Arnauld E. Nicogossian · Richard S. Williams Carolyn L. Huntoon · Charles R. Doarn James D. Polk · Victor S. Schneider *Editors*

Space Physiology and Medicine

From Evidence to Practice Fourth Edition

EXTRAS ONLINE

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Arnauld E. Nicogossian • Richard S. Williams Carolyn L. Huntoon • Charles R. Doarn James D. Polk • Victor S. Schneider Editors

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From Evidence to Practice

Fourth Edition

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We dedicate this book to our families whose encouragements guided us through the writing process. We are grateful to the astronauts and the men and women of NASA whose dedicated work and selfless contributions generated the knowledge used in this book. John Glenn's [1921–2016] encouragements inspired the editors during the preparation of this textbook.

Foreword

This Fourth Edition provides a unique perspective into the principles of the expanding space medicine practice, drawing on the underpinning knowledge gathered over the last six decades of research and clinical practice in support of human space exploration. This textbook serves as a ready reference for space medicine specialists, engineers, and educators alike. Built upon the success of the first three editions, this fourth edition brings to the reader the evidence necessary to plan for human health of future space explorers. Since my last space flight on the Space Shuttle on October 29, 1998, on Discovery mission STS-95: the U.S. Space Shuttles docked with the *Mir* Space Station; the ISS was built using the Space Shuttle, which was retired in 2011; commercial companies began to launch spacecraft for logistic and resupply to the ISS; and China joined the U.S. and Russia as the third nation with piloted space mission capabilities. And last but not least, the exciting discovery of exoplanets and the abundance of water and life building blocks in the universe.

My reason for receiving the assignment to go up again on the Space Shuttle was to investigate the striking similarities between what happens to all astronauts in all extended space flight and how it compares to some of the natural aging processes right here on Earth. If breakthroughs can be made in our knowledge barrier, perhaps we can provide for longer term space flight and cut out or reduce many of the frailties of old age right here on Earth.

The materials presented in this textbook not only unravel in an elegant and understandable manner our current clinical knowledge but also point to the major knowledge gaps to be addressed to ensure future crews' health and safety while launching bold missions into the solar system. The wealth of knowledge gathered from space research and practice has also benefitted the life on Earth. I was instrumental in bringing together the best minds of NIH and NASA to work on the problems of aging. Healthy and productive aging reduces healthcare costs and minimizes disparities in future generations. My two space flights demonstrated that people do survive and work productively in space, but also more importantly age is not a limiting factor to space travel. My experiences do pave the way to future space tourism as the information provided in this textbook will undoubtedly contribute to safe space travel.

I am pleased to introduce this book and commend NASA and its medical community for their contributions to the future of human space travel.

Columbus, OH, USA John Glenn

Preface

Built upon the success of the first three editions, this fourth edition brings to the reader currency of space physiology and medicine. It has been more than 20 years since the publication of the third edition. In that time, U.S. Space Shuttles docked with the *Mir* Space Station, the International Space Station (ISS) was built using the Space Shuttles and Russian launch capabilities, the *Mir* Space Station was deorbited into the Pacific Ocean, the Space Shuttle Program was retired, commercial companies began to launch spacecraft for logistic and resupply to the ISS, and countries such as China and India have initiated programs that launched space assets into low Earth orbit. In the case of China, there have been a number of Chinese crew members who have been launched into space and safely returned to the Earth. The knowledge gained from the third edition (1994) to the fourth edition (2016) has been significant.

The growth in knowledge is attributable to international life sciences' research, significantly more complex space-based systems, technology in medical monitoring as well as computers and telecommunications, and the number of "man" hours in space. The construction of the ISS required an increase in the duration of an individual's time in space, a significant increase in the number of extravehicular activities, as well as advancements in environmental control and life support systems. With each successive program from the Mercury Program in the U.S. and the Voshkod in the Soviet Union to the ISS, our ability to understand the nuances and capabilities of providing healthcare in support of astronauts and cosmonauts during all phases of flight has progressively evolved. The current ISS Program, and the opportunity it provides for research and significantly longer stays in space, concomitant with commercial capabilities, has set the stage for exploration of other celestial bodies such as Mars and our own moon.

This fourth edition provides a foundation for those interested in space physiology and medicine practice and research. It is intended to be a teaching textbook with accompanying teaching materials to help the educator and student alike. Through 19 chapters, it provides a comprehensive review of space medicine, spacecraft environments, adaptation and rehabilitation in response to space flight, occupational health and safety issues, and ground-based testbeds and training of physicians and other personnel to support space medicine. The textbook is unique in distilling currently published clinical evidence for the benefit of the busy practitioner and researcher.

As editors, we are pleased to provide clinicians the needed practice tools at the time of expanding commercial interests in space.

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Introduction

Parts

- 1. Introduction to Space Medicine
- 2. Spacecraft Environments
- 3. Space Flight and Crew Health: Adaptation, Pathophysiology, Rehabilitation, and Countermeasures
- 4. Occupational Health and Safety Issues in Space Flight
- 5. Ground-Based and Academic Training Programs
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Optional Teaching Tools (Web-Based)

- 1. Syllabus
- 2. Teaching aids

On April 13, 1960, a meeting was held at the Aviation Medical Acceleration Laboratory, WDAC, Johnsville, Pennsylvania, to examine and recommend biomedical information to be collected from astronauts and the Mercury spacecraft.¹ Fifty-six years later the editors and chapter contributors, in cooperation with Springer, are delighted to introduce the fourth edition of *Space Medicine and Physiology*. This revised textbook is intended for teachers, students, and practitioners interested and engaged in this rapidly evolving discipline. The knowledge

¹Space medicine in Project Mercury Chap. [7.](http://dx.doi.org/10.1007/978-1-4939-6652-3_7) history.nasa.gov/SP-4003/ch7-3.htm*.*

gained over the past five decades is reflected in this textbook. A common outline provides the reader with a ready cross-referencing between different chapters. The authors have provided their expertise from a variety of disciplines. Each has contributed in significant ways. All of the authors have been part of the space program in the U.S. over the course of their careers, and their experiences helped shape the narrative of NASA's space medicine activities.

This fourth edition builds on the foundation of the previous three editions, adding new information on relevant legislation, medical policy, and ethics. A syllabus and a set of teaching materials are made available for academic purposes.

The standard chapter outline (see Box) contains Case Studies, which add an important element for the reader to connect the evidence to clinical practice. The information contained in the chapters reflects the evidence in a point of time, which is subject to potential change based on new information obtained from research and observations conducted on the International Space Station (ISS). The Self-Study Questions and Key Points summarize the knowledge underpinning space medicine standards of practice and the remaining uncertainties to be addressed. Supplemental information from biomedical research based on human surrogate subjects and biological specimens are also included as appropriate.

Many of the research findings and publications from the Skylab and Apollo programs remain current and have been retained in this textbook.

Highly cited NASA experts at the forefront of space physiology and medicine contributed to this textbook. This is especially true with Part II, which includes Chaps. [8–](http://dx.doi.org/10.1007/978-1-4939-6652-3_8)[14](http://dx.doi.org/10.1007/978-1-4939-6652-3_14) on a variety of human systems and the impact space flight has on them. Each of the 19 chapters is either new or rewritten to reflect knowledge gained from the 30 years of research conducted in the Space Shuttle Program, the Shuttle/*Mir* Program, and the ISS Program. In addition, groundbased research, a critical component of space flight activities, has also been updated.

This textbook describes space medicine from the U.S. perspective and relies on the peerreviewed literature and government sources divided into six parts and 19 chapters. A short overview of each part is described as follows:

Part 1, entitled *"Introduction to Space Medicine*,*"* presents a discussion on the evolution of human capabilities and space medicine, the environment of space exploration, and the clinical implications of the adaptation to space flight.

Chapter [1](http://dx.doi.org/10.1007/978-1-4939-6652-3_1) provides a historical context for the human spacecraft design evolution, robotic tools, and biomedical research intended to protect the crew health and safety. It also dwells on the nascent commercial human space flight initiatives, the evolution of medically relevant legislation, policies, and standards, and ethical dilemma in the conduct of space exploration.

Chapter [2](http://dx.doi.org/10.1007/978-1-4939-6652-3_2) describes the space exploration environment of Low Earth Orbit (LEO) and beyond. Most recent information on the Solar System and astrobiology, gathered from ground and space telescopes and robotic planetary rovers and probes, is presented. Robotic explorers are pathfinders for future bold human initiatives and provide invaluable information on potential resources, and possible health risks from physical and perhaps biological threats. Life's building blocks, the availability of water and other energy sources, are summarily addressed for individual planets and their satellites, including asteroids and comets. Planetary protection and associated policies are also discussed.

Chapter [3](http://dx.doi.org/10.1007/978-1-4939-6652-3_3) is an overview of the bioastronautics of space flight and clinical implications to the human living and working in space. This chapter addresses astronaut demographics, epidemiological findings, and crew health maintenance in space and post-flight. The chapter also informs on microgravity as an analog to aging on Earth. Human factors influencing crew performance and the status of current countermeasures are concisely summarized. This chapter sets the stage for Parts II and III.

Part II, entitled *"Spacecraft Environments,"* provides an in-depth review of spacecraft internal environments that impact crew health, including toxicology, microbiology, immunology, acoustics, audition, and radiation. These four chapters provide a thorough review of how spacecraft systems may impact the human in the system. While research continues to be done

in these disciplines, the literature, while current, requires additional information to enhance clinical practice, through the development of spacecraft standards and systems.

Chapter [4](http://dx.doi.org/10.1007/978-1-4939-6652-3_4) provides an overview of spacecraft atmosphere contaminants, particulates, and other chemicals. In addition, monitoring and management of the toxicological threats to crew health and performance are addressed, including the handling of emergency events such as a fire or leaking coolant. This chapter also discusses accepted environmental hazards exposure limits for astronauts living in closed environments in missions in excess of 1 year.

Chapter [5](http://dx.doi.org/10.1007/978-1-4939-6652-3_5) is focused on microbiology and infectious disease that may result from crew exposures to microorganisms present in the spacecraft environment, including the air, water, and surfaces. Environmental systems of the Space Shuttle, Shuttle/*Mir*, ISS, and management of health risks are discussed.

Chapter [6](http://dx.doi.org/10.1007/978-1-4939-6652-3_6) describes health problems related to noise onboard the spacecraft. Attributes, characteristics, and mechanisms of acoustics and auditory response are described. Hearing conservation and principles for monitoring hearing thresholds and mitigating impact to crew health and performance are discussed.

Chapter [7](http://dx.doi.org/10.1007/978-1-4939-6652-3_7) provides a thorough review of radiation exposure, including types and location of spacecraft related to the Earth and the Sun. In addition, acute and chronic effects of radiation exposure on crew members are discussed as well as countermeasures and prevention to minimize impact to crew health and performance. Radiation protection standards and clinical management of radiation-related health risks are presented.

Part III, entitled *"Space Flight and Crew Health: Adaptation, Pathophysiology, Rehabilitation, and Countermeasures,"* covers major body systems and the impact space flight has on them. There are seven comprehensive chapters, including cardiopulmonary, neurology, regulatory, metabolism and nutrition, clinical pharmacology and therapeutics, musculoskeletal, and behavioral health and performance. The chapter authors bring forward highly relevant findings and clinical evidence of the impact to the space traveler and the adaptive response to short-duration and long-duration LEO missions. These chapters have been refined with major updates from the third edition. Thirteen chapters of the third edition were consolidated into seven to minimize duplications.

Chapter [8](http://dx.doi.org/10.1007/978-1-4939-6652-3_8) reviews aeromedical considerations and the cardiopulmonary system's response to space flight. An examination of the evidence supporting cardiopulmonary system function during all phases of flight is provided. This includes a discussion on cardiac physiology, orthostatic tolerance, pulmonary response, and countermeasures to maintain physical fitness and aerobic capacity.

Chapter [9](http://dx.doi.org/10.1007/978-1-4939-6652-3_9) is a thorough review of the acute and chronic responses of the neurosensory and motor functions during space flight and post-flight health risks. Clinical findings presented include sensation, vision, cerebellar/vestibular function, spatial orientation, space motion sickness, and post-flight control of the circulatory system. Visual changes and risks due to altered cerebrovascular circulation are addressed.

Chapter [10](http://dx.doi.org/10.1007/978-1-4939-6652-3_10) is focused on regulatory physiology, including health implications of space flight impacts on the endocrine, fluid, electrolyte, and hematological systems. Endocrine and biochemical functions as well as hematologic alterations in plasma volume, red cell mass production, and destruction are discussed. Renal stone formation risks and the impact of the iodine, used as a potable water biocidal treatment, on the thyroid function, and interventions to minimize health risk are presented.

Chapter [11](http://dx.doi.org/10.1007/978-1-4939-6652-3_11) provides a summary of metabolism and nutrition in human space flight. A historical review of the development of food systems for space is presented as well as a discussion of the current system used on the ISS. Nutritional requirements for crew members are provided, which include palatability and cultural considerations. Food as a countermeasure for mental health and physical deconditioning is discussed.

The authors of Chap. [12](http://dx.doi.org/10.1007/978-1-4939-6652-3_12) present a review of the clinical pharmacology and therapeutics concerns in human space flight. The principles of space pharmacology are addressed and the

use and efficiency of medications in the space environment are discussed. Testbeds for the study of space pharmacokinetics as well as future directions of the use of medication in space flight are presented.

Chapter [13](http://dx.doi.org/10.1007/978-1-4939-6652-3_13) is focused on the musculoskeletal adaptation to space flight. The mechanisms for musculoskeletal adaptation to microgravity, the health risks association with this adaptation, and effective countermeasures are discussed in detail. The outcomes of the use of animal models both in-flight and ground-based analogs and simulated conditions are provided.

Chapter [14](http://dx.doi.org/10.1007/978-1-4939-6652-3_14) provides an updated review of behavioral health and performance and how space flight affects them and how these effects can be ameliorated. A discussion on the psychological adaptation factors, human-to-human system interface factors, sleep, and circadian factors is provided. Examples of pre-, in-, and post-flight and ground-based activities are used to illustrate the challenges for exploration missions.

Part IV, entitled *"Occupational Health Safety Issues in Space Flight,"* is focused on crew health monitoring and the international aspects of space medicine. This includes healthcare delivery systems and its challenges, telemedicine, and the development of a multinational approach to health through interaction with international partners and the experience gained between the U.S. and the USSR/Russia beginning in the 1960s.

Chapter [15](http://dx.doi.org/10.1007/978-1-4939-6652-3_15) explores the principles of crew health monitoring and care. This includes a description of astronaut medical certification, pre-flight, in-flight, and post-flight crew biomedical monitoring, and use of countermeasures, psychosocial support, and in-flight environmental monitoring; and post-flight rehabilitation is discussed. Health and Medical Technical Authority and medical policy development are presented for both NASA and international partners.

Chapter [16](http://dx.doi.org/10.1007/978-1-4939-6652-3_16) is focused on a number of issues related to the international aspects of space medicine. These include historical perspectives dating back to U.S./USSR relations in the late 1960s and early 1970s, the foundations for multilateral medical operations in support of international initiatives, the development of selection standards for the ISS era, and a look at the future of space medicine in a multicultural and technologically diverse environment.

Part V, entitled *"Ground-Based and Academic Training Programs,"* includes a concise summary of simulations and analogs as well as training in space medicine. Both of these are necessary elements for research and practice in space medicine.

Chapter [17](http://dx.doi.org/10.1007/978-1-4939-6652-3_17) describes the development and utilization of a wide variety of testbeds and analogs used to conduct clinical supporting research and training. Testbeds discussed range from simulations to analogs that provide geographic and physical similarities to space flight with regard to isolation, limited communications, and time delays in reaching definitive care.

Chapter [18](http://dx.doi.org/10.1007/978-1-4939-6652-3_18) addresses the current academic and professional skill training offered to space medicine practitioners as these individuals prepare to support human space flight activities with a focus on its development and evolution. In addition to academic training, other modes of training in space medicine are detailed, including international collaborations in preparing flight surgeons for duty. The chapter also highlights training efforts in a number of other spacefaring nations.

The book closes out with *"Future Perspectives,"* which provides a conclusion and next steps as space medicine moves forward. Chapter [19](http://dx.doi.org/10.1007/978-1-4939-6652-3_19) discusses national plans and commercial endeavors as human space flight and the launch systems that support LEO operations are being conducted by nongovernment entities. This includes a summary support for NASA-specific missions and emerging technological designs as well as efforts for commercial tourism. Commercial entities are discussed with regard to their progress and collaboration with NASA and its partners.

The operational necessities of space missions require that the clinical knowledge base is continuously updated to ensure that proper medical policies and standards are incorporated in a timely manner into space medicine practice. Recent reviews of biomedical publications suggest that the volume of research literature has been on the rise, but most of the epidemiological and clinical information is still of a descriptive nature. A scientometric and bibliometric analysis of the space medicine literature from major databases, such as the ISI Thompson Web of

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Science, PubMed/Medline, and archives specific or relevant to this textbook, was conducted by the George Mason University faculty,² validating high relevance materials are cited in this textbook. When compared to the wealth of the world biomedical literature, the number of clinical publications in space medicine is quite modest. Only 15% of the space medicine literature provides robust knowledge readily translatable into medical policies, practice, and standards. This does not differ from other biomedical disciplines, and especially in the field of occupational and environmental health [1–4]. It is also worth mentioning that this analysis revealed that while the U.S. remains the leading contributor to the space medicine and physiology knowledge base, the People's Republic of China has an increasing presence in this discipline, primarily in simulations and ground-based research.

This textbook will be updated on a regular basis to reflect new knowledge and challenges as they are made available from the ISS research and clinical observations. This text contains an extensive amount of information, and the editors and authors took utmost care to ensure accuracy and minimize potential errors or omissions. The authors are especially grateful to NASA for the use of the archives, narratives, and illustrations. In many instances those were the only sources of information available at this time as the NASA-supported research continues.

Finally, the editors will be more than happy to consider requests for tutorials on the subject matter.

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²Professor Laurie Schintler, Rajendra Kulkarni, Research Instructor, and Kingsley Haynes, Eminent Scholar, University Professor Emeritus at George Mason University (supported by the NASA Research Grant # NNX12AK32G).

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The editors acknowledge the efforts of a large number of individuals who contributed to this book. The fourth edition of *Space Physiology and Medicine: Evidence and Practice* has been completely revised and covers a wide range of topics that have relevancy to the practice of and research in space medicine. Numerous experts within academia and private organizations, especially the NASA Lyndon B. Johnson Space Center, played a part in early drafts of chapters as well as rewrites and refinement of the text. These contributions, whether small or large, helped the chapter authors develop the evidence and practices reflected in this textbook.

While it would be appropriate to list all the individuals that played some role, we would assuredly omit some. So we choose to acknowledge the men and women who have been associated with NASA over the past 20 years since the release of the third edition. There are two individuals who were selfless in their contributions, Dr. Sam Pool and Dr. Lakshmi Putcha. The space medicine community lost them both in the past few years. Each spent most of their careers conducting medical research and operations at the NASA Lyndon B. Johnson Space Center.

The editors wish to extend their appreciation to the astronauts who contributed so much as researchers and experiment participants to collect the information contained in this book. Special thanks are offered to Professor Laurie Schintler and Mr. Rajendra Kulkarni, Schar School of Policy and Government, George Mason University, who lent their expertise and time to help with the bibliometric analysis of the literature cited in this book.

As editors, we appreciate the dedication of the editorial staff and the publisher for keeping us on our toes and accurate in our summary of space medicine.

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Part I Introduction to Space Medicine

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 Part I

 Introduction to Space Medicine

Evolution of Human Capabilities and Space Medicine

Arnauld E. Nicogossian, Charles R. Doarn, and Yinyue Hu

Chapter Overview

NASA is the world's largest civilian space and aviation engineering research agency and a showcase of U.S. technological advances. It is the true birthplace of modern space medicine, which continues to be primarily influenced by engineering requirements. This chapter brings together the historical evolution of space medicine and human factors driven by technological development and political imperatives of human space exploration. Sustaining life, minimizing health risks and chances of injury have been and continues to be the primary goals for space medicine practice. In the sixteenth century, Ramazzini (Bernardino Ramazzini. *De Morbis Artificum Diatriba*. Apud Guilielmum van de Water Academii \bar{x} Typographaphum [Publisher]. Geneva, Switzerland. 1703), the father of occupational medicine, observed that sailors on long voyages of exploration did not fare as well as those on land when afflicted by chronic disorders. His observations still apply to modern space medicine practice, which is rooted in the principles of preventive medicine. Thus, space medicine practitioners' primary focus is on life support, food and water production systems, selection of cabin atmosphere and gas composition, hygiene, space habitat toxicology, radiation protection, and preventing infections. The principles of astronaut medical selection are to ensure

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"healthy" and disease-free candidates, while medical retention standards (annual clinical evaluations) and care ensures career longevity. Depending on the type and length of the space mission, certain medical conditions are considered compatible with the ability to perform assigned mission duties and a medical waiver is issued. Space medicine draws heavily on 50 years of aviation medicine knowledge and continues to be the central focus of today's practice of "personalized medicine." Traditionally, the knowledge underpinning space medicine practice lagged behind operational needs and remained largely empirical, relying on data from terrestrial analogs and simulations. Historically, extremely complex short duration missions to the Moon, followed by long duration low Earth orbit missions, did not permit adequate time for a systematic acquisition of biomedical knowledge base. Clinical and psychological research remained resource constrained for access to space, funding, and sufficient sample size of astronauts (the astronaut community constituted the astronaut sample size. Astronaut exposure to the space environment precluded a meaningful selection of a control group) due to political pressures of the "space race" and mounting costs from unexpected technological challenges. The national debate on the future of the space program, following the Apollo 17 mission, coupled with federal deficits due to the Vietnam War, resulted in significant reduction to NASA budgets, and termination of follow-on missions to the Moon. Excess Apollo program hardware was used to deploy the first U.S. space station (Skylab) and to conduct the first U.S.- Soviet collaboration in space: docking an Apollo and a Soyuz spacecraft. The U.S. investment in Skylab produced an unprecedented amount of data on the human responses to orbital long-duration space flight. The three Skylab missions produced the most comprehensive and fundamental seminal knowledge used by all space-faring nations in designing medical support and habitability systems for human space flight. Despite the many operational and logistic challenges, and occasional in-flight crew illness, no U.S. missions resulted in an unscheduled termination, or loss of life, due to medical conditions. This in itself is a testimonial to the soundness and

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efficiency of the clinical infrastructure and space medicine skills and expertise evolved since the early 1960s.

Budgetary and political imperatives led to periodic NASA management and programmatic reorganizations often affecting medical staff and projects. The number of NASA space medicine physicians (flight surgeons), remains small, and between 1960 and 1990 reached its peak of 35 individuals (not including astronaut physicians). Today, this small federal workforce is supplemented by military detailees and supported by contractors in the many demanding duties outlined throughout this textbook.

The experimental nature of spacecraft designed by the U.S. and other space-faring nations are briefly detailed in the context of space medicine. The interaction of the environment and spacecraft design, leading to potential health risk(s) are summarily reviewed. Extravehicular systems and robotics, intended to minimize unsafe exposures, while enhancing human performance, are briefly discussed. Space tourism and evolving commercial infrastructure and the potential for space medicine practice expansion are also presented. Finally the socioeconomic, cultural, and health care impacts of space exploration are briefly addressed.

Learning Objectives

- 1. Review major historical events framing the evolution of space medicine and exploration technology.
- 2. Discuss international collaboration in space medicine and contributions to the peaceful uses of space exploration by humans.
- 3. Become familiar with the evolving legislation, policies, and ethics guiding space medicine practice.

Introduction

"…chronic diseases also attack them (sailors), but they do not suffer from them as long as do those whose occupation is on land, for a ship is not a good place to ministering to chronic diseases...'

Bernardino Ramazzini: *De Morbis Artificum Diatriba* 1703 *[Diseases of the Workers]*

For millennia, humans have watched in awe the unending spectacle of the night skies. Ancient civilizations relied on the sun, moon, planets, and stars for time keeping and navigation, exploring the far horizons of our spaceship Earth. Over time, astronomy became intimately involved with religion, science, health, medicine, politics, philosophy, and war (Box 1.1). Changes in seasons, appearance of floods, and agriculture were predicated on the astronomical observations. These observations were essential to ensure the health security of the communities and mitigate disasters, plagues, and food shortages.

Box 1.1

The roots of space flight can be traced to the introduction of rocketry more than 2000 years ago with the invention of gunpowder in China. By the time of Genghis Khan's reign, gunpowder in the form of firecrackers and rudimentary rockets had become an integral part of Chinese defenses.

Box 1.2

Shortly before World War II, Oberth joined his former pupil, Wernher von Braun, in Germany's secret facility at Peenemünde Army Research Center (Heeresversuchsanstalt Peenemünde in German), located on the northern peninsula of the Baltic Island of Usedom, where they both worked on the development of the V-2 rocket [\[1](#page--1-0)]. This weapon was responsible for the death of 9000 civilians and military personnel and more than 12,000 slave laborers used to build them.

In the late nineteenth and beginning of the twentieth centuries, and purportedly inspired by the Jules Verne novel, *From the Earth to the Moon*, theoreticians, Konstantin Tsiolkovsky in Russia and Hermann Oberth in Germany published mathematical theories and calculations of speeds required to escape the Earth's gravitational force (Box 1.2) [[1\]](#page--1-0). Tsiolkovsky was primarily interested in space travel and was the first to describe the concepts of artificial gravity to protect the health of future explorers.

The era of modern rocketry truly began with Goddard's rocket launch on March 16, 1926, followed by an instrumented rocket in 1929. German engineers at the Peenemünde Army Research Center carefully monitored his experiments, and incorporated his discoveries into the design of Germany's offensive weapons. Under the direction of Wernher von Braun, the liquid-fueled *Aggregat-4* (A4) or V-2 became the first offensive ballistic missile of World War II. In 1945, after the defeat of Germany, V2 technology, together with surviving engineering expertise, was absorbed into the U.S. and Soviet Union military and space programs.

The Origins of Space Medicine in the United States

Interest in human space flight grew rapidly among a small circle of talented biomedical scientists—most of whom were initially from military aeromedical research laboratories.

Box 1.3

Since its inception, space medicine developed interdependencies with astronautics, human factors, habitability engineering, and biomedical research. Scientists and physicians soon recognized the need for an organization to coordinate and exchange information concerning space medicine research and practice. In 1951, the *Space Medicine Association* became the first constituent of the *Aerospace Medical Association*. In May 1978, NASA physicians P. Buchanan, J. Buhaine, J. DeGioanni, R. Hessberg, W. Hoffler, and A. Nicogossian established the *Society of NASA Flight Surgeons* as a constituent member of the *Aerospace Medical Association*.

This interest was also fueled in part by concern over the health and safety of pilots involved in supersonic test flights. Many conditions faced by space crews during launch and landing were similar to those encountered by test pilots in supersonic flights.

In 1948, a U.S. Air Force (USAF) physician, Colonel Harry G. Armstrong (1899–1983), organized a meeting at the USAF School of Aviation Medicine to discuss aeromedical problems of space travel [[2\]](#page--1-0). Participants included biologist Hubertus Strughold¹ (1898–1986) and astrophysicist Heinz Haber (1913–1990) [[3\]](#page--1-0). This meeting saw the birth of a new aerospace medicine discipline within the field of preventive medicine (Box 1.3) [\[2](#page--1-0)]. Space medicine is considered a product of the Cold War, borrowing applicable standards and knowledge from occupational, environmental, and aviation medicine [[4\]](#page--1-0).

Sustaining life and productive human function during space flight presented unique technological challenges requiring innovation in distance health monitoring, medical care, and life support. Concerns with physiological responses to weightlessness led both Soviet and U.S. pioneers in space medicine to use high-altitude balloons, Earth suborbital rockets, and orbital spacecraft carrying a variety of living organisms to study responses to the stresses of flight and validating engineering systems design [\[5](#page--1-0)]. In 1957, the Soviets flew invertebrates and vertebrates, including dogs. The *Sputnik 2* carried a dog named Laika into space on a

Box 1.4 Early NASA Biomedical Research Partnerships

The newly formed NASA research and operations field centers in Texas and California, as well as select institutes and universities—such as Lovelace and Mayo Clinics, University of New York at Rochester, The Ohio State University in Columbus, the Veterans Administration Hospital in San Francisco, Houston Medical Center in Texas, and others—began accelerated ground-based research and technology development programs in space human factors. These institutions in partnership with the aerospace industry developed a ground-based knowledge and technology base and successfully incorporated this information into future spacecraft development for the exploration of the Moon and low Earth orbit (LEO).

non-retrievable platform [[6\]](#page--1-0). Subsequent missions returned their canine passengers safely back to Earth.

To learn more about how the body would adapt to space flight, the U.S. launched two primates into space on board V-2 rockets by 1950. Although neither animal survived, these early flights demonstrated the need for reliable life-support systems and began the long process of requirements definition for the protection of mammals against the rigors and stresses of flight into space [\[7](#page--1-0)]. Early practitioners were trained in aviation medicine by the U.S. Navy (USN) and USAF. Beginning in the 1950s, these two organizations expanded their curricula to include space flight. These developments were reflected by new organizational designations: the Air Force Aviation Medical Facility in San Antonio, Texas became the School of Aerospace Medicine and the Navy's Aviation School at Pensacola, Florida became the Naval Aerospace Medical Institute. The Schools of Public Health at Johns Hopkins University, Harvard University, and The Ohio State University, which cooperated with military organizations in providing residency training, also reflected the changing focus in their curricula (Box 1.4). These training programs eventually led to the development of Aerospace Medicine residencies at Wright State University and the University of Texas Medical Branch (UTMB) (see Chap. [18](http://dx.doi.org/10.1007/978-1-4939-6652-3_18)) [\[8\]](#page--1-0).

Politics and Space Medicine

The October 4, 1957 launch of the Sputnik marked the Union of Soviet Socialist Republics (USSR) and U.S. engaging in a "space race" that allowed little time for biomedical research and medical practice to conduct a systematic indepth program of investigations into the health risks of space

¹Hubertus Strughold, PhD, was brought from Germany to the U.S. by the U.S. Army, together with other German physicians and researchers. They brought with them results of aeromedical and physiological research conducted in Germany up to the end of World War II. Dr. Strughold continued his research at the USAF Brooks School of Aerospace Medicine, which contributed significantly to the U.S. space medicine successes. Controversy and suspicion surrounding his participation in Nazi medical experiments resulted in several federal investigations and a tarnished reputation [[3\]](#page--1-0).

flight. Anecdotal reports, ground-based studies, and clinical observations, together with existing aviation databases became the foundation of the nascent space medicine program. This sense of urgency had a profound impact on the progress of space physiology and medicine, whereas clinical problems were handled empirically and research was conducted addressing issues after they were identified in space missions.

Specialized space biomedical research laboratories, established within the U.S. Air Force (USAF) and the U.S. Navy (USN), supported National Advisory Committee for Aeronautics (NACA) installations (NASA absorbed the NACA installations in 1958) as early as 1955. Some of the needed systems such as the USN's full-pressure suit used for high-altitude flights were adapted for space flight needs. Despite political differences, the U.S. and Soviet space medicine specialists met and exchanged knowledge at scientific meetings. After the Apollo Soyuz Test Project (ASTP), collaboration intensified, and the sides began planning for more ambitious activities using analogs and Soviet/Russian Cosmos missions for joint biological experiments (Box 1.5).

In the 1990s, some of the biomedical experiments using non-human primates came under attack from the People for the Ethical Treatment of Animals (PETA). NASA's use of animal test subjects was an attractive and highly visible target for PETA's goal to eliminate biomedical research using animal test subjects, especially non-human primates, in the U.S. The publicity surrounding the Bion 11 flight resulted in several Congressional hearings and further intensified postflight with the death of one of the primates, and ultimately led to the cancellation of U.S. funding for the remainder of the program. NASA physicians and biomedical scientists worked with many influential professional and international associations to ensure that ethical and scientific research priorities remained in the domain of the biomedical community and minimized political imperatives. The Bion 11 mission provided valuable inputs into space medicine, but unfortunately it severely impacted the Russian space biomedical community's ability to continue with non-human primate missions, especially at a time of severely constrained funding in the "post-Perestroika" era.

Despite these challenges, space biomedical collaboration between the U.S. and the Soviet Union/Russia continued to flourish, surviving political turmoil. Academicians Oleg G. Gazenko and Anatoli I. Grigoriev remained at the forefront of such collaboration for at least four decades and were joined by other nations.

Historical Demographics

The majority of professional space travelers are supported by the space agencies from the U.S., USSR/Russia, European Union, and Peoples Republic of China. Russia was the first

Box 1.5 NASA's Research Using Non-Human Primates in Space

The U.S. developed a series of biosatellites to initially fly invertebrates and rodents, and later to orbit nonhuman primates for several days in LEO. The last spacecraft in this series, *Biosatellite III*, was launched on June 28, 1969. On board was a single, male primate, *Macaca nemestrina*, named Bonnie, weighing 6 kg, for a planned 30-day mission. The mission objective was to investigate the effect of long-term space flight on behavior, performance, cardiovascular, and fluid and electrolyte metabolism. Bonnie was over instrumented and became sick after several days on orbit. The mission was terminated short of 9 days. Bonnie died 8 h after recovery from dehydration and other medical complications. The U.S. Congress terminated the program by reducing NASA life sciences funding. A similar Russian-funded research effort *Bion 11* also had issues. One of the primates died on the second post-flight day following anesthesia for tissue, muscle, and bone biopsies.

A NASA/Russian review committee under the chairmanship of Dr. Ronald Merrell found that the post-flight dehydration, cardiovascular compromise, and poor body temperature regulation contributed to the death of the non-human primate. The review committee characterized the immediate post-flight period to a *high risk perioperative category of patients* (American Society of Anesthesiologists [ASA] III/IV Class). NASA dropped out of participation in a planned *Bion 12* mission due to the inability to adhere to the approved study protocols based on the probability of high mortality risks associated with early post-flight procedures. Following several hearings the U.S. Congress disapproved further participation and funding for the Bion 12 mission.

country to provide access to paying space tourists. Tables [1.1,](#page-23-0) [1.2,](#page-23-0) and [1.3](#page-23-0) summarize these human space flight demographics [\[9](#page--1-0)]. By October 2016, only 551 individuals (491 men, 60 women) had spent a combined total of more than 135 human years in space missions.2 Sending humans into space is still the domain of the U.S., Russia, and China, and as of June 2015, with the retirement of the NASA Space Shuttle, the U.S. has been relying on Russian Soyuz spacecrafts to deliver astronauts to the International Space Station (ISS), while developing a new exploration class spacecraft and relying on the private sector to develop a "space taxi" capability for astronauts to travel to and from the ISS.

²This number varies depending on the three existing definitions of astronauts based on altitude reached.

Table 1.1 Number of months logged in space by individual crew members

Adapted from <http://www.astronautix.com/articles/aststics.htm>[[9\]](#page--1-0)

Table 1.2 Actual number of missions flown by individual crew member and by country of origin

Missions							
United States	102	103	58	58	21		
Russia/USSR	51	29	23			$\overline{4}$	
China						Ω	
Europe	29	15				Ω	Ω
Asia	Ω	6			Ω	θ	Ω
Canada						θ	Ω
Other countries					Ω	Ω	Ω
Total men	188	140	81	60	19	Ω	
Total women	19	18					

Adapted from http://www.astronautix.com/articles/aststics.htm [[9\]](#page--1-0)

Table 1.3 Space tourists by countries of origin (paying space explorers)

	Country of origin Number of paying tourists	Vehicle	Country of origin
United States	.5	Soyuz	One Iranian-American and one Hungarian-American, one British-American
Japan	(1)		Did not fly due to medical reasons
Iran		Soyuz	Iranian-American
Canada		Soyuz	
Hungary			Hungarian-American
United Kingdom	\Box	Soyuz and <i>Mir</i>	A woman chemist flew under the project Juno
South Africa		Soyuz	South African-British
Total women	2		
Total men	6		

Medical Diplomacy

Medicine is an integral element of all human space missions, both human and robotic. Sustaining humans, searching for habitable planets or extraterrestrial life, is of interest and involves flight surgeons and medical personnel. Humans can present a threat to extraterrestrial life and the reverse is also true. Planning human exploration into the Solar System also requires a good understanding of the operational environments, determining potential health threats and hazards, and devising appropriate tools and countermeasures (Box [1.6](#page-24-0) [\[7](#page--1-0)]). International collaboration in space is also viewed as a showcase and demonstration of a national scientific capability. Space exploration is considered a contributor to knowledge and a better life on Earth. Most of the international

collaborative human space missions are expensive, use unique attributes of space to solve space medicine problems, and offer a promise for potential return on the investment.

NASA medical personnel expertise and innovative practices are routinely sought to address earthbound medical problems. Technological demonstrations such as telemedicine, remote sensing and vector borne infections, natural disaster warnings and mitigations, and miniaturized and compact health care systems for home use were transferred to the private sector by the sponsoring individual space agencies [\[10](#page--1-0)]. It is estimated that biomedical space research has contributed significantly to the private sector enterprise [\[11](#page--1-0)]. These real and perceived benefits or "spinoffs" are the basis for a vigorous outreach program, often with the participation

Before the formation of NASA, most of the NACA functions were located at the Langley Research Center in Hampton, Virginia. It is in this location that the majority of the Department of Defense (DoD) detailed medical specialists began their work on Project Mercury [\[7](#page--1-0)].

Initially, the Office of Advanced Research and Technology was responsible for both the biotechnology and human research, and institutionally for the biological activities at the NASA Ames Research Center.

NASA followed the military model for its space medicine program. The purpose was, and remains, to ensure crew health and safety and to capitalize on relevant biomedical knowledge gathered by other federal or private agencies and organizations.

With time, a second major activity was developed, mostly under the auspices of the physical sciences, addressing the understanding of the origins, evolution, and destiny of life in the universe—or astrobiology.

Though both programs remained separate for decades, the human exploration of the solar system is bound to bring their research and operational interests together, notably in planetary protection, preventing back contamination and the hazards associated with the search for alien life forms—mostly bacteria (see Chap. [2](http://dx.doi.org/10.1007/978-1-4939-6652-3_2)).

The NASA chief health and medical officer (CHMO) is responsible for coordinating the planetary protection policies with the human solar system planning activities.

of physicians, to ensure the diffusion of space research into terrestrial health programs.

For many years, Dr. Oleg Gazenko, a physician and statesman, and the director of the USSR Institute of Biomedical Problems, led the Soviet delegations to the United Nations (UN) Committee on the Peaceful Uses of Outer Space (COPUOS). He established annual special sessions at the UN to showcase health benefits from applications of space research and technology. He ensured that NASA and other space-faring nations participated in these sessions [[12\]](#page--1-0). NASA routinely testifies to the U.S. Congress on its technology transfer programs, especially in disaster mitigation and humanitarian help, as required by the Space Act of 1958 [\[13](#page--1-0), [14\]](#page--1-0). Unlike with other U.S. agencies currently, medical diplomacy is not part of the official space medicine training curriculum and is usually acquired over time as part of NASA career opportunities.

Space Medicine in the United States

From NASA's beginning, the responsibilities for life sciences research and space medicine remained diffused and fragmented. NASA's expectation of space medicine is to ensure crew health and mission safety. Most of the knowledge used to plan for the mission was derived from the aviation medicine and ground analogs. The National Research Council Committee on Bioastronautics (1958–1960) was asked to predict possible health risks to astronauts. These risks are detailed in Table [1.4](#page-25-0) [[15\]](#page--1-0). The predictions were quite accurate and helped focus the clinical research and development approach [[15\]](#page--1-0).

Traditionally, NASA management assigned a high priority to practice and research with relationship to health and safety of the space crews. NASA did seek support for fundamental biological research through cooperation with the international community and especially the U.S. National Institutes of Health (NIH). Flight access was given priority to those activities that were judged to have a direct benefit to human safety, or for the "improvement of life on Earth." This philosophy led to tensions between field centers, clinical (operational) personnel, and academic bioscientists competing for limited funds. In the early 2000s, it became necessary to address these conflicts, real or apparent, by separating organizationally clinical programs from the biomedical research activities within NASA. An office of the Chief Health and Medical Officer (OCHMO) was established, reporting directly to the NASA Administrator. Reorganization of the biomedical programs within NASA continues on a decadal basis and is driven by the availability of resources and human exploration priorities as set by the U.S. president through revisions of the National Space Policy.

NASA has sponsored specific high-priority biomedical research consistent with mission needs during the Gemini, Skylab, and Space Shuttle in anticipation of the ISS program. Such an approach does delay the acquisition of a systematic space medicine knowledge base. Major events shaping the NASA space medicine program, organizations, and relevant policies are outlined in Table [1.5](#page-26-0) [[16–22\]](#page--1-0).

Project Mercury (May 5, 1961 to May 16, 1963)3

In 1960, NASA Administrator T. Keith Glennan established a life sciences program under the direction of Clark T. Randt, a prominent physician and bioscientist. This move was designed to ensure collaboration with the larger community of biologists and clinicians. NASA's intent was to harness the existing

³The dates reflect the launch dates of the first and last missions.

Predicted by the NRC 1958	Observed as of August 2015
Anorexia	Only in association with space motion sickness
Nausea	With space motion sickness
Disorientation	During initial adaptation to the space environment
Sleepiness	With severe space motion sickness
Sleeplessness	Throughout space mission duration
Fatigue	Poor workload scheduling
Restlessness	None
Euphoria	None
Hallucinations	None
Decreased g tolerance	post-flight
Urinary retention	Affects less than 1% of crews
Diuresis	None measured
Muscular incoordination	Gait disturbances post mission
Muscle atrophy	Documented
Demineralization of bones	Documented
Renal calculi	Pre-mission increased risk
Motion sickness	In most crew members with varying degree. Repeat exposure, reduces the incidence
Pulmonary atelectasis	None
Tachycardia	Rare and during high metabolic load activities
Hypertension	None
Hypotension	Post-flight and transient
Cardiac arrhythmias	Documented in few instances. Pre-existing rhythm abnormalities not aggravated by space flight
Post-flight syncope	Documented
Decreased exercise capacity	Post-flight
Reduced blood volume	Documented post-flight
Reduced plasma volume	Documented post-flight
Dehydration	Observed post-flight due to decreased fluid intake and mild.
Weight loss	Variable
Infectious illness	Minor infections treated in space. Increased shedding and reactivation of HSV attributed both to stress and possible decrease immune response (see Chap. 5)
Not predicted	Visual changes and pathological findings

Table 1.4 Predicted and observed medical problems associate with space missions

NRC National Research Council, *HSV* Herpes simplex virus

knowledge to be applied to the space program needs and not necessarily to spin a major life sciences research enterprise. After 1 year in office, Randt felt that his efforts in establishing a strong medical presence in NASA were blocked, and tendered his resignation. This led to a fragmentation of the life sciences organization in 1962, with the Biosciences Programs, including the Biosatellite Office, being established in the Office of Space Sciences; NASA Ames Research Center, at Moffett Field, California (former Navy Air Station) was designated as the laboratory responsible for biological research including the development of the Biosatellite project to explore the physiological responses of primates, and other living organisms, to extended duration space missions [\[23](#page--1-0)]. The space medicine program was transferred to the human space flight organization, to better integrate and address engineering and health safety risk management. The new organizations captured many of these early leaders in space medicine [[19](#page--1-0)]. By 1962, some of the team members were transferred to the Office of Advance Research and Technology, the Space

Medicine Directorate, Office of Manned Space Flight, at NASA Headquarters or to the Manned Spacecraft Center (MSC) located in Houston Texas (Box [1.7\)](#page-28-0). These individuals took on the responsibilities at NASA Headquarters and the MSC for developing requirements and systems to support NASA and the nation's program of getting to the Moon and back, including development of the clinical research and designing the Lunar Receiving Operations, following return from the Moon missions.

In the late 1950s, the selection process began with Eisenhower's direction that all astronaut candidates be recruited from the ranks of military test pilots. As a group, military test pilots were required to demonstrate many abilities deemed to be crucial for an astronaut: good judgment, decision making under stress and in threatening situations, quick reaction time, and refined motor skills. Of the first group of applicants, 100 test pilots were given interviews, psychiatric examinations, and a complete medical evaluation that included medical stress tests (Box [1.8](#page-28-0)) [[24](#page--1-0), [25\]](#page--1-0). The main

(11981–2007–2007) NASA-DOS (1998–1998–1998) ISS (1998–1998) ISS (1998–1998–present) ISS (1998–1998–present) IS •••• Increased neurosensory and gaze control, target requires three times the Increased neurosensory Radiation levels higher and gaze control, target with the nearing of the requires three times the length of time spent in Increased rest periods, performance improves Radiation levels higher NASAMIR (1992-1998) • Bone and muscle loss changes (e.g., posture length of time spent in following exposure to with the nearing of the • Bone and muscle loss Risk of kidney stones Risk of kidney stones changes (e.g., posture following exposure to Increased rest periods performance improves increases with length Tilt position counters maintaining physical Tilt position counters redistribution during communication with maintaining physical increases with length long-duration flights redistribution during • Recovery post-flight communication with equal length of time relaxation time, and long-duration flights Recovery post-flight equal length of time relaxation time, and mission completion continues unabated mission completion continues unabated family and friends orbit for bone and orbit for bone and family and friends • Increased cardiac Increased cardiac >70 days in-flight >70 days in-flight acquisition) with acquisition) with microgravity for performance and performance and microgravity for and neuromotor and neuromotor dysrhythmia at dysrhythmia at Good nutrition Good nutrition improves crew improves crew than expected necessary for necessary for than expected rapid blood rapid blood safetyCrew safetyCrew strength muscles of flight deorbit •••••••landing; some astronauts unable to Salt and water drink most effective • Decreased sensorimotor control at landing; some astronauts unable to inflation, for post-flight orthostatic Salt and water drink most effective Midodrine potential for preventing Midodrine potential for preventing • Decreased sensorimotor control at walk unaided or make rapid egress inflation, for post-flight orthostatic walk unaided or make rapid egress thermal load, with 33% incidence thermal load, with 33% incidence suit hard torso during underwater Fluid loading before deorbiting a Risk of renal stone and proposed Simulation of pre-flight sleeping Fluid loading before deorbiting a Shoulder injuries from the space suit hard torso during underwater orthostatic intolerance post-flight orthostatic intolerance post-flight Urinary retention relieved by the Air cooled escape suit increased Risk of renal stone and proposed Urinary retention relieved by the Increased risk of virus shedding Increased risk of virus shedding Simulation of pre-flight sleeping arrangements (bed sack, pillow) arrangements (bed sack, pillow) Air cooled escape suit increased Shoulder injuries from the space post-flight, reduced to 3% with Increased use of analgesics and post-flight, reduced to 3% with Increased use of analgesics and moderately effective for space together with anti-g suit early together with anti-g suit early moderately effective for space Space shuttle/spacelab/spacehab Caloric intake improved with Space shuttle/spacelab/spacehab (immunocompromised) with Caloric intake improved with (immunocompromised) with Women more susceptible to • Women more susceptible to helps counter sleeplessness promising countermeasure, helps counter sleeplessness promising countermeasure, increasing flight duration of orthostatic intolerance of orthostatic intolerance increasing flight duration Injectable promethazine Injectable promethazine Transient low back pain wider dietary selections • Transient low back pain use of urethral catheters use of urethral catheters wider dietary selections potassium citrate as a orthostatic intolerance orthostatic intolerance liquid cooled garment liquid cooled garment potassium citrate as a sleeping medications sleeping medications training sessions training sessions countermeasure motion sickness motion sickness countermeasure oral solution oral solution intolerance intolerance $(1981 - 2011)$ in-flight •••••••••••••• Ocular foreign body (pencil conducting in-flight clinical conducting in-flight clinical Rigorous data collection to physician
Ocular foreign body (pencil sufficient to protect against • Rigorous data collection to due to unscheduled ground due to unscheduled ground potential countermeasures corrected by the astronaut superficial lacerations and sufficient to protect against potential countermeasures superficial lacerations and corrected by the astronaut cardiovascular, bone, and deconditioning, using the systematic investigations Crew and ground control cardiovascular, bone, and deconditioning, using the Dispelled the low caloric missions reported during systematic investigations into man's physiological Crew and ground control lead tip) removed by the missions reported during Crew schedule overload lead tip) removed by the • Dry skin and occasional Dispelled the low caloric into man's physiological Crew schedule overload • Inflation of anti-gravity Dry skin and occasional for post-flight physical Subluxation of a finger for post-flight physical Inflation of anti-gravity suits during post-flight Subluxation of a finger suits during post-flight treadmill on Skylab 4, requirements in space scientific experiments requirements in space scientific experiments treadmill on Skylab 4 Aerobic exercise not Aerobic exercise not Thornton-Whitmore Extensive testing of Extensive testing of boils, responding to boils, responding to Thornton-Whitmore First U.S. astronaut control requests for astronaut physician control requests for • First U.S. astronaut meet experimental astronaut physician Introduction of the • Introduction of the physician in space physician in space responses to flight meet experimental previous missions responses to flight bicycle ergometer bicycle ergometer Skylab (1973-1974) evaluations. First protected against previous missions (1958–1963) Gemini (1961–1966) Apollo (1961–1972) Skylab (1973–1974) evaluations. First protected against musculoskeletal deconditioning musculoskeletal deconditioning deconditioning deconditioning antibiotics antibiotics protocols tensions landing •••••••••••orthostatic tolerance orthostatic tolerance bigemini) attributed • Reduced post-flight bigemini) attributed Reduced post-flight mass $(2-10\%)$ and (recovery within 1 arrhythmias (a run Decreased red cell mass $(2-10\%)$ and food consumption (recovery within 1 exercise tolerance Apollo 15 cardiac • Apollo 15 cardiac arrhythmias (a run Decreased red cell Less than optimal Less than optimal food consumption negative pressure exercise tolerance to potassium loss) to potassium loss) Apollo (1961-1972) $(1260 - 2903 \text{ kcal})$ (1260–2903 kcal/ negative pressure dehydration and dehydration and (tilt/lower body (tilt/lower body plasma volume plasma volume (first 3 days) disturbances disturbances (first 3 days) weight loss of 22 nodal day)
Post-flight weight loss of 22 nodal • Vestibular Vestibular • Post-flight Decreased • Decreased post-flight $(4 - 9\%)$ week) tests) ••••••Gemini (1961-1966) mission of 14 days mission of 14 days (medical focus), in (medical focus), in preparation for the preparation for the post-flight medical post-flight medical capacity compared bone density (7% capacity compared bone calcium and space rendezvous Second astronaut metabolic cost of during the first in • Second astronaut bone density (7 $%$ Sustained loss of bone calcium and during the first in space rendezvous Loss of os calcis metabolic cost of Apollo program Loss of exercise Loss of *os calcis* Sustained loss of New Nine" and New Nine" and Apollo program Loss of red cell Loss of exercise muscle nitrogen comprehensive muscle nitrogen selection "The comprehensive Loss of red cell $100\,\%$ of crews selection "The 100% of crews with pre-flight from baseline) mass (ranging with pre-flight extravehicular extravehicular mass (ranging intolerance in intolerance in from baseline) $5-20\%$ from 5–20% from performance performance and docking evaluations documented documented Higher than • Higher than and docking the longest evaluations the longest • Post-flight orthostatic orthostatic Post-flight Sustained Sustained baseline) predicted pre- and baseline activity First ••••••••and without major and without major body functions for body functions for periods up to 34 h screening focused demonstrated that periods up to 34 h • Medical selection Clinical Findings • Medical selection on psychological, function properly screening focused demonstrated that on psychological. function properly deteriorations of deteriorations of laboratory tests space missions laboratory tests space missions of weightless of weightless Six Mercury Six Mercury stress, and *man* could stress, and endurance endurance man could $(1958 - 1963)$ untoward untoward physical Programs^a Mercury Mercury flight •Clinical Findings Programs^a Elements

Table 1.5 Major medical and select policy milestones in the United States space program [16-22] **Table 1.5** Major medical and select policy milestones in the United States space program [[16–22](#page--1-0)]

reduction countermeasure reduction countermeasure • Validation of the risk • Validation of the risk ISS (1995-present) for renal stones for renal stones

resistive exercise protocols resistive exercise protocols • Validation of the most Validation of the most effective aerobic and effective aerobic and for musculoskeletal or musculoskeletal

(primarily males) suffer from physiological and anatomical (primarily males) suffer from physiological and anatomical visual problems in long visual problems in long $20\,\%$ of astronauts \bullet 20% of astronauts deconditioning deconditioning

 Some astronauts reaching Some astronauts reaching career limits of radiation career limits of radiation duration missions duration missions

exposures exposures

"The program dates refer to the year of authorization (start) and termination aThe program dates refer to the year of authorization (start) and termination

Authority

Box 1.7

The Houston MSC was renamed in honor of President Lyndon B. Johnson on February 19, 1973. It is the premier facility for astronaut training, care and preparation for space missions. This NASA facility has and is hosting many international crews, flight surgeons, and biomedical researchers and is the home of the Mission Control Center for space operations. The center also developed and hosts linguistic and cultural suitability training and flight surgeon training programs (see Chap. [16](http://dx.doi.org/10.1007/978-1-4939-6652-3_16)).

Box 1.8

The purpose of these extensive evaluations was to discover any hidden medical problems, to establish baseline levels of physical fitness, and perhaps most important, to compile a medical database for each individual against which any changes brought about by later space missions might be measured and quantified.

goal was to identify individuals in good health and able to withstand extremes of physical stresses, isolation, and sensory deprivation, without psychological or physiological consequences. This type of testing remained in effect until the end of the Apollo era, and was modified over time as new knowledge about health effects of space flight became available.

Following the selection of the Mercury 7 astronauts, Dr. W. Randolph Lovelace, on his own initiative, began the medical evaluation of a group of women aviators who could at a later date fly in space [[26](#page--1-0), [27](#page--1-0)]. Unfortunately, Lovelace's vision would not be fulfilled for decades, until the Space Shuttle era. The First Ladies Astronaut Trainees Project fell victim to political and Cold War space race rivalries. The "Women in Space" training was terminated shortly after its initiation. It was the Soviets who launched the first female cosmonaut, Valentina Tereshkova, on the Vostok 6 spacecraft on June 16, 1963.

In 1958, the USAF "Man in Space Soonest" program was transferred to NASA and it became the foundation of the Mercury project. The human requirements for radiation protection, atmospheric pressure and gas composition, food and water, and thermal control had to be established within the constraints on system failure tolerance, size, weight, power, and operation under conditions of thermal extremes, acceleration, and weightlessness [[28\]](#page--1-0). Almost from the onset of human space flight, space medicine physicians and engineers were often at odds on design requirements. The astronauts and physicians insisted that there be a window in the Mercury

Fig. 1.1 Alan Shephard getting ready for launch (Courtesy of NASA)

capsule so the astronaut could look out the window. This addition, notwithstanding the cost and delays, proved a valuable tool for human direct observations of the flight environment, navigation, and Earth observations.

The primary goal of Mercury to launch and recover a person was reached with Alan Shepard's flight in May 1961 (Fig. 1.1), and in all, two sub orbital and four orbital Mercury missions were flown, including one that lasted for 34 h and accomplished 22 orbits around the Earth. All 6 Mercury astronauts returned to Earth in satisfactory physical condition [\[29](#page--1-0)]. All Mercury astronauts went on to fly on Gemini, Apollo, and Skylab Missions.

However, a seventh astronaut, Donald "Deke" Slayton, was disqualified from flight duties due to cardiac arrhythmia (slow atrial fibrillation), which did not prevent him from flying 20 years later on July 15, 1975 on the first international "détente" mission, Apollo–Soyuz Test Project (ASTP). In the intervening period, Slayton served as the head of the Astronaut Office. He continued to seek reinstatement to flight duties and was seen repeatedly by preeminent cardiologists including Drs. Eugene Braumwald and Dudley White. Through the efforts of Dr. Charles Berry, he was treated with anti-arrhythmia medications, resigned from the USAF, and was put back on aviation duties in the late 1960s.

Box 1.9

Like Yuri Gagarin of the Soviet Union, John Glenn was considered a national hero, and NASA did not allow him to fly again until 1998, following his retirement as a U.S. Senator from Ohio.

At this time, many U.S. scientists and engineers did not see the value of having the human in the system.

John Glenn became a national hero following his first orbital mission on February 20, 1962. He returned to space on October 29, 1998 as a payload specialist aboard the STS-95 (Box 1.9).

The early space missions were valuable for both dispelling and validating numerous medical concerns (Box 1.10). The principal findings of human adaptation to space flight were weight loss, resulting primarily from dehydration, and some impairment of cardiovascular function. Cardiovascular data from the final and longest Mercury flight showed postflight orthostatic intolerance, dizziness on standing, and hemoconcentration [[30\]](#page--1-0). From a behavioral perspective, astronauts performed well under conditions of weightlessness and stress. The program had succeeded in accomplishing its purposes: to successfully orbit a man in space, to explore human ability of tracking and control, and to learn about microgravity and other biomedical problems associated with space flight.

Gemini Program (March 23, 1965 to November 15, 1966)

Planning for the Gemini Program began in May 1961, just after the successful completion of the first sub-orbital Mercury mission. The 2-man Gemini capsule was based upon the experience of Project Mercury and was designed to demonstrate new capabilities, such as extravehicular activities (EVAs), while providing NASA with the necessary experience in conducting extended space missions. The program also allowed the biomedical community to delineate the physiological limits of astronaut endurance, an essential step for planning future missions of greater complexity.

Gemini successfully completed ten manned space missions, with many notable accomplishments (Box 1.11). The program itself was a resounding success in advancing the science of space technology. Fifty-two different experiments were performed during its ten missions. The Gemini achievements were a litany of precedents and records: the

Box 1.11 Gemini Program Objectives:

- 1. Demonstrate the feasibility of space flight lasting long enough to complete a lunar landing;
- 2. Perfect the techniques and procedures for orbital rendezvous and docking of two spacecraft;
- 3. Achieve precisely controlled Earth reentry and landing;
- 4. Establish capability for extravehicular activity; and
- **Box 1.10** 5. Enhance the flight and ground crew proficiency.

first U.S. extravehicular activity during Gemini-4 (Fig. [1.2](#page--1-0)); the first rendezvous and docking maneuver during Gemini-8; and the 14-day Gemini-7 mission, dedicated to biomedical studies. The question remained, however, whether the observed cardiovascular deconditioning was a self-limiting problem.

For the first time, slow and sustained tumbling was experienced by the crew of Gemini 8 during the docked phase with the Agena spacecraft [\[31](#page--1-0)]. Physiological and technical risks led to premature undocking of the two spacecraft. Nevertheless significant knowledge on manual piloting in space under off-nominal conditions was acquired, to be applied during future missions. Once the uncoupling was accomplished, astronaut Neil Armstrong was able to regain control of the tumbling spacecraft and return safely to the Earth.

The Gemini missions reinforced the medical conclusion that humans could live and work in space and could certainly do so for the duration required for the forthcoming Apollo missions (Box [1.12\)](#page--1-0). A number of new responses to the space flight environment, such as bone mineral loss, were noted; however, none were considered of real consequence for missions lasting 2 weeks or less. While bringing new issues and concerns to light, Gemini left other medical questions unresolved [\[32](#page--1-0)]. The program's biomedical findings nonetheless served to structure and guide studies to be designed for later, longer missions (Box [1.13](#page--1-0)) [\[33](#page--1-0), [34\]](#page--1-0). Such experiments would be needed to determine the basis and time course of the observed physiological changes.

Mercury and Gemini projects attracted new physicians and researchers into the space program. Practitioners in space medicine began the formulation of the operational aeromedical support for space missions and provision of astronaut care. These events led to the expansion of the scientific data base in space physiology: Drs. Craig Fischer and Phillip Johnson were first to describe "space anemia," fluid loss and head ward fluid shifts as a response to exposure to microgravity [\[30](#page--1-0), [35](#page--1-0)]. Dr. Carolyn Leach described fluid, electrolyte, and endocrine changes [\[36](#page--1-0)], while Dr. Steve Kimzey began research into the causes of space anemia,