



Monitoring astronaut health in space

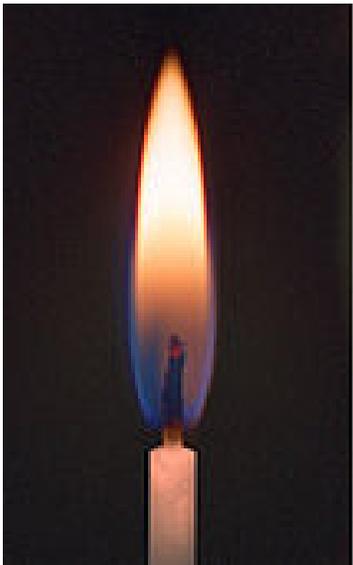
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Pharmacology Discipline Lead, NASA Johnson Space Center

5 May 2014

Sensing Technologies for Global Health, Military Medicine and
Environmental Monitoring
Baltimore, MD

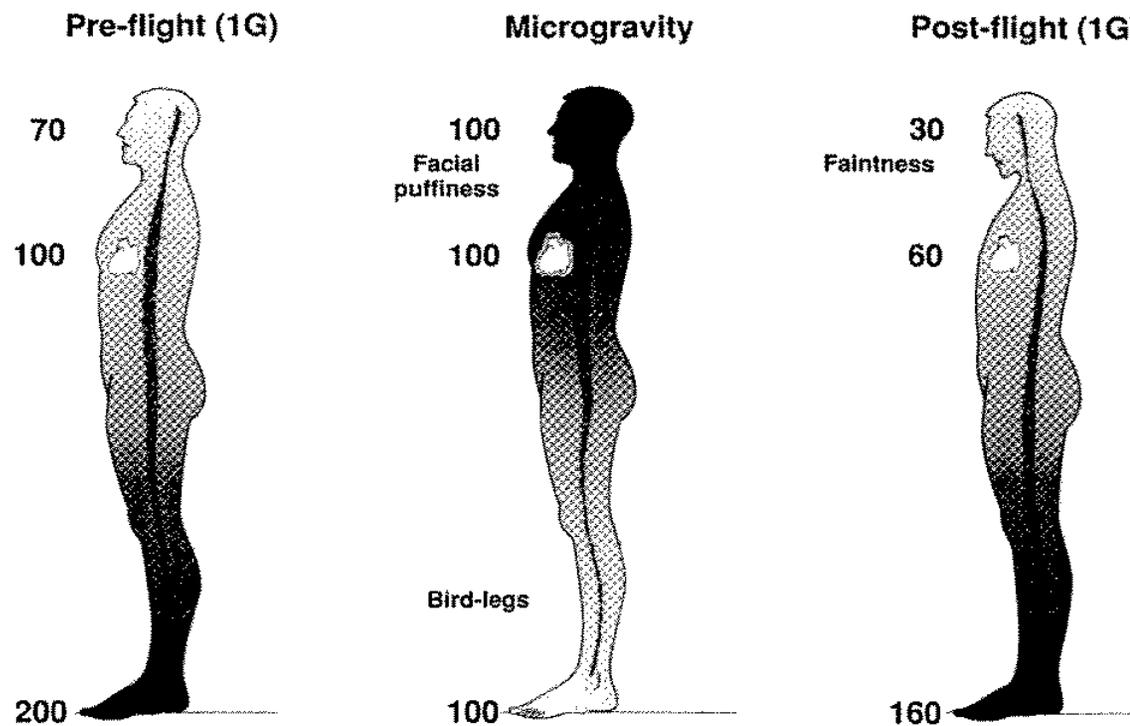
Things are different in microgravity





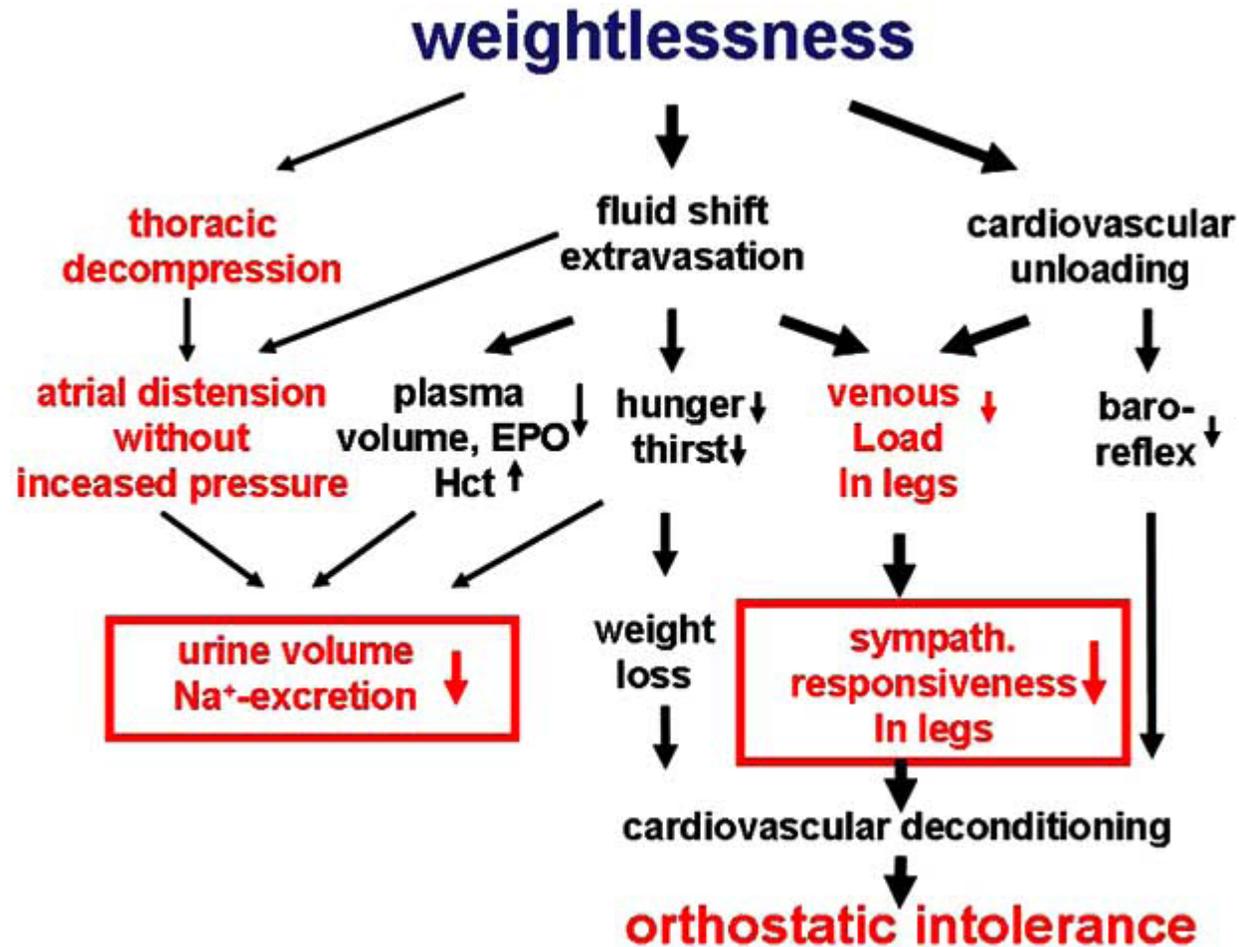
Even people are different in microgravity

Decreased gravity makes body fluids shift upward



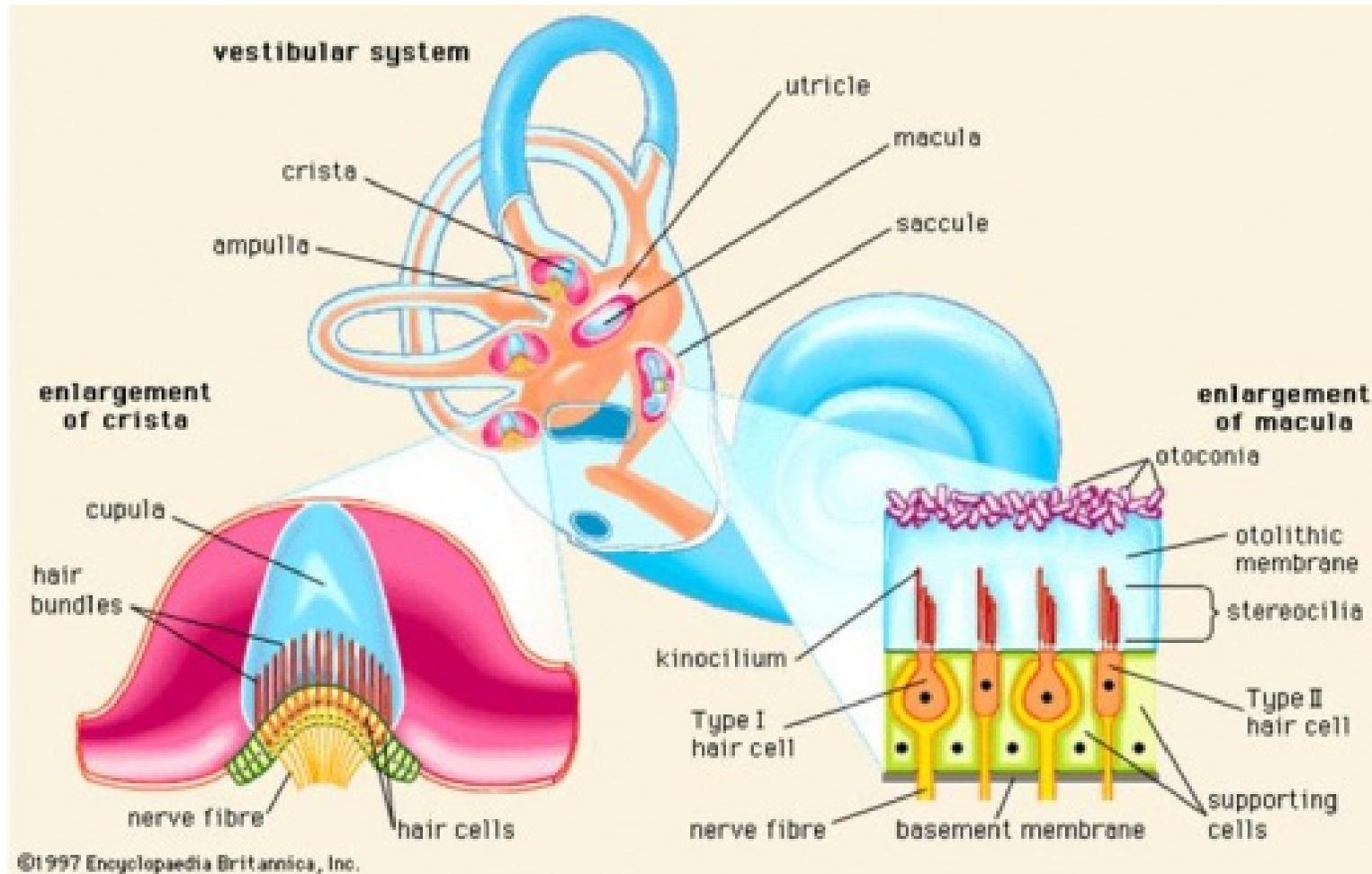
Cardiovascular adaptations, fluid shifts, and countermeasures related to space flight*

Alan R. Hargens^{a,*}, Sara Richardson^b *Respiratory Physiology & Neurobiology* 169S (2009) S30–S33



Regulation of Body Fluid and Salt Homeostasis – from Observations in Space to New Concepts on Earth. R. Gerzer and M. Heer *Current Pharmaceutical Biotechnology*, 2005, 6, 299-304 299

Decreased gravity disrupts the sense of balance



[http://www.skybrary.aero/index.php/Vestibular_System_and_Illusions_\(OGHFA_BN\)](http://www.skybrary.aero/index.php/Vestibular_System_and_Illusions_(OGHFA_BN))

Medical Complaints Associated with Spaceflight



Weight loss

Bone loss

Muscle atrophy

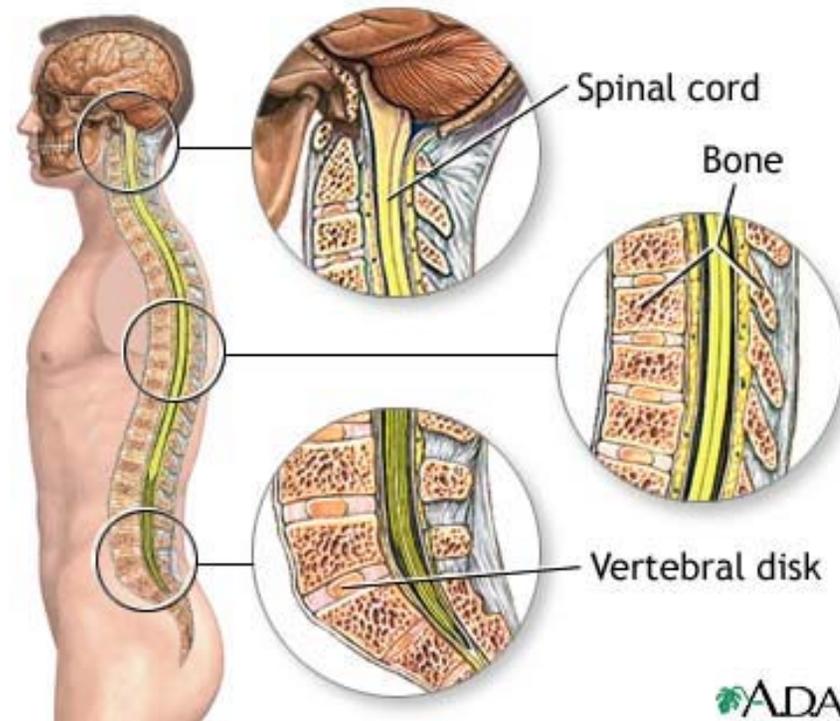
Loss of Bone Mineral Density



Russian ground personal members and doctors carry Italian ESA astronaut Roberto Vittori to the medical tent upon his arrival to the town of Arkalyk, northern Kazakhstan, early Monday, April 25, 2005. [AP]

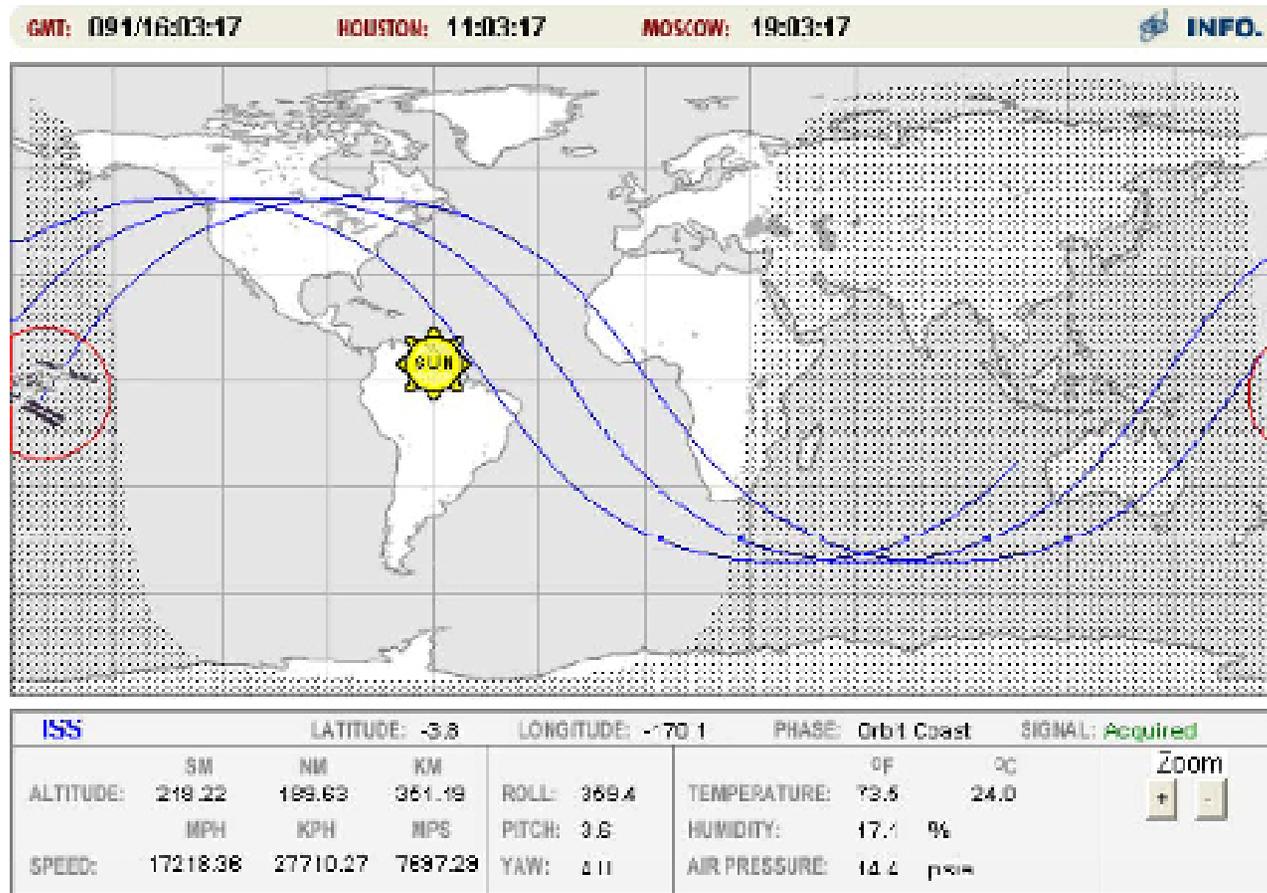


Body Pain





Circadian Rhythm Disruption



Medical Complaints in Space



Based on ISS Missions:

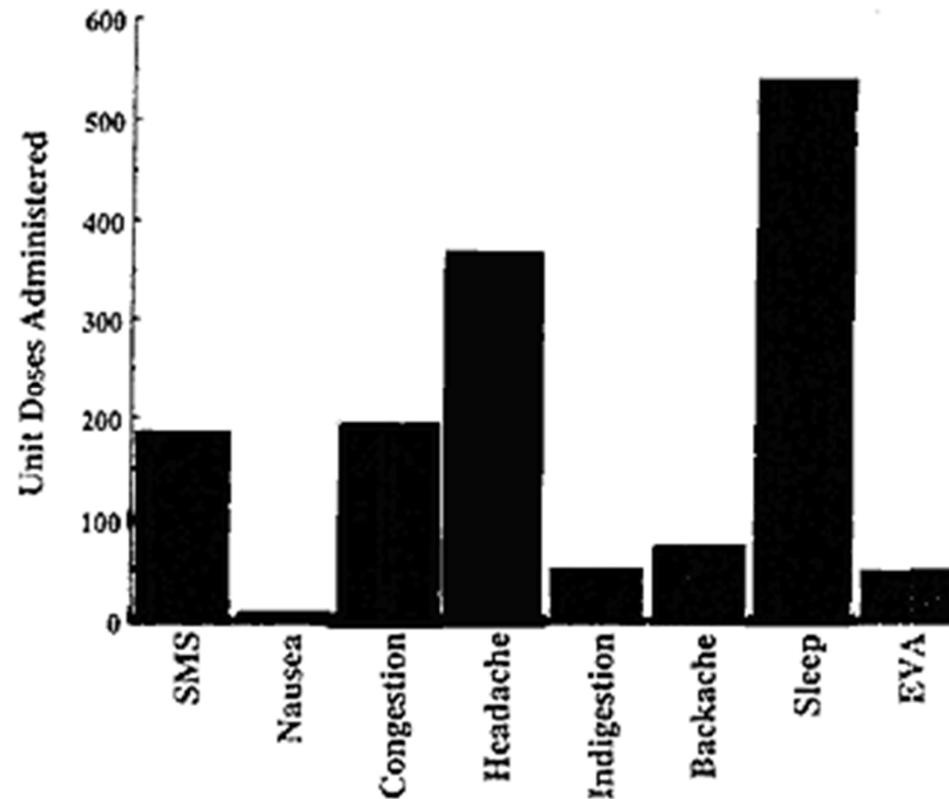
Anorexia
Space motion sickness
Fatigue
Insomnia
Dehydration
Dermatitis
Back pain
Upper respiratory infection
Conjunctival irritation
Subungual hemorrhage
Urinary tract infection
Cardiac arrhythmia
Headache
Muscle strain
Diarrhea
Constipation

From Clement , Fundamentals of Space Medicine, 2003

Based on Space Shuttle, 1988- 1995

Facial Fullness
Headache
Sinus congestion
Dry skin, irritation, rash
Eye irritation, dryness, redness
Foreign body in eye
Sneezing/coughing
Sensory changes
Upper respiratory infection
Back muscle pain
Leg/foot muscle pain
Cuts
Shoulder/trunk muscle pain
Hand/arm muscle pain
Anxiety/annoyance
Contusions
Ear problems (usu. Pain)
Neck muscle pain
Stress/tension
Muscle cramp
Abrasions
Fever, chills
Nosebleed
Psoriasis, folliculitis, seborrhea
Low heart rate
Myoclonic jerks

Pharmaceutical Use on Shuttle



PUTCHA L, BERENS KL, MARSHBURN TH, ORTEGA HJ, BILICA RD.
Pharmaceutical use by U.S. astronauts on space shuttle missions.
Aviat Space Environ Med 1999; 70:705-8.

Our Mission at the JSC Pharmacology Lab...



...is to ensure that flight surgeons have good information about how administered pharmaceuticals will work in the extreme conditions of spaceflight

...which means that we have to understand the physiological changes caused by living in the spaceflight environment

...as well as the effect of the spaceflight environment on the stored drugs themselves

...as well as the pharmaceuticals' mechanism of action

How and where we do our Research



In order to conduct the kind of studies that can help improve the safety and productivity of space travel, the HRP has established a number of experimental facilities known as “analog” that mimic environmental conditions in space.

- NASA Extreme Environment Mission Operations (NEEMO)
- Haughton Mars Project
- Desert Research and Technology Studies (Desert RATS)
- Bedrest Studies
- Human-Related Altitude Chamber Complex
- Antarctica



Limitations of Spaceflight Experiments:

- Non-invasive methods best
- Non-toxic
- Lightweight and small equipment
- No degassing, explosion or fire risk
- Low power consumption
- Low impact on crew schedule
- N will be small (~400 people have flown to space)

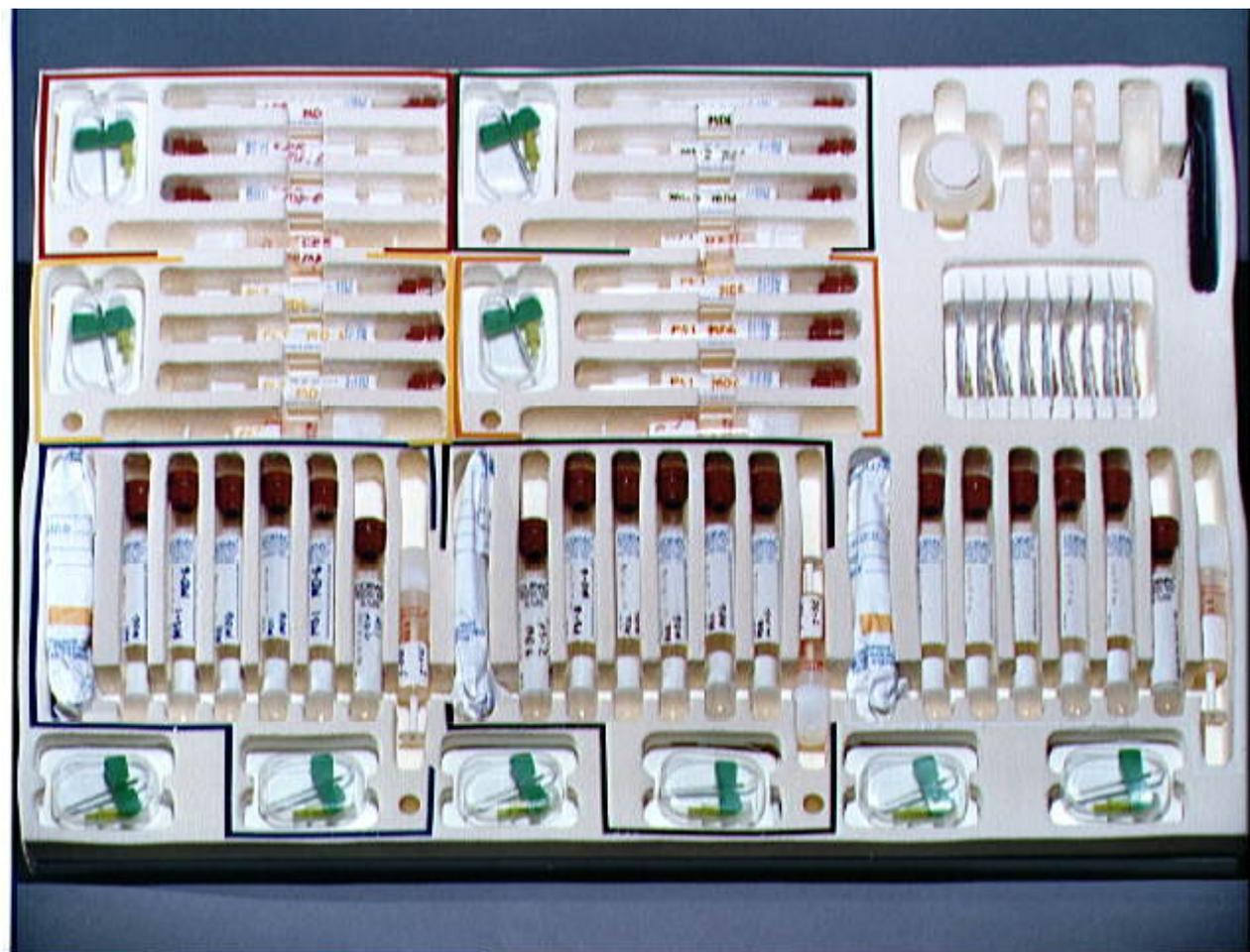
Current model: return samples to Earth for analysis



Inside the science module aboard the Earth-orbiting Space Shuttle Columbia, Astronaut David A. Wolf draws blood from payload specialist Martin J. Fettman, DVM. Blood samples from crew members are critical to Life Sciences investigations, but must be returned to Earth for analyses.



Blood collection kit for Space Lab 1 1981



-80 Freezer is available



ISS013E64639

Some tests can be performed inflight, with data transmitted to ground



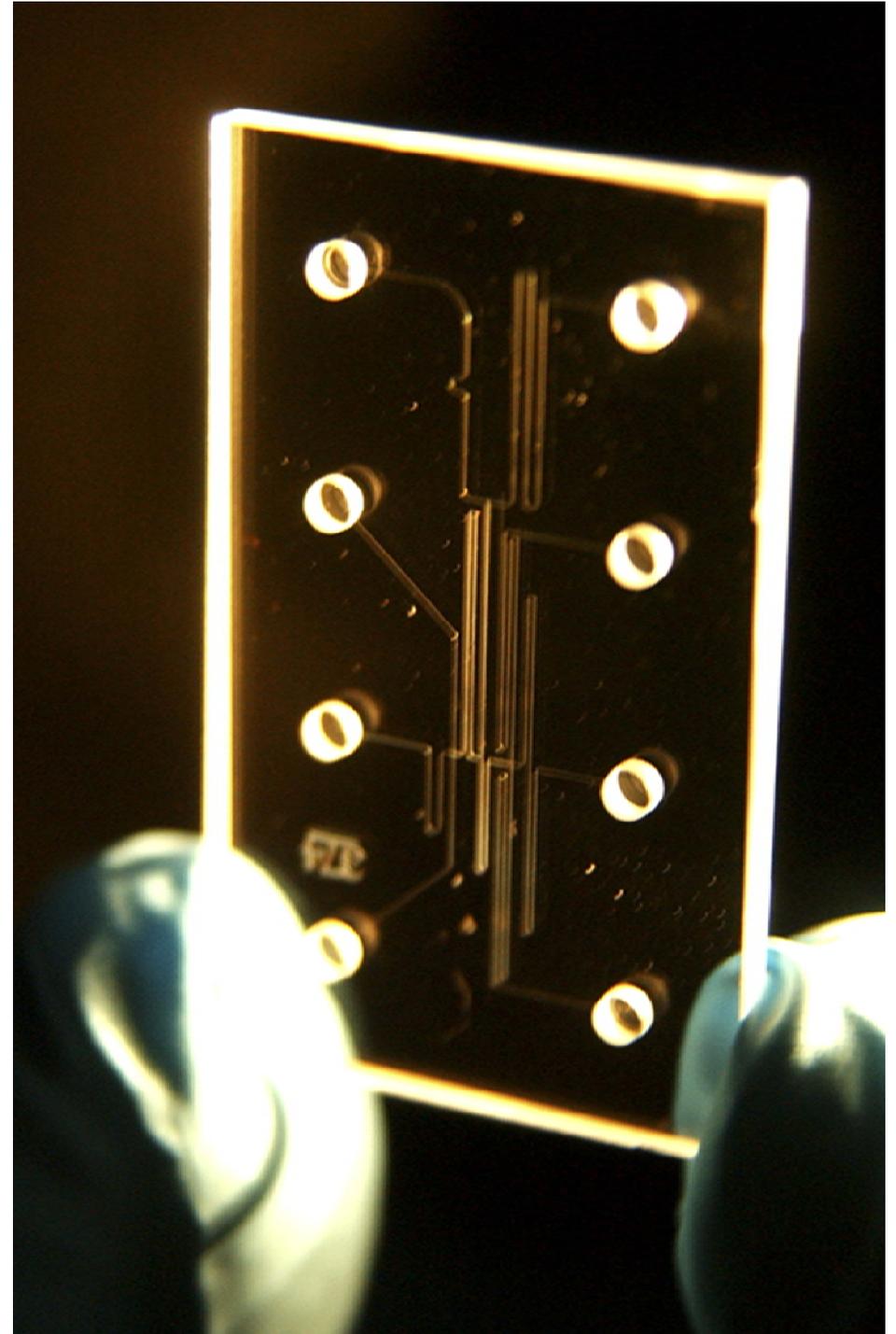
ISS commander and science officer Leroy Chiao performs an ultrasound scan on the eye of flight engineer Salizhan Sharipov during ISS Expedition 10. Crewmembers may be guided real-time by flight surgeons and/or an experienced ultrasound technician.



Lab on a chip may enable inflight analysis

The eight holes on this chip are ports that can be filled with fluids or chemicals. Tiny valves control the chemical processes by mixing fluids that move in the tiny channels that look like lines, connecting the ports.

Scientists at NASA's Marshall Space Flight Center in Huntsville, Alabama designed this chip to grow biological crystals on the ISS. These chips, the size of dimes, could be loaded on a rover looking for biosignatures of past or present life. Other types of chips could be placed in handheld devices used to monitor microbes in water or to quickly conduct medical tests on astronauts. (NASA/MSFC/D.Stoffer)



Human Research Roadmap



A web tool that is an excellent source of information on each of the identified HRP Risks to human spaceflight.

<http://humanresearchroadmap.nasa.gov/>

<http://humanresearchroadmap.nasa.gov/>



Risks ⓘ

Risks 1 - 32 (of 32)

Risk Factor of Inadequate Nutrition

Risk of Acute and Late Central Nervous System Effects from Radiation Exposure

Risk of Acute Radiation Syndromes Due to Solar Particle Events (SPEs)

Risk of Adverse Behavioral Conditions and Psychiatric Disorders

Risk of Adverse Health Effects Due to Alterations in Host-Microorganism Interactions

Risk of Adverse Health Effects of Exposure to Dust and Volatiles During Exploration of Celestial Bodies

Risk of an Incompatible Vehicle/Habitat Design

Risk of Bone Fracture

Risk of Cardiac Rhythm Problems

Risk of Clinically Relevant Unpredicted Effects of Medication

Risk of Compromised EVA Performance and Crew Health Due to Inadequate EVA Suit Systems

Risk of Crew Adverse Health Event Due to Altered Immune Response

Risk of Decompression Sickness

Risk Of Degenerative Tissue Or Other Health Effects From Radiation Exposure

Risk Of Early Onset Osteoporosis Due To Spaceflight

Risk of Impaired Control of Spacecraft, Associated Systems and Immediate Vehicle Egress Due to Vestibular/Sensorimotor Alterations Associated with Space Flight

Risk of Impaired Performance Due to Reduced Muscle Mass, Strength and Endurance

Risk of Inadequate Critical Task Design

Risk of Inadequate Design of Human and Automation/Robotic Integration

Risk of Inadequate Human-Computer Interaction

Risk of Injury from Dynamic Loads

Risk of Intervertebral Disk Damage

Risk of Orthostatic Intolerance During Re-Exposure to Gravity

Risk of Performance Decrement and Crew Illness Due to an Inadequate Food System

Risk of Performance Decrements Due to Inadequate Cooperation, Coordination, Communication, and Psychosocial Adaptation within a Team

First pharmaceuticals in US spaceflight

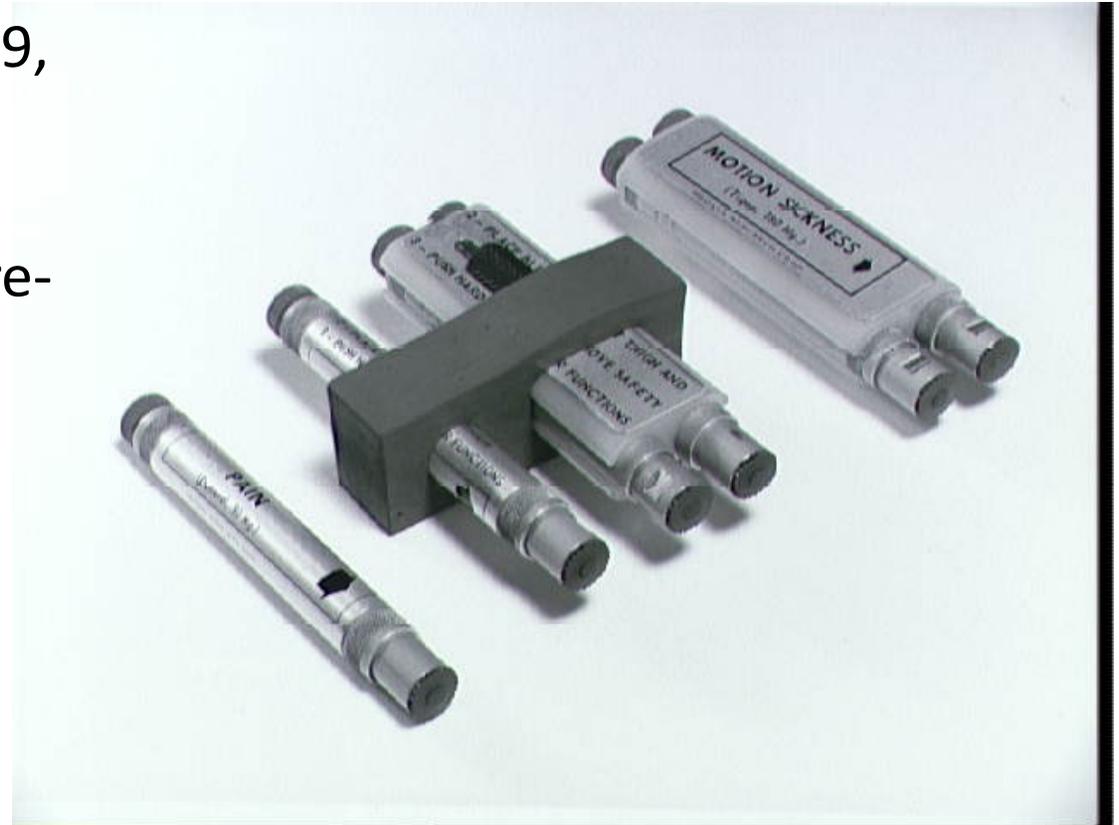


In 1963 on Mercury Atlas 9,
22 Earth orbits, 35 hours

Gordon Cooper carried pre-
loaded drug injectors in
space suit pocket

Demerol – pain relief

Tigan - motion sickness



Before 1988, there were no pharmacological countermeasures-
just exercise, fluid loading and g suits.

Shuttle missions lasted no more than 7 days.

In 1988 Congress approved funds to expand missions to 16 days.

Countermeasure development began in earnest.

Currently, 6 months on the ISS is routine and the first 1 year
mission will begin later this year.

We are planning exploration missions that may last 3 years and
the countermeasures (many of them pharmaceutical) that will be
required to maintain crew health over periods of years.

Research in JSC Pharmacology

Pharmaceuticals

Usage tracking

Stability

Pharmacokinetics

Absorption/Distribution

Metabolism/Excretion

Pharmacodynamics

all the indications that people treat



Evidence: Flight studies

Cintron & Putcha acetaminophen case studies, 1987

Salivary acetaminophen measured over time after oral administration of tablets (acetaminophen is the gold standard for examination of oral absorption, but substitution of saliva concentrations for plasma not established)

3 individuals - each panel shows data from a single individual. Use of other medications is unspecified.

Two individuals show reduced peak concentration for FD1-2 (middle and lower)

One person shows slower absorption preflight, but this value is much slower than average ground values from the literature (0.8 h).

Paucity of data and variability in data preclude definitive conclusion.

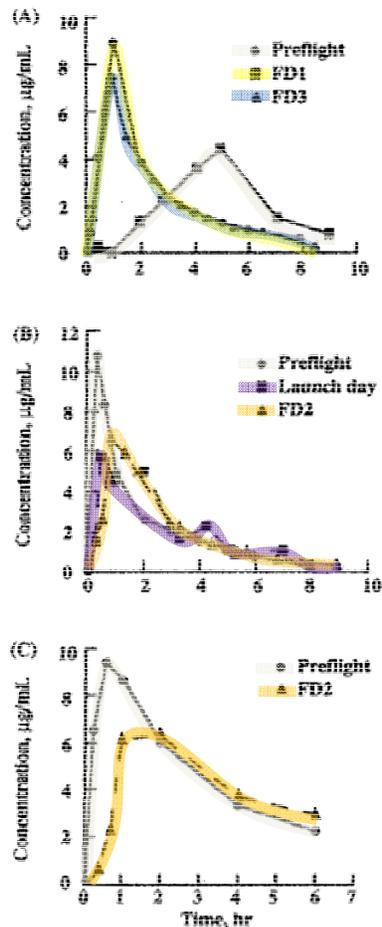
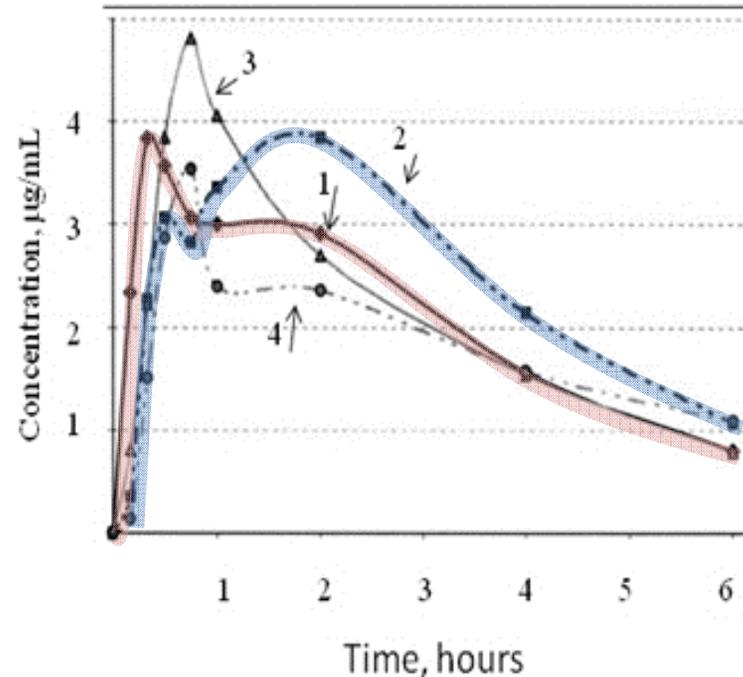


Fig. 2 Saliva concentrations of acetaminophen over time after oral administration of a 650 mg dose to crew members.



Evidence: Flight studies

Average pharmacokinetic curves after acetaminophen tablet administration.



1. ~~tablets, on ground;~~
2. ~~tablets during spaceflight;~~
3. capsules, ground;
4. capsules during spaceflight.

AUC is the same, peak looks slower (but not significantly so)

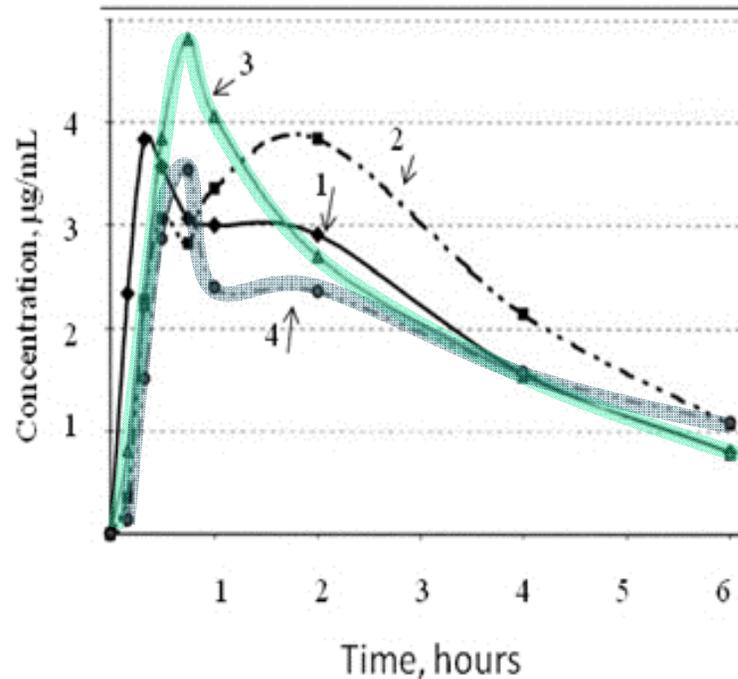
Long duration ISS crew (sampling date unspecified), males, N= 5, free of other medications, salivary sampling. The absence of error bars on this graph exaggerates differences between curves.

Kovachevich *et al.*,2009



Evidence: Flight studies

Average pharmacokinetic curves after acetaminophen capsule administration.



1. tablets, on ground;
2. tablets during spaceflight;
3. capsules, ground;
4. capsules during spaceflight.

Time to peak is the same, but the half-life is significantly longer.

Long duration ISS crew (sampling date unspecified), males, N= 5 (different 5 than tablet group), free of other medications, salivary sampling. The absence of error bars on this graph makes comparisons difficult.

Kovachevich *et al.*,2009

PK- Does the spaceflight environment (radiation, microgravity, etc) alter ADME?

Use of salivary sampling for PK studies would improve feasibility inflight

Inflight analysis would remove the requirements for sample storage (in costly frozen conditions) and sample return



Salivary sampling use in spaceflight

- Collection of saliva is one such technique that has been frequently used for pharmacologic, metabolic, circadian and immune studies.
- About 40 studies, on Apollo, Mir, Apollo-Soyuz Test Project, Shuttle and ISS

Hardware – Saliva Collection Kit

NEW SUPPLIES: USED SUPPLIES:

CREWMEMBER 1:

CREWMEMBER 2:

CREWMEMBER 3:



Dry saliva collection book (2 per crewmember, 6 in kit)

Clipped liquid saliva preservative bags (7 count per crewmember, 21 total in kit)

Collection swabs for liquid saliva (7 count in one bag per crewmember, 3 bags/21 swabs in kit)





Need more ?

General NASA info:

<http://www.nasa.gov>

Human Research at NASA:

<http://humanresearch.jsc.nasa.gov/>

NASA Research Grants:

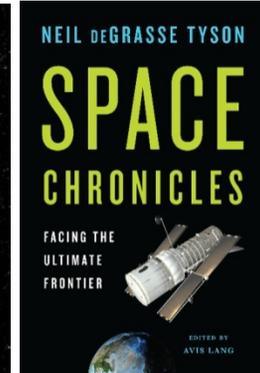
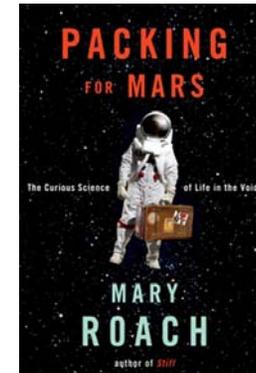
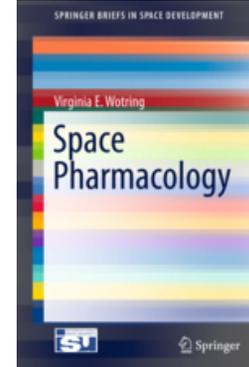
<http://nspires.nasaprs.com/external/>

Don Pettit's Blog

<http://blogs.airspacemag.com/pettit/>

Ginger

virginia.e.wotring@nasa.gov





The NASA budget is about \$17B.

Half a penny of every tax dollar goes to NASA.

NIH budget is about \$30B.

Americans spend more on pizza (\$27B) than on the space program.

