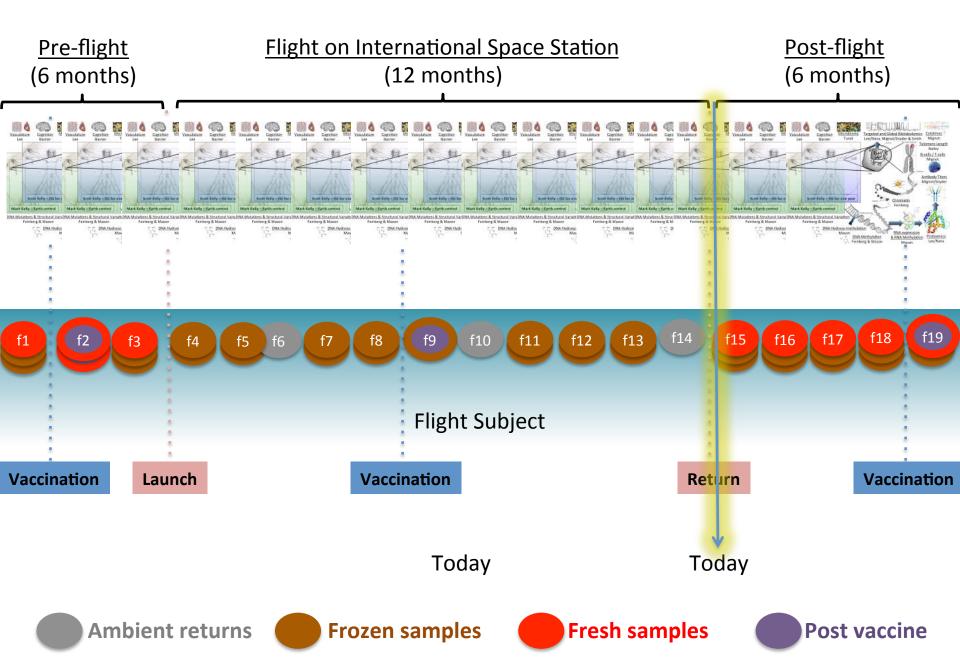
<u>Cytokines</u>

Targeted and Global Metabolomics

(large and small)



### Sample collections for the NASA Twins Study (2014-2016)



### **Protocol Planning**

NASA asks, "What's your terrestrial protocol?"

## Two Options:

- 1. Bring Samples Back
  - -Frozen
  - -Ambient

2. Sequence in Space



http://www.youtube.com/watch?v=0ODYdyLy9Tg





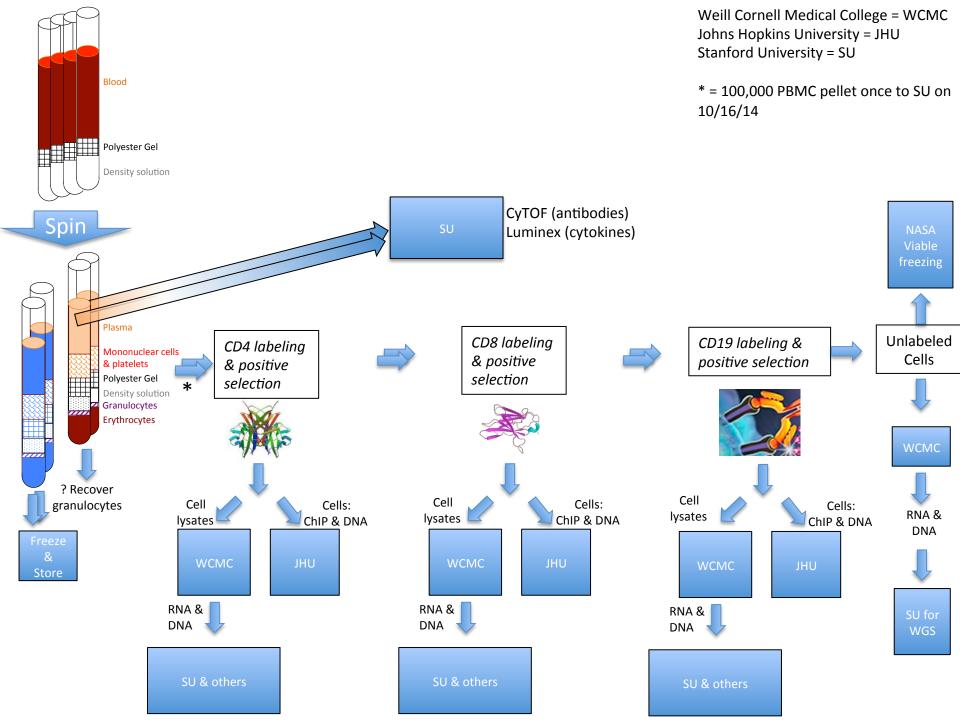




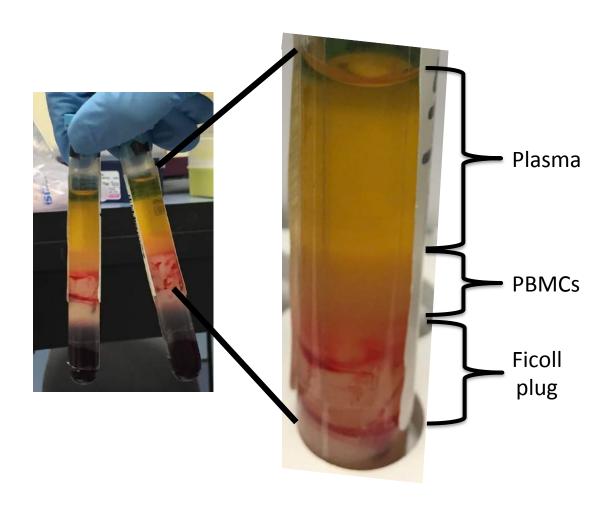
http://www.youtube.com/watch?v=VmhEtPN5u9Y



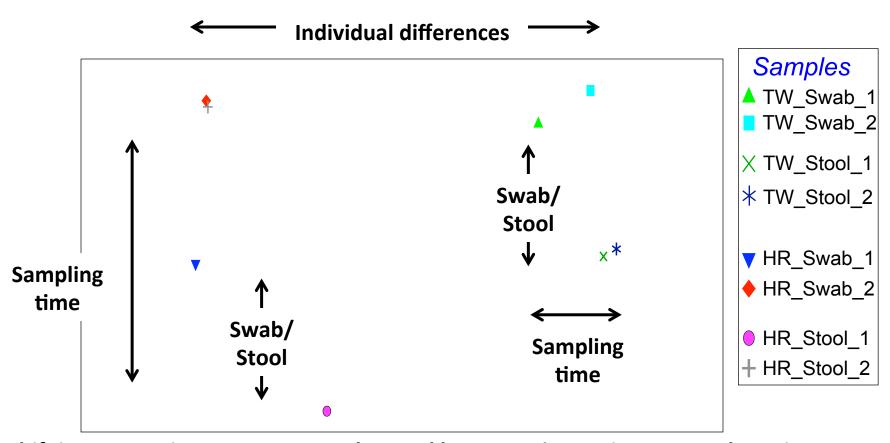




### Only 36 hours prior – in space!



## We can already see differences in the twins



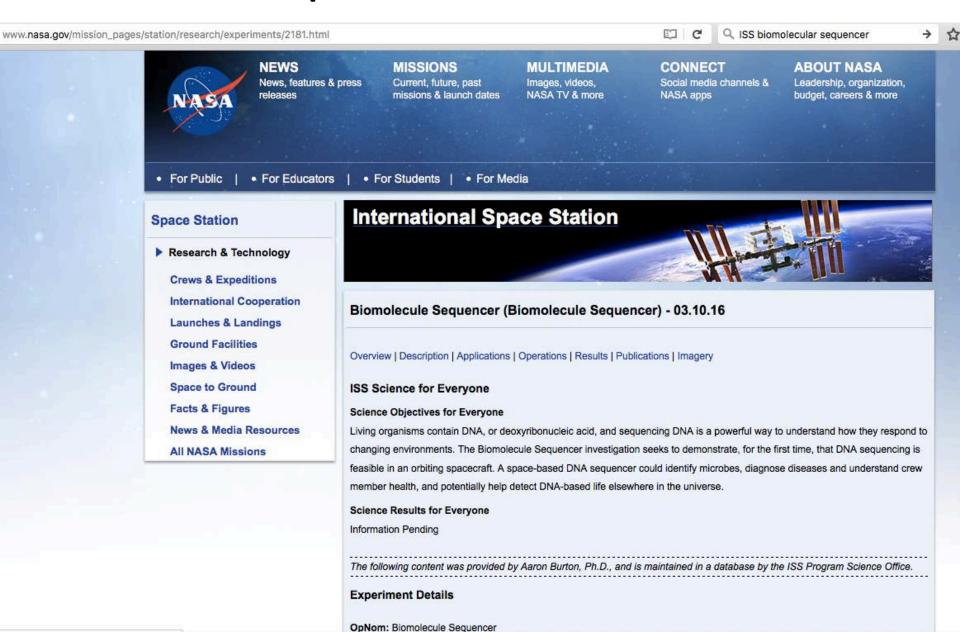
 A shift in community structure was observed between time points one and two in both individuals.

## Two Options:

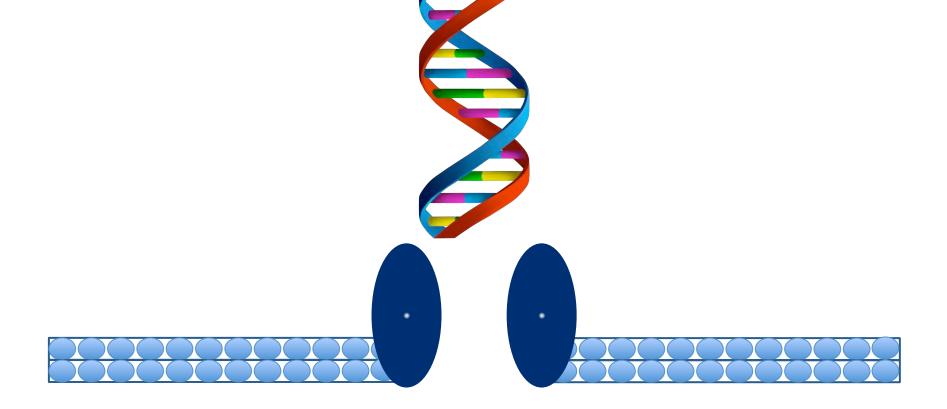
- 1. Bring Samples Back
  - -Frozen
  - -Ambient

Sequence in Space MinION sequencer (v5 chemistry)

### Launch planned for June 2016







## Zero-G Pipetting: Hardest Lab Job Ever







NATURE | NEWS







### Zero-gravity genomics passes first test

Two experiments demonstrate sample transfer and sequencing in a low-gravity environment.

#### **Chris Cesare**

13 October 2015



Rights & Permissions

After 160 swoops in NASA's zero-gravity aeroplane, researchers have the first evidence that genetic sequencing can be done in space.





### **EDUCATION**

**ABOUT US** 

MEDICAL EDUCATION

BASIC SCIENCE & CLINICAL RESEARCH

PATIENT CARE

DEPARTMENTS

ACADEMIC AFFAIRS

ADMISSIONS

CURRICULUM

FACILITIES

STUDENT AFFAIRS

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General Info

For Students

For Faculty

Curriculum Retreats

Med Ed Webinars

### Selected Areas of Concentration (AOC) Summaries and Contact Information

Listed below are some selected Areas of Concentration that have been established. Many others are in different stages of development by the faculty. Students are encouraged to consider developing their own Area of Concentration.

Updated January 26, 2016

Addiction and Trauma in Special Populations

Addiction Medicine

Aerospace Medicine, Space Genetics and Technology

Autoimmune and Inflammatory Disease

Cancer Biology and Therapy

Cardiometabolic Disease and Prevention

Community and Public Health

AOC Home

List of AOCs

Scholarly Project

Core Activities

Timelines

AOC Canvas Page

Learning Objectives

People and Contacts

FAOs

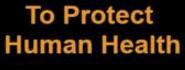


### Consortium for Space Genetics HARVARD MEDICAL SCHOOL

https://spacegenetics.hms.harvard.edu/

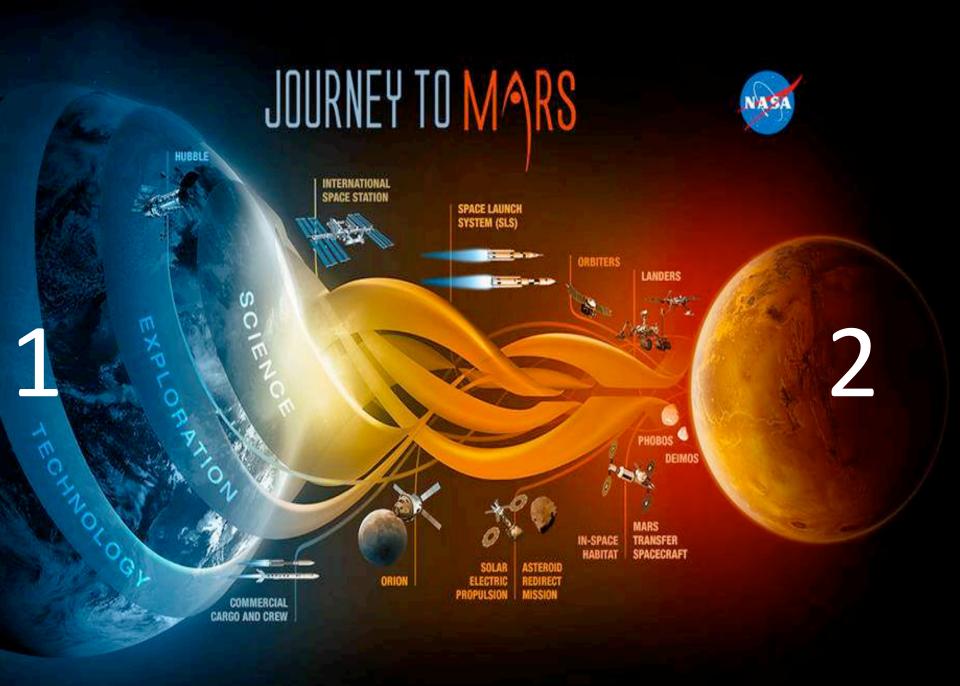
FACULTY CONTACT US





To Promote **Exploration** 





https://www.nasa.gov/content/nasas-journey-to-mars

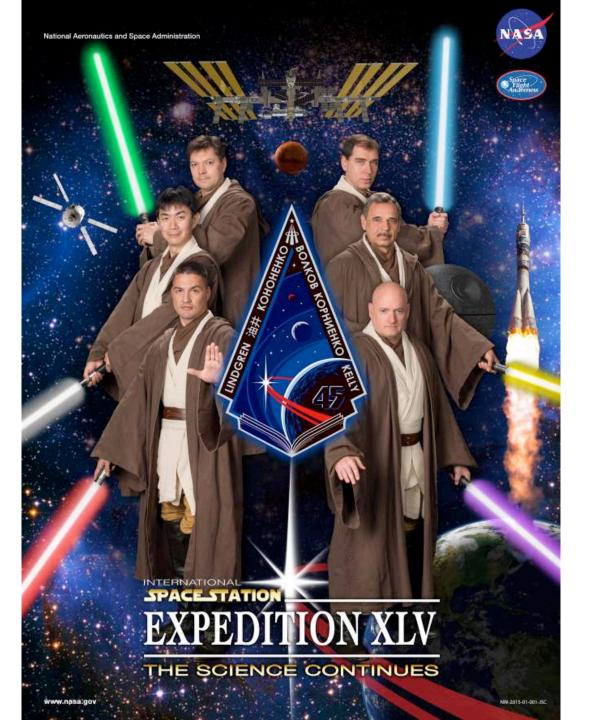
### **Conclusions**

- The subway is safe
- Forensics applies across all kingdoms of life
- Multi-disciplinary science is fantastic
- Space is Hard
- First demonstration of zero-gravity sequencing
  - Some optimization of the current and input is needed, but indeed we could potentially sequence on Mars or the ISS



### These People are Awesome





### Thanks to the Swabbing Teams! www.pathomap.org/people/



#### Weill Cornell Medical College















Queens College









New York University



York College





John Jay College





Fordham University





Brooklyn College





State University of New York (SUNY) Downstate

City University of New York (CUNY)

### Gratitude to Many People and Places



Mason Lab

Ebrahim Afshinnekoo

Sofia Ahsanuddin

Noah Alexander

Pradeep Ambrose

Marjan Bozinoski

Dhruva Chandramohan

Sagar Chhangawala

Shanin Chowdhury

Jorge Gandara

Francine Garrett-Bakelman

Elizabeth Hénaff

Sheng Li

Alexa McIntyre

Cem Meyden

Lenore Pipes

Darryl Reeves

Yogesh Saletore

Priyanka Vijay

HudsonAlpha Shawn Levy

NYU Jane Carlton

Julia Maritz



Icahn/MSSM

Eric Schadt, Andrew Kasarskis,

Joel Dudley, Ali Bashir,

Bobby Sebra

<u>UMMS</u>

Maria E Figueroa

Horner Lab

Yale University

Nenad Sestan

Sherman Weissman

Stacy Horner

<u>ABRF</u>

George Grills

Scott Tighe Don Baldwin

> Univ. Chicago Yoav Gilad

Illumina

Gary Schroth Marc Van Oene

Baylor Jeff Rogers

MSKCC
Christina Leslie
Ross Levine

AMNH
George Amato
Mark Sidall



Cornell/WCMC

Jason Banfelder

Scott Blanchard Selina Chen-Kiang

Olivier Elemento

Yariv Houvras

Samie Jaffrey

Ari Melnick

Margaret Ross

Adam Siepel

**Epigenomics Core** 

FDA/SEQC/Fudan Univ.

Leming Shi

NIH/UDP/NCBI

Jean & Danielle Thierry-Mieg



Nati Neu

FOUNDATION

National Institute of Neurological Disorders and Stroke

STARR CANCER CONSORTIUM

