Exh. CAC-1T Witness: Charles A. Czeisler

BEFORE THE WASHINGTON UTILITIES AND TRANSPORTATION COMMISSION

WASHINGTON UTILITIES AND TRANSPORTATION COMMISSION, Complainant,

Docket TP-

v.

PUGET SOUND PILOTS,

Respondent.

TESTIMONY OF

CHARLES A. CZEISLER

ON BEHALF OF PUGET SOUND PILOTS

JUNE 29, 2022

Haglund Kelley, LLP 2177 SW Broadway Portland, OR 97201 Tel: (503) 225-0777 / Fax: (503) 225-1257

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1	1		
^))	

IDENTIFICATION OF WITNESS I.

2 3

O:

Please state your name, occupation and business address.

4 A: My name is Dr. Charles A. Czeisler. I am currently the Frank Baldino, Jr., Ph.D. 5 Professor of Sleep Medicine, Professor of Medicine and Director of the Division of Sleep 6 Medicine at the Harvard Medical School, and I am Chief of the Division of Sleep and Circadian 7 Disorders in the Departments of Medicine and Neurology and Director of the Sleep Matters 8 Initiative, both at Brigham and Women's Hospital in Boston, Massachusetts. My business 9 address is 221 Longwood Ave., BLI 438, Boston, MA 02115-5817. 10

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O:

Please describe your educational background.

13 A: I received my bachelor's degree from Harvard College in biochemistry and molecular 14 biology magna cum laude in 1974 and was inducted into Phi Beta Kappa at Harvard College in 15 1999. I received the degree of Doctor in Philosophy (Ph.D.) in Neuro- and Biobehavioral 16 Sciences in 1978 from Stanford University and the degree of Medical Doctor (M.D.) from 17 Stanford University in 1981. 18

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22

20 **O**: Please describe your experience in the physiology of the human circadian timing 21 system and its relationship to the sleep-wake cycle.

A: I have more than four decades of experience in the field of basic and applied research on 23 the physiology of the human circadian timing system and its relationship to the sleep-wake cycle. 24 Together with my collaborators, I have done significant study and research into the areas of sleep 25 and fatigue. This research has resulted in 286 original reports in peer-reviewed journals and 118 26

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review articles/proceedings of meetings, as well as five books/monographs/theses and numerous abstracts of research.

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2	abstracts of research.
2	I am a Diplomate of the American Board of Sleep Medicine, a Fellow of the American
4	Academy of Sleep Medicine, a Fellow of the American Physiological Society and an elected
5	Fellow of the Royal College of Physicians (London). I have also served as President of the Sleep
6	Research Society; Chairman of the Board of Trustees of the National Sleep Foundation;
7	Chairman of the National Institutes of Health Sleep Disorders Research Advisory Board of the
8	National Center on Sleep Disorders Research in the National Heart, Lung and Blood Institute;
9	Chair, Steering Committee, Academic Alliance for Sleep Research; and Chair, Steering
10 11	Committee, Sleep Research Network. For ten years, I served as the Team Leader of the Human
12	Performance Factors, Sleep and Chronobiology Team of NASA's National Space Biomedical
13	Research Institute in Houston, Texas.
14	I have also been elected as a Fellow of the American Society for Clinical Investigation, a
15	
15	Member of the Association of American Physicians, a Member of the International Academy of
16	Member of the Association of American Physicians, a Member of the International Academy of Astronautics and a Member of the National Academy of Medicine (formerly called the Institute
16 17	
16 17 18	Astronautics and a Member of the National Academy of Medicine (formerly called the Institute
16 17	Astronautics and a Member of the National Academy of Medicine (formerly called the Institute of Medicine of the National Academies). I served on the faculty of the World Economic Forum
16 17 18 19	Astronautics and a Member of the National Academy of Medicine (formerly called the Institute of Medicine of the National Academies). I served on the faculty of the World Economic Forum in Davos, Switzerland in 2014, the Aspen Ideas Festival Spotlight Health in Aspen, Colorado in
16 17 18 19 20	Astronautics and a Member of the National Academy of Medicine (formerly called the Institute of Medicine of the National Academies). I served on the faculty of the World Economic Forum in Davos, Switzerland in 2014, the Aspen Ideas Festival Spotlight Health in Aspen, Colorado in 2016 and 2017, and Aspen Brain Health in 2018.
16 17 18 19 20 21	Astronautics and a Member of the National Academy of Medicine (formerly called the Institute of Medicine of the National Academies). I served on the faculty of the World Economic Forum in Davos, Switzerland in 2014, the Aspen Ideas Festival Spotlight Health in Aspen, Colorado in 2016 and 2017, and Aspen Brain Health in 2018. In addition, I serve or have served as a committee member on a number of national and
 16 17 18 19 20 21 22 	Astronautics and a Member of the National Academy of Medicine (formerly called the Institute of Medicine of the National Academies). I served on the faculty of the World Economic Forum in Davos, Switzerland in 2014, the Aspen Ideas Festival Spotlight Health in Aspen, Colorado in 2016 and 2017, and Aspen Brain Health in 2018. In addition, I serve or have served as a committee member on a number of national and international advisory panels in the public and private sectors related to sleep and circadian

Assessment, U.S. Congress; the Work Hours, Sleepiness and Accidents Consensus Conference, 26

4	National Institute for Psychosocial Factors and Health, Department of Clinical Neuroscience,
1 2	Karolinska Institute, Stockholm, Sweden; the Advisory Committee on Night Operations and
2	Human Chronobiology, Life and Environmental Sciences Division, Air Force Office of
4	Scientific Research; the Panel on Workload Transition, Committee on Behavioral and Social
5	Sciences and Education, National Research Council; the Research Briefing Panel on Basic Sleep
6	Research, Division of Health Science Policy, Institute of Medicine, National Academy of
7	Sciences; Plenary Address, American Trucking Association Foundation Conference on
8	Managing Fatigue in Transportation; the NASA Advisory Panel on the Pre-flight Circadian
9 10	Shifting of Shuttle Flight Crews, Space and Life Sciences Directorate, National Aeronautics and
11	Space Administration; the Biological Rhythms Task Force, Mental Health Research Network I,
12	John D. and Catherine T. MacArthur Foundation; the External Advisory Committee, National
13	Science Foundation Center for Biological Timing at the University of Virginia; External
14	Advisory Committee, University of Wisconsin Sleep Center; External Advisory Committee,
15	Neuroscience Institute, Morehouse School of Medicine; the Boards of Trustees/Directors of the
16	National Sleep Foundation, the Association for Patient-Oriented Research, the Institute for
17 18	Experimental Psychiatry Research Foundation, the Sleep Research Society and the Sleep
10	Research Society Foundation; Circadian Rhythm Sleep Disorders Advisory Board, Takeda
20	Pharmaceutical Company, Inc.; Sleep-Wake Scientific Advisory Board, Cephalon, Inc.;
21	Scientific Advisory Board, Hypnion, Inc.; Executive Committee, Chair, Research Committee,
22	Chair, Presidential Task Force on Sleep and Public Policy, and President, Sleep Research
23	Society; President, Sleep Research Society Foundation; Chair, Scientific Advisory Panel, Mars
24	Exploration Rover Surface Operations Program, NASA Jet Propulsion Laboratory, California
25 26	Institute of Technology; Scientific Advisory Board, Air Transport Association of America; the

	Neuroscience Psychiatry Advisory Board, Pfizer, Inc; Chair, Scientific Advisory Board, Vanda
1 2	Pharmaceuticals, Inc.; Scientific Advisory Board, Zeo, Inc. (formerly Axon Labs, Inc.); Expert
3	Consultant, Committee on Sleep Medicine and Research, Institute of Medicine, National
4	Academy of Sciences; Member, External Advisory Panel, International Space Station and
5	Shuttle Utilization Reinvention Team, NASA; Member, Medical Expert Panel on Sleep Apnea
6	and Commercial Truck Driving, Federal Motor Carrier Safety Administration (FMCSA), US
7	Department of Transportation (DOT); Expert Consultant, Committee on Optimizing Graduate
8	Medical Trainee (Resident) Hours and Work Schedules to Improve Patient Safety, Institute of
9	Medicine, National Academy of Sciences; External Advisory Committee, Wisconsin Sleep
10 11	Center, University of Wisconsin; Panelist, Fatigued Driving Committee, National Highway
12	Traffic Safety Administration's (NHTSA) Office of Behavioral Safety Research; Member,
13	Expert Panel on Obstetrics, Staffing and Communication Task Group, Betsy Lehman Center for
14	Patient Safety, Boston, MA; Member, Steering Committees, Academic Alliance for Sleep
15	Research; Member and Chair, Sleep Disorders Research Advisory Board, National Heart, Lung,
16	and Blood Institute; Member, Advisory Board on Insomnia, Novartis Consumer Health; Member
17 18	and Chair, Steering Committee, Sleep Research Network, Clinical and Translational Science
18	Award (CTSA) institutions; and Member, Drowsy Driving Commission for the Commonwealth
20	of Massachusetts.
21	In December 2011, I provided a briefing to the FMCSA's Motor Carrier Safety Advisory
22	Committee (MCSAC) and the FMCSA's Medical Review Board on "Addressing Obstructive
23	Sleep Apnea in Commercial Motor Vehicle Drivers." In 2014–2015, I served as a Task Force

Sleep Apnea in Commercial Motor Vehicle Drivers." In 2014–2015, I served as a Task Force
Member of a Panel on Truck Safety, Hours of Service, and Fatigue for the National Research
Council of the National Academy of Sciences (NAS) [3]; in 2018-2019, I have served as a

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member of an Expert Panel for the National Highway Transportation Safety Administration and currently chair the National Sleep Foundation Sleep Timing and Variability Consensus Panel.

4	
3	At the Brigham and Women's Hospital, Partners HealthCare and the Harvard Medical
4	School, my research and educational programs, and those of the Divisions that I direct, have
5	been supported by a number of federal agencies, public charities and foundations, and industry.
6	These include: Agency for Healthcare Research and Quality (AHRQ); Air Force Office of
7	Scientific Research (AFOSR); Alza Corporation; Apria Healthcare; Brigham and Women's
8	Hospital; Brigham Health; Bristol Myers-Squibb Company; Beth Israel Deaconess Medical
9	Center, Boston; Centers for Disease Control (CDC); Cephalon, Inc.; City of
10 11	Philadelphia/Fraternal Order of Police (Lodge 5); Dayzz Live Well Ltd.; Defense Advanced
12	Research Projects Agency (DARPA); Department of Defense (DoD); Falck Foundation; Federal
13	Air Marshal Service (FAMS), Transportation Security Administration (TSA), Department of
14	Homeland Security (DHS); Federal Aviation Administration (FAA); Federal Emergency
15	Management Agency (FEMA), Department of Homeland Security (DHS); Harvard Medical
16	School; Harvard University; Helena Rubinstein Foundation; Jazz Pharmaceuticals, Inc.; Josiah
17 18	Macy Foundation; Koninklijke Philips Electronics, N.V.; March of Dimes Birth Defect
10	Foundation; Mary Ann & Stanley Snider via Combined Jewish Philanthropies; Merck Research
20	Laboratories; National Aeronautics and Space Administration (NASA); Harvard Catalyst;
21	Lighting Science Group; National Center for Advancing Translational Sciences (NCATS);
22	National Center for Research Resources (NCRR); National Center for Complementary and
23	Alternative Medicine (NCCAM); National Football League (NFL) Charities; National
24	Geographic Society; National Heart, Lung and Blood Institute (NHLBI); National Institute of
25 26	Child Health and Human Development (NICHD); National Institute of Environmental Health

	Sciences (NIEHS); National Institute of General Medical Sciences (NIGMS); National Institute
1 2	of Justice (NIJ), Department of Justice (DOJ); National Institute of Mental Health (NIMH);
2	National Institute of Neurological and Communicative Disorders and Stroke (NINCDS);
4	National Institute on Aging (NIA); National Institute of Occupational Health and Safety
5	(NIOSH); National Space Biomedical Research Institute (NSBRI); Office of Naval Research
6	(ONR); Optum; Partners HealthCare; Peter Bent Brigham Hospital Biomedical Research Support
7	Grant (BRSG); Peter Brown and Margaret Hamburg; Pfizer, Inc.; Regeneron Pharmaceuticals,
8 9	Inc.; ResMed Foundation; Respironics, Inc.; Rx Foundation; Sanofi Aventis, Inc.; Sanofi S.A.;
9 10	San Francisco Bar Pilots; Schering Plough Pharmaceutical Corporation; Schneider National,
11	Inc.; Sepracor, Inc.; Simmons Bedding Company; Sleep HealthCenters, Inc.; Sysco; Takeda
12	Pharmaceuticals North America, Inc.; Tempur-Pedic, Inc.; Teva Pharmaceutical Industries, Ltd.;
13	United States Olympic Committee; Vanda Pharmaceuticals, Inc.; W.K. Kellogg Foundation;
14	Wake Up Narcolepsy; and Whoop, Inc.
15	In addition to my employment by the Brigham and Women's Hospital, which is a
16	Founding Member of Mass General Brigham, I founded the Center for Design of Industrial
17 18	Schedules, which I directed for over a decade, and which consulted with businesses on the
18	application of sleep and circadian principles in the design of work schedules, pioneering the field
20	of fatigue risk management consulting.
21	I serve or have served as a consultant for: A2Z Development Center, Accelerator Corp.;
22	Actelion, Ltd.; Accreditation Council of Graduate Medical Education; Air Transport Association
23	of America; Alfresa Pharmaceutical Company; Amazon.com, Inc., American Academy of
24	Allergy and Infectious Disease; American Academy of Sleep Medicine; Association of
25 26	University Anesthesiologists; Aventis, Inc.; Avera Pharmaceuticals, Inc.; Axis Healthcare, Inc.;

	Bose Corporation; Boston Bruins; Boston Celtics; Boston Edison, Inc.; Boston Red Sox; Bristol-
1 2	Myers Squibb; Center for Design of Industrial Schedules; Cephalon, Inc.; Chevron Chemicals,
2	Inc.; Citgo, Inc.; Cleveland Browns; Columbia River Bar Pilots; CME Outfitters; Cornell
4	Medical College; M. Davis and Company; Department of Homeland Security, U.S. Fire
5	Administration; Eli Lilly and Co.; Exxon Chemical Co.; FedEx Kinko's; Federal Motor Carriers
6	Safety Administration, U.S. Department of Transportation; Fraternal Order of Police, Lodge 5;
7	Fusion Medical Education, LLC; Garda Inspectorate, Republic of Ireland; General Electric;
8	Gerson Lehman Group, Great Salt Lake Minerals and Chemical Co.; Harvard College; Harvard
9 10	Medical International; Health Science Communications, Inc.; Hypnion, Inc. (acquired by Eli
10	Lilly and Co. in April 2007); Innovative Medical Technologies, Inc.; Institute of Digital Media
12	and Child Development; Institute of Medicine; Institute of Sleep Health Promotion; Jack
13	Mattson Group; Japan National Railroad; Jet Propulsion Laboratory, California Institute of
14	Technology; Jazz Pharmaceuticals, Inc.; Koninklijke Philips Electronics, N.V.; Light Sciences,
15	Inc.; Lifetrac Systems, Inc.; Massachusetts Institute of Technology; Medical Consulting;
16	Medical Science Partners; Merck and Co.; Merck Sharpe and Dohme, Inc.; Michelin Tire Co.;
17 18	Minnesota Timberwolves; 3M; Montefiore Hospital and Medical Center; Morgan Stanley; MPM
18	Asset Management; the National Academies; National Broadcasting Company (NBC); National
20	Research Council, the National Academies; National Space Biomedical Research Institute; New
21	South Wales Clinical Excellence Commission; Neurocrine, Inc.; North Pacific Paper Co.;
22	Northeast Utilities, Inc.; Novartis Consumer Health, S.A.; Novartis, Inc.; Office of Technology
23	Assessment, U.S. Congress; Oxford Biosignals; Pfizer, Inc.; Philips Respironics, Inc.; the
24	Portland Trailblazers; Primary Research, LLC; Purdue Pharma; Quest Diagnostics, Inc.; Rohm
25 26	and Haas; Respironics, Inc.; Rockpointe, Inc.; Saatchi and Saatchi, Inc.; Samsung Electronics

_	Co., Ltd.; San Francisco Bar Pilots; Sanofi Aventis, Inc.; Sanofi Synthelabo, Inc.; Sepracor, Inc.;
1 2	ShiftWork Systems, Inc.; Sleep Multimedia, Inc.; Society for Neurological Surgeons; Somnus
2	Therapeutics, Inc.; Swiss Air; Tanabe Seiyaku Co., Ltd.; Takeda North America
4	Pharmaceuticals, Inc.; Teva, Inc.; Tokyo Electric Power Co., Inc.; Unilever, Inc.; United
5	Airlines, Inc.; U.S. Air Force; U.S. Central Intelligence Agency; U.S. National Aeronautics and
6	Space Administration (NASA); U.S. National Institutes of Health; U.S. Navy; U.S. Nuclear
7	Regulatory Commission; U.S. Olympic Committee; U.S. Secret Service; University of Virginia
8	National Science and Technology Center, National Science Foundation; Wisconsin Sleep Center,
9 10	University of Wisconsin; Valero, Inc.; Vanda Pharmaceuticals, Inc.; Vital Issues in Medicine;
10	Warburg-Pincus; Washington Board of Pilotage Commissioners; With Deep, Inc.; and Zeo, Inc.
12	(formerly Axon Labs, Inc.). Since 1985, I have also served as an expert witness/consultant on a
13	number of civil matters, criminal matters, and arbitration cases, including those involving the
14	following commercial and government entities: Advanced Power Technologies; Alvarado
15	Hospital, LLC; Amtrak; Aransas-Corpus Christi Pilots; Bombardier, Inc.; Burlington Northern
16	Railroad; Casper Sleep, Inc.; Catapult Energy Services Group, LLC; Celadon Group Inc.; CEVA
17 18	Logistics; Charlotte, North Carolina Police Department; CITGO Petroleum Corporation; City of
18	Albany, New York; C&J Energy Services; Clyde Machines, Inc.; Columbia River Bar Pilots;
20	Complete General Construction Company; Covenant Testing Technologies, LLC; Crete Carrier
21	Corporation; Covenant Testing Technologies, LLC; Dallas Police Association; Dallas Police
22	Department; Delta Airlines/Comair; EAN Holdings LLC, d/b/a Enterprise Rent-A-Car;
23	Fédération des Médecins Résidents du Québec (FMRQ); FedEx; Greyhound Lines, Inc./Motor
24	Coach Industries/Firstgroup America; Fraternal Order of Police, Lodge 5; H.G. Energy LLC;
25 26	Maricopa County, Arizona Sheriff's Office; Miami Truck Leasing, Inc.; Municipal Light and

Power, City of Anchorage, Alaska; Murrieta Valley Unified School District; Pomerado Hospital,
Palomar Health District; Philadelphia Electric; Philadelphia Police Department; Puckett EMS;
Purdue Pharma; South Carolina Central Railroad Company, LLC.; State of New Hampshire;
Steel Warehouse, Inc.; St. Louis University; Tideport Distributing, Inc.; Town of Natick,
Massachusetts; Union Pacific Railroad; United Parcel Service; Vanda Pharmaceuticals, Inc.;
Valero Energy Corporation; and the United States of America.

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Q:

What awards and recognition have you received for your work?

I received the William C. Dement Academic Achievement Award from the American 10 Academy of Sleep Medicine; the Lifetime Achievement Award from the National Sleep 11 Foundation; the Lord Adrian Gold Medal from the Royal Society of Medicine (London); the 12 Distinguished Scientist Award from the Sleep Research Society; the Senator Mark O. Hatfield 13 14 Public Policy Award from the American Academy of Sleep Medicine; the Mary A. Carskadon 15 Outstanding Educator Award from the American Academy of Sleep Medicine; the National 16 Institute for Occupational Safety and Health (NIOSH) Director's Award for Scientific 17 Leadership in Occupational Safety and Health; "Aschoff's Rule" International Award in 18 Circadian Biology; the NASA Johnson Space Center Director's Innovation Award from National 19 Aeronautics and Space Administration; a 2013 Major League Baseball World Championship 20 21 Ring from the Boston Red Sox; the 2018 Green Cross for Safety Innovation Award from the 22 National Safety Council; the Harriet Hardy Award from the New England College of 23 Occupational and Environmental Medicine; the Healthy Sleep Community Award (for research 24 done by the Harvard Work Hours, Health and Safety Group on the occupational safety impact of 25 the work hours of resident physicians) from the National Sleep Foundation; the E.H. Ahrens, Jr. 26

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	Award from the Association for Patient Oriented Research; the Gordon Wilson Lectureship		
1	Award from the American Clinical and Climatological Association; the Robert R. J. Hilker		
2 3	Award by the Central States Occupational Medical Association; the Honorary Membership and		
4	Keynote Lectureship Award from the American Academy of Dental Sleep Medicine; the Golden		
5	Mind-Body Medicine Lectureship from the University of Buffalo; the Distinguished Speakership		
6	from the Walter Reed Army Institute for Research; the Michael S. Aldrich Commemorative		
7	Lectureship from the University of Michigan; the 2018 Jürgen Walther Ludwig Aschoff,		
8	Ph.D.—Colin Stephenson Pittendrigh, Ph.D. Keynote Lectureship at the bi-annual meeting of the		
9 10	Society for Research on Biological Rhythms; the 2018 Distinguished Lectureship from the		
10	Office of Naval Research; the 2019 Bernese Sleep Award from the University of Bern in Bern,		
12	Switzerland; the 2019 Peter C. Farrell Prize in Sleep Medicine from the Division of Sleep		
13	Medicine at the Harvard Medical School; the 2019 J.E. Wallace Sterling Lifetime Achievement		
14	Award in Medicine from the Stanford Medical Alumni Association; and the 2022 Thomas Roth,		
15	Ph.D. Keynote Lectureship of Excellence on Sleep Education from the Associated Professional		
16	Sleep Societies.		
17 18	A copy of my curriculum vitae is Exhibit CAC-02.		
18			
20	II. <u>PURPOSE OF TESTIMONY</u> .		
21			
22	Q: What is the purpose of your testimony?		
23	A: My testimony addresses the following topics:		
24	1. A summary of my experience advising U.S. maritime pilot groups and other		
25	transportation sector workers on the importance of sound fatigue risk management practices;		
26			

1	2.	A review of the factors influencing alertness, cognitive performance, risk of error/accident and health of night shift workers;	
2	3.	Influence of circadian phase (biological time of day);	
3	4.	The sleep homeostat;	
4	5.	The impact of chronic sleep curtailment;	
5	6.	Drowsy driving and sleep-related motor vehicle crashes;	
6 7	7.	Interruptions of sleep and sleep inertia;	
8	8.	Night shift work induces misalignment between the phase of circadian pacemaker and the work sleep schedule;	
9	9.	Night shift work leads to sleep deprivation;	
10	10.	Worker survey data on the effects of shift work;	
11			
12	11.	Sleep disorders and crash risk;	
13	12.	A summary of the evaluation performed by my colleagues and I at the Brigham and Women's Hospital Division of Sleep and Circadian Disorders evaluating Puget	
14		Sound Pilot's scheduling efficiency and making recommendations for improvement.	
15		improvement.	
16			
17	А.	<u>Summary of Experience Advising U.S. Maritime Pilot Groups and Other</u> Transportation Sector Workers on Fatigue Risk Management Practices	
18	Q: Plea	se describe your involvement in the Columbia River Bar Pilots' 2004 rate	
19			
20	proceeding	•	
21	A: In Ju	ane 2004 I was contacted by Attorney Michael E. Haglund in preparation for the	
22	Columbia River Bar Pilots' 2004 Rate Proceedings, requesting my assistance as an expert in		
23	sleep medicine to evaluate the existing schedule of the Columbia River Bar Pilots and to		
24	recommend	a new schedule and appropriate staffing levels to manage fatigue in the Pilots'	
25	safety-cenci	tive jobs. On November 15, 2004, I provided a report to Haglund, Kelley, Horngren,	
26	5arety-501151	are jobs. On revenuer 19, 2007, i provided a report to fragiund, Keney, fioffigien,	

Jones & Wilder, LLP on the matter. On January 18, 2005, I conducted a site visit in Astoria, 1 during which time I met with the Columbia River Bar Pilots and provided them with an 2 educational session about sleep, circadian rhythms, performance and safety. That night, I crossed 3 the Columbia River Bar with Captain Roger Nelson via the Chinook pilot boat, climbed aboard 4 the Alam Mesra with Captain Nelson at midnight using a pilot rope ladder, and observed Captain 5 6 Nelson pilot the Alam Mesra across the Columbia River Bar to Astoria, Oregon. On February 28, 7 2005, I testified before Judge Allan Arlow at a rate proceeding hearing of the Board of Maritime 8 Pilots of the State of Oregon. In February 2010, I was contacted again by Attorney Haglund in 9 connection with the then-current pending rate proceeding and asked to review the current status 10 of the Columbia River Bar Pilots. I again reviewed the work schedules of the Columbia River 11 Bar Pilots and interviewed Captain Gary Lewin and various other pilots. On March 29, 2010, I 12 provided Declaration Testimony before the Board of Maritime Pilots of the State of Oregon in 13 14 the matter of the petition of the Columbia River Bar Pilots for a change in pilotage rates. 15

Q: Please describe your involvement with the shipping industry in Corpus Christi, Texas and the Aransas-Corpus Christi Pilots Association.

A: In June 2013, I was contacted by Attorney Keith Letourneau, Attorney at Law, of 19 Houston, Texas, on behalf of Valero Energy Corporation, CITGO Petroleum Corporation and 20 21 other shippers whose ships are guided to and from the harbor in Corpus Christi, Texas by 22 members of the Aransas-Corpus Christi Pilots Association. I spoke with Attorney Letourneau, 23 who asked me to review materials related to the work-rest schedules of the pilots and to render 24 an expert opinion regarding best practices related to fatigue risk management in such a round-25 the-clock transportation operation. Every year, vessels and barges from around the world 26

transport about 80 million tons of cargo on about 6,000 vessels and barges through Corpus 1 Christi Port, which is the fifth largest port in the United States as measured by tonnage. Each of 2 these arriving and departing vessels and barges is piloted through the Corpus Christi Port harbor 3 by the harbor pilots of the Aransas-Corpus Christi Pilots Association, which has indicated that 4 safety and protection of the environment are its highest priorities. I requested and received from 5 6 Ms. Lorène Rothschild, of Compton & Wendler, P.C., detailed information regarding the work 7 schedules of four Aransas-Corpus Christi pilots throughout the 2012 calendar year. I also 8 reviewed a copy of the July 29, 2013, Order of The Board of Pilot Commissioners for the Port of 9 Corpus Christi Authority Regarding Pilotage Rules, Regulations and Rates and correspondence 10 regarding Aransas-Corpus Christi Pilots Fatigue Guidelines. Based on my analysis of these 11 materials, on October 6, 2013, I developed and provided a set of recommendations for the 12 Aransas-Corpus Christi Pilots designed to reduce the risk of fatigue-related errors and accidents. 13 14 15 **O**: Please discuss your relationship with the San Francisco Bar Pilots and the study 16

that you formed for them in 2013.

On or about August 16, 2013, I was contacted by Captain Joe Long of the San Francisco 18 Bar Pilots, who explained that the Board of Pilot Commissioners governing the San Franciso Bar 19 Pilots had been charged by the California legislature to contract with an independent entity to 20 21 conduct a study of the effects of work and rest periods on psychological ability and safety for 22 pilots. The legislature specified that the study "shall evaluate sleep- and human-related factors 23 for pilots, and shall include information and recommendations on how to prevent pilot fatigue 24 and ensure the safe operation of vessels." Cal. Harb. & Nav. Code § 1196.5(a). In preparation for 25 that initiative, the San Francisco Bar Pilots requested that I initiate a study, which I did with Dr. 26

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	Laura Barger, to assess the impact of the hours worked by the San Francisco Bar Pilots and mak	
1 2	recommendations on how best to mitigate the impact of fatigue on the health of their pilots and	
3	safety of their operations. In order to gather information about the history and operations of the	
4	San Francisco Bar Pilots, we reviewed a Manalytics study that had been conducted in 1986 and	
5	then interviewed almost half of the San Francisco Bar Pilots to explore their preferences and	
6	perspectives. We also reviewed the current work rules governing San Francisco Bar Pilots'	
7	schedules. To analyze the San Francisco Bar Pilots' schedules for factors that affect	
8	physiological measures of alertness and performance, scheduling data from a full year was	
9 10	reviewed. On May 7, 2014, I accompanied Captain Joe Long on an operational night shift during	
10	which we departed Pier 9 at night by maritime pilot boat and were transferred to the offshore	
12	station boat, 11 miles out at sea. Thereafter, Captain Long and I used a pilot rope ladder to climb	
13	aboard the container vessel Oakland Express during the night, which Captain Long then piloted	
14	across the San Francisco Bay and docked shortly after dawn at the Port of Oakland. That	
15	experience confirmed what I had learned from working with the Columbia River Bar Pilots about	
16	the intense and demanding nature of the work that maritime pilots perform, and the challenging	
17 18	conditions under which they must carry out their safety-sensitive operations. The importance of	
10	instituting a comprehensive fatigue management program for the San Francisco Bar Pilots was	
20	reaffirmed. This evaluation of the work hours of the San Francisco Bar Pilots enabled Dr. Laura	
21	Barger and I to develop and provide recommendations for a Fatigue Risk Management Program	
22	there. We also conducted an education session for the San Francisco Bar Pilots. On October 24,	
23	2014, Dr. Laura Barger and I provided a report to the San Francisco Bar Pilots on the results of	
24	our evaluation and recommendations to improve the health and safety of the San Francisco Bar	
25 26	Pilots. On April 7, 2016, I made an invited presentation on Pilotage, Sleep and Circadian	

Rhythms to the 2016 West Coast Pilots' Conference held at the Barbey Maritime Center/Columbia Maritime Museum in Astoria, Oregon, at which both the Columbia River Bar Pilots and the San Francisco Bar Pilots were represented, along with many other pilot groups. At that time, I also made a separate presentation on fatigue risk management training to the Columbia River Bar Pilots.

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⁷ Q: Please describe the work that you performed for the Washington State Board of ⁸ Pilotage Commissioners in 2017.

A: On June 8, 2017, Sheri J. Tonn, Chair of the Washington State Board of Pilotage 10 Commissioners contacted me, indicating that the Board wanted to gather information about 11 current best practices in sleep science and wanted an overview of best practices with other 12 pilotage programs around the US and abroad, an assessment of how the Puget Sound Pilots 13 14 fatigue management program is functioning, an evaluation of the importance of measuring and 15 accounting for "time on task" vs "bridge time" and recommendations for any improvements that 16 could be made to the existing program. The Board had invited the then-president of Puget Sound 17 Pilots to give a presentation at the July 13, 2017, meeting of the Board. Board Chair Tonn 18 indicated that the Board was seeking the assistance of a sleep professional because information 19 about fatigue management would be an important factor in determining the number of licensed 20 21 pilots. Chair Tonn indicated that the Washington State Legislature was reviewing the Pilotage 22 Commission for best practices, and the Board planned to have a final report for the legislature for 23 the 2018 legislative session that began in January 2018. Therefore, I met in Seattle with 24 Chairperson Tonn and Interim Executive Director of the Board Jaimie Bever regarding the issue 25 of fatigue and safety on September 21, 2017. Following the meeting, I was contracted by the 26

Washington State Board of Pilotage Commissioners to make a Fatigue Management presentation 1 to the Board at the regular public meeting on December 14, 2017 in which I (a) outlined the 2 science of and current best practices for fatigue management; (b) explained how I view the 3 Board's then-current rest rules (RCW 88.16.103 and WAC 363-116-081); (c) explained how I 4 view the pilot's rest rules (including assignment time vs. bridge time); (d) compared BPC/PSP 5 6 rest rules; and I recommended any improvements. In my December 14, 2017, presentation to the 7 Washington State Board of Pilotage Commissioners, I critiqued the Agency's then-current Rest 8 Rules, which excluded travel time in hours of service, potentially allowing unsafe, extended 9 duration work shifts. I also critiqued the then-current Agency rules, which did not limit duration 10 of work shifts and therefore allowed unsafe, extended duration work shifts. As written at that 11 time, a 6.9-hour pilotage assignment could be followed by an assignment of 22 or more hours, 12 resulting in a 29-hour work shift, including travel time. The then-current Agency rules 13 14 promulgated by the Washington State Board of Pilotage Commissioners provided inadequate 15 time for rest between work shifts, potentially creating an unsafe condition. I indicated that seven 16 hours of off-duty time is inadequate for maritime pilots to fulfill their daily sleep need, inducing 17 sleep deficiency that could cause fatigue. The then-current Agency rules promulgated by the 18 Washington State Board of Pilotage Commissioners failed to ensure that pilots were provided 19 with 34 consecutive hours of uninterrupted rest, including two nights between midnight and 6 20 21 am, within every running 7-day interval. Fortunately, the Puget Sound Pilots had promulgated 22 rules that addressed some of the deficiencies in the Agency rules. However, as I pointed out at 23 the time, voluntary inclusion of travel time in work hours restrictions by the Puget Sound Pilots, 24 for example, was not sufficient to substitute for regulatory action by the Washington State Board 25 of Pilotage Commissioners. After reviewing the existing regulations promulgated by the 26

Washington State Board of Pilotage Commissioners, I recommended a series of 10

improvements to the existing program, as requested by the Board. The complete slides from my 2

presentation to the Washington State Board of Pilotage Commissioners, which include a

4 summary of each of those ten recommendations, are included as Exhibit CAC-03.

5

1

6	Q: Have You Worked Directly with the Puget Sound Pilots Before?		
7	A: Yes. On March 12, 2021, Captain Ivan Carlson, President of the Puget Sound Pilots,		
8	contacted Dr. Laura Barger requesting consultation regarding the recommendation by the Utility		
9 10	and Transportation Commission for Puget Sound Pilots to improve their efficiency without		
10	sacrificing safety or prudent fatigue mitigation efforts. Subsequent discussions led the Puget		
12	Sound Pilots to commission the Sleep Matters Initiative in the Division of Sleep and Circadian		
13	Disorders to conduct a quality improvement program through the Brigham and Women's		
14	Physicians Organization in which Dr. Barger and I evaluated the Puget Sound Pilots' scheduling		
15	efficiency and made recommendations for improvement, with a final report delivered on June 1,		
16	2022.		
17			
18 19	B. <u>A Review of the Factors Influencing Alertness, Cognitive Performance, Risk</u> of Error/Accident and Health of Night Shift Workers		
20	Q: The Puget Sound Pilots regularly work night shifts yet perform a high-risk service		
21	that requires constant alertness and mental fortitude. In light of that, what factors		
22	influence night shift workers' ability to sustain effective waking neurocognitive		
23	performance?		
24			
25	A: Multiple factors influence the ability to sustain effective waking neurocognitive		
26	performance in healthy individuals not taking medications. These include biological time of day		

	(i.e., circadian phase); the length of prior wakefulness; nightly sleep duration; and the recency of			
1 2	the last sleep episode (sleep inertia). While the effects of these circadian and homeostatic sleep			
2	regulatory processes can be modified by environmental conditions, physical activity and			
4	pharmacological agents (i.e., caffeine and/or nicotine during night wakefulness or hypnotics			
5	during day sleep), they cannot consistently overcome the impact of adverse circadian phase			
6	and/or sleep deprivation on performance. The effect of these four factors (misalignment of			
7	circadian phase and work/sleep schedule, cumulative sleep deprivation, lengthy prior			
8	wakefulness and/or recent awakening) can create an imposing biological force that can			
9	overpower a worker's ability to remain awake and attentive while at work and during the			
10 11	commute to and from work [2, 4-18]. This can lead to impaired neurocognitive performance			
12				
13				
14	to sustain attention for a continuous duration of time (e.g., for 10-20 minutes or more) or when			
15	performing a routine, highly overlearned task, such as driving, for 10–20 minutes or more.			
16				
17 18	Q: What are some possible effects that these factors have on worker performance?			
10				
20	A: Sleep-related performance decrements can lead to impaired job performance and higher			
20 21	rates of errors, accidents and injuries. In fact, sleep deficiency and sleep disorders cost U.S.			
21	businesses an estimated \$411 billion per year in absenteeism, workplace accidents, lost			
23				
24	awareness and impaired performance, is estimated to be a possible contributing factor in nearly			
25	30% of railway collisions [21, 22]. The National Transportation Safety Board, which is charged			
26	with conducting root-cause investigations on major transportation accidents in the United States, TESTIMONY OF CHARLES A. CZEISLER Exh. CAC-1Tr Page 18			

	has identified fatigue as a probable cause, a contributing factor or a finding in 17% of its railroa			
1 2	crash investigations, 23% of its major aviation investigations and nearly 40% of its highway			
2	crash investigations [23]. Moreover, sleep deficiency and circadian disruption have a major			
4	impact on safety in the general population. Every year, motor vehicle crashes worldwide cause			
5	20-50 million injuries and cost more than a half-trillion dollars. Road Traffic Injuries, World			
6	Health Organization (June 20, 2022), https://www.who.int/news-room/fact-sheets/detail/road-			
7	traffic-injuries. At least twenty percent of crashes, serious injuries and deaths occurring in road			
8	crashes are due to driver sleepiness [1, 24, 25]. In fact, when 9 hours of time in bed per night is			
9	used as the referent, data from a recent prospective cohort study reveals that up to forty percent			
10 11	of motor vehicle crashes in the general population are attributable to sleep deficiency and sleep			
11	disorders [25].			
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15				
13				
	Q: What effect do sleep disorders have on sleep deficiency and other health risks?			
14	Q: What effect do sleep disorders have on sleep deficiency and other health risks?A: The prevalence of sleep disorders, which are a major cause of sleep deficiency, has			
14 15				
14 15 16	A: The prevalence of sleep disorders, which are a major cause of sleep deficiency, has			
14 15 16 17	A: The prevalence of sleep disorders, which are a major cause of sleep deficiency, has increased substantially in recent years. In addition to impairment of neurocognitive performance,			
14 15 16 17 18	A: The prevalence of sleep disorders, which are a major cause of sleep deficiency, has increased substantially in recent years. In addition to impairment of neurocognitive performance, chronic sleep deficiency and recurrent circadian disruption have adverse health effects. The			
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 14 15 16 17 18 19 20 21 22 23 24 	A: The prevalence of sleep disorders, which are a major cause of sleep deficiency, has increased substantially in recent years. In addition to impairment of neurocognitive performance, chronic sleep deficiency and recurrent circadian disruption have adverse health effects. The Institute of Medicine concluded that "50 to 70 million Americans chronically suffer from disorders of sleep and wakefulness, hindering daily functioning and adversely affecting health and longevity" [1]. The Institute of Medicine further concluded that "the cumulative long-term effects of sleep loss and sleep disorders have been associated with a wide range of deleterious health consequences, including an increased risk of hypertension, diabetes, obesity, depression,			
 14 15 16 17 18 19 20 21 22 23 	A: The prevalence of sleep disorders, which are a major cause of sleep deficiency, has increased substantially in recent years. In addition to impairment of neurocognitive performance, chronic sleep deficiency and recurrent circadian disruption have adverse health effects. The Institute of Medicine concluded that "50 to 70 million Americans chronically suffer from disorders of sleep and wakefulness, hindering daily functioning and adversely affecting health and longevity" [1]. The Institute of Medicine further concluded that "the cumulative long-term effects of sleep loss and sleep disorders have been associated with a wide range of deleterious			

Research on Cancer has classified night shift work as being "probably carcinogenic to humans" [27].

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C. Influence of Circadian Phase (Biological Time of Day)

5 Q: What are circadian rhythms?

6 A: Circadian rhythms, i.e., biological rhythms oscillating with an approximate period of 7 twenty-four hours (from the Latin words: *circa* [about] and *dies* [day]), are present at all levels of 8 biological complexity from unicellular organisms to humans. Circadian rhythms are endogenous 9 (i.e., internally generated), self-sustaining oscillations that persist in the absence of periodic 10 external time cues. In humans, many physiological processes, including the body temperature 11 cycle, endocrine patterns, renal and cardiac function, subjective alertness, sleep-wake behavior, 12 performance and the risk of sleep-related attentional failures and involuntary micro-sleep 13 14 episodes vary according to the time of day [2, 28-46]. 15 16 **Q**: Generally, how does the circadian rhythm relate to variations in alertness and 17 performance? 18 19 A: The circadian contribution to variations in alertness and performance is generated by a 20 light-sensitive circadian pacemaker that also drives the circadian rhythms of core body 21 temperature, plasma cortisol and plasma melatonin [44, 47-61]. The endogenous circadian clock 22 is a major determinant of the timing and internal architecture of sleep in humans [37, 42, 44, 60-23 65]. Experimental studies have demonstrated that spontaneous sleep duration, sleepiness, REM 24 25 sleep propensity, and both the ability and the tendency to sleep vary markedly with circadian 26

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Exh. CAC-1Tr Page 20 phase or biological time of day and interact with a homeostatic process to regulate sleep propensity and daytime alertness and neurocognitive performance [33, 37, 42, 44, 62, 66-72].

3

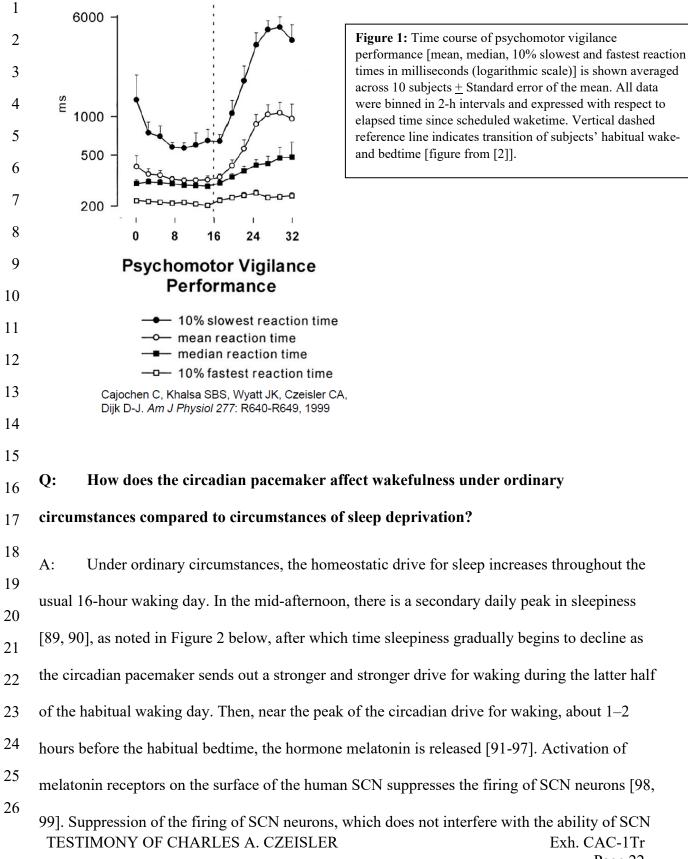
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4 Q: What is the circadian pacemaker, and how does it affect alertness and cognitive 5 performance?

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Nearly fifty years ago the central neuroanatomic structures responsible for both the 7 generation of endogenous circadian rhythms and their synchronization with the 24-hour day were 8 9 identified [73, 74]. Deep within the brain, two bilaterally paired clusters of hypothalamic neurons 10 comprising the suprachiasmatic nuclei (SCN) act as the central neural pacemaker for the 11 generation and/or synchronization of circadian rhythms [44, 75-85]. This endogenous circadian 12 pacemaker is a major determinant of daily variations in subjective alertness and cognitive 13 performance [2, 15, 30, 42, 44, 47, 70, 86, 87]. These and other studies have shown that, as 14 illustrated in Figure 1 below, there is a prominent circadian variation in objective and subjective 15 measures of alertness, performance (psychomotor, vigilance, memory) and attention or ability to 16 17 concentrate, with a primary peak in the daily rhythm of sleepiness in the latter half of the usual 18 sleep time, just before our usual wake time. Similarly, the peak drive for waking, emanating 19 from the hypothalamic circadian pacemaker, occurs a couple of hours before our habitual 20 bedtime [2, 33, 88]. This paradoxical relationship between the output of the circadian pacemaker 21 and the timing of the sleep-wake cycle is thought to facilitate consolidation of sleep and 22 wakefulness in humans [33, 42, 68]. 23 24

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Page 22

to oscillate with a near-24-hour period [100], is thought to quiet the wake-promoting signal 1 emanating from the SCN. Melatonin thereby suppresses the wake-promoting signal emanating 2 from the SCN, allowing us to fall asleep at our habitual hour without stopping the circadian 3 oscillation. Similarly, the SCN sends out a strong drive promoting sleep in the latter half of the 4 night, which helps to consolidate sleep once the pressure for deep slow-wave sleep is satiated in 5 6 the first half of the night [33, 42]. The latter half of the night is richest in REM sleep, which has a 7 very prominent circadian rhythm in its propensity [42, 67, 68, 101]. 8 In the absence of sleep, in the latter half of the night near the habitual wake time, elevated 9 homeostatic drive for sleep interacts with the circadian peak of sleep propensity to create a 10 critical zone of vulnerability [2, 42]. This is a time of increased risk for both motor vehicle 11 crashes [89, 102-106] and train crashes [107] (Figure 2). The magnitude of these circadian 12 variations in sleep propensity is greatly amplified when the homeostatic drive for sleep is 13 14 elevated [13, 108, 109]. 15

Time of Day

Peak drowsy driving danger times:

 5 am – 8 am (primary nighttime peak)
 3 pm – 5 pm (secondary mid-day peak)

 Size of the Time of Day peaks are INCREASED by sleep deficiency
 Size of the Time of Day peaks are DECREASED when sleep is sufficient

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Figure 2. Impact of time of day on risk of fatigue-related crashes [22]. TESTIMONY OF CHARLES A. CZEISLER

Q: What is the period of a typical circadian rhythm, and what environmental factors influence the circadian pacemaker?

A: In the absence of periodic time cues, the period of the human circadian pacemaker is slightly longer than 24 hours [94, 110-112]. In order for the biological clock to coordinate its function with the timing of events in the external world that operates on a 24-hour schedule, daily information from the environment must therefore reach the circadian pacemaker. The circadian pacemaker is essentially reset by a small amount each day by this external stimulus in order to maintain synchrony with the 24-hour day [44, 49, 60, 110].

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Light is the principal environmental synchronizer of the mammalian SCN [44]. The SCN 10 11 is connected to the retina via a monosynaptic pathway called the retinohypothalamic tract (RHT), 12 the presumed conduit by which information from the external light dark cycle reaches the 13 circadian clock. Research conducted in my laboratory at the Brigham and Women's 14 Hospital/Harvard Medical School has demonstrated that properly timed exposure to bright light 15 and darkness can rapidly shift the phase of the endogenous circadian pacemaker in humans [31, 16 44, 47-59, 113, 114]. Both the magnitude and direction of the phase shifts induced by light are 17 dependent on the timing, duration, intensity and wavelength of the light exposure [44, 48-50, 55-18 19 58, 114-122], together with the prior light exposure history [44, 61, 123]. Exposure to blue-20 enriched light from a light-emitting eReader can suppress melatonin, shift circadian phase and 21 affect sleep architecture [124, 125]. On average, light exposure occurring during the first half of 22 the biological night resets the circadian clock to a later hour; light received in the last quarter of 23 the biological night resets the circadian clock to an earlier hour [44, 49, 61, 114]. Thus, the 24 circadian pacemaker of an individual living on a conventional schedule of day work and night 25 26

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Exh. CAC-1Tr Page 24 sleep is synchronized by the naturally occurring light dark cycle to oscillate at the same period as the Earth's solar day, i.e., 24 hours.

23

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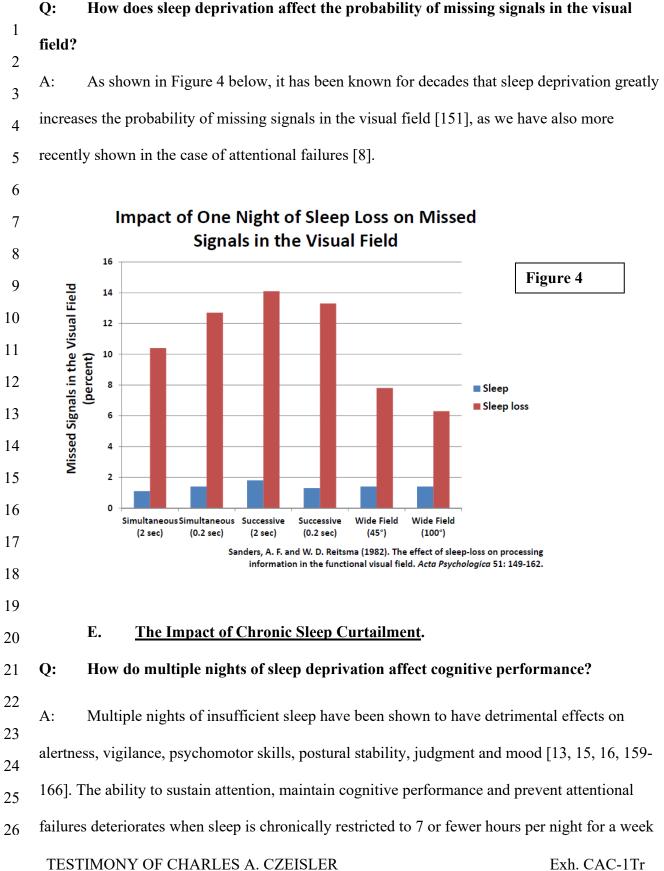
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D. The Sleep Homeostat

Q: What is the range of senses, traits, and behaviors that sleep deprivation can affect, and how does that compare to alcohol's effects?

- A: Without sleep, alertness and neurocognitive performance exhibit a steady deterioration 7 attributable to sleep loss, onto which a rhythmic circadian variation is superimposed [2, 12, 30, 8 9 33, 38, 42, 44, 60, 62, 68, 70, 71, 86, 87, 108, 126-132]. During sustained wakefulness, 24 hours 10 of sleep deprivation has been shown to greatly impair neurobehavioral reaction time performance 11 [2, 133, 134], to an extent that is comparable to a level of 0.10 percent blood alcohol 12 concentration [135-140]. The sedative effects of alcohol can persist for more than six hours after 13 alcohol ingestion, even after blood alcohol concentration is no longer elevated [141]. As noted in 14 Figure 3 below, acute sleep deprivation impairs judgment [142-144]; decision making [145]; 15 cognitive performance [15, 146-148]; memory [149]; reaction time [2, 14, 128, 150]; visual-16 17 perceptual ability [9, 41, 150-154]; sensorimotor gating [155]; distractibility [150, 156] and 18 ability to focus attention [149, 150], and increases the instability of waking neurobehavioral 19 functions [146, 147, 150] and the probability of eyelid closure and the risk of loss of situational 20 awareness, even when the eyes remain open [150]. 21 22 23 24 25
- 26

1	Sleep deficiency		
2	– Slows reaction time		
3	 Raises risk: attentional failures & sleep attacks 		
4	 Impairs judgment, increases risk taking 		
5	 Makes people more distractible, fast & sloppy 		
6	 Hinders perception of objects in the visual field 		
7	 Increases eyelid closure; risk of falling asleep 		
8	- Degrades cognitive performance		
9	 Interferes with memory formation 		
10	 Induces loss of situational awareness, even when eyes are open 		
11	- Slows thinking; can induce automatic behavior		
12	clove inning, can induce automatic senarior		
13	Figure 3. Sleep deficiency, which can be the result of many factors, impairs neurobehavioral performa	nce [22].	
14	Q: How does sleep deprivation affect the likelihood of making mistakes and t	aking	
15	risks?		
16	A: Paradoxically, instead of slowing response times to preserve accuracy, researc	h from my	
17	laboratory and from others has demonstrated that sleep-deprived participants increase	speed at	
18			
19	the expense of making more mistakes (i.e., becoming "fast and sloppy"), taking greate	_	
20	157, 158], and making hasty decisions based on inadequate information [41]. This res	ults in	
21	increased rates of errors on selective attention tasks that require a search for targets in	the visual	
22	field [41]. Commonly used stimulants, such as caffeine, modafinil and amphetamines, all fail to		
23	reverse the impairment of decision making and increased risk-taking behavior induced by sleep		
24	deficiency, whereas both are restored once recovery sleep is taken [158].		
25			
26			



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or longer [1, 16, 163]. This effect is even seen in the absence of extended wakefulness [167].
Subjective measures of stress, tiredness, sleepiness, irritability, hostility, distractibility and the
frequency of complaints increase, while sociability and optimism decrease in the face of chronic
sleep loss, which significantly increases psychometric measures of total mood disturbance and
impairs judgment [143, 145, 156, 168, 169]. Objective measures of performance, including
reaction time and memory, worsen with each successive day that insufficient sleep is obtained
[16, 163] (see Figure 6 below).

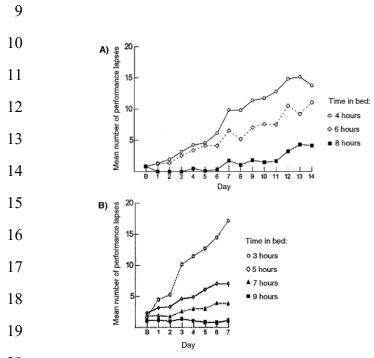


Figure 6. Repeated nights of sleep loss result in cumulative cognitive impairment. Higher number of performance lapses indicate poorer performance and more unstable alertness. Note: B, baseline day.

Figure and legend reprinted from *Sleep Deprivation and Sleep Disorders: An Unmet Public Health Problem*, a report of the Institute of Medicine Committee on Sleep Medicine and Research Sleep Disorders (2006) [1].



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Q:

How does increasing chronic sleep deprivation affect the frequency of symptoms,

22 and how long do those take to correct?

A: The effects of sleep deprivation are not overcome with a single night of sleep but can
 carry over several days. Increasing chronic sleep deprivation leads to an increased probability of
 experiencing lapses of attention and falling asleep involuntarily in inappropriate or dangerous

situations. In a condition of chronic sleep deprivation, even if work were scheduled during an appropriate circadian phase, the probability of falling asleep while working is markedly increased.

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Q: Can chronic sleep curtailment progressively deteriorate performance with additional weeks of insufficient sleep?

A: Yes. Recent evidence suggests that even after a single night of 8 to 12 hours of recovery 8 9 sleep, individuals with a history of recent exposure to chronic sleep loss may be more vulnerable 10 to the effects of re-exposure to sleep restriction, exhibiting nearly double the deterioration in 11 performance on a vigilance task upon acute sleep restriction to 4 hours of time in bed as 12 compared with their response to the same challenge when their recovery sleep was not preceded 13 by exposure to chronic sleep debt [170]. Thus, work schedules that induce chronic sleep 14 curtailment may generate a deterioration of performance that becomes progressively greater with 15 additional weeks of insufficient sleep. Despite intermittent opportunities for recovery sleep, 16 17 individuals exposed to such schedules become increasingly vulnerable to the adverse effects of 18 sleep loss on performance. We recently demonstrated in my laboratory that when individuals are 19 chronically exposed to a schedule that affords inadequate time for sleep (i.e., 5.6 hours available 20 for sleep each day), then the adverse effects of misalignment of circadian phase and acute sleep 21 loss are an order of magnitude greater, even after participants were provided a 10-hour sleep 22 opportunity [13]. We found that even a 10-hour sleep opportunity in bed in a dark and quiet 23 room is insufficient to reverse the increased vulnerability to sleep loss associated with a week of 24 25 chronic sleep restriction to 5.6 hours of time-in-bed per 24 hours [13]. Moreover, we have found 26

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1

4 **O**: How exactly does increased and chronic sleep deprivation affect performance and 5 the probability of performance failure?

6

Increasing sleep deprivation leads to cognitive slowing, an increased probability of A: 7 experiencing lapses of attention, episodes of automatic behavior [171, 172] and/or falling asleep 8 9 while attempting to remain awake at work or on the commute to or from work. In a condition of 10 chronic sleep deprivation, even if work were scheduled during an appropriate circadian phase, 11 the probability of a sleep-related attentional failure or neurobehavioral performance failure while 12 awake is markedly increased [15, 16, 163]. In fact, six hours of time in bed per night for a week 13 or two brings the average young adult to the same level of neurobehavioral impairment as 24 14 hours of wakefulness, whereas 4 hours of time in bed per night induces that same level of 15 impairment in four to six days and, within a week or two, induces a level of impairment 16 17 comparable with 48 hours of wakefulness (i.e., two consecutive days and nights without sleep). 18 19 20

21

O: Can you give examples of some other problems associated with sleep deprivation?

A: Young adults are particularly vulnerable to the adverse impact of acute and chronic sleep 22 deficiency on neurobehavioral performance and fatigue-related motor vehicle crash risk [106, 23 173, 174]. Metabolic studies have demonstrated that such sleep curtailment also has adverse 24 effects on the metabolic and immune systems, leading to alterations in glucose metabolism and 25 26 increasing the probability of weight gain [175-180]. As with alcohol intoxication, chronically **TESTIMONY OF CHARLES A. CZEISLER** Exh. CAC-1Tr Page 30

sleep-deprived individuals tend to underestimate the extent to which their performance is impaired, failing to judge accurately the impact of sleep deficiency on their functioning and alertness, despite increasing impairment evident in objective recordings of the rate of lapses of attention (see Figure 7 below from [16]).

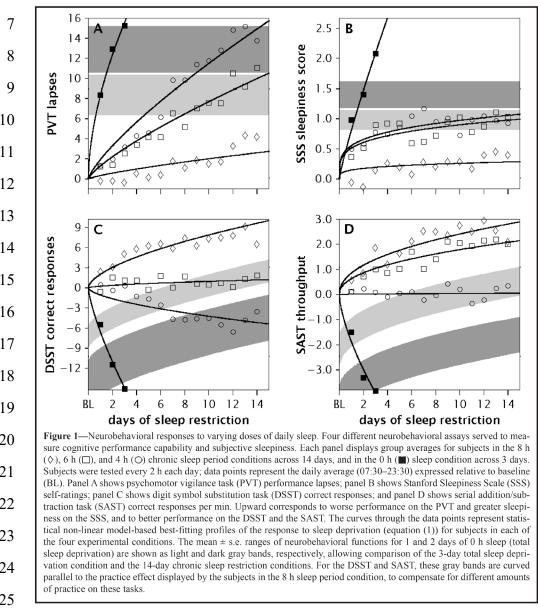


Figure 7.

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F.

Sleep-wake Transitions, Sleep Attacks and Automatic Behavior

Q: Are individuals able to force themselves to stay awake when struggling with elevated 3 sleep pressure?

A: Importantly, individuals struggling to stay awake in the face of elevated sleep pressure— 5 whether due to chronic sleep restriction or circadian misalignment-are not always able to do so. 6 Sleep deprivation greatly increases the risk that an individual will succumb to the increased sleep 7 pressure, and that the brain's "sleep switch" located in the ventrolateral preoptic (VLPO) area of 8 9 the hypothalamus [64] will initiate an involuntary transition from wakefulness to sleep. Due to 10 the influence of the circadian timing system, such an involuntary transition from wakefulness to 11 sleep is most probable in the latter half of the night near the habitual wake time, which coincides 12 with the morning commute, and in the mid-afternoon.

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Q: Can you give one common example of where the consequences of sleep deprivation are especially evident?

17

These classic consequences of chronic partial sleep deprivation are particularly evident A: 18 while doing a routine task like driving. In a condition of chronic sleep deprivation, even when 19 20 work (or wakefulness) is scheduled during an appropriate circadian phase, the probability of a 21 sleep-related neurobehavioral performance failure while working and/or driving is markedly 22 increased. Of course, once an individual has lost the struggle to stay awake and makes the 23 transition from wakefulness to sleep-however briefly-driving performance is much worse 24 than that of a drunk driver, as the individual is unresponsive to the environment throughout the 25 duration of the micro-sleep episode or the sleep attack. Moreover, when work (or wakefulness) is 26

scheduled during an inappropriate circadian phase, the probability of a sleep-related neurobehavioral performance failure while commuting home from work after the overnight shift is also markedly increased.

4 In addition, sometimes drowsy individuals linger in an intermediate state between sleep 5 and wakefulness. The operator of a motor vehicle in this sleep-related condition, which probably 6 represents a transitional state in which part of the brain is locally asleep while part of the brain 7 remains awake, may maintain full pressure on the accelerator pedal and proceed for a 8 considerable distance, even negotiating gradual turns and exhibiting goal-directed behavior, but 9 fail to heed stop signals or respond appropriately to traffic conditions in a timely manner. This 10 11 intermediate state, which has been termed automatic behavior syndrome, is characterized by 12 retention of the ability to maintain pressure on the accelerator, to turn the steering wheel and to 13 carry out rudimentary tasks and to provide semi-automatic responses to stimuli without 14 appropriate cortical integration, often resulting in a complete loss of situational awareness and 15 judgment [171, 181, 182]. In the case of a motor vehicle driver, this could involve driving toward 16 the flashing hazard lights of disabled vehicles in a highway breakdown lane rather than steering 17 clear of those vehicles. 18

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Q: Can you give another example where automatic behavior syndrome has been documented?

A: Another example of automatic behavior syndrome has been repeatedly documented in
 trains equipped with alerter devices that are designed to stop the train if an engineer were to fall
 asleep or otherwise be incapacitated. Unfortunately, as Oman and Liu reported in 2007,

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	"automatic resetting behavior among engineers ultimately defeats the purpose [of the
1 2	alerter]." [Oman CM, Liu AM. Locomotive In-Cab Alerter Technology Assessment. In
3	Development of Alternative Locomotive In-Cab Alerter Technology: Final Technical Report
4	DOT Volpe PR#79-3389, DTS-79, Volpe National Transportation Systems Center, Cambridge,
5	MA, 30 November, 2006]. One stakeholder in the rail industry provided an apt description of the
6	impact of fatigue on the effectiveness of the alerter devices: "Engineers can become inattentive
7	and keep hitting the alerter even when nodding off. 'Alerter naps' has become a commonly used
8 9	term." Oman and Liu report that: "Accident investigations show that fatigued engineers are able
10	continue their automatic resetting behavior even while in a very low level of physiological
11	arousal, including brief episodes of micro-sleep, thus defeating the original purpose of the
12	alerter." And that "The NTSB has noted that current alerter designs foster the development of
	automatic pre-emptive resetting behavior by drowsy engineers"
13	automatic pre-emptive resetting behavior by drowsy engineers
14	automatic pre-emptive resetting behavior by drowsy engineers
14 15	G. <u>Drowsy Driving and Sleep-related Motor Vehicle Crashes</u>
14 15 16	G. <u>Drowsy Driving and Sleep-related Motor Vehicle Crashes</u>
14 15	
14 15 16 17	G. <u>Drowsy Driving and Sleep-related Motor Vehicle Crashes</u>
14 15 16 17 18	 G. <u>Drowsy Driving and Sleep-related Motor Vehicle Crashes</u> Q: How frequently do Americans drive drowsy and fall asleep behind the wheel?
14 15 16 17 18 19	 G. <u>Drowsy Driving and Sleep-related Motor Vehicle Crashes</u> Q: How frequently do Americans drive drowsy and fall asleep behind the wheel? A: National survey data reveal that 56 million Americans drive while drowsy each month,
14 15 16 17 18 19 20	 G. <u>Drowsy Driving and Sleep-related Motor Vehicle Crashes</u> Q: How frequently do Americans drive drowsy and fall asleep behind the wheel? A: National survey data reveal that 56 million Americans drive while drowsy each month, and an estimated 8 million Americans fall asleep at the wheel each month—with half of those
 14 15 16 17 18 19 20 21 22 23 	 G. <u>Drowsy Driving and Sleep-related Motor Vehicle Crashes</u> Q: How frequently do Americans drive drowsy and fall asleep behind the wheel? A: National survey data reveal that 56 million Americans drive while drowsy each month, and an estimated 8 million Americans fall asleep at the wheel each month—with half of those who do so drifting into another lane or onto the rumble strip and 10 percent running off the road
 14 15 16 17 18 19 20 21 22 	 G. <u>Drowsy Driving and Sleep-related Motor Vehicle Crashes</u> Q: How frequently do Americans drive drowsy and fall asleep behind the wheel? A: National survey data reveal that 56 million Americans drive while drowsy each month, and an estimated 8 million Americans fall asleep at the wheel each month—with half of those who do so drifting into another lane or onto the rumble strip and 10 percent running off the road [183]. Remarkably, many drivers continue to operate their vehicles in a nether world between

a video recording, when the husband of a motorist who was nearly driven off the highway by a drowsy driver called 911 to report the drowsy driver and then videotaped the drowsy driver as she drove for 30 minutes on U.S. Interstate 25 in Denver. In that 30-minute video, the driver can be seen with her head leaning back against the headrest, as shown in Figure 8.



Figure 8. An admittedly drowsy driver, Ms. Karyn Steinert, is shown leaning against the headrest while driving on I-25 in Denver at speeds of up to 70 mph. She reportedly caught herself nodding off at the wheel frequently during this drive and told the Colorado State Trooper who ticketed her for investigation of reckless driving that she only had 2 hours of sleep the night beforehand. The incident was documented on the morning of October 15, 2007 by Christian Pruett and his wife, who was driving on Interstate 25 in Denver, when they both noticed a white sport utility vehicle next to them drifting from lane to lane, and nearly sideswiping their car. Mr. Pruett grabbed their video camera and photographed the sleeping woman with her head back as her vehicle moved along I-25.

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12 Figure 9. Screen capture from a video of a white sport utility vehicle (right) driven by an admitted drowsy driver, Ms. Karyn Steinert. In this image, the white SUV driven by Ms. Steinert is shown drifting across the white line on 13 the left side of the middle lane in which she is driving, forcing the dark-colored pickup truck in the left passing lane over the fog line and the rumble strip and onto the shoulder of the interstate highway. Ms. Steinert admitted that she 14 had caught herself nodding off at the wheel frequently during this 30-minute videotaped drive, in which she similarly caused a tanker truck to go onto the shoulder. The incident was documented on the morning of October 15, 15 2007 by Christian Pruett and his wife, who was driving on Interstate 25 in Denver, when they both noticed a white sport utility vehicle next to them drifting from lane to lane, and nearly sideswiping their car. After calling the police, 16 Mr. Pruett videotaped and photographed the sleeping woman with her head back as her vehicle moved along I-25. 17 18 **O**: What percentage of automobile accidents are caused by sleep deficiency and sleep

- ¹⁹ disorders?
- 20

A: Sleep deprivation degrades reaction time, impairs judgment, memory and vigilance,

reduces attention span, increases distractibility, and raises the risk of attentional failures,

23 automatic behavior, falling asleep at the wheel and motor vehicle crashes, with sleep deficiency

²⁴ and sleep disorders accounting for a population-attributable risk of at least 20 percent of all

²⁵ motor vehicle crashes [25]. This estimate is consistent with the conclusion of the Institute of

26 Medicine and the AAA Foundation for Traffic Safety that sleep deficiency and sleep disorders TESTIMONY OF CHARLES A. CZEISLER Exh. CAC-1Tr Page 36

account for 20 percent of serious crash injuries and deaths, and with the National Center for
 Statistics and Analysis that "Drowsy/Asleep/Inattentive" drivers account for 20 percent of fatal
 two-vehicle truck crashes [1, 22, 24, 156, 184, 185]. Moonesinghe, et al., *An Analysis of Fatal Large Truck Crashes*, Mathematical Analysis Division, National Center for Statistics and
 Analysis, National Highway Traffic Safety Administration, U.S. Dept. of Transp., Report No.
 DOT HS 809 (Mar. 2003).
 It's official: Distracted drivers are dangerous

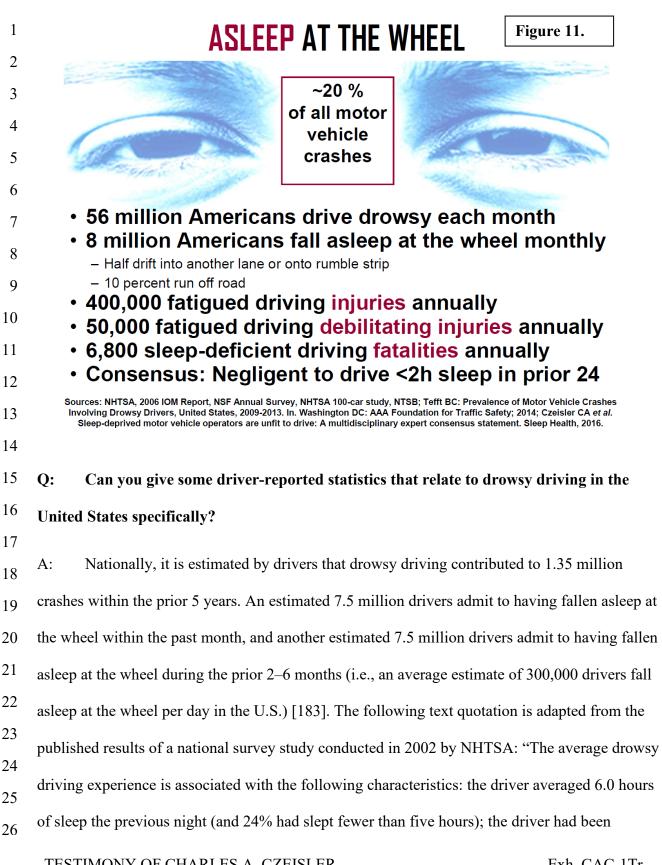
(CNN) -- A new study lends scientific credence to what many aready suspect: Drivers dabbing on makeup, chatting on cell phones or eating breakfast are three times as likely to be involved in a crash as more attentive motorists.

A new study of car crashes and near-misses identified leading contributing factors in drivers' behavior. 10 11 22% of all actual and near-miss 22.16 % Drowsiness MV crashes due to drowsiness 12 3.58 % **Dialing hand-held device** (NHTSA 100-car study, 2006) Fatal-to-driver truck crashes: 1/3 3.56 % Talking/listening to a hand-held device 13 are fatigue-related—equal drug 2.85 % Reading and alcohol-related crashes 14 combined (NTSB, 1990) 2.15 % Eating 15 Drowsy driving crashes account 1.41 % Applying makeup for 20% of serious MVC injuries 16 (Institute of Medicine, 2006) 1.23 % **Reaching for object** 17 1.11 % Reaching for a moving object 18 0.91 % Looking at external object 0.35 % Insect in vehicle 19 Source: National Highway Safety Administration, Virginia Tech Tra 20

20 Drowsiness is also a problem, the researchers found. They said drowsy drivers are four times as likely to have a crash or near-crash.

The conclusion that drowsy driving is a causal factor in 20 percent of all serious injuries resulting from motor vehicle crashes in the U.S. [1, 184] is consistent with earlier reports [186] and with estimates in the U.K. [105]. As illustrated in Figure 11 below, sleep-related crashes therefore result in nearly 55,000 debilitating injuries and 6,400 to 6,800 deaths annually [1, 24]

	(Figure 12). These estimates are also consistent with data from a recent study conducted by
1	researchers at Virginia Tech that was funded by the U.S. Department of Transportation, in which
2 3	100 automobiles were equipped with multiple video monitors, and the driving of all of the
4	individuals driving those cars were monitored over a year-long duration [187]. Sleep deprivation
5	renders drivers unfit to drive [185]. Video camera recording of the driver's face was
6	continuously recorded and scored for distractions, such as cell phone use, eating, personal
7	grooming, conversations with passengers, operating a GPS, etc., and drowsiness. The
8	automobiles were instrumented to capture actual and near-miss crash incidents. Drowsiness was
9	by far the most common contributing factor to these incidents, accounting for 22 percent of them
10 11	[187, 188], equal to the contribution frequency of all other distracters combined (see Figure 10
12	above). Legislation is needed at the state level to address the epidemic of drowsy driving [184].
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1	driving for an average of 2.9 hours (but 22% had been driving for more than four hours); the
1 2	incident occurred while driving on an interstate type highway with posted speeds of 55 mph or
3	higher (59%); and nearly half (48%) of the drivers reported that they nodded off between 9 p.m.
4	and 6 a.m. [183]. The overwhelming majority (92%) of drivers who nodded off while driving
5	within the past six months reported that they were startled awake. However, 33% drifted into
6	another lane or onto the shoulder, and 19% admit that they crossed the centerline. In 10% of
7	cases, the driver reportedly ran off the road. An estimated 292,000 drivers are involved in some
8	type of crash within the past six months as a result of nodding off at the wheel." [183]. In 2009,
9 10	the Commonwealth of Massachusetts issued the report of the Massachusetts Commission on
11	Drowsy Driving, on which I served as a member, to address this important public safety issue in
12	Massachusetts [184] I also served as Chair of a National Sleep Foundation Task Force to develop
13	a multidisciplinary expert consensus statement on the impact of sleep deficiency on driver fitness
14	[185].
15	
16	Q: Why are the consequences of sleep deficiency so evident in a routine task like
17 18	driving?
18 19	uning.
20	A: These widely recognized consequences of sleep deficiency are particularly evident while
20	These what is recognized consequences of steep denotency are particularly evident white
21	doing a routine task like driving because operating a motor vehicle is a demanding, vigilant task
21 22	
	doing a routine task like driving because operating a motor vehicle is a demanding, vigilant task
22	doing a routine task like driving because operating a motor vehicle is a demanding, vigilant task that is uniquely vulnerable to a momentary lapse of attention or slowed reaction time. Moreover,
22 23	doing a routine task like driving because operating a motor vehicle is a demanding, vigilant task that is uniquely vulnerable to a momentary lapse of attention or slowed reaction time. Moreover, driving is a routine, highly over-learned task with minimal novelty that occurs while drivers are

1	Drowsiness is Special Problem for
2	Motor Vehicle Operators Figure 12.
3	 Operating motor vehicles is a routine, highly-
4	over-learned task with minimal novelty
5	 Drivers are usually in a sedentary position
6	 At night, motor vehicle drivers are usually in
7	dim light or near darkness
8	Task is uniquely vulnerable to momentary lapse
9	of attention or slowed reaction time
10	 Drunk Driving ~10,800 deaths annually
11	 Drowsy Driving ~ 6,800 deaths annually
12	 Distracted Driving ~ 5,500 deaths annually
13	
14 15	Figure 12 . Operating a motor vehicle is a demanding vigilance task that is uniquely vulnerable to sleep deficiency and circadian misalignment and is associated with nearly twenty fatalities in the United States each day [22].
16	
17	Q: How does chronic sleep deprivation effect driving compared driving under the
18	influence of alcohol?
19 20	A: In a condition of chronic sleep deprivation, even when work (or wakefulness) is
20 21	scheduled during an appropriate circadian phase, the probability of a sleep-related
21	neurobehavioral performance failure while working or driving is markedly increased. Once an
22	individual has lost the struggle to stay awake and makes the transition from wakefulness to
24	sleep—however briefly—driving performance is much worse than that of a drunk driver, as the
25	individual is unresponsive to the environment throughout the duration of a sleep-related
26	attentional failure, such as a sleep-related lapse of attention, a micro-sleep episode or a sleep TESTIMONY OF CHARLES A. CZEISLER Exh. CAC-1Tr Page 41

attack. In 2004, Lamond and colleagues published a study in which they quantified the 1 deterioration in neurobehavioral performance that occurs at the end of the night shift in 2 comparison with that induced by alcohol intoxication [189]. In this laboratory-based study 3 simulating night shift work, individuals were scheduled to work in the laboratory on a series of 4 consecutive 8-hour shifts and scheduled to sleep up to 11 hours per day in a sound-attenuated, 5 6 dark room during the daytime. Neurobehavioral performance in those participants in the night 7 shift work condition was compared with an alcohol intoxication condition in which those same 8 participants were required to consume alcoholic beverage "consisting of 40% vodka and a non-9 caffeinated soft drink mixer, at hourly intervals." Notwithstanding the greater opportunity for 10 sleep between shifts afforded by an 8-hour as compared to a > 12-hour shift, after the third 11 consecutive 8-hour overnight shift, "impairment equivalent to that caused by a BAC [blood 12 alcohol concentration] of 0.10% was observed for lapse frequency [i.e., attentional failures] at 13 14 the 8th hour [i.e., at 7 am]" [189]. As with alcohol intoxication, chronically sleep deprived 15 individuals tend to underestimate the extent to which their performance is impaired, despite 16 increasing impairment evident in objective recordings of the rate of lapses of attention or 17 attentional failures [16]. Moreover, most drivers who cause sleep related motor vehicle crashes 18 are often reportedly unaware that they fell asleep or had a sleep related attentional failure that 19 caused the crash [190]. Sleep deficiency, circadian disruption, sleep-promoting drugs or foods, 20 21 and medical illness can all increase the risk of impairment by fatigue (Figure 13). Soporific 22 environmental conditions can exacerbate this risk. 23 24 25

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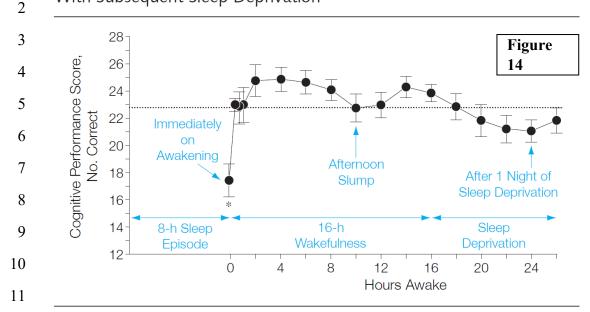
Multiple pathways can lead to Figure 13. 1 operator impairment by fatigue 2 Sleep deficiency 3 - Insufficient Sleep (acute or chronic) 4 Recreational sleep loss 5 Occupational sleep loss - Sleep Disorder 6 Circadian disruption 7 Drug- or alcohol-induced 8 Secondary to other medical condition 9 Drowsy drivers seek distractions to stay awake 10 - Sleep deficiency increases distractibility 11 12 Figure 13. Many different factors can result in the fatigue-related impairment [22]. 13 14 15 H. Interruptions of sleep and sleep inertia 16 **O**: How do sleep interruptions affect alertness and what are some compounding 17 factors? 18 19 A: Repeated interruptions of sleep, such as is experienced by employees when they are on 20 call, degrade the restorative quality of sleep, compared to an equal amount of consolidated sleep. 21 Such repeated interruptions of sleep are thought to contribute to the excessive daytime sleepiness 22 associated with sleep disordered breathing, which induces many brief arousals during the night. 23 Interestingly, just being on call itself disturbs sleep, even when the individual is not called [191, 24 192]. 25 26

Q:

What is sleep inertia and how does it relate to cognitive performance?

A: Cognitive performance is markedly degraded during the transition from sleep to wakefulness [11, 193-204]. The extent to which this phenomenon, which is called sleep inertia, interferes with neurobehavioral performance is related to the depth of the prior sleep episode [200]. Thus, agents that interfere with sleep, such as caffeine, can mute the effect of sleep inertia [205]. The adverse impact of sleep inertia on neurobehavioral performance can exceed the impact of total sleep deprivation, as shown in Figure 14 from our publication in the Journal of the American Medical Association (JAMA)[11]. Individuals who are subjected to acute total sleep deprivation or chronic sleep restriction often experience very deep sleep, which together with an adverse circadian phase will increase the effects of sleep inertia [10, 11, 200, 206, 207]. Upon awakening from sleep, individuals will often be confused and may even lack situational awareness.

Figure. Cognitive Performance on Awakening From Sleep Compared With Subsequent Sleep Deprivation



12 The group mean (horizontal dotted line) across the sleep inertia period and 26hour awake period has been added to the deviation from the mean scores for the 13 number of correct responses in 2 minutes to permit overall assessment of the magnitude of performance impairment. Error bars indicate SEM; asterisk, differs from 14 all subsequent time points at $P \leq .01$.

Q: You performed research that documents the effects that sleep inertia can have on cognitive performance. Would you please briefly describe that study?

17

18 A: Yes. As shown in Figure 15 below from research performed in my laboratory by Dr. SE

19 McCormack in collaboration with Drs. K Kwong and BR Rosen at MIT, sleep inertia following a

²⁰ 45-minute sleep episode in an MRI brain scanner at 4 am is associated with local loss of

21 functional Magnetic Resonance (brain) Imaging activation in response to an attention task. (SE

- McCormack, D Aeschbach, K Kwong, BR Rosen, CA Czeisler, Presented at the 17th Congress 23
- of the European Sleep Research Society, P087, October 9, 2004, Prague, Czech Republic).
- 25

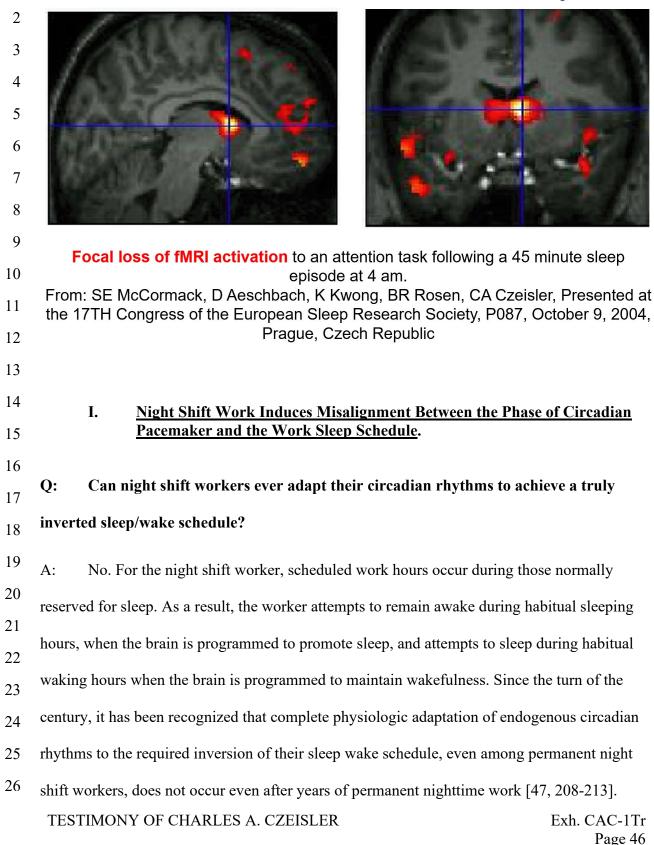
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SLEEP INERTIA: Thalamus and frontal cortex are particularly slow to recover in the first 15-30 minutes after awakening

Figure 15.

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	Maladaptation of night shift workers' circadian system to their required work schedule
1	results from their exposure to conflicting environmental and behavioral synchronizers. For a
2	results from their exposure to conflicting environmental and behavioral synchronizers. For a
3	commercial motor vehicle driver, this is particularly true on days that they are driving, since
4	drivers are exposed to the solar light-dark cycle to a much greater extent than a person working
5	in an indoor environment. This light exposure synchronizes the biological clock to the solar day,
6	reinforcing a conventional schedule of day activity and night sleep. Therefore, a maladaptive
7	circadian phase relationship with the outside world is maintained for the night commercial motor
8	vehicle driver, promoting sleep at night and wakefulness during the day. This environmental
9	synchronizer is also reinforced by exposure to social and behavioral influences promoting day
10	
11	activity and night sleep, such as the often day-oriented schedules of pickup and drop-off
12	locations. These influences often lead the night shift worker to postpone, interrupt or truncate
13	daytime sleep. Exposure to these environmental, social and behavioral synchronizers can
14	overcome the potential effect of the inverted work-rest schedule on the circadian clock, resulting
15	in misalignment between the biological time programmed into the brain's circadian clock and the
16	work rest schedule required of the night shift worker [47]. Misalignment of the circadian
17	
18	pacemaker with the desired sleep wake schedule is thus a chronic consequence of night shift
19	work, even among permanent night shift workers.
20	

Q: Is there a period during the usual sleep time when night shift workers are especially vulnerable to impaired cognitive performance?

23

24 A: Numerous laboratory studies, including those in my own laboratory, have shown that

²⁵ there is a prominent circadian variation in objective and subjective measures of alertness,

performance (psychomotor, vigilance, memory) and attention or ability to concentrate with a TESTIMONY OF CHARLES A. CZEISLER
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	nadir i	n the latter half of the usual sleep time [4, 30, 66, 86, 210, 214-226]. Because of the failure
1	of the	circadian pacemaker to adapt to the inverted work schedule in most night shift workers,
2		
3	the pe	ak drive for sleep continues to occur during the latter half of the night and into the morning
4	hours	while the employee is at work and/or commuting home from work. In 1995, Pack et al.
5	noted	that the peak of fatigue-related motor vehicle crashes occurs in the early morning hours
6	betwee	en 7–8 a.m. [190]. Extended duration work shifts, in which night workers are also
7	schedu	led to work during daytime hours can further increase sleep pressure and lead to increased
8	risk of	sleep-related neurobehavioral performance failure [227].
9		
10		
11	Q:	Do individuals differ in their ability to maintain performance during night work?
12	A:	Yes, there are inter-individual differences in the ability to maintain performance during
13	night v	work. Increased vulnerability to attentional failure during acute sleep deprivation in women
14	•	
15	depen	ds on menstrual phase [228].
16		
17	Q:	What is Shift Work Disorder and are there effective treatments?
18	A:	Excessive sleepiness during night work or insomnia during day sleep are the cardinal

A: Excessive sleepiness during night work or insomnia during day sleep are the cardinal symptoms of Shift Work Disorder [229], which recent research indicates may have a genetic basis in some individuals [230]. A number of countermeasures and treatments can be effective in treating the symptoms of Shift Work Disorder, including improvements in work scheduling, use of properly timed use of light, administration of nutritional supplements such as caffeine and melatonin, and use of pharmacologic agents such as modafinil, as we have demonstrated in my laboratory and our field study investigations [8, 43, 47, 231-236].

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Night Shift Work Leads to Sleep Deprivation.

Q: How does night shift work lead to sleep deprivation?

Several mechanisms are responsible for the sleep deprivation associated with night shift 4 A: 5 work. The first and most direct effect is that the employees are now awake and working during 6 the hours at which they ordinarily sleep. This can occur when employees work throughout the 7 time of their usual sleep episode (e.g., an overnight work shift), during the first half of the usual 8 sleep episode (e.g., a late evening shift), or during the final half of their usual sleep episode (e.g., 9 an early morning shift). Second, abnormalities in the internal architecture and consolidation of 10 sleep result from the misalignment of their attempted sleep time with the output of their 11 12 biological clock, causing sleep deprivation, even when there is ample time available for sleep 13 between shifts. Laboratory studies that I and others have conducted have demonstrated 14 conclusively that the length of a sleep episode is dependent on the phase of the circadian system 15 at which the sleep occurs, even when participants are studied in sound-attenuated rooms and 16 shielded from external time cues [28, 29, 66, 67, 237]. Studies have demonstrated that sleep 17 duration is reduced during daytime hours as compared to the night [238, 239]; daytime sleep is 18 accompanied by more frequent arousals and changes in intra-sleep architecture. Even in a 19 20 soundproofed environment and/or one without time cues, sleep is quantitatively and qualitatively 21 different from that obtained at night, with a deficit of both minutes and types of sleep. Daytime 22 sleep episodes are shorter in duration than nighttime sleep episodes because the night shift 23 worker is attempting to sleep just as the internal biological alarm clock sounds to wake him or 24 her up, a consequence of a misalignment between the circadian pacemaker and the external 25 environment. For example, the daytime sleep of train drivers scheduled to work at night is cut 26

short when sleep is attempted during the daytime at an adverse circadian phase [Foret J and 1 Lantin G, in Aspects of Human Efficiency, W. P. Colquhoun, Ed. (English Univ. Press, London, 2 1970), pp. 273-282] [66]. The daytime sleep of night shift workers worsens with advancing age 3 [238-240]. When an inadequate amount of time is available between shifts, sleep deprivation will 4 further alter sleep stages. Sleep disruption is more pronounced in middle aged shift workers than 5 6 in younger shift workers [239, 241]. Nurses commonly admit to falling asleep while working at 7 night, with almost two thirds (65%) of the nurses working the night shift reporting struggling to 8 stay awake at work and 20% falling asleep at work [242]. 9 10 11 **Q**: How do environmental disturbances and social norms contribute to night shift 12 worker sleep deprivation? 13 A: Environmental disturbances further interrupt the daytime sleep of the night shift worker. 14 Noise from children, road traffic, telephones, doorbells, and other family members are the most 15 commonly reported causes of sleep interruption of the night shift worker [243, 244]. In one 16 17 survey of 9,000 shift workers, children's noise was mentioned in 77.9 percent of the complaints, 18 road traffic noise in 63.2 percent and telephone ringing in 54.5 percent [243]. Field studies of 19 shift workers worldwide indicate that the majority of workers whose schedule includes night 20 work (midnight to 7 am) experience disturbed day sleep [243, 244]. 21 Furthermore, night workers are often attempting to sleep at a time when not only 22 environmental activities, but also social activities are at a peak. Therefore, many night shift 23 24 workers voluntarily elect to shorten their sleep in order to attend to social and family obligations. 25 In fact, night shift workers are much more likely than day workers to set alarms to interrupt sleep 26 [245], notwithstanding the increasing sleep debt they are accumulating. Thus, pressure from the **TESTIMONY OF CHARLES A. CZEISLER** Exh. CAC-1Tr Page 50

demands of society often leads night shift workers to interrupt and even prematurely truncate the sleep that they would be able to achieve during the day. Employees working on call and/or irregular schedules also suffer from poor quality sleep. i.e., sleep that is less restorative for optimal work time functioning.

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6 Q: How much less sleep do night shift workers typically get during the day compared to 7 at night?

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A: Overall, night shift workers sleep an average of one to two hours less during the day than
at night [47, 239, 244], accumulating approximately one lost full night of sleep for every week of
night work. Furthermore, on individual days they are often able to sleep only 1 to 3 hours [244],
resulting in acute sleep deprivation. Subjective sleep quality and quantity is also poorer during
day sleep [246-248].

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Data from field studies collected on shift workers are thus consistent with the conclusion 15 that rotating shift workers are both chronically and acutely sleep deprived when working at night 16 and attempting to sleep during the day. These problems with shorter and poorer quality sleep 17 18 often persist despite government or industry polices that mandate rest time. Unless sufficient 19 duration and quality sleep occurs, rest alone will have no significant impact on promoting 20 alertness and performance during subsequent working hours. For this reason, it is critical that 21 employers in safety-sensitive industries require employees in general, and night workers in 22 particular, to notify the employer and be relieved of any scheduled work assignment if, for 23 whatever reason, the employee is not fit for duty due to insufficient sleep duration or quality. 24

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K.

Worker Survey Data on the Effects of Shift Work

Q: What research have you performed that catalogs the prevalence of sleeping and 2 3 inattentiveness among shift workers? 4 A: My own research in the field has provided compelling evidence of the prevalence of 5 sleeping and inattentiveness among shift workers. My colleagues and I have worked with 6 companies from a variety of industries nationwide to evaluate the shift work schedules of 7 workers at those companies using questionnaires designed to evaluate shift work schedules, 8 9 employee preferences and sleep disorders. 10 The questionnaires sought information on worker satisfaction, health practices, and 11 adaptation to their existing shift schedules. The questionnaires specifically asked how frequently 12 the employees "nodded off" or fell asleep while at work. Steps were taken to protect worker 13 confidentiality and encourage participation. 14 15 **Q**: What were the results of these questionnaires? 16 17 The analysis presented below represents results of surveys administered to a total of A: 18 2,583 workers from 12 companies. Only those workers who indicated that they currently worked 19 a 3-shift rotation, including day, evening, and night shifts, were included in the analysis. The 12 20 companies are from a variety of industries and include an urban police department, a paper 21 company, a plant producing rubber for the tire industry, a potash harvesting plant, a tire 22 manufacturing company and a utility with both nuclear and non-nuclear facilities. 23 Summarized below are the results of a series of questions related to attentiveness and the 24 25 ability to remain awake at work. The survey data show falling asleep at work and loss of 26 alertness to be common features of shift work. Moreover, the risk of falling asleep at work and **TESTIMONY OF CHARLES A. CZEISLER** Exh. CAC-1Tr

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loss of alertness is elevated during the night shift. These results are consistent with those obtained by other researchers using similar methods [209, 221, 223, 226, 249-254].

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When the shift workers were asked how many times per week of working on the day shift 3 they usually nodded off or fell asleep while at work, 21.6% of the 2402 workers who responded 4 said they fell asleep at least once each week on the day shift, with 2% of those indicating that 5 6 they fell asleep more than five times per week. When working evening shifts, 15.1% of the 2392 7 workers surveyed said they nodded off or fell asleep one or more times per week, with 0.7% 8 indicating that they fell asleep more than five times per week. When asked about working the 9 night shift, 55.7% of the 2424 respondents said they fell asleep at least once per week, with 10 12.1% responding that they fell asleep more than 5 times per week. These inter-individual 11 differences in the severity of symptoms of Shift Work Disorder derived from workers 12 participating in field studies are consistent with laboratory investigations that have revealed 13 14 considerable differential vulnerability to the deterioration in neurobehavioral performance 15 associated with sleep loss and night shift work [255-260]. 16

Overall, 57.0% of 2363 workers reported falling asleep at least once during a full shift rotation. While results varied by company, at least 40% of the workers from each and every company reported nodding off or falling asleep on the night shift at least once per week.

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Q: Can you describe your findings with respect to sleepiness among Nuclear Powered Electric Power Generation plant workers on 8-hour Shifts?

A: There is a common misconception that in critical situations, where individuals have great
 responsibility for human life, they can voluntarily overcome the effects of sleep loss and
 misalignment of circadian phase. Our survey data did not support this hypothesis. Even nuclear TESTIMONY OF CHARLES A. CZEISLER
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power plant workers experienced the same debilitating effects of night shift work on alertness and performance.

As with shift workers in the other industries we surveyed, the nuclear plant personnel 3 reported a significant drop in their level of alertness on the night shift. This drop was reported by 4 both licensed and non-licensed nuclear plant personnel. In addition, both licensed and non-5 6 licensed personnel reported that they were most tired on the night shift. In fact, 49.4% of the 7 nuclear power plant workers reported that they fell asleep at least once per week on the night 8 shift, with 30.5% reporting that they fell asleep 2 to 5 times per week, and 5.2% reporting 6 to 10 9 times per week. As in other industries, nodding off or falling asleep at work occurred to a lesser 10 extent on the day and evening shifts. The predominance of reported sleeping on the night shift is 11 consistent with the fact that 71% of the 2,498 workers responding indicated that they were most 12 tired on the night shift. 17.3% indicated they were most tired on the day shift, 4.2% responded 13 14 that they were most tired on the evening shift, and 7.5% said that they were not more tired on 15 any shift.

16 Similarly, when asked to rate their alertness on the VAS, the average + S.E.M. alertness 17 score of 2,392 workers on the night shift was 42.0 + 0.54; the average alertness score of 2,322 18 workers on the day shift was 66.5 ± 0.47 ; and the average alertness score of 2,366 workers on the 19 evening shift was 73.0 + 0.42; and the average alertness score of 2,180 respondents on their days 20 21 off was 76.9 + 0.43. When asked if they thought that their schedule makes them sleepy, a 22 majority of workers (54.1% of 2,342) answered YES. On a separate question asking if they felt 23 too sleepy during work, 36.8% of 799 workers who were asked this question responded 24 affirmatively.

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Q: Can describe the questionnaires' results regarding attentional failures and occupational safety?

2 A: The impact of the profound fatigue associated with night shift work is highlighted by the 3 responses from employees reporting instances of falling asleep driving to or from work: 17.6% 4 of 1,270 shift workers reported that they had fallen asleep while driving to or from work in the 5 6 last year and 30.4% of 2,368 shift workers reporting that they had at least one near miss or actual 7 automobile accident due to sleepiness in the past year. A subset of these workers was asked more 8 detailed questions regarding the circumstances of these accidents. 3.6% of all workers 9 responding reported having an actual automobile accident due to sleepiness. Of these 15 10 workers, ten had accidents on a workday. Two reported accidents that happened while on 11 evening shift rotation, and eight occurred to workers on the night shift. 12 Of the 428 workers responding to a question about near-miss automobile accidents due to 13 14 sleepiness, 146 (34.1%) reported having at least one actual or near-miss automobile accident due 15 to sleepiness in the past year. When asked about procedural errors at work due to sleepiness, 16 19.7% of 233 workers responding reported at least one procedural error, and 33.2% of 214 17 workers responding reported near-miss procedural errors at work due to sleepiness. When 18 questioned about personal injury accidents at work due to sleepiness, 6.9% of 245 workers 19 responding reported actual personal injury accidents, and 18.4% of 228 workers reported near-20 21 miss personal injury accidents due to sleepiness. 22 23 24

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O:

There are many studies regarding attentional failures and occupational safety. Can you please briefly describe some of those studies and their results?

Yes. First, in 1992, we published data on a study of 876 nurses. We found that 32.4 A: 3 percent of those who always worked nights reported falling asleep during the night shift at least 4 once a week [261]. Furthermore, we demonstrated that nurses who worked the night shift had 5 6 2.24-fold greater odds of a motor vehicle crash and nearly double the risk of a near-crash [261]. 7 In 1996, Novak and Auvil-Novak published data from a cohort of night-working nurses, 8 95.5 percent of whom report that they had experienced at least one accident or near-accident 9 driving home from the night shift [262]. In 1999, Stutts et al. reported that working the night 10 shift increases the odds of a sleep-related crash by nearly six times and [263]. In 2007, in a study 11 of nurses, Scott et al. reported that working at night increased the odds of a drowsy driving 12 incident four-fold, such that 79.5 percent of nurses who worked only the night shift had an 13 14 episode of drowsy driving within a 4-week study interval [264]. Moreover, 3.3 percent of those 15 nurses reported experiencing drowsy driving following every shift worked. In a 2008 study of 16 nurses, Dorrian et al. [265] reported that "nearly half of extreme drowsiness and near-accident 17 reports occurred between 0700 and 0900h. This coincides with the end of the night shift" [265]. 18 These data are consistent with the conclusions from the published literature published by 19 Lyznicki et al. for the Council on Scientific Affairs of the American Medical Association, by 20 21 Horne [190, 266], by Scott et al. [264], and by Dorrian et al. [265], who concluded that night 22 shift work was associated with a markedly increased risk of fatigue-related motor vehicle 23 crashes. In 2007, Scott et al. concluded: "Given the large numbers of nurses who reported 24 struggling to stay awake when driving home from work and the frequency with which nurses 25 reported drowsy driving, greater attention should be paid to increasing nurse awareness of the 26

risks and to implementing strategies to prevent drowsy driving episodes to ensure public safety. Without mitigation, fatigued nurses will continue to put the public and themselves at risk" [264].

1 2

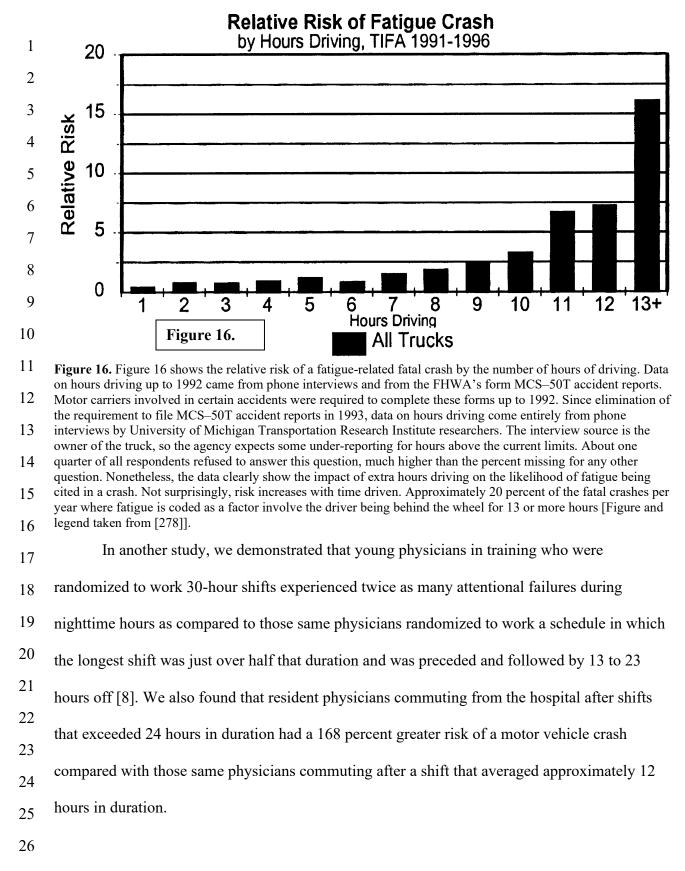
The work schedule itself has a significant effect on the risk of errors and accidents. The 3 number of consecutive work hours; the number of consecutive days of work; the time of day and 4 the frequency of rest breaks have all been found to influence the risk of error and accident [267-5 6 270]. A meta-analysis of a number of research studies has revealed that the relative risk of Injury 7 or accident is more than doubled after 12 hours of work [267], comparable to the results of 8 Haneke et al., who reported in 1998 that the relative accident risk increased exponentially after 9 the 9th hour of work to more than double the risk of an 8-hour shift [271]. Rogers and her 10 colleagues found that the risk of error was three times higher when nurses worked shifts of 12.5 11 hours or longer [270]. 12

When workers experienced with both 8- and 12-hour night shifts were asked which shift 13 14 resulted in fatigue that decreased their performance at work, 80 percent of respondents felt that 15 the 12-hour workday resulted in the least efficient performance due to fatigue [272]. In a 16 modeling study entitled "The 12-hour Shift Revisited: Recent Trends in the Electric Power 17 Industry" conducted by Ontario Hydro in Toronto, Canada, it was estimated that human error by 18 operators would be doubled by a change from 8-hour to 12-hour shifts, due to fatigue and 19 misalignment of circadian phase. This resulted in a 70 percent increase in their risk assessment of 20 21 a public accident [273-275]. In addition to (1) night shifts posing a greater risk of injuries and 22 accidents than day shifts, (2) 12-hour shifts posing a greater hazard than 8-hour shifts, (3) the 23 later hours of the night shift posing a much greater hazard than the initial hours of the night shift, 24 and (4) the circadian peak of sleep-related errors and accidents occurring between 6 a.m. to 9 25 a.m., the number of consecutive days of lengthy shifts also affects the risk of errors and 26

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	accidents. In 2006, Folkard and Lombardi [267] reported that "there was a significant trend [in
1 2	the risk of accidents] across four successive night shifts [such that] risk was about 6% higher
2	on the second night, 17% higher on the third night, and 36% higher on the fourth night."
4	In 1998, the National Highway Transportation Safety Administration concluded, based
5	on evidence from crash reports and self-reports of sleep behavior and driving performance, that
6	one of the three top population groups at highest risk of drowsy driving crashes was: "Shift
7	workers whose sleep is disrupted by working at night or working long or irregular hours" [276].
8	The National Highway Transportation Safety Administration noted that "periods of work longer
9 10	than 8 hours have been shown to impair task performance and increase [the risk of motor
10	vehicle] crashes"[276]. As reported by the Department of Transportation, the relative risk of a
12	fatigue-related crash among truck drivers-which is the leading cause of fatal-to-the-driver truck
13	crashes [102, 277]—increases sharply after 10 hours of driving, such that there is a fifteen-fold
14	increase in the risk of a fatigue-related crash in the group driving more than 13 hours [278]
15	(Figure 16).
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_	As Harris observed more than forty years ago, truck driving in particular is one of the
1 2	most demanding vigilance tasks, in which a momentary lapse of attention has the potential for
3	disastrous consequences [104]. Fatigue-related crashes in commercial motor vehicles occur most
4	commonly during night-time hours [103, 104, 279]. Sleep-related fatigue is a major cause of
5	crashes involving heavy trucks due to fatigue-related impairment of driving skills (poor
6	judgment, slowed reaction times, decreased awareness and increased distractibility) [102, 277,
7	280]. Sleep-related fatigue can be caused by long work hours and lack of sleep and is strongly
8 9	influenced by time of day [with increased risk in the latter half of the habitual sleep episode and
9 10	in the mid-afternoon between about 1-5 p.m. [89]]. In May of 1994, two parents in Maine whose
11	son and three of his friends were killed by a truck driver who the parents believe fell asleep at the
12	wheel formed an organization they call "Parents Against Tired Truckers," an organization with
13	the stated goal of reducing heavy truck crashes resulting from truck driver fatigue.
14	
15	Q: Can you please describe the study that you performed regarding sleepiness among
16 17	urban police officers working 8-hour Shifts?
18	A: Thirty years ago, I conducted an evaluation of the impact of the shift work schedule on
19	the alertness and performance of the officers in the Philadelphia Police Department. In order to
20	do so, a survey questionnaire was administered to a representative sample of patrolmen and
21 22	detectives from all districts and divisions in the City of Philadelphia to obtain information from a
	detectives from an districts and divisions in the City of Finadelpina to obtain mornation from a
23	cross section of the officers (a horizontal sample). Simultaneously, a single district of patrolmen
23 24	

Exh. CAC-1Tr Page 60 collected from members of the 35th District and the Northwest Detectives in the City of Philadelphia, for a total of 375 respondents.

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2 I found that 54.3 percent of the officers reported that they received an inadequate amount 3 of sleep. In fact, the only time that the officers averaged eight hours of sleep per night was during 4 a series of days off. When working, the officers' average daily sleep times were 6.7 hours per 24 5 6 hours while on day shift; 7.2 hours while on the evening shift; and 6.3 hours when on the night 7 shift. Significantly, more than 50 percent of the officers reported a moderate to severe problem 8 with poor quality sleep. More than 70 percent of the officers admitted falling asleep on the job 9 (intentional or unintentional) during the night shift; one third of the officers reported falling 10 asleep on the job when working the day shift; and one in seven officers reported falling asleep at 11 work while working the evening shift. Officers who admitted falling asleep at work during the 12 night shift estimated that it happened an average of more than three times per week of night 13 14 shifts. Officers on the night shift averaged 3.8 of such on-the-job sleep incidents per week, even 15 though falling asleep at work was considered grounds for disciplinary action and/or termination. 16 Furthermore, 41.3 percent of the officers reported feeling tired frequently or most of the time 17 both on and off the job while on the night shift. In fact, one out of every ten officers reported 18 incidents when they actually fell asleep while driving to or from work during the prior year. 19 Nearly one in four officers reported an actual or near miss automobile accident due to sleepiness 20 21 in the prior year. Thirteen percent of the officers reported three or more such incidents in the 22 prior year. Finally, I found that 75 percent of the officers reported that their schedule did not 23 provide them enough time with their children. Nearly 80 percent reported that their families were 24 dissatisfied with their work schedule. Only 16 percent reported that their families were satisfied 25 with their work schedule. 26

Q: There have been surveys regarding sleepiness among air traffic controllers on a 2 rapidly rotating shift-work schedules. Can you please provide some of the data from the Federal Aviation Administration's survey?

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5 A: Yes. Survey data reveal that sleep disruption and fatigue are serious problems among air 6 traffic controllers. Data from a Congressionally mandated, FAA-sponsored survey distributed 7 from 1999 to 2000 to all 23,500 FAA personnel with an ATCS designation revealed that 62% of 8 the 4,112 enroute and terminal ATCS respondents reported difficulty sleeping before the 9 midnight shift. In fact, according to a pamphlet published by the FAA in 2001, "67% of ATCS 10 11 shift workers reported having trouble sleeping because of shift work [and] 46% of ATCS shift 12 workers indicated that they often fall asleep unintentionally." [281]. Reported sleep quality was 13 rated most poorly (averaging only 1.3 on a 1-5 scale in which 5 was high) on the sleep between 14 the day shift and the night shift on the FAA's counterclockwise, rapidly rotating schedule 15 worked by 92 percent of the respondents [281], and the respondents felt least rested (average 16 rating of 1) after the midnight shifts and the least mentally sharp in terms of alertness and 17 memory at the end of the midnight shift, consistent with data from our studies in the laboratory 18 19 [2]. As the FAA reported in 2001, "Mental sharpness is lowest during the midnight shift because 20 at this time, shift workers must deal with the circadian low point for energy and alertness levels, 21 and the effects of poor quality daytime sleep." [281]. In fact, 77% of the FAA's air traffic 22 controllers on the counterclockwise, rapidly rotating shift work schedule with midnight shifts 23 reported that they had caught themselves about to doze off while at work; 79 percent of the air 24 traffic controllers had sometimes, frequently or always had lapses of attention and 36 percent had 25 actually fallen asleep at the wheel while commuting on the midnight shift [281]. This most often 26

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occurred on the commute home from the midnight shift. When asked if the air traffic controllers
had any other comments or observations relating to their sleep and fatigue, the most common
category of response was: "Quick turn-arounds between shifts cause fatigue and/or are
dangerous." This spontaneous response was written in by 164 air traffic controllers. The second
most common write-in response category was that "shift work in general causes fatigue, is
dangerous, and does not allow a sleep routine," which was noted by 115 respondents.

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Q: Can you please provide some objective field data on the effects of shift work across various industries?

11 A: Yes. The self-reported rate of nodding off or falling asleep at work among nuclear power 12 plant control room operators and police officers seen in our survey data is consistent with survey 13 data on severe sleepiness of train drivers working an irregularly scheduled night shift [282], and 14 with data recently obtained using continuous electroencephalographic (EEG) recordings of 15 power plant operators in Sweden. Dr. Kerstin Dahlgren of the Swedish Nuclear Inspectory has 16 recorded ambulatory polysomnograms on 20 nuclear power plant workers on a rotating shift 17 work schedule and has found that approximately 25 percent of them fall asleep while operating 18 19 the plant, mainly while on the night shift (Dr. Dahlgren, personal communication). Dr. Dahlgren 20 notes that the operators not only napped when she left them on their own, but they progressed to 21 slow wave sleep, according to the EEG recordings (Dr. Dahlgren, personal communication), a 22 progression which our laboratory studies indicate requires an average of 18.0 + 8.6 minutes 23 (range 9.5 to 33.5 minutes). During her study, she has further personally observed situations in 24 which all five nuclear power plant operators were simultaneously asleep. 25

26

Objective field studies have also discovered sleeping among workers subject to circadian 1 rhythm disruptions and/or sleep deprivation in other industries where worker alertness is 2 commonly viewed as a necessary element of successful job performance. For example, studies of 3 airline pilots have revealed instances of pilots asleep while on duty on the flight deck [222, 283] 4 and physicians falling asleep while caring for patients in Intensive Care Units [8]. EEG 5 6 recordings of the pilots have shown patterns consistent with sleep or inattention. Observers in the 7 cockpit have also seen airline crews take voluntary and involuntary naps. In some cases, the crew 8 discussed who would nap, despite regulations prohibiting such behavior. In fact, a NASA/FAA 9 sponsored study revealed that the pilots of commercial airlines were napping approximately 11% 10 of the time on long haul flights (average flight duration, 10.4 hours) with nap episodes averaging 11 46 minutes in duration, occurring throughout the flights despite the presence of NASA observers 12 and the knowledge that their EEG patterns were being monitored [284]. 13 14 Studies of train drivers in the field have shown similar results: EEG and EOG changes 15 consistent with inattention, microsleep, and/or sleep and observer (in the locomotive) recorded 16 instances of inattention or missed signals and other severe performance lapses [252, 254, 285]. 17 One study [252] found that of 2,290 train trips, 1.5% (34 cases) had actual operational misses 18 involving drowsing or napping; 79% of these occurred between midnight and 6 a.m. and 12% 19 between 10 p.m. and midnight. The NASA/FAA field study revealed that alertness of pilots from 20 21 top of descent to landing can be improved by the use of scheduled cockpit naps [222], similar to 22 the results of a CAMI study. However, we have found that brief (<4 hr) sleep episodes are 23 insufficient for restoring performance in first-year resident physicians working overnight 24 extended-duration work shifts [286]. 25

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When the work to be performed is relatively monotonous and routine, then sleepiness 1 may be even more difficult to overcome [226, 287]. Dr. Dahlgren found that workers at the 2 power plants were generally inactive and that there was even less activity on the night shift. The 3 biological factors causing sleep and inattentiveness in shift workers would therefore be even 4 more difficult to overcome in a relatively non-stimulating environment, such as that of 5 6 monitoring a control room in an electric power plant, driving a motor vehicle on a limited access 7 highway, or piloting a vessel at night from the darkened bridge of a slowly moving cargo ship. 8 As illustrated in Figure 17, continuous polysomnographic monitoring of 80 licensed 9 commercial drivers without OSA during 7,500 hours of long-haul trucking operations revealed 10 that more than half of the long-haul truck drivers experienced episodes of drowsy driving, mostly

during night driving [288]. Predictably, these episodes of drowsy driving were not distributed evenly across the 80 drivers. In fact, more than half of the drowsy driving episodes occurred in just 8 (10%) of the 80 drivers [288], revealing the differential vulnerability of this subset of drivers to the sleep deprivation and circadian misalignment associated with night driving. This is consistent with the differential vulnerability to nocturnal sleep loss observed in laboratory studies [255-260].

Countermeasures such as the use of light exposure of optimal wavelength and intensity to suppress endogenous melatonin secretion, enhance alertness and reset the circadian clock of night shift workers [44, 47, 120, 289], the use of wake-promoting therapeutics during night work [233, 234, 290] and the use of sleep-promoting therapeutics during day sleep [43, 291-293] hold promise for use in overcoming the propensity for the occurrence of lapses of attention and unintended sleep episodes during night shift work.

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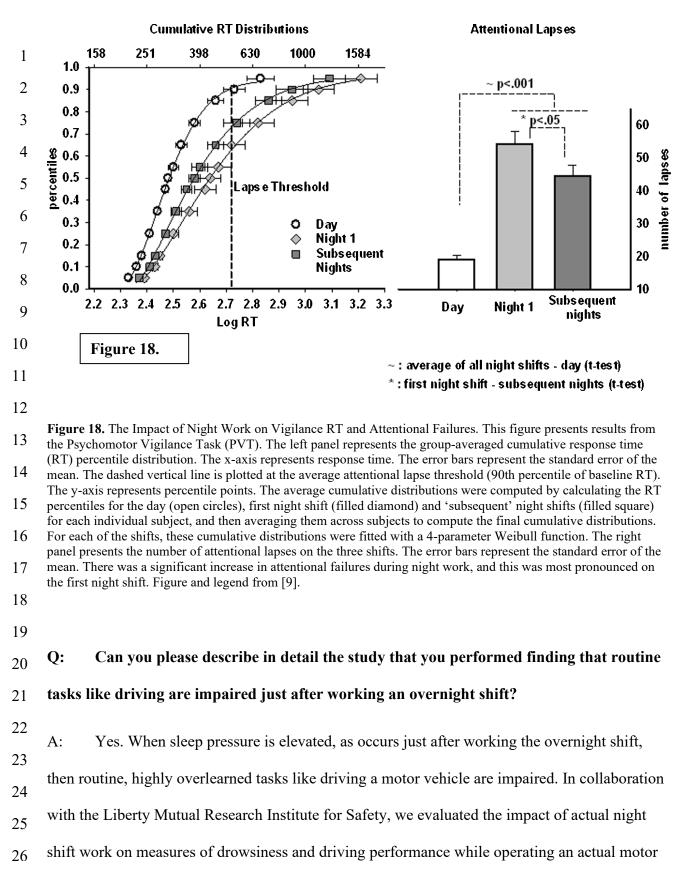
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2	Overnight Trucking and Drowsy Driving Figure 17.
3 4 5	Polysomnographic recordings reveal that truck drivers working between midnight and noon
6	sleep an average of only 3.8 hours per day ^{1,2}
7	 Videos recordings revealed that truck drivers are often drowsy while driving
8	Forty-five drivers (56 percent) had at least 1 six-minute interval of drowsiness while driving
9	_ 1067 of the 1989 six-minute segments (54 percent) that showed
10	drowsy drivers involved just eight drivers
11	¹ Wylie et al., Commercial Motor Vehicle Driver Fatigue and Alertness Study: Technical Summary. Report No. FHWA-MC-97-001, National Technical
12	Information Service, Springfield, VA, November, 1996. ² Mitler, M. M., Miller, J. C., Lipsitz, J. J., Walsh, J. K., and Wylie, C. D. The sleep of
13	long-haul truck drivers. New England Journal of Medicine 337 , 755-761. 1997.
14	
15	As noted in the 1975 FAA study carried out at the FAA Civil Aeromedical Institute in
16	Oklahoma City, Oklahoma that was entitled "The Effects of a 12-Hour Shift in the Wake-Sleep
17	Cycle on Physiological and Biochemical Responses and on Multiple Task Performance," the
18	interaction of sleep loss with misalignment of circadian phase may explain why performance is
19	so significantly degraded at the end of the first night shift [294]. As Higgins and his colleagues at
20	the FAA Civil Aeromedical Institute concluded from their study, "The implications of these
21	findings are as follows: (1) Individuals making a 12-hour alteration in the wake-sleep cycle
22	
23	should not perform critical tasks during the first awake period following the change.
24	Performance decrements observed during this first period might be due to either the sleep loss or
25	the initial adjustment to the new schedule, or both factors could be contributory. (2) After the
26	first full sleep period following the change, subjects appeared to perform well even though the
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physiological and biochemical parameters measured were still adjusting to the change. ..." [294] (pp. 23-24).

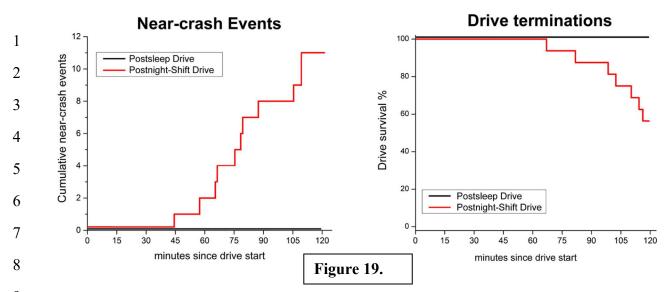
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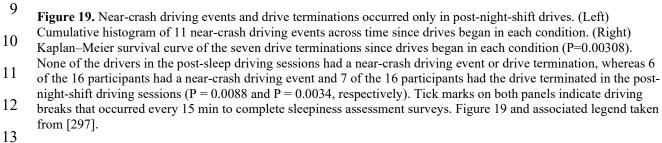
2	(pp. 23-24).
3	More than 30 years later, we have published a paper based on studies sponsored by the
4	National Institute of Occupational Health in my laboratory at the Harvard Medical
5	School/Brigham and Women's Hospital that comes to a similar conclusion, as shown in Figure
6	18, which is from our publication entitled "Acute Sleep Deprivation and Circadian Misalignment
7	Associated with Transition onto the First Night of Work Impairs Visual Selective Attention" [9].
8	Moreover, we have demonstrated that the sleep deprivation associated with night shift work
9 10	resulted in individuals becoming "fast and sloppy" when doing visual searches [41]-with
10	impairment of decision making [295]—and that sleep deprivation interferes with the search for
12	rare targets [296]. Excessive sleepiness is also known to enhance distractibility when performing
13	a routine task [156]. We have also found the degradation of performance on a spatial
14	configuration visual search task to be significantly worse on the first night shift as compared to
15	subsequent night shifts, with participants exhibiting a significantly greater number of attentional
16	failures on the first night of an overnight work episode [9].
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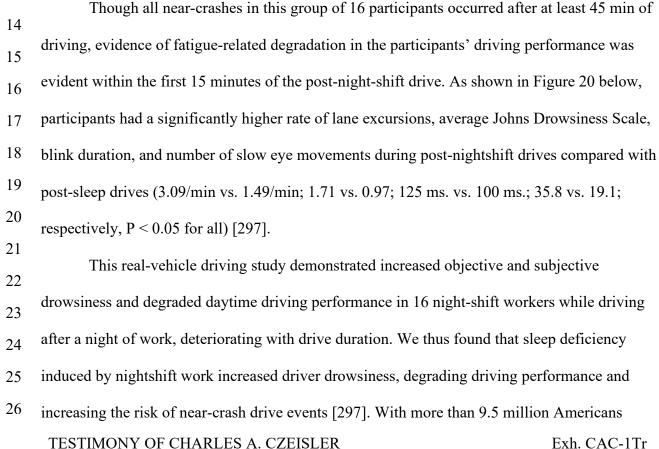


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	vehicle in sixteen night-shift workers driving during the daytime [297]. Each of those workers
1	completed two 2-hour daytime driving testing sessions on a closed driving track operated by the
2 3	Liberty Mutual Research Institute for Safety in Hopkinton, Massachusetts. The sessions all
4	began between 9:30 a.m. and 2:30 P.M. and were time-matched for each individual to control for
5	potential time-of-day effects. In one instance, the drive testing session occurred after a prior
6	night of sleep averaging 7.6 ± 2.4 h in duration, without having worked on the night shift (the
7	post-sleep condition). In another instance, the drive testing session occurred after a prior night of
8	work averaging 8.3 ± 4.1 h in duration (the post-night-shift condition). Participants in the post
9	night-shift condition reported obtaining an average (\pm SD) of 0.4 \pm 1.1 h of sleep between the
10 11	start of their overnight shift and the start of their post-night-shift drive (in three cases this sleep
12	occurred during the night shift, whereas in one case it occurred between the end of the night shift
13	and the start of the post-night shift-drive). As shown in the Figure 19 below, we found that
14	eleven near-crashes occurred in 6 of 16 post night shift drives (37.5%), and that 7 of 16 post
15	night-shift drives (43.8%) were terminated early for safety reasons, compared with zero near-
16	crashes or early drive terminations during 16 post sleep drives (Fishers exact: P=0.0088 and
17	P=0.0034, respectively). No near-crashes occurred during driving after a night of sleep; 11
18 19	occurred during driving after night-work [297] (Figure 19).
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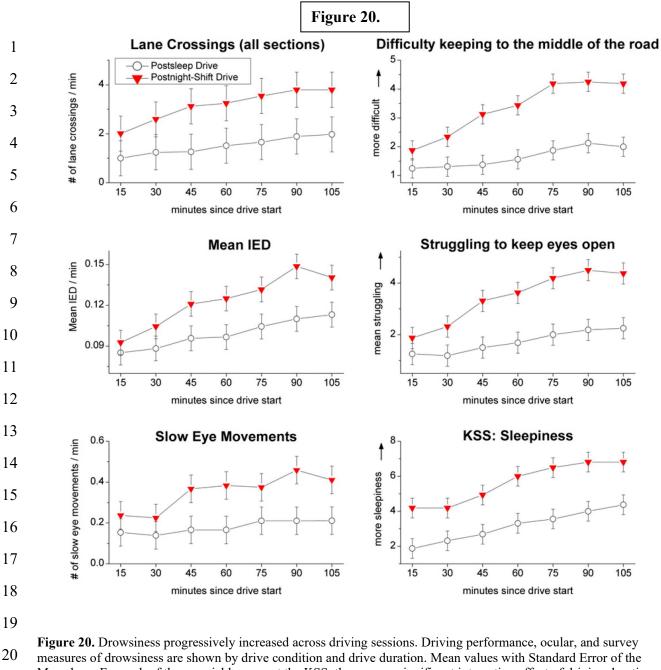




working overnight or rotating shifts and one-third of United States commutes exceeding 30 min, these results have implications for traffic and occupational safety.

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3	In 2012, while we were in the process of analyzing the data that we had collected in 2011
4	for the Lee et al. study that we published in 2016 [297], ABC News requested that we test the
5	driving performance of ABC News journalist and correspondent Mr. Ron Claiborne after a night
6	without sleep using the same recording methods for the driving test sessions that we used in
7	2011, to evaluate the impact of increased homeostatic sleep pressure on his ability to drive using
8	the same methods that we used in the experiment described in [297] and that are illustrated in
9	Figure 20 below.
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Mean bars. For each of these variables except the KSS, there was a significant interaction effect of driving duration (driving block 1–7) by condition (P < 0.05). Condition alone had a significant effect on the KSS (P < 0.05). Tick marks indicate driving breaks, as in Figure 19. Figure 20 and associated legend taken from [297].

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Figure 21. ABC News correspondent Mr. Ronald driving a specially instrumented van during a test driving session while outfitted for polysomnographic recording and eye tracking equipment. Investigator-observer in the front passenger seat can activate emergency braking maneuvers to prevent a crash. Seated in the second row, Dr. Charles Czeisler and Dr. Michael Lee are monitoring Mr. Claiborne's electroencephalogram and electrooculogram during the drive.

15 As shown in the video report that he filed on ABC News Nightline on December 3, 2012,

16 ABC News, *Sleepy Drivers Can Doze Unknowingly*, YouTube (Dec. 3, 2012),

¹⁷ <u>https://www.youtube.com/watch?v=JwCnJZyU70M</u>, and on ABC World News Tonight with

18 Diane Sawyer, *Micro-sleep May be to Blame for Accidents*,

20 <u>https://abcnews.go.com/WNT/video/micro-sleep-blame-accidents-17871728</u>, (accessed May 26,

2018), Mr. Claiborne was very impaired when driving after he had missed a night of sleep. Like

- 22 most sleep-deprived individuals, Mr. Claiborne reported that he thought he could handle the
- 23 driving after missing a night of sleep. However, about a half-hour into the drive, Mr. Claiborne
- ²⁴ fell asleep at the wheel and drove completely off the roadway on the closed track (see Figures 22

²⁵ and 23 below).

26

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Figure 22. Photograph of ABC News correspondent Mr. Ronald Claiborne when he was asleep at the wheel while driving an instrumented van occupied by himself and three other people on a test track in Hopkinton, MA, while he was being recorded in a driving test session after missing one night sleep.



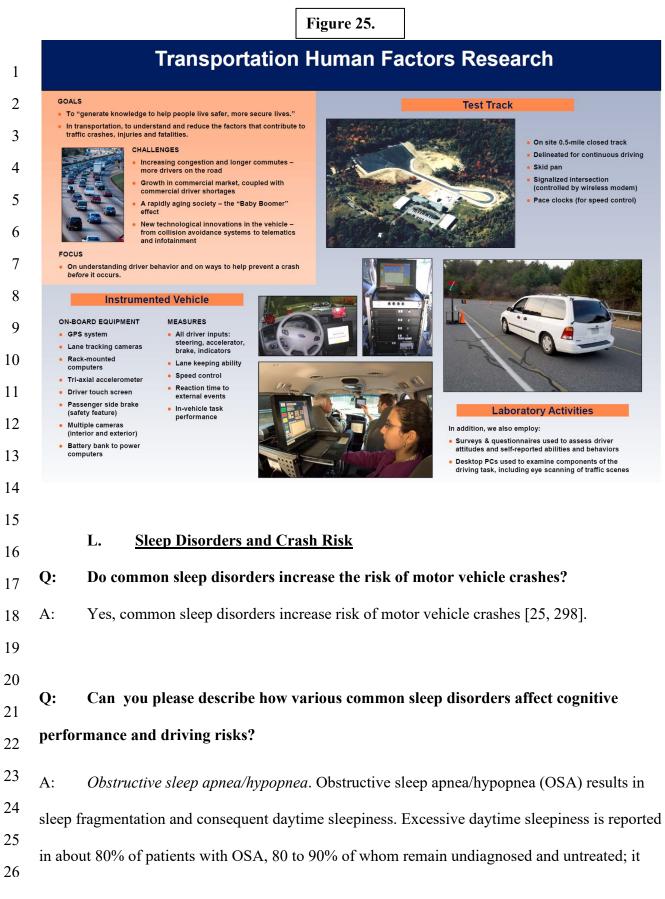
Figure 23. Photograph from the instrumented van driven by ABC News correspondent Mr. Ronald Claiborne and occupied by himself and three other people as it drove off the roadway completely on a test track in Hopkinton, Massachusetts, while Mr. Claiborne was being recorded by Dr. Charles Czeisler and colleagues in a driving test session after a night without sleep.

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Recognizing that it was unsafe to continue driving, Mr. Claiborne decided to terminate 1 the 2-hour drive testing session early. After terminating the drive testing session, Mr. Claiborne 2 forgot to put the vehicle in "Park," and had to be reminded to do so by an investigator before 3 leaving the driver's seat of the vehicle. 4 During Mr. Claiborne's half-hour drive testing session, polysomnographic (PSG) 5 6 recording-including electroencephalographic ("brain wave" or EEG) recording-using 7 electrodes attached to the face and scalp (illustrated in Figure 24 below) revealed that Mr. 8 Claiborne experienced 22 attentional failures and/or micro-sleep episodes, though he 9 remembered only two of them. 10 11 Sleepy Drivers Can Dose Unknowingly Figure 24. 12 13 14 15 16 17 18 19 20 abc N EWS 21 2:16 / 4:48 22 Figure 24. Polysomnographic (PSG) recording equipment used to monitor the electroencephalogram (EEG) and 23 electrooculogram (EOG) of ABC News Correspondent Mr. Ronald Claiborne. 24 25 26

	Thus, despite being hooked up to all of our monitoring equipment, and despite the
1	presence of television news crews filming at multiple positions along the track, and despite the
23	presence of scientist observers in the car, including myself, Mr. Claiborne recapitulated all of the
4	findings that we observed in the 16 participants that we had carefully evaluated in the
5	randomized study that we later published [297]. The Liberty Mutual Research Institute for Safety
6	test track, located in Hopkinton, Massachusetts, on which we conducted the driving test sessions
7	is illustrated in Figure 25 below.
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often takes several years from the time a patient presents to a physician with symptoms until the 1 correct diagnosis is made [1]. Performance related to executive functions such as verbal fluency, 2 planning and sequential thinking-the purview of the frontal cortex, an area of the brain that 3 imaging studies has revealed to be exquisitely sensitive to sleep loss-is very adversely affected 4 by OSA [299]. Vigilance and the ability to sustain attention are also degraded in patients with 5 6 OSA. Reaction times on a sustained attention task in patients with mild to moderate OSA have 7 been reported to be comparable to or worse than those of a young adult with a blood alcohol 8 concentration of 0.080 g/dL [300].

Untreated patients with OSA perform much more poorly in a driving simulator (in terms 10 of lane deviations, tracking errors, off-road events and collisions with obstacles) and have a 11 markedly increased in risk for having a motor vehicle crash [25, 301-314]. Two systematic 12 reviews reported that the majority of studies found that patients with obstructive sleep apnea had 13 14 a motor vehicle crash risk that was 2 to 3 times higher than those without obstructive sleep apnea 15 [315, 316], with individual studies estimating the crash risk to be more than 6-fold higher in 16 patients with obstructive sleep apnea [311]. We found that nonadherence with an employer-17 mandated sleep apnea treatment greatly increased the risk of serious truck crashes [317], and that 18 implementation of a sleep health education and sleep disorders screening and referral for 19 treatment program significantly reduced injury rate and reduced disability day usage by 46% 20 21 [318, 319]. 22

Insomnia. Insomnia is the most common sleep disorder, with symptoms of disturbed
 sleep affecting most adults at some time in their lives [1]. The clinical definition of insomnia is a
 chronic (> 6 months) complaint of latency to sleep onset or awakenings from sleep of at least a

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half hour occurring at least three times per week. The prevalence of insomnia associated with a 1 daytime complaint of increased sleepiness or fatigue is 8% to 18% [320]. Broadly, insomnia may 2 be psychophysiologic or secondary to the sleep environment, an irregular sleep schedule [321], a 3 sleep-related movement disorder, a psychiatric condition, a medical condition, a medication or 4 substance, or a circadian rhythm sleep disorder such as shift work disorder [1]. Insomnia in non-5 6 depressed adults is a risk factor for depression [322]. Insomnia syndrome, which includes 7 increased daytime sleepiness, is associated with an increased rate of work absenteeism, reduced 8 productivity and increased risk of accidents [1, 323].

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Shift Work Sleep-Wake Disorder. As reviewed above, night shift work is often associated with insomnia during daytime sleep. Excessive sleepiness during night work or insomnia during 12 day sleep are the cardinal symptoms of a condition known as Circadian Rhythm Sleep Disorder 13 14 (CRSD), Shift Work Type (Shift Work Disorder, SWD) [229], which recent research indicates 15 may have a genetic basis in some individuals [230]. While most employees nod off or fall asleep 16 regularly while working at night or while commuting to or from night work [233], about 5–10% 17 of such workers suffer chronically from moderate to severe SWD [324]. One-third of nurses who 18 work at night report that they fall asleep in the hospital every week of night work (302), with half 19 admitting that they fall asleep at the wheel while driving to or from the hospital; twice as many 20 21 night-working nurses reported making medication errors or having motor vehicle crashes as 22 compared to those who worked the day and evening shifts, but did not work at night [261]. Long 23 work weeks and extended duration work shifts that do not allow sufficient time for sleep can 24 elicit attentional failures during work in most if not all employees [8], effectively inducing SWD 25 in all workers in the same way that depriving people of food will eventually induce malnutrition 26

in all people who are starved. Exposure to less extreme work schedules reveals considerable variability in the ability of individuals to cope with the demands of work schedules that require employees to work during times that they would otherwise be asleep.

A number of countermeasures and treatments can be effective in treating the symptoms of Shift Work Disorder, including improvements in work scheduling, use of properly timed use of light, administration of nutritional supplements such as caffeine and melatonin, and use of pharmacologic agents such as modafinil, as we have demonstrated in my laboratory and our field study investigations [8, 43, 47, 231-236], although none eliminate them altogether.

There are a number of other factors that can have an impact on alertness and 10 performance. These include the length of time on task, the level of environmental stimulation, 11 the level of physical activity, posture, the level of task stimulation/novelty, many other medical 12 conditions, and the use of pharmacologic agents with stimulant or hypnotic properties. Caffeine 13 14 administration can reduce the adverse impact of misalignment of circadian phase as well as 15 increased homeostatic pressure on neurobehavioral performance [234]. It should be noted that 16 caffeine, which has a 6- to 9-hour half-life, may interfere with recovery sleep. Neither caffeine 17 nor other wake-promoting therapeutics are a substitute for sleep. In a study of fatal-to-the-driver 18 truck crashes, the National Transportation Safety Board (NTSB) found that plasma caffeine 19 levels were highest in drivers involved in fatigue-related crashes [102, 277]. The NTSB 20 21 interpreted high plasma caffeine data from those drivers as indicating that the drivers were taking 22 caffeine to try to combat their fatigue. Unfortunately, even high levels of caffeine were 23 insufficient to save those drivers from the effects of fatigue, which the NTSB found to be the 24 leading cause of fatal crashes in those trucker drivers, accounting for 31 percent of fatalities, 25

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equal to the fraction of fatal truck crashes caused by both drugs and alcohol combined [102, 277].

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M. A Summary of the Evaluation Performed at the Brigham and Women's Hospital Division of Sleep and Circadian Disorders Evaluating Puget Sound **Pilots' Scheduling Efficiency and Making Recommendations for Improvement**

6 **O**: Please describe the scope of work performed by you and your colleagues at Brigham 7 and Women's Physicians Organization for the Puget Sound Pilots.

8 A: Over the course of one year that began with our engagement on June 1, 2021, we 9 reviewed the previous studies of pilot fatigue issues involving the Puget Sound pilots, the PSP 10 Operating Rules, conducted interviews of PSP dispatchers and a series of pilot focus groups 11 involving 68% of the PSP pilot corps, examined large data files provided by PSP, transformed 12 that data into usable databases for analysis and queried PSP management when questions arose 13 14 regarding the data. During the course of our work, PSP implemented multiple new efficiency 15 measures between the fall of 2021 and spring of 2022. Ultimately, we produced a report with 11 16 recommendations, all but three of which have already been implemented. 17

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Q: What previous studies had been performed related to fatigue risk management for 19 the Puget Sound Pilots prior to beginning your most recent work for PSP in June 2021? 20 21 A: There were two. We reviewed the San Jose State Foundation/NASA Ames Research 22 Center Fatigue Countermeasure Laboratory Puget Sound Pilot Fatigue Study Report (2019) and 23 the previous presentation that I made to the Washington Board of Pilotage Commissioners in 24

2017. A copy of the 2017 presentation to the Board is Exhibit CAC-03.

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O: How would you describe the nature of the overall effort by the Puget Sound Pilots to 1 address the directive from the Washington Utilities and Transportation Commission to 2 thoroughly examine their scheduling practices in order to improve overall work efficiency? 3 Based upon the time that PSP leadership and its members devoted to examining all of the A: 4 potential means of improving efficiency and the group's implementation to date of no less than 5 6 eight efficiency measures, I would characterize PSP's response to the UTC directive as both 7 robust and clearly in good faith.

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Q: In an overview fashion, how would you describe the nature of the efficiency measures both recommended and now adopted by PSP?

The efficiency measures adopted by PSP over a period of approximately 10 months range A: 12 from modest incremental efficiency gains to highly significant improvement of non-watch 13 14 efficiency. The first recommendation in our report is an example of a modest efficiency gain. In 15 this instance, PSP changed its rule requiring rest following attendance at a meeting. The new 16 operating rule allows for direct dispatch of a pilot to a pilotage assignment following the meeting 17 where the combination of the meeting and assignment can be concluded within 13 hours from 18 the start of the meeting. Further, in order to increase the efficiency gain associated with this rule 19 change, PSP began greater utilization of virtual meetings and established hard stops for meetings 20 21 when possible to facilitate dispatcher awareness of when pilots involved in a meeting will be 22 available for efficient assignment to a follow-on job. An example of a major operating rule 23 change that we believe will deliver significant efficiencies is the adoption by PSP of a rolling 24 start for pilots coming on and going off their watch intervals. For decades, PSP used the 25 scheduling approach that is common among pilot groups to divide their pilot workforce into two 26

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watches that changeover on a single day. In other words, one watch of half the pilot corps that is 1 on-duty for a specified period is replaced by the other watch of half the pilot corps that has been 2 off-duty on a single changeover day. Following extensive work back and forth between PSP and 3 me and my colleagues, PSP created and implemented a five-watch rolling start schedule with 4 starts on Tuesdays and Thursdays. Our evaluation of this new rolling start schedule shows that it 5 6 increases on-watch efficiency significantly. As we note in the report: "On this schedule, there are 7 never fewer people scheduled to be on duty than in the standard non-rolling schedule. We 8 estimate on-duty capacity increases by at least one schedule pilot for 99% of days and by two or 9 more schedule pilots for 29% of days." 10 11 **Q**: The last three recommendations in your report have not yet been implemented. 12 These include 72-hour no assignment shoulders on each side of a pilot's on-watch interval 13 14 and the pilot's next rolling start, an annual cap on the maximum number of callback jobs 15 and limiting pilot workload by increasing the number of licensed pilots. What is your 16 understanding about why these recommendations have not been implemented? 17 A: Two of these recommendations—72-hour no assignment shoulders at the beginning and 18 end of each pilot's off-watch interval and placing an annual cap on the number of callbacks any 19 pilot can perform—are both potential new operating rules that PSP could implement, but I am 20 21 advised by PSP leadership that it is not possible to do so at this time. This is 22 because implementing one or both of these new rules would significantly increase ship delays. 23 At present, PSP is experiencing high levels of callbacks, utilizing off-duty pilots to work in order 24 to avoid ship delays. During the two years preceding the pandemic, callbacks as a percentage of 25 total assignments were at 18% in 2018 and 19.7% in 2019. Callbacks as a percentage of total 26

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assignments moderated with lower traffic levels in 2020 and 2021 due to Covid-19, but are now 1 back up with rebounding traffic in 2022 to levels above 15% during the first five months of this 2 year, which I consider to be unsafe from a fatigue risk management standpoint. Given the 3 transportation-critical nature of PSP pilot work, the unpredictable nature of assignments during a 4 pilot's on-watch interval and the prevalence of night work throughout that interval, it is 5 6 important for each PSP pilot to use their off-watch interval for what it is called "respite" by PSP. 7 In my opinion, the number of callbacks that a pilot group like PSP should experience throughout 8 the year should average 5% or less in order to maximize the level of alertness and rest necessary 9 to perform this challenging work. 10 11 **Q**: Please explain why extensive use of callbacks utilizing off-watch pilots is not 12 acceptable from a fatigue risk best practices standpoint. 13 14 A: Over the past decade, research has demonstrated that an extended duration of time off 15 duty, without scheduled work time, is required to recover from the chronic sleep deficiency and 16 disruption of circadian rhythms induced by the types of erratic and demanding schedules 17 required of the Puget Sound Pilots. For example, as demonstrated in 2011 in a study conducted 18 by Van Dongen et al. [325] from the Sleep and Performance Research Center at Washington 19 State University in Spokane and published in the peer-reviewed journal *Sleep*, a 34-hour restart 20 21 break, which was efficacious for maintaining neurobehavioral functioning from a pre-restart duty 22 cycle to a post-restart duty cycle after daytime work shifts, was not sufficient or efficacious for 23 maintaining neurobehavioral reaction-time performance and other objective neurobehavioral 24 functioning profiles from one duty cycle to the next after nighttime work shifts. Moreover, the 25 scientists found that subjective sleepiness did not reliably track objective neurobehavioral 26

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deficits observed in the nighttime duty condition. Thus, extensive use of callbacks utilizing offwatch pilots—especially when this *averages* 15% across more than 50 pilots and is not capped,
such that some pilots may be working multiple times during their off-duty recovery intervals—
will result in conditions in which maritime pilots will **begin** their next work-shift interval preloaded with fatigue, lacking resilience for coping with the very demanding schedule that each
sequence of shifts requires. This will increase the risk of a fatigue-related adverse safety event.

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Q:

Did you receive any comments from PSP regarding your final report?

A: Yes. PSP leadership questioned the accuracy of the following sentence on page 5 of our 10 report: "Given that the PSP management affirmed that one type of assignment is not inherently 11 more dangerous than another, the work hour limit does not need to limit multiple shifts to just 12 harbor assignments." I concur that the reference to one type of assignment not being inherently 13 14 more dangerous than another is ambiguous. In reality, however, the challenge of any particular 15 pilotage assignment is a function of many different factors such as ship type, weather, current, 16 proximity to other vessel traffic, etc. Hence, the difficulty level of any particular pilotage 17 assignment in Puget Sound is also a function of these many factors. The same holds true for the 18 difficulty that an airline pilot faces on any given flight. There is a significant difference between 19 a flight in perfect weather compared to one with weather or wind factors which substantially 20 21 increase the difficulty of the pilot's job safely completing a particular flight. 22

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III. CONCLUSION.

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Q: Does this conclude your testimony?

26 A: Yes.

IV. <u>REFERENCES</u>.

- 1. Institute of Medicine: **Sleep Disorders and Sleep Deprivation: An Unmet Public Health Problem**. Washington, D.C.: Institute of Medicine of the National Academies; The National Academies Press; 2006.
- 2. Cajochen C, Khalsa SBS, Wyatt JK, Czeisler CA, Dijk DJ: **EEG and ocular correlates** of circadian melatonin phase and human performance decrements during sleep loss. *American Journal of Physiology* 1999, 277:R640-R649.
- 3. Stern HS, Blower D, Cohen ML, Czeisler CA, Dinges DF, Greenhouse JB, Guo F, Hanowski RJ, Hartenbaum NP, Krueger GP *et al*: **Data and methods for studying commercial motor vehicle driver fatigue, highway safety and long-term driver health**. *Accident; analysis and prevention* 2018.
- 4. Rutenfranz J, Aschoff J, Mann H: **The effects of a cumulative sleep deficit, duration of preceding sleep period and body-temperature on multiple choice reaction time**. In: *Aspects of Human Efficiency: Diurnal Rhythm and Loss of Sleep*. edn. Edited by Colquhoun WP. London: The English Universities Press Ltd; 1972: 217-229.
- 5. Tilley AJ, Wilkinson RT, Warren PSG, Watson B, Drud M: **The sleep and performance** of shift workers. *Human Factors* 1982, **24**:629-641.
- 6. Tomasi D, Wang RL, Telang F, Boronikolas V, Jayne MC, Wang GJ, Fowler JS, Volkow ND: Impairment of attentional networks after 1 night of sleep deprivation. *Cereb Cortex* 2009, **19**(1):233-240.
- 7. Barger LK, Ayas NT, Cade BE, Cronin JW, Rosner B, Speizer FE, Czeisler CA: **Impact** of extended-duration shifts on medical errors, adverse events, and attentional failures. *PLoS Med* 2006, **3**(12):e487.
- 8. Lockley SW, Cronin JW, Evans EE, Cade BE, Lee CJ, Landrigan CP, Rothschild JM, Katz JT, Lilly CM, Stone PH *et al*: **Effect of reducing interns' weekly work hours on sleep and attentional failures**. *NEnglJMed* 2004, **351**(18):1829-1837.
- 9. Santhi N, Horowitz TS, Duffy JF, Czeisler CA: Acute sleep deprivation and circadian misalignment associated with transition onto the first night of work impairs visual selective attention. *PLoS ONE* 2007, **2**(11):e1233.
- 10. Scheer FA, Shea TJ, Hilton MF, Shea SA: An Endogenous Circadian Rhythm in Sleep Inertia Results in Greatest Cognitive Impairment upon Awakening during the Biological Night. *JBiol Rhythms* 2008, **23**(4):353-361.
- 11. Wertz AT, Ronda JM, Czeisler CA, Wright Jr. KP: Effects of sleep inertia on cognition. *Journal of the American Medical Association* 2006, **295**(2):163-164.
- Jewett ME, Wyatt JK, Ritz-De Cecco A, Khalsa SB, Dijk DJ, Czeisler CA: Time course of sleep inertia dissipation in human performance and alertness. *J Sleep Res* 1999, 8:1-8.
- 13. Cohen DA, Wang W, Wyatt JK, Kronauer RE, Dijk DJ, Czeisler CA, Klerman EB: Uncovering residual effects of chronic sleep loss on human performance. *Science Translational Medicine* 2010, **2**(14):14ra13.

- 14. Lim J, Dinges DF: Sleep deprivation and vigilant attention. *Ann N Y Acad Sci* 2008, 1129:305-322.
- 15. Durmer JS, Dinges DF: Neurocognitive consequences of sleep deprivation. SeminNeurol 2005, 25(1):117-129.
- 16. Van Dongen HPA, Maislin G, Mullington JM, Dinges DF: The cumulative cost of additional wakefulness: Dose-response effects on neurobehavioral functions and sleep physiology from chronic sleep restriction and total sleep deprivation. *Sleep* 2003, **26**(2):117-126.
- 17. Czeisler CA: Medical and genetic differences in the adverse impact of sleep loss on performance: ethical considerations for the medical profession. *Trans Am Clin Climatol Assoc* 2009, **120**:249-285.
- 18. Czeisler CA: **Sleep Deficit. The Performance Killer**. In: *Harvard Business Review*. Edited by Fryer B, vol. 84; 2006: 53-59.
- National Commission on Sleep Disorders Research: Wake up america: A national sleep alert. In. Palo Alto, CA: Stanford University Sleep Disorders Clinic and Research Center; 1992: 1-18.
- 20. Hafner M, Stepanek M, Taylor J, Troxel WM, van Stolk C: Why sleep matters the economic costs of insufficient sleep: A cross-country comparative analysis. In. Santa Monica, CA; 2016.
- 21. Collision Analysis Working Group, Federal Rail Administration(CAWG): **65 Main-Track Train Collisions, 1997 through 2002: Review, Analysis, Findings, and Recommendations**. In., vol. August 2006: Federal Railroad Administration; 2006.
- 22. Duffy JF, Zitting KM, Czeisler CA: The Case for Addressing Operator Fatigue. *Rev Hum Factors Ergon* 2015, **10**(1):29-78.
- 23. Marcus JH, Rosekind MR: Fatigue in transportation: NTSB investigations and safety recommendations. *Injury Prevention* 2016.
- 24. Tefft BC: Prevalence of Motor Vehicle Crashes Involving Drowsy Drivers, United States, 2009-2013. In. Washington DC: AAA Foundation for Traffic Safety; 2014.
- 25. Gottlieb DJ, Ellenbogen JM, Bianchi MT, Czeisler CA: Sleep deficiency and motor vehicle crash risk in the general population: a prospective cohort study. *BMC Med* 2018, **16**(1):44.
- 26. Barger LK, Rajaratnam SMW, Cannon CP, Lukas MA, Im K, Goodrich EL, Czeisler CA, O'Donoghue ML: Short Sleep Duration, Obstructive Sleep Apnea, Shiftwork, and the Risk of Adverse Cardiovascular Events in Patients After an Acute Coronary Syndrome. J Am Heart Assoc 2017, 6(10).
- 27. World Health Organization International Agency fo Research on Cancer: IARC Monographs on the Evaluation of Carcinogenic Risk to Humans. 2007.
- 28. Czeisler CA: Human circadian physiology: internal organization of temperature, sleep-wake, and neuroendocrine rhythms monitored in an environment free of time cues. *Dissertation*. Stanford University; 1978.
- 29. Czeisler CA, Jewett ME: Human circadian physiology: Interaction of the behavioral rest activity cycle with the output of the endogenous circadian pacemaker. In: *Handbook of Sleep Disorders*. edn. Edited by Thorpy MJ. New York: Marcel Dekker, Inc.; 1990: 117-137.

- 30. Johnson MP, Duffy JF, Dijk DJ, Ronda JM, Dyal CM, Czeisler CA: Short-term memory, alertness and performance: A reappraisal of their relationship to body temperature. *J Sleep Res* 1992, 1:24-29.
- 31. Allan JS, Czeisler CA: **Persistence of the circadian thyrotropin rhythm under constant conditions and after light-induced shifts of circadian phase**. *J Clin Endocrinol Metab* 1994, **79**:508-512.
- 32. Waldstreicher J, Duffy JF, Brown EN, Rogacz S, Allan JS, Czeisler CA: Gender differences in the temporal organization of prolactin (PRL) secretion: Evidence for a sleep-independent circadian rhythm of circulating PRL levels a clinical research center study. J Clin Endocrinol Metab 1996, 81(4):1483-1487.
- 33. Dijk DJ, Czeisler CA: **Paradoxical timing of the circadian rhythm of sleep propensity** serves to consolidate sleep and wakefulness in humans. *Neurosci Lett* 1994, **166**(1):63-68.
- 34. El-Hajj Fuleihan G, Klerman EB, Brown EN, Choe Y, Brown EM, Czeisler CA: The parathyroid hormone circadian rhythm is truly endogenous-a general clinical research center study. *J Clin Endocrinol Metab* 1997, **82**:281-286.
- 35. Czeisler CA, Klerman EB: Circadian and sleep-dependent regulation of hormone release in humans. *Recent Prog Horm Res* 1999, **54**:97-132.
- 36. Dijk DJ, Neri DF, Wyatt JK, Ronda JM, Riel E, Ritz-De Cecco A, Hughes RJ, Elliott AR, Prisk GK, West JB et al: Sleep, performance, circadian rhythms, and light-dark cycles during two space shuttle flights. Am J Physiol Regul Integr Comp Physiol 2001, 281:R1647-R1664.
- 37. Khalsa SBS, Conroy DA, Duffy JF, Czeisler CA, Dijk DJ: **Sleep- and circadian - dependent modulation of REM density**. *J Sleep Res* 2002, **11**:53-59.
- 38. Cajochen C, Wyatt JK, Czeisler CA, Dijk DJ: Separation of circadian and wake duration-dependent modulation of EEG activation during wakefulness. *Neurosci* 2002, **114**(4):1047-1060.
- 39. Dijk DJ, Neri DF, Wyatt JK, Ronda JM, Riel E, Ritz-De Cecco A, Hughes RJ, Czeisler CA: Sleep, circadian rhythms, and performance during space shuttle missions. In: *The Neurolab spacelab mission: Neuroscience research in space*. edn. Edited by Buckey Jr JC, Homick JL. Houston, Texas: NASA, Lyndon B. Johnson Space Center; 2002: 211-221.
- 40. Van Cauter E, Buxton OM: **Circadian modulation of endocrine secretion**. In: *Handbook of Neurophysiology*. edn. Edited by Takahashi J, Turek FW, Moore RY; 2001: 685-714.
- 41. Horowitz TS, Cade BE, Wolfe JM, Czeisler CA: Searching night and day: a dissociation of effects of circadian phase and time awake on visual selective attention and vigilance. *Psychol Sci* 2003, 14(6):549-557.
- 42. Czeisler CA, Buxton OM, Khalsa SBS: **The human circadian timing system and sleepwake regulation**. In: *Principles and Practices of Sleep Medicine. Volume 4th*, edn. Edited by Kryger MH, Roth T, Dement WC. Philadelphia: W.B.Saunders Company; 2005: 375-394.
- 43. Wyatt JK, Dijk DJ, Ritz-De Cecco A, Ronda JM, Czeisler CA: Sleep facilitating effect of exogenous melatonin in healthy young men and women is circadian-phase dependent. *Sleep* 2006, **29**:609-618.

- 44. Czeisler CA, Gooley JJ: **Sleep and circadian rhythms in humans**. *Cold Spring Harb Symp Quant Biol* 2007, **72**:579-597.
- 45. Scheer FA, Hilton MF, Mantzoros CS, Shea SA: Adverse metabolic and cardiovascular consequences of circadian misalignment. *Proc Natl Acad Sci U S A* 2009, **106**(11):4453-4458.
- 46. Zitting KM, Vujovic N, Yuan RK, Isherwood CM, Medina JE, Wang W, Buxton OM, Williams JS, Czeisler CA, Duffy JF: **Human resting energy expenditure varies with circadian phase**. *Current Biology* 2018, **28**(22):3685-3690 e3683.
- 47. Czeisler CA, Johnson MP, Duffy JF, Brown EN, Ronda JM, Kronauer RE: **Exposure to bright light and darkness to treat physiologic maladaptation to night work**. *N Engl J Med* 1990, **322**:1253-1259.
- 48. Czeisler CA: **The effect of light on the human circadian pacemaker**. In: *Circadian Clocks and Their Adjustment*. edn. Edited by Waterhouse JM. Chichester (Ciba Found. Symp. 183): John Wiley and Sons, Inc.; 1995: 254-302.
- 49. Czeisler CA, Wright Jr. KP: **Influence of light on circadian rhythmicity in humans**. In: *Neurobiology of Sleep and Circadian Rhythms*. edn. Edited by Turek FW, Zee PC. New York: Marcel Dekker, Inc.; 1999: 149-180.
- 50. Czeisler CA, Kronauer RE, Allan JS, Duffy JF, Jewett ME, Brown EN, Ronda JM: Bright light induction of strong (type 0) resetting of the human circadian pacemaker. *Science* 1989, **244**:1328-1333.
- 51. Jewett ME, Kronauer RE, Czeisler CA: Light-induced suppression of endogenous circadian amplitude in humans. *Nature* 1991, **350**:59-62.
- 52. Czeisler CA, Chiasera AJ, Duffy JF: **Research on sleep, circadian rhythms and aging:** Applications to manned spaceflight. *Experimental Gerontology* 1991, **26**:217-232.
- 53. Shanahan TL, Czeisler CA: Light exposure induces equivalent phase shifts of the endogenous circadian rhythms of circulating plasma melatonin and core body temperature in men. *J Clin Endocrinol Metab* 1991, **73**:227-235.
- 54. Duffy JF, Kronauer RE, Czeisler CA: **Phase-shifting human circadian rhythms:** Influence of sleep timing, social contact and light exposure. *J Physiol (Lond)* 1996, **495**(1):289-297.
- 55. Jewett ME, Kronauer RE, Czeisler CA: **Phase-amplitude resetting of the human** circadian pacemaker via bright light: A further analysis. *JBiol Rhythms* 1994, **9**(3-4):295-314.
- 56. Jewett ME, Rimmer DW, Duffy JF, Klerman EB, Kronauer RE, Czeisler CA: Human circadian pacemaker is sensitive to light throughout subjective day without evidence of transients. *American Journal of Physiology Regulatory, Integrative, and Comparative Physiology* 1997, **273**:R1800-R1809.
- 57. Shanahan TL, Zeitzer JM, Czeisler CA: **Resetting the melatonin rhythm with light in** humans. *JBiol Rhythms* 1997, **12**(6):556-567.
- 58. Boivin DB, Czeisler CA: Resetting of circadian melatonin and cortisol rhythms in humans by ordinary room light. *Neuroreport* 1998, **9**(5):779-782.
- 59. Shanahan TL, Kronauer RE, Duffy JF, Williams GH, Czeisler CA: Melatonin rhythm observed throughout a three-cycle bright-light stimulus designed to reset the human circadian pacemaker. *JBiol Rhythms* 1999, 14:237-253.

- 60. Czeisler CA, Khalsa SBS: **The human circadian timing system and sleep-wake regulation**. In: *Principles and Practice of Sleep Medicine. Volume 3rd*, edn. Edited by Kryger MH, Roth T, Dement WC. Philadelphia: W.B. Saunders Company; 2000: 353-375.
- 61. Czeisler CA, Buxton OM: **The Human Circadian Timing System and Sleep-Wake Regulation**. In: *Principles and Practices of Sleep Medicine. Volume 5th*, edn. Edited by Kryger MH, Roth T, Dement WC: Elsevier; 2010: 402-419.
- 62. Czeisler CA, Dijk DJ: **Human circadian physiology and sleep-wake regulation**. In: *Handbook of Behavioral Neurobiology: Circadian Clocks*. edn. Edited by Takahashi JS, Turek FW, Moore RY. New York: Plenum Publishing Co.; 2001: 531-569.
- 63. Münch M, Silva EJ, Ronda JM, Czeisler CA, Duffy JF: **EEG sleep spectra in older** adults across all circadian phases during NREM sleep. *Sleep* 2010, **33**(3):389-401.
- 64. Saper CB, Scammell TE, Lu J: **Hypothalamic regulation of sleep and circadian rhythms**. *Nature* 2005, **437**(7063):1257-1263.
- 65. Dijk DJ, Lockley SW: Integration of human sleep-wake regulation and circadian rhythmicity. *Journal of Applied Physiology* 2002, **92**:852-862.
- Czeisler CA, Weitzman ED, Moore-Ede MC, Zimmerman JC, Knauer RS: Human sleep: Its duration and organization depend on its circadian phase. Science 1980, 210:1264-1267.
- 67. Czeisler CA, Zimmerman JC, Ronda JM, Moore-Ede MC, Weitzman ED: **Timing of REM sleep is coupled to the circadian rhythm of body temperature in man**. *Sleep* 1980, **2**:329-346.
- 68. Dijk DJ, Czeisler CA: Contribution of the circadian pacemaker and the sleep homeostat to sleep propensity, sleep structure, electroencephalographic slow waves, and sleep spindle activity in humans. *Journal of Neuroscience* 1995, **15**(5):3526-3538.
- 69. Dijk DJ, Shanahan TL, Duffy JF, Ronda JM, Czeisler CA: Variation of electroencephalographic activity during non-rapid eye movement and rapid eye movement sleep with phase of circadian melatonin rhythm in humans. J Physiol (Lond) 1997, 505.3:851-858.
- 70. Wyatt JK, Ritz-De Cecco A, Czeisler CA, Dijk DJ: Circadian temperature and melatonin rhythms, sleep, and neurobehavioral function in humans living on a 20-h day. *Am J Physiol Regul Integr Comp Physiol* 1999, **277**:R1152-R1163.
- 71. Dijk DJ, Duffy JF, Riel E, Shanahan TL, Czeisler CA: Ageing and the circadian and homeostatic regulation of human sleep during forced desynchrony of rest, melatonin and temperature rhythms. *J Physiol (Lond)* 1999, **516.2**:611-627.
- 72. Van Dongen HPA, Dinges DF: Circadian Rhythms in Sleepiness, Alertness, and Performance. In: Principles and Practices of Sleep Medicine. Volume Fourth, edn. Edited by Kryger MH, Roth T, Dement WC. Philadelphia: W.B. Saunders; 2005: 435-443.
- 73. Moore RY, Eichler VB: Loss of a circadian adrenal corticosterone rhythm following suprachiasmatic lesions in the rat. *Brain Research* 1972, **42**:201-206.
- 74. Stephan FK, Zucker I: Circadian rhythms in drinking behavior and locomotor activity of rats are eliminated by hypothalamic lesions. *PNAS* 1972, **69**(6):1583-1586.
- 75. Ralph MR, Foster RG, Davis FC, Menaker M: **Transplanted suprachiasmatic nucleus** determines circadian period. *Science* 1990, **247**:975-978.

- 76. Edgar DM, Dement WC, Fuller CA: Effect of SCN lesions on sleep in squirrel monkeys: Evidence for opponent processes in sleep-wake regulation. *Journal of Neuroscience* 1993, **13(3)**:1065-1079.
- 77. Klein DC, Moore RY, Reppert SM: **Suprachiasmatic nucleus: The mind's clock**. New York: Oxford University Press; 1991.
- 78. Silver R, Lesauter J, Tresco PA, Lehman MN: A diffusible coupling signal from transplanted suprachiasmatic nucleus controlling circadian locomotor rhythms. *Nature* 1996, **382**:810-813.
- 79. Weaver DR: The suprachiasmatic nucleus: A 25-year retrospective. *JBiol Rhythms* 1998, **13**(2):100-112.
- 80. Welsh DK, Logothetis DE, Meister M, Reppert SM: Individual neurons dissociated from rat suprachiasmatic nucleus express independently phased circadian firing rhythms. *Neuron* 1995, 14:697-706.
- 81. Moore RY: **Organization of the mammalian circadian system**. In: *Circadian clocks and their adjustment*. edn. Edited by Waterhouse JM. Chichester (Ciba Foundation Symp 183): John Wiley and Sons, Inc.; 1995: 88-99.
- 82. Moore RY, Speh JC, Leak RK: **Suprachiasmatic nucleus organization**. *Cell TissueRes* 2002, **309**(1):89-98.
- 83. Kuhlman SJ: Biological Rhythms Workshop IB: Neurophysiology of SCN Pacemaker Function. Cold Spring Harb Symp Quant Biol 2007, 72:21-33.
- 84. Kalsbeek A, Palm IF, la Fleur SE, Scheer FA, Perreau-Lenz S, Ruiter M, Kreier F, Cailotto C, Buijs RM: **SCN outputs and the hypothalamic balance of life**. *JBiol Rhythms* 2006, **21**(6):458-469.
- 85. Lee ML, Swanson BE, de la Iglesia HO: Circadian timing of REM sleep is coupled to an oscillator within the dorsomedial suprachiasmatic nucleus. *Curr Biol* 2009, **19**(10):848-852.
- 86. Dijk DJ, Duffy JF, Czeisler CA: **Circadian and sleep/wake dependent aspects of subjective alertness and cognitive performance**. *J Sleep Res* 1992, **1**:112-117.
- 87. Czeisler CA, Dijk DJ, Duffy JF: Entrained phase of the circadian pacemaker serves to stabilize alertness and performance throughout the habitual waking day. In: *Sleep Onset: Normal and Abnormal Processes.* edn. Edited by Ogilvie RD, Harsh JR. Washington, D.C.: American Psychological Association; 1994: 89-110.
- 88. Strogatz SH, Kronauer RE, Czeisler CA: Circadian pacemaker interferes with sleep onset at specific times each day: Role in insomnia. *American Journal of Physiology* 1987, **253**:R172-R178.
- 89. Mitler MM, Carskadon MA, Czeisler CA, Dement WC, Dinges DF, Graeber RC: Catastrophes, sleep, and public policy: Consensus report. *Sleep* 1988, 11:100-109.
- 90. Bes F, Ing MJ, Schulz H: Modeling napping, post-lunch dip, and other variations in human sleep propensity. *Sleep* 2009.
- 91. Scheer FA, Cajochen C, Turek FW, Czeisler CA: **Melatonin in the regulation of sleep and circadian rhythms**. In: *Principles and Practice of Sleep Medicine. Volume 4th*, edn. Edited by Kryger MH, Roth T, Dement WC. Philadelphia: W.B. Saunders; 2005: 395-404.
- 92. Scheer FA, Czeisler CA: Melatonin, sleep, and circadian rhythms. *Sleep Med Rev* 2005, **9**(1):5-9.

- 93. Czeisler CA: The Gordon Wilson Lecture: Work hours, sleep and patient safety in residency training. *Trans Am Clin Climatol Assoc* 2006, **117**:159-189.
- 94. Duffy JF, Cain SW, Chang AM, Phillips AJK, Münch MY, Gronfier C, Wyatt JK, Dijk D, Wright KP, Czeisler CA: Sex difference in the near-24-hour intrinsic period of the human circadian timing system. *Proc Nat Acad Sci* 2011, **108**(suppl. 3):15602-15608.
- 95. Gronfier C, Wright Jr. KP, Kronauer RE, Czeisler CA: Entrainment of the human circadian pacemaker to longer-than-24-h days. *Proceedings of the National Academy of Sciences USA* 2007, 104(21):9081-9086.
- 96. Cain SW, Dennison CF, Zeitzer JM, Guzik AM, Khalsa SS, Santhi N, Schoen M, Czeisler CA, Duffy JF: **Sex differences in phase angle of entrainment and melatonin amplitude in humans**. *JBiol Rhythms* 2010, **25**(4):288-296.
- 97. Gooley JJ, Chamberlain K, Smith KA, Khalsa SB, Rajaratnam SM, Van RE, Zeitzer JM, Czeisler CA, Lockley SW: **Exposure to Room Light before Bedtime Suppresses Melatonin Onset and Shortens Melatonin Duration in Humans**. *J Clin EndocrinolMetab* 2011, **96**(3):E463-E472.
- Barinaga M: How jet-lag hormone does double duty in the brain. Science 1997, 277(5325(25)):480.
- 99. Liu C, Weaver DR, Jin X, Shearman LP, Pieschi RL, Gribkoff VK, Reppert SM: Molecular dissection of two distinct actions of melatonin on the suprachiasmatic circadian clock. *Neuron* 1997, **19**:91-102.
- 100. Schwartz WJ, Gross RA, Morton MT: **The suprachiasmatic nuclei contain a tetrodotoxin-resistant circadian pacemaker**. *PNAS* 1987, **84**:1694-1698.
- 101. Czeisler CA, Guilleminault C: **REM sleep: Its temporal distribution**. New York: Raven Press; 1980.
- 102. National Transportation Safety Board: Safety Study: Fatigue, Alcohol, Other Drugs, and Medical Factors in Fatal-to-the-Driver Heavy Truck Crashes (Volume 1). In., vol. NTSB/SS-90/01. Washington, DC: National Transportation Safety Board; 1990: 1-181.
- 103. Langlois PH, Smolensky MH, Hsi BP, Weir FW: **Temporal patterns of reported** single-vehicle car and truck accidents in Texas, USA during 1980-1983. *Chronobiology International* 1985, 2:131-140.
- 104. Harris W: Fatigue, circadian rhythm, and truck accidents. In: *Vigilance Theory, Operational Performance, and Physiological Correlates.* edn. Edited by Mackie R. New York: Plenum; 1977: 133-146.
- 105. Horne JA, Reyner LA: Sleep related vehicle accidents. BMJ 1995, 310:565-567.
- 106. Pack AI, Pack AM, Rodgman E, Cucchiara A, Dinges DF, Schwab CW: Characteristics of crashes attributed to the driver having fallen asleep. *Accident Analysis & Prevention* 1995, **27**(6):769-775.
- 107. Resources, Community and Economic Development Division, General Accounting Office: **Railroad Safety: Engineer Work Shift Length and Schedule Variability**. In.: United States General Accounting Office; Report RCED-92-122, 1992.
- 108. Jewett ME: Models of circadian and homeostatic regulation of human performance and alertness. *Thesis*. Harvard University; 1997.
- 109. Jewett ME, Kronauer RE: Interactive mathematical models of subjective alertness and cognitive throughput in humans. *JBiol Rhythms* 1999, 14(6):588-597.

- 110. Czeisler CA, Duffy JF, Shanahan TL, Brown EN, Mitchell JF, Rimmer DW, Ronda JM, Silva EJ, Allan JS, Emens JS *et al*: **Stability, precision, and near-24-hour period of the human circadian pacemaker**. *Science* 1999, **284**:2177-2181.
- 111. Wright Jr. KP, Hughes R, Kronauer RE, Dijk DJ, Czeisler CA: Intrinsic near-24-h pacemaker period determines limits of circadian entrainment to a weak synchronizer in humans. *Proc Natl Acad Sci USA* 2001, **98**(24):14027-14032.
- 112. Scheer FA, Wright Jr. KP, Kronauer RE, Czeisler CA: **Plasticity of the intrinsic period** of the human circadian timing system. *PLoS ONE* 2007, **2**(8):e721.
- 113. Czeisler CA, Allan JS, Strogatz SH, Ronda JM, Sánchez R, Ríos CD, Freitag WO, Richardson GS, Kronauer RE: **Bright light resets the human circadian pacemaker independent of the timing of the sleep-wake cycle**. *Science* 1986, **233**:667-671.
- 114. Khalsa SBS, Jewett ME, Cajochen C, Czeisler CA: A phase response curve to single bright light pulses in human subjects. *Journal of Physiology (London)* 2003, 549(Pt 3):945-952.
- 115. Boivin DB, Duffy JF, Kronauer RE, Czeisler CA: **Sensitivity of the human circadian** pacemaker to moderately bright light. *JBiol Rhythms* 1994, **9**(3-4):315-331.
- 116. Boivin DB, Duffy JF, Kronauer RE, Czeisler CA: **Dose-response relationships for** resetting of human circadian clock by light. *Nature* 1996, **379**:540-542.
- 117. Zeitzer JM, Dijk DJ, Kronauer RE, Brown EN, Czeisler CA: Sensitivity of the human circadian pacemaker to nocturnal light: Melatonin phase resetting and suppression. *J Physiol* 2000, **526**(3):695-702.
- 118. Shanahan TL, Czeisler CA: **Physiological effects of light on the human circadian pacemaker**. *Semin Perinatol* 2000, **24** 299-320.
- 119. Klerman EB, Shanahan TL, Brotman DJ, Rimmer DW, Emens JS, Rizzo JF, III, Czeisler CA: Photic resetting of the human circadian pacemaker in the absence of conscious vision. *JBiol Rhythms* 2002, **17**(6):548-555.
- 120. Lockley SW, Brainard GC, Czeisler CA: **High sensitivity of the human circadian** melatonin rhythm to resetting by short wavelength light. *J Clin Endocrinol Metab* 2003, **88**(9):4502-4505.
- 121. Zeitzer JM, Khalsa SB, Boivin DB, Duffy JF, Shanahan TL, Kronauer RE, Czeisler CA: **Temporal dynamics of late-night photic stimulation of the human circadian timing system**. *Am J Physiol Regul Integr Comp Physiol* 2005, **289**(3):R839-R844.
- 122. Zaidi FH, Hull JT, Peirson SN, Wulff K, Aeschbach D, Gooley JJ, Brainard GC, Gregory-Evans K, Rizzo III JF, Czeisler CA *et al*: Short-wavelength light sensitivity of circadian, pupillary, and visual awareness in humans lacking an outer retina. *Curr Biol* 2007, 17(24):2122-2128.
- 123. Smith KA, Schoen MW, Czeisler CA: Adaptation of human pineal melatonin suppression by recent photic history. *J Clin Endocrinol Metab* 2004, **89**:3610-3614.
- 124. Chang A, Aeschbach D, Duffy JF, Czeisler CA: Impact of evening use of light-emitting electronic readers on circadian timing and sleep latency. *Sleep* 2012, **35**:A206.
- 125. Chinoy ED, Duffy JF, Czeisler CA: Unrestricted evening use of light-emitting tablet computers delays self-selected bedtime and disrupts circadian timing and alertness. *Physiological Reports* 2018, **6**(10):e13692.

- 126. Dijk DJ, Duffy JF, Czeisler CA: **Contribution of circadian physiology and sleep homeostasis to age-related changes in human sleep**. *Chronobiol Int* 2000, **17**(3):285-311.
- Wright Jr. KP, Hull JT, Czeisler CA: Relationship between alertness, performance, and body temperature in humans. Am J Physiol Regul Integr Comp Physiol 2002, 283:R1370-R1377.
- 128. Dorrian J, Rogers NL, Dinges DF: Psychomotor Vigilance Performance: Neurocognitive Assay Sensitive to Sleep Loss. In: Sleep Deprivation Clinical Issues, Pharmacology, and Sleep Loss Effects. edn. Edited by Kushida CA. New York: Marcel Dekker; 2005: 39-70.
- 129. Klein T, Martens H, Dijk DJ, Kronauer RE, Seely EW, Czeisler CA: Chronic non-24hour circadian rhythm sleep disorder in a blind man with a regular 24-hour sleepwake schedule. *Sleep* 1993, 16(4):333-343.
- 130. Boivin DB, Czeisler CA, Dijk DJ, Duffy JF, Folkard S, Minors DS, Totterdell P, Waterhouse JM: Complex interaction of the sleep-wake cycle and circadian phase modulates mood in healthy subjects. Arch Gen Psychiatry 1997, 54(2):145-152.
- 131. Jewett ME, Borbély AA, Czeisler CA: **Biomathematical modeling workshop. May** 118-21, 1999. *JBiol Rhythms* 1999, 14(6):429-630.
- 132. Chee MW, Tan JC, Zheng H, Parimal S, Weissman DH, Zagorodnov V, Dinges DF: Lapsing during sleep deprivation is associated with distributed changes in brain activation. *Journal of Neuroscience* 2008, **28**(21):5519-5528.
- 133. Venkatraman V, Chuah YM, Huettel SA, Chee MW: Sleep deprivation elevates expectation of gains and attenuates response to losses following risky decisions. *Sleep* 2007, **30**(5):603-609.
- 134. Bocca ML, Denise P: Total sleep deprivation effect on disengagement of spatial attention as assessed by saccadic eye movements. *Clin Neurophysiol* 2006, **117**(4):894-899.
- Dawson D, Reid K: Fatigue, alcohol and performance impairment. *Nature* 1997, 388(6639):235.
- 136. Williamson AM, Feyer AM: Moderate sleep deprivation produces impairments in cognitive and motor performance equivalent to legally prescribed levels of alcohol intoxication. *OccupEnvironMed* 2000, **57**:649-655.
- 137. Lamond N, Dawson D: Quantifying the performance impairment associated with fatigue. *J Sleep Res* 1999, **8**(4):255-262.
- 138. Roehrs T, Burduvali E, Bonahoom A, Drake C, Roth T: **Ethanol and sleep loss: a** "dose" comparison of impairing effects. *Sleep* 2003, **26**(8):981-985.
- 139. Falleti MG, Maruff P, Collie A, Darby DG, McStephen M: Qualitative similarities in cognitive impairment associated with 24 h of sustained wakefulness and a blood alcohol concentration of 0.05%. J Sleep Res 2003, 12(4):265-274.
- 140. Powell NB, Schechtman KB, Riley RW, Li K, Troell R, Guilleminault C: The road to danger: the comparative risks of driving while sleepy. *Laryngoscope* 2001, 111(5):887-893.
- 141. Walsh JK, Humm T, Muehlbach MJ, Sugerman JL, Schweitzer PK: Sedative effects of ethanol at night. *Journal of Studies on Alcohol* 1991, **52**(6):597-600.

- 142. Babkoff H, Zukerman G, Fostick L, Ben Artzi E: Effect of the diurnal rhythm and 24 h of sleep deprivation on dichotic temporal order judgment. *J Sleep Res* 2005, 14(1):7-15.
- 143. Killgore WD, Killgore DB, Day LM, Li C, Kamimori GH, Balkin TJ: **The effects of 53** hours of sleep deprivation on moral judgment. *Sleep* 2007, **30**(3):345-352.
- 144. Anderson C, Dickinson DL: Bargaining and trust: the effects of 36-h total sleep deprivation on socially interactive decisions. *J Sleep Res* 2009, 19:54-63.
- 145. Killgore WDS, Balkin TJ, Wesensten NJ: Impaired decision making following 49 h of sleep deprivation. *J Sleep Res* 2006, **15**(1):7-13.
- 146. Doran SM, Van Dongen HP, Dinges DF: Sustained attention performance during sleep deprivation: evidence of state instability. *Arch Ital Biol* 2001, **139**(3):253-267.
- 147. Goel N, Rao H, Durmer JS, Dinges DF: Neurocognitive consequences of sleep deprivation. *Semin Neurol* 2009, **29**(4):320-339.
- 148. Ratcliff R, Van Dongen HPA: Sleep deprivation affects multiple distinct cognitive processes. *Psychon Bull Rev* 2009, **16**(4):742-751.
- 149. Turner TH, Drummond SPA, Salamat JS, Brown GG: Effects of 42 hr of total sleep deprivation on component processes of verbal working memory. *Neuropsychology* 2007, 21(6):787-795.
- 150. Anderson C, Wales AWJ, Horne JA: **PVT lapses differ according to eyes open, closed, or looking away**. *Sleep* 2010, **33**(2):197-204.
- 151. Sanders AF, Reitsma WD: The effect of sleep-loss on processing information in the functional visual field. *Acta Psychologica* 1982, **51**:149-162.
- 152. Russo MB, Kendall AP, Johnson DE, Sing HC, Thorne DR, Escolas SM, Santiago S, Holland DA, Hall SW, Redmond DP: **Visual perception, psychomotor performance, and complex motor performance during an overnight air refueling simulated flight**. *AviatSpace Environ Med* 2005, **76**(7 Suppl):C92-103.
- 153. Kendall AP, Kautz MA, Russo MB, Killgore WDS: Effects of sleep deprivation on lateral visual attention. *Int J Neurosci* 2006, **116**(10):1125-1138.
- 154. Rogé J, Pébayle T, El Hannachi S, Muzet A: Effect of sleep deprivation and driving duration on the useful visual field in younger and older subjects during simulator driving. *Vision Research* 2003, **43**:1465-1472.
- 155. Petrovsky N, Ettinger U, Hill A, Frenzel L, Meyhofer I, Wagner M, Backhaus J, Kumari V: Sleep deprivation disrupts prepulse inhibition and induces psychosis-like symptoms in healthy humans. *The Journal of neuroscience : the official journal of the Society for Neuroscience* 2014, **34**(27):9134-9140.
- 156. Anderson C, Horne JA: Sleepiness enhances distraction during a monotonous task. *Sleep* 2006, **29**(4):573-576.
- McKenna BS, Dickinson DL, Orff HJ, Drummond SPA: The effects of one night of sleep deprivation on known-risk and ambiguous-risk decisions. J Sleep Res 2007, 16(3):245-252.
- 158. Killgore WD, Grugle NL, Balkin TJ: **Gambling when sleep deprived: don't bet on stimulants**. *Chronobiology International* 2012, **29**(1):43-54.
- 159. Wilkinson RT: **Sleep deprivation**. In: *The Physiology of Human Survival*. edn. Edited by Edholm OG, Bacharach AL. New York: Academic Press; 1965: 399-430.

- 160. Taub JM, Berger RJ: **Performance and mood following variations in the length and timing of sleep**. *Psychophysiology* 1973, **10**:559-570.
- Johnson LC: Sleep deprivation and performance. In: *Biological Rhythms, Sleep, and Performance*. edn. Edited by Webb WB. New York: John Wiley & Sons Ltd.; 1982: 111-141.
- 162. Naitoh P: Sleep deprivation in human subjects: A reappraisal. *Waking and Sleeping* 1976, 1:53-60.
- 163. Belenky G, Wesensten NJ, Thorne DR, Thomas ML, Sing HC, Redmond DP, Russo MB, Balkin TJ: Patterns of performance degradation and restoration during sleep restriction and subsequent recovery: A sleep dose-response study. J Sleep Res 2003, 12:1-12.
- 164. Czeisler CA: Quantifying consequences of chronic sleep restriction. *Sleep* 2003, **26** (3):247-248.
- 165. Van Dongen HPA, Dinges DF: Sleep debt and cumulative excess wakefulness. *Sleep* 2003, **26** (3):249.
- 166. Mander BA, Reid KJ, Baron KG, Tjoa T, Parrish TB, Paller KA, Gitelman DR, Zee PC: **EEG measures index neural and cognitive recovery from sleep deprivation**. *Journal of Neuroscience* 2010, **30**(7):2686-2693.
- 167. McHill AW, Hull JT, Wang W, Czeisler CA, Klerman EB: Chronic sleep curtailment, even without extended (>16-h) wakefulness, degrades human vigilance performance. Proceedings of the National Academy of Sciences of the United States of America 2018, 115(23):6070-6075.
- 168. Dinges DF, Pack F, Williams K, Gillen KA, Powell JW, Ott GE, Aptowicz C, Pack AI: Cumulative sleepiness, mood disturbance, and psychomotor vigilance performance decrements during a week of sleep restricted to 4-5 hours per night. *Sleep* 1997, 20(4):267-277.
- 169. Haack M, Mullington JM: Sustained sleep restriction reduces emotional and physical well-being. *Pain* 2005, **119**(1-3):56-64.
- 170. Banks S, Van Dongen H, Dinges D: Response to sleep restriction depends upon preexisting sleep debt. *Sleep* 2007, Abstract Supplement(30):A119.
- 171. Guilleminault C, Phillips R, Dement WC: A syndrome of hypersomnia with automatic behavior. *ElectroencephalogrClin Neurophysiol* 1975, **38**:403-413.
- 172. Guilleminault C, Montplaisir J, Billiard M, Dement WC: Automatic behavior syndrome in patients with excessive daytime somnolence. *ProcWestern EEG Soc* 1975:432-433.
- Duffy JF, Willson HJ, Wang W, Czeisler CA: Healthy older adults better tolerate sleep deprivation than young adults. *Journal of the American Geriatric Society* 2009, 57(7):1245-1251.
- 174. Zitting KM, Münch MY, Cain SW, Wang W, Wong A, Ronda JM, Aeschbach D, Czeisler CA, Duffy JF: Young adults are more vulnerable to chronic sleep deficiency and recurrent circadian disruption than older adults. *Sci Rep* 2018, **8**(1):11052.
- 175. Spiegel K, Leproult R, Van Cauter E: **Impact of sleep debt on metabolic and endocrine function**. *Lancet* 1999, **354**:1435-1439.
- 176. Mander BA, Colecchia EF, Spiegel K, Van Cauter E: **Short Sleep: A risk factor for insulin resistance and obesity**. *Sleep* 2001, **24**(Supp):A74-A75.

- 177. Spiegel K, Sheridan JF, Van Cauter E: Effect of sleep deprivation on response to immunization. *Journal of the American Medical Association* 2002, **288**(12):1471-1472.
- 178. Spiegel K, Tasali E, Penev P, Van Cauter E: **Brief communication: Sleep curtailment** in healthy young men is associated with decreased leptin levels, elevated ghrelin levels, and increased hunger and appetite. *Annals of Internal Medicine* 2004, 141(11):846-850.
- 179. Van Cauter E, Holmback U, Knutson K, Leproult R, Miller A, Nedeltcheva A, Pannain S, Penev P, Tasali E, Spiegel K: **Impact of sleep and sleep loss on neuroendocrine and metabolic function**. *Horm Res* 2007, **67 Suppl 1**:2-9.
- 180. Knutson KL, Spiegel K, Penev P, Van Cauter E: The metabolic consequences of sleep deprivation. *Sleep Med Rev* 2007, **11**(3):163-178.
- 181. Guilleminault C, Billiard M, Montplaisir J, Dement WC: Altered states of consciousness in disorders of daytime sleepiness. *Journal of the Neurological Sciences* 1975, 26:377-393.
- 182. **State of Florida v. Jorge Rosario**. In.: District Court of Appeal of Florida, Second District; 2004.
- 183. Royal D: National survey of distracted and drowsy driving attitudes and behavior: 2002. In., vol. DOT HS 809 566. Washington, D.C.: National Highway Traffic Safety Administration; 2003: 1-61.
- 184. Moore RT, Kaprielian R, Auerbach J: Asleep at the wheel. Report of the Massachusetts commission on drowsy driving. In.; 2009.
- 185. Czeisler CA, Wickwire EM, Akerstedt T, Barger LK, Dement W, Gamble K, Hartenbaum N, Ohayon MM, Phillips B, Strohl K *et al*: **Sleep-deprived motor vehicle operators are unfit to drive: A multidisciplinary expert consensus statement on drowsy driving**. *Sleep Health* 2016, **2**:94-99.
- 186. Garbarino S, Nobili L, Beelke M, De Carli Phy F, Ferrillo F: **The contributing role of** sleepiness in highway vehicle accidents. *Sleep* 2001, 24(2):203-206.
- 187. Dingus TA, Klauer SG, Neale VL, Petersen A, Lee SE, Sudweeks J, Perez MA, Hankey J, Ramsey D, Gupta S *et al*: The 100-car naturalistic driving study; Phase II-Results of the 100-car field experiment. In., vol. DOT HS 810 593. Washington, DC: National Highway Traffic Safety Administration; 2006: 1-352.
- 188. Klauer SG, Dingus TA, Neale VL, Sudweeks JD, Ramsey DJ: **The impact of driver inattention on near-crash/crash risk: an analysis using the 100-car naturalistic driving study data**. In., vol. HS810594. Washington, D.C.: National Highway Traffic Safety Administration; 2006: 1-192.
- 189. Lamond N, Dorrian J, Burgess H, Holmes A, Roach G, McCulloch K, Fletcher A, Dawson D: Adaptation of performance during a week of simulated night work. *Ergonomics* 2004, 47(2):154-165.
- 190. Horne J, Reyner L: Vehicle accidents relating to sleep: A review. *OccupEnvironMed* 1999, **56**:289-294.
- 191. Torsvall L, Åkerstedt T: Disturbed sleep while being on call: an EEG study of ships' engineers. *Sleep* 1988, 11:35-38.
- 192. Richardson GS, Wyatt JK, Sullivan JP, Orav E, Ward A, Wolf MA, Czeisler CA:
 Objective assessment of sleep and alertness in medical house-staff and the impact of protected time for sleep. *Sleep* 1996, 19:718-726.

- 193. Langdon DE, Hartman B: **Performance upon sudden awakening**. *School of Aerospace Medicine* 1961, **62-17**:1-8.
- 194. Hartman BO, Langdon DE: A second study on performance upon sudden awakening. In., vol. TR-65-61. Brooks AFB, TX: U.S. Air Force; 1965: 1-10.
- 195. Hartman BO, Langdon DE, Mc Kenzie RE: A third study on performance upon sudden awakening. In., vol. TR-65-63. Brooks,TX: U.S. Air Force; 1965: 1-4.
- 196. Koulack D, Schultz KJ: Task performance after awakenings from different stages of sleep. *Percept Mot Skills* 1974, **39**:792-794.
- 197. Dinges DF, Orne MT, Orne EC: Assessing performance upon abrupt awakening from naps during quasi-continuous operations. *Behav Res Methods, Instruments & Computers* 1985, 17:37-45.
- 198. Balkin TJ, Badia P: Relationship between sleep inertia and sleepiness: Cumulative effects of four nights of sleep disruption/restriction on performance following abrupt nocturnal awakenings. *Biol Psychol* 1988, **27**:245-258.
- 199. Balkin TJ, Braun AR, Wesensten NJ, Jeffries K, Varga M, Baldwin P, Belenky G, Herscovitch P: The process of awakening: A PET study of regional brain activity patterns mediating the re-establishment of alertness and consciousness. *Brain* 2002, 125:2308-2319.
- 200. Dinges DF: **Sleep Inertia**. In: *Encyclopedia of Sleep and Dreaming*. edn. Edited by Carskadon MA. New York: Macmillillan Publishing Company; 1993: 553-554.
- 201. Ferrara M, De Gennaro L: The sleep inertia phenomenon during the sleep-wake transition: Theoretical and operational issues. *AviatSpace Environ Med* 2000, 71(8):843-848.
- 202. Bruck D, Pisani D: The effects of sleep inertia on decision-making performance. J Sleep Res 1997, 8:95-103.
- 203. Achermann P, Werth E, Dijk DJ, Borbély AA: **Time course of sleep inertia after nighttime and daytime sleep episodes**. *Archives Italiennes de Biologie* 1995, **134**:109-119.
- 204. Naitoh P, Kelly T, Babkoff H: Sleep inertia: Best time not to wake up. *Chronobiol Int* 1993, **10**(2):109-118.
- 205. Van Dongen HPA, Price NJ, Mullington JM, Szuba MP, Kapoor SC, Dinges DF: Caffeine eliminates psychomotor vigilance deficits from sleep inertia. Sleep 2001, 24(7):813-819.
- 206. Burke TM, Scheer FA, Ronda JM, Czeisler CA, Wright KP, Jr.: Sleep inertia, sleep homeostatic and circadian influences on higher-order cognitive functions. *J Sleep Res* 2015, **24**(4):364-371.
- 207. Silva EJ, Duffy JF: Sleep inertia varies with circadian phase and sleep stage in older adults. *Behav Neurosci* 2008, **122**(4):928-935.
- 208. Åkerstedt T, Patkai P, Dahlgren K: Field studies of shiftwork: II. Temporal patterns in psychophysiological activation in workers alternating between night and day work. *Ergonomics* 1977, **20**:621-631.
- 209. Benedict FG: **Studies in body-temperature. I. Influence of the inversion of the daily routine; the temperature of night-workers**. *American Journal of Physiology* 1904, **11**:145-169.

- Patkai P, Åkerstedt T, Pettersson K: Field studies of shiftwork: I. Temporal patterns in psychophysiological activation in permanent night workers. *Ergonomics* 1977, 20:611-619.
- 211. Bjerner B, Holm A, Swensson A: Diurnal variation in mental performance a study of three-shift workers. *British Journal of Industrial Medicine* 1955, **12**:103-110.
- 212. Waldhauser F, Vierhapper H, Pirich K: Abnormal circadian melatonin secretion in night-shift workers. (Letter to the Editor). *NEnglJMed* 1986:1614-1614.
- 213. Smith MR, Cullnan EE, Eastman CI: Shaping the light/dark pattern for circadian adaptation to night shift work. *Physiol Behav* 2008, **95**(3):449-456.
- 214. Carskadon MA, Dement WC: Effects of total sleep loss on sleep tendency. *Perceptual and Motor Skills* 1979, **48**:495-506.
- 215. Colquhoun WP: **Rhythms in performance**. In: *Handbook of Behavioral Neurobiology: Biological Rhythms*. edn. Edited by Aschoff J. New York: Plenum Press; 1981: 333-348.
- 216. Folkard S, Knauth P, Monk TH: **The effect of memory load on the circadian variation in performance efficiency under a rapidly rotating shift system**. *Ergonomics* 1976, **19**:479-488.
- 217. Colquhoun WP, Blake MJF, Edwards RS: Experimental studies of shift-work II: Stabilized 8-hour shift systems. *Ergonomics* 1968, 11:527-546.
- 218. Colquhoun WP, Blake MJF, Edwards RS: **Experimental studies of shift work**. *Studia Laboris et Saluds* 1969, **4**:47-63.
- 219. Colquhoun WP, Blake MJF, Edwards RS: Experimental studies of shift-work I: A comparison of 'rotating' and 'stabilized' 4-hour shift systems. *Ergonomics* 1968, 11:437-453.
- 220. Colquhoun WP, Blake MJF, Edwards RS: Experimental studies of shift-work III: Stabilized 12-hour shift systems. *Ergonomics* 1969, 12:865-882.
- 221. Coleman RM, Dement WC: Falling asleep at work: A problem for continuous operations. *Sleep Research* 1986, 15:265-265.
- 222. Rosekind MR, Gander PH, Dinges DF: Alertness management in flight operations: strategic napping. *Aerospace Technology Conference and Exposition,Long Beach CA* 1991:1-12.
- 223. Dahlgren K: Adjustment of circadian rhythms and EEG sleep functions to day and night sleep among permanent night workers and rotating shiftworkers. *Psychophysiol* 1981, **18**:381-391.
- 224. Dahlgren K: Long-term adjustment of circadian rhythms to a rotating shiftwork schedule. *Scandinavian Journal of Work, Environment and Health* 1981, 7:141-151.
- 225. Tilley AJ, Wilkinson RT, Drud M: Night and day shifts compared in terms of the quality and quantity of sleep recorded in the home and performance measured at work: A pilot study. In: Night and Shift Work, Biological and Social Aspects. edn. Edited by Reinberg A, Vieux N, Andlauer P. Oxford: Pergamon Press; 1981: 187-196.
- 226. Kogi K, Ohta T: Incidence of near accidental drowsing in locomotive driving during a period of rotation. *Journal of Human Ergology (Tokyo)* 1975, 4:65-76.
- 227. Anderson C, Ftouni S, Ronda JM, Rajaratnam SMW, Czeisler CA, Lockley SW: Selfreported drowsiness and safety outcomes while driving after an extended duration work shift in trainee physicians. *Sleep* 2018.

- 228. Vidafar P, Gooley JJ, Burns AC, Rajaratnam SMW, Rueger M, Van Reen E, Czeisler CA, Lockley SW, Cain SW: Increased vulnerability to attentional failure during acute sleep deprivation in women depends on menstrual phase. *Sleep* 2018, **41**(8).
- 229. American Academy of Sleep Medicine: The International Classification of Sleep Disorders: Diagnostic and Coding Manual, vol. Second Edition, 2nd edn. Westchester, IL: American Academy of Sleep Medicine; 2005.
- 230. Viola AU, Archer SN, James LM, Groeger JA, Lo JCY, Skene DJ, von schantz M, Dijk DJ: **PER3 polymorphism predicts sleep structure and waking performance**. *Current Biology* 2007, **17**:613-618.
- Czeisler CA, Moore-Ede MC, Coleman RM: Rotating shift work schedules that disrupt sleep are improved by applying circadian principles. Science 1982, 217:460-463.
- 232. Czeisler CA, Dijk DJ: Use of bright light to treat maladaptation to night shift work and circadian rhythm sleep disorders. J Sleep Res 1995, 4(Suppl 2):70-73.
- 233. Czeisler CA, Walsh JK, Roth T, Hughes RJ, Wright Jr. KP, Kingsbury L, Arora S, Schwartz JRL, Niebler GE, Dinges DF: **Modafinil for excessive sleepiness associated** with shift work sleep disorder. *N Engl J Med* 2005, **353**:476-486.
- 234. Wyatt JK, Cajochen C, Ritz-De Cecco A, Czeisler CA, Dijk DJ: Low-dose repeated caffeine administration for circadian-phase-dependent performance degradation during extended wakefulness. *Sleep* 2004, **27**(3):374-381.
- 235. Landrigan CP, Rothschild JM, Cronin JW, Kaushal R, Burdick E, Katz JT, Lilly CM, Stone PH, Lockley SW, Bates DW *et al*: **Effect of reducing interns' work hours on** serious medical errors in intensive care units. *NEnglJMed* 2004, **351**(18):1838-1848.
- 236. Barger LK, Cade BE, Ayas NT, Cronin JW, Rosner B, Speizer FE, Czeisler CA: Extended work shifts and the risk of motor vehicle crashes among interns. N Engl J Med 2005, 352:125-134.
- 237. Strogatz SH, Kronauer RE, Czeisler CA: Circadian regulation dominates homeostatic control of sleep length and prior wake length in humans. *Sleep* 1986, 9:353-364.
- 238. Torsvall L, Åkerstedt T, Gillberg M: Age, sleep and irregular workhours: A field study with electroencephalographic recordings, catecholamine excretion and self-ratings. Scandinavian Journal of Work, Environment and Health 1981, 7:196-203.
- 239. Åkerstedt T, Gillberg M: Sleep disturbances and shift work. In: *Night and Shift Work: Biological and Social Aspects*. edn. Edited by Reinberg A, Vieux N, Andlauer P. Oxford: Pergamon Press; 1981: 127-137.
- 240. Åkerstedt T, Torsvall L: Age, sleep and adjustment to shiftwork. In: *Sleep 1980.* edn. Edited by Koella WP. Basel: S. Karger; 1981: 190-195.
- 241. Åkerstedt T, Torsvall L: Shift-dependent well-being and individual differences. *Ergonomics* 1981, **24**:265-273.
- 242. Scott LD, rogers AE, Hwang WT, Zhang Y: Effects of critical care nurses' work hours on vigilance and patients' safety. *AmJ CritCare* 2006, **15**(1):30-37.
- Rutenfranz J, Colquhoun WP, Knauth P, Ghata JN: Biomedical and psychosocial aspects of shift work. Scandinavian Journal of Work, Environment and Health 1977, 3:165-182.
- 244. Rutenfranz J, Knauth P, Angersbach D: **Shift work research issues**. In: *The Twenty-Four Hour Workday: Proceedings of a NIOSH Symposium on Variations in Work-Sleep*

Schedules. Volume NIOSH Publication 81-127, edn. Edited by Johnson LC, Tepas DI, Colquhoun WP, Colligan MJ. Cincinnati: U.S. Department of Health and Human Services; 1981: 221-259.

- 245. Mahan RP, Carvalhais AB, Queen SE: Sleep reduction in night-shift workers: is it sleep deprivation or a sleep disturbance disorder? *Percept Mot Skills* 1990, **70**:723-730.
- 246. Torsvall L, Åkerstedt T: A diurnal type scale. Scandinavian Journal of Work, Environment and Health 1980, 6:283-290.
- 247. Rutenfranz J, Knauth P: Hours of work and shiftwork. Ergonomics 1976, 19:331-340.
- 248. Vokac Z, Lund L: Patterns and duration of sleep in permanent security night guards. *Journal of Human Ergology (Tokyo)* 1982, **11**:311-316.
- 249. Åkerstedt T, Torsvall L, Fröberg J: A questionnaire study of irregular work hours and sleep/wake disturbances. *Sleep Research* 1983, **12**:358-358.
- 250. Folkard S, Condon R: Night shift paralysis in air traffic control officers. *Ergonomics* 1987, **30**:1353-1363.
- 251. Foret J, Lantin G: The sleep of train drivers: An example of the effects of irregular work schedules on sleep. In: *Aspects of Human Efficiency Diurnal Rhythm and Loss of Sleep.* edn. Edited by Colquhoun WP. Cambridge,UK: English Universities Press Ltd; 1972: 273-282.
- 252. Kogi K: Repeated short indifference periods in industrial vigilance. Journal of Human Ergology (Tokyo) 1972, 1:111-121.
- 253. Torsvall L, Åkerstedt T: Sleepiness on the job: Continuously measured EEG changes in train drivers. *ElectroencephalogrClin Neurophysiol* 1987, 66:502-511.
- 254. Torsvall L, Åkerstedt T, Gillander K, Knutsson A: Sleep on the night shift: 24-Hour EEG monitoring of spontaneous sleep/wake behavior. *Psychophysiol* 1989, 26:352-358.
- 255. Van Dongen HPA, Baynard MD, Maislin G, Dinges DF: Systematic interindividual differences in neurobehavioral impairment from sleep loss: evidence of trait-like differential vulnerability. *Sleep* 2004, **27**:423-433.
- 256. Van Dongen HPA, Maislin G, Dinges DF: **Dealing with inter-individual differences in the temporal dynamics of fatigue and performance: importance and techniques.** *AviatSpace Environ Med* 2004, **75**(3 Suppl):A147-A154.
- 257. Van Dongen HPA, Vitellaro KM, Dinges DF: Individual differences in adult human sleep and wakefulness: leitmotif for a research agenda. *Sleep* 2005, **28**(4):479-496.
- 258. Van Dongen HPA: Brain activation patterns and individual differences in working memory impairment during sleep deprivation. *Sleep* 2005, **28**(4):386-388.
- 259. Van Dongen HPA, Caldwell JA, Jr., Caldwell JL: Investigating systematic individual differences in sleep-deprived performance on a high-fidelity flight simulator. *Behav* Res Methods 2006, **38**(2):333-343.
- 260. Van Dongen HPA: Shift work and inter-individual differences in sleep and sleepiness. *Chronobiology International* 2006, **23**(6):1139-1147.
- 261. Gold DR, Rogacz S, Bock N, Tosteson TD, Baum TM, Speizer FE, Czeisler CA: **Rotating shift work, sleep, and accidents related to sleepiness in hospital nurses**. *American Journal of Public Health* 1992, **82**:1011-1014.

- 262. Novak RD, Auvil-Novak SE: Focus group evaluation of night nurse shiftwork difficulties and coping strategies. *Chronobiol Int* 1996, **13**(6):457-463.
- 263. Stutts JC, Wilkins JW, Vaughn GM: **Why do people have drowsy driving crashes? Input from drivers who just did**. In., vol. Nov 1999. Washington, DC: AAA Foundation for Traffic Safety; 1999: 1-85.
- Scott LD, Hwang WT, rogers AE, Nysse T, Dean GE, Dinges DF: The relationship between nurse work schedules, sleep duration, and drowsy driving. *Sleep* 2007, 30(12):1801-1807.
- 265. Dorrian J, Tolley C, Lamond N, van den HC, Pincombe J, rogers AE, Drew D: Sleep and errors in a group of Australian hospital nurses at work and during the commute. *Appl Ergon* 2008, **39**(5):605-613.
- 266. Lyznicki JM, Doege TC, Davis RM, Williams MA: **Sleepiness, driving, and motor** vehicle crashes. *Journal of the American Medical Association* 1998, **279**:1908-1913.
- 267. Folkard S, Lombardi DA: Modeling the impact of the components of long work hours on injuries and "accidents". *Am J Ind Med* 2006, **49**:953-963.
- 268. Dembe AE, Erickson JB, Delbos RG, Banks SM: The impact of overtime and long work hours on occupational injuries and illnesses: new evidence from the United States. *OccupEnvironMed* 2005, **62**(9):588-597.
- 269. Loomis D: Long work hours and occupational injuries: New evidence on upstream causes. *OccupEnvironMed* 2005, **62**(9):585.
- 270. Rogers AE, Hwang WT, Scott LD, Aiken LH, Dinges DF: **The working hours of hospital staff nurses and patient safety**. *Health Aff (Millwood)* 2004, **23**(4):202-212.
- 271. Hänecke K, Tiedemann S, Nachreiner F, Grzech-Sukalo H: Accident risk as a function of hour at work and time of day as determined from accident data and exposure models of the German working poulation. Scand J Work Environ Health 1998, 24(Suppl 3):43-48.
- 272. Colligan MJ, Tepas DI: **The stress of hours of work**. *American Industrial Hygiene Association Journal* 1986, **47**(11):686-695.
- 273. Kelly RJ, Schneider MF: The twelve-hour shift revisited: Recent trends in the electric power industry. *Journal of Human Ergology (Tokyo)* 1982, 11:369-384.
- 274. Rosa RR, Colligan MJ: Extended workdays: effects of 8-hour and 12-hour rotating shift schedules on performance, subjective alertness, sleep patterns, and psychosocial variables. *Work and Stress* 1989, **3**:21-32.
- 275. Lewis PM, Swaim DJ, Rosa RR, Colligan MJ, Booth RJ, Swint MJ: Evaluation of the 12-hour shift schedule at the Fast Flux Test Facility. In.: U.S. Department of Energy; 1986.
- 276. Strohl KP, Blatt J, Council F, Georges K, Kiley J, Kurrus R, McCartt AT, Merritt SL, Pack AI, Rogus S et al: Drowsy driving and automobile crashes: Reports and recommendations. In., vol. DOT HS 808 707. Washington, DC: U.S. Department of Transportation, National Highway Traffic Safety Administration; U.S. Department of Health and Human Services, Public Health Service, National Institutes of Health, National Heart, Lung, and Blood Institute; 1998: III-30.
- 277. National Transportation Safety Board: Factors that affect fatigue in heavy truck accidents. Volume 1: Analysis. In., vol. NTSB/SS-95/01. Washington, D.C.; 1995.

- 278. Department of Transportation: **Hours of Service of Drivers; Driver Rest and Sleep for Safe Operations.** In. Edited by Federal Motor Carrier Safety Administration, vol. 65. Washington, DC; 2000.
- 279. Czeisler CA, Kennedy WA, Allan JS: Circadian rhythms and performance decrements in the transportation industry. In: *Proceedings of a Workshop on the Effects of Automation on Operator Performance*. edn. Edited by Coblentz AM. Paris: Universite Rene Descartes; 1986: 146-171.
- 280. National Transportation Safety Board: Factors That Affect Fatigue in Heavy Truck Accidents Volume 2: Case Summaries. In., vol. NTSB/SS-95/02. Washington, DC: National Transportation Safety Board; 1995: 1-227.
- 281. Federal Aviation Administration, Terminal, Enroute: **Shiftwork Survey Results**. Human Resources Research Organization; Federal Aviation Administration; 2001.
- 282. Harma M, Sallinen M, Ranta R, Mutanen P, Muller K: The effect of an irregular shift system on sleepiness at work in train drivers and railway traffic controllers. *J Sleep Res* 2002, **11**(2):141-151.
- 283. Gander PH, Myhre G, Graeber RC, Andersen HT, Lauber JK: Adjustment of sleep and the circadian temperature rhythm after flights across nine time zones. *AviatSpace Environ Med* 1989, **60**:733-743.
- 284. Gander PH, Graeber RC, Connell LJ: Crew factors in flight operations II: Psychophysiological responses to short-haul air transport operations. In., vol. Tech.Memo.108856. Washington, D.C.: NASA; 1994: 1-60.
- 285. Åkerstedt T: Sleepiness as a consequence of shift work. Sleep 1988, 11:17-34.
- 286. St Hilaire MA, Anderson C, Anwar J, Sullivan JP, Cade BE, Flynn-Evans EE, Czeisler CA, Lockley SW, Harvard Work Hours H, Safety G: Brief (<4 hr) sleep episodes are insufficient for restoring performance in first-year resident physicians working overnight extended-duration work shifts. *Sleep* 2019, 42(5).
- 287. Endo T, Kogi K: Monotony effects of the work of motormen during high-speed train operation. *Journal of Human Ergology (Tokyo)* 1975, **4**:129-140.
- 288. Mitler MM, Miller JC, Lipsitz JJ, Walsh JK, Wylie CD: The sleep of long-haul truck drivers. *NEnglJMed* 1997, **337**(11):755-761.
- 289. Lockley SW, Evans EE, Scheer FAJL, Brainard GC, Czeisler CA, Aeschbach D: Shortwavelength sensitivity for the direct effects of light on alertness, vigilance, and the waking electroencephalogram in humans. *Sleep* 2006, **29**(2):161-168.
- 290. Muehlbach MJ, Walsh JK: The effects of caffeine on simulated night-shift work and subsequent daytime sleep. *Sleep* 1995, **18**(1):22-29.
- 291. Walsh JK, Muehlbach MJ, Schweitzer PK: Acute administration of Triazolam for the daytime sleep of rotating shift workers. *Sleep* 1984, 7:223-229.
- 292. Walsh JK, Muehlbach MJ, Schweitzer PK: Hypnotics and caffeine as countermeasures for shiftwork-related sleepiness and sleep disturbance. *J Sleep Res* 1995, 4(Suppl. 2):80-83.
- 293. Walsh JK, Schweitzer PK, Anch AM, Muehlbach MJ, Jenkins NA, Dickins QS: Sleepiness/alertness on a simulated night shift following sleep at home with triazolam. *Sleep* 1991, 14(2):140-146.
- 294. Higgins E, Chiles WD, McKenzie JM, Iampietro PF, Winget CM, Funkhouser GE, Burr MJ, Vaughan JAJ, Jennings A: **The effects of a 12-hour shift in the wake-sleep cycle**

on physiological and biochemical responses and on multiple task performance. In., vol. DOT/FAA/AM-75-10. Washington,DC: Federal Aviation Administration, Office of Aviation Medicine; 1975.

- 295. Santhi N, Horowitz TS, Czeisler CA: Night shift work impairs decision-making in visual search. *Sleep* 2005, 28:A64.
- 296. Santhi N, Horowitz T, Wolfe J, Czeisler CA: Sleep deprivation impairs search for rare targets. *Sleep* 2007, **30**(Abstract Supplement):A22.
- 297. Lee ML, Howard ME, Horrey WJ, Liang Y, Anderson C, Shreeve MS, O'Brien CS, Czeisler CA: High risk of near-crash driving events following night-shift work. Proceedings of the National Academy of Sciences of the United States of America 2016, 113(1):176-181.
- 298. Barger LK, Rajaratnam SM, Wang W, O'Brien CS, Sullivan JP, Qadri S, Lockley SW, Czeisler CA: **Common Sleep Disorders Increase Risk of Motor Vehicle Crashes and Adverse Health Outcomes in Firefighters**. *Journal of Clinical Sleep Medicine* 2015, 11(3):233-240.
- 299. Beebe DW, Gozal D: Obstructive sleep apnea and the prefrontal cortex: towards a comprehensive model linking nocturnal upper airway obstruction to daytime cognitive and behavioral deficits. *J Sleep Res* 2002, **11**(1):1-16.
- 300. Powell NB, Riley RW, Schechtman KB, Blumen MB, Dinges DF, Guilleminault C: A comparative model: Reaction time performance in sleep-disordered breathing versus alcohol-impaired controls. *Laryngoscope* 1999, **109**:1648-1654.
- 301. George CF, Nickerson PW, Hanly PJ, Millar TW, Kryger MH: Sleep apnoea patients have more automobile accidents. *Lancet* 1987, **2**:447.
- 302. Findley LJ, Bonnie RJ: Sleep apnea and auto crashes: What is the doctor to do? *Chest* 1988, **94**(2):225-226.
- 303. Findley LJ, Unverzagt ME, Suratt PM: Automobile accidents involving patients with obstructive sleep apnea. *Am Rev Respir Dis* 1988, **138**:337-340.
- 304. Findley LJ, Fabrizio M, Thommi G, Suratt PM: Severity of sleep apnea and automobile crashes. *N Engl J Med* 1989, **320**(13):868-869.
- 305. Findley LJ, Weiss JW, Jabour ER: Drivers with untreated sleep apnea: A cause of death and serious injury. *Arch Intern Med* 1991, **151**:1451-1452.
- Haraldsson P-O, Carenfelt C, Tingvall C: Sleep apnea syndrome symptoms and automobile driving in a general population. *Journal of Clinical Epidemiology* 1992, 45(8):821-826.
- 307. Findley LJ, Levinson MP, Bonnie RJ: **Driving performance and automobile accidents** in patients with sleep apnea. *Clinics in Chest Medicine* 1992, **13**(3):427-435.
- 308. Gurubhagavatula I, Maislin G, Nkwuo JE, Pack AI: Occupational screening for obstructive sleep apnea in commercial drivers. *Am J RespirCrit Care Med* 2004, 170(4):371-376.
- 309. George CF, Boudreau AC, Smiley A: Simulated driving performance in patients with obstructive sleep apnea. *Am J RespirCrit Care Med* 1996, **154**(1):175-181.
- 310. Findley L, Unverzagt M, Guchu R, Fabrizio M, Buckner J, Suratt P: Vigilance and automobile accidents in patients with sleep apnea or narcolepsy. *Chest* 1995, 108(3):619-624.

- 311. Teran-Santos J, Jimenez-Gomez A, Cordero-Guevara J: The association between sleep apnea and the risk of traffic accidents. Cooperative Group Burgos-Santander. *NEnglJMed* 1999, **340**(11):847-851.
- 312. George CF: Sleep. 5: Driving and automobile crashes in patients with obstructive sleep apnoea/hypopnoea syndrome. *Thorax* 2004, **59**(9):804-807.
- 313. Hack MA, Choi SJ, Vijayapalan P, Davies RJ, Stradling JR: Comparison of the effects of sleep deprivation, alcohol and obstructive sleep apnoea (OSA) on simulated steering performance. *Respiratory Medicine* 2001, **95**(7):594-601.
- 314. Pack AI, Maislin G, Staley B, Pack FM, Rogers WC, George CF, Dinges DF: Impaired performance in commercial drivers: role of sleep apnea and short sleep duration. *Am J Respir Crit Care Med* 2006, **174**(4):446-454.
- 315. Ellen RLB, Marshall SC, Palayew M, Molnar FJ, Wilson KG, Man-Son-Hing M: Systematic review of motor vehicle crash risk in persons with sleep apnea. *J Clin Sleep Med* 2006, **2**(2):193-200.
- 316. Tregear S, Reston J, Schoelles K, Phillips B: **Obstructive sleep apnea and risk of motor vehicle crash: systematic review and meta-analysis**. *Journal of Clinical Sleep Medicine* 2009, **5**(6):573-581.
- 317. Burks SV, Anderson JE, Bombyk M, Haider R, Ganzhorn D, Jiao X, Lewis C, Lexvold A, Liu H, Ning J *et al*: Nonadherence with Employer-Mandated Sleep Apnea Treatment and Increased Risk of Serious Truck Crashes. *Sleep* 2016, **39**(5):967-975.
- 318. Sullivan JP, O'Brien CS, Barger LK, Rajaratnam SM, Czeisler CA, Lockley SW: Randomized, prospective study of the impact of a sleep health program on firefighter injury and disability. *Sleep* 2017.
- 319. Barger LK, O'Brien CS, Rajaratnam SM, Qadri S, Sullivan JP, Wang W, Czeisler CA, Lockley SW: **Implementing a Sleep Health Education and Sleep Disorders Screening Program in Fire Departments: A Comparison of Methodology**. *Journal of Occupational and Environmental Medicine* 2016, **58**(6):601-609.
- 320. Kryger MH, Roth T, Dement WC. **Principles and Practice of Sleep Medicine**, Vol. 4. Philadelphia: W.B. Saunders; 2005.
- 321. Phillips AJK, Clerx WM, O'Brien CS, Sano A, Barger LK, Picard RW, Lockley SW, Klerman EB, Czeisler CA: Irregular sleep/wake patterns are associated with poorer academic performance and delayed circadian and sleep/wake timing. *Sci Rep* 2017, 7(1):3216.
- 322. Nutt D, Wilson S, Paterson L: Sleep disorders as core symptoms of depression. *Dialogues Clin Neurosci* 2008, **10**(3):329-336.
- 323. Daley M, Morin CM, LeBlanc M, Gregoire JP, Savard J, Baillargeon L: Insomnia and its relationship to health-care utilization, work absenteeism, productivity and accidents. *Sleep Med* 2008.
- 324. Drake CL, Roehrs T, Richardson G, Walsh JK, Roth T: Shift work sleep disorder: prevalence and consequences beyond that of symptomatic day workers. *Sleep* 2004, 27(8):1453-1462.
- 325. Van Dongen HP, Belenky G, Vila BJ: The efficacy of a restart break for recycling with optimal performance depends critically on circadian timing. *Sleep* 2011, 34(7):917-929.