DRAFT

APPENDICES TO THE REPORT FROM THE

PEDIATRIC MILD TRAUMATIC BRAIN INJURY GUIDELINE WORKGROUP

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Appendix A: Roster

Appendix A Roster

Workgroup Members

CHAIR: Shelly D. Timmons, MD, PhD, FACS, FAANS Professor of Neurosurgery Vice Chair for Administration

Director of Neurotrauma Department of Neurosurgery Penn State University Milton S. Hershey Medical Center

Edward C. Benzel, MD

Chairman Cleveland Clinic Department of Neurosurgery

Catherine Broomand, PhD, ABPP-CN

Past President of MindGame/Sacramento Valley Concussion Care Consortium

Former Director of Sports Concussion Clinic Kaiser Permanente

Neuropsychologist Center for Neuropsychological Services Kaiser Permanente (Retired)

Linda Ewing-Cobbs, PhD

Professor of Pediatrics Harriet and Joe Foster Chair in Cognitive Neuroscience

Director Dan L. Duncan Children's Neurodevelopmental Clinic Department of Pediatrics and Children's Learning Institute University of Texas Health Science Center at Houston Gerard Gioia, PhD

Division Chief of Pediatric Neuropsychology Children's National Health System

Professor Departments of Pediatrics, Psychiatry & Behavioral Sciences George Washington University School of Medicine

Christopher C. Giza, MD

Professor of Pediatric Neurology and Neurosurgery Brain Injury Research Center University of California, Los Angeles

Director Steve Tisch BrainSPORT Sports Concussion - Mild TBI Program University of California, Los Angeles

Medical Director Operation MEND - Wounded Warrior Project TBI Program Interdepartmental Programs for Neuroscience and Biomedical Engineering Mattel Children's Hospital University of California, Los Angeles

Wayne A. Gordon, PhD, ABPP/Cn

Jack Nash Professor and Vice Chair Department of Rehabilitation Medicine Icahn School of Medicine at Mount Sinai

Kevin Guskiewicz, PhD, ATC

Kenan Distinguished Professor of Exercise and Sport Science University of North Carolina at Chapel Hill

Dean, College of Arts & Sciences University of North Carolina at Chapel Hill

Mark E. Halstead, MD

Associate Professor Departments of Pediatrics and Orthopedics St Louis Children's Hospital Washington University School of Medicine

Stanley A. Herring, MD

Clinical Professor, Departments of Rehabilitation Medicine, Orthopaedics and Sports Medicine and Neurological Surgery University of Washington

Director University of Washington Medicine Sports Health and Safety Institute Seattle, Washington

Barbara Holshouser, PhD

Medical Physicist & Professor Division of Magnetic Resonance Imaging Department of Radiology Loma Linda University Medical Center

Madeline Matar Joseph, MD, FACEP, FAAP

Chair American College of Emergency Physicians Pediatric Committee

Vice President Florida Chapter of the American Academy of Pediatrics

Professor of Emergency Medicine and Pediatrics Assistant Chair of Pediatric Emergency Medicine Quality Department of Emergency Medicine University of Florida College of Medicine-Jacksonville

Angela Lumba-Brown, MD, FAAP

Assistant Professor of Pediatrics Pediatric Emergency Medicine Washington University School of Medicine St. Louis Children's Hospital

Robert E. O'Connor, MD, MPH

Professor and Chair of Emergency Medicine Professor of Public Health Sciences University of Virginia School of Medicine

Rosemarie Scolaro Moser, PhD, ABN, ABPP-RP

Director Sports Concussion Center of New Jersey

Adjunct Professor of Clinical Psychology Graduate Program Widener University

Theodore J. Spinks, MD, FAANS

Clinical Assistant Professor Health Science Center Texas A&M

Attending Staff St. David's Hospitals

Stacy Suskauer, MD

Director of Brain Injury Rehabilitation Programs Kennedy Krieger Institute

Associate Professor of Physical Medicine & Rehabilitation and Pediatrics Johns Hopkins University School of Medicine

Michael Turner, MD

Neurosurgeon (Retired) Goodman Campbell Brain & Spine Department of Neurosurgery Indiana University

Barbara Weissman, MD

Associate Professor of Pediatrics Division of Child Neurology Department of Pediatrics Emory University School of Medicine

David W. Wright, MD, FACEP

Associate Professor Vice Chair for Research

Director Emergency Neurosciences

Director Injury Prevention Research Center EmoryDepartment of Emergency Medicine Emory University School of Medicine

Keith Owen Yeates, PhD, ABPP

Ronald and Irene Ward Chair in Pediatric Brain Injury Professor of Psychology University of Calgary

Ad-Hoc Experts

Katrina Altenhofen, MPH, Paramedic, CME, CCPSTI

Council Member National EMS Advisory Council Emergency Medical Educational Services

James M. Callahan, MD

Division of Emergency Medicine Department of Pediatrics The Children's Hospital of Philadelphia

Professor of Clinical Pediatrics Perelman School of Medicine at The University of Pennsylvania

Meeryo C. Choe, MD

Assistant Clinical Professor Division of Pediatric Neurology Mattel Children's Hospital University of California, Los Angeles

Associate Director Steve Tisch BrainSPORT Program Brain Injury Research Center University of California, Los Angeles

Cindy W. Christian, MD

Anthony A. Latini Endowed Chair in Child Abuse and Neglect Prevention The Children's Hospital of Philadelphia

Professor of Pediatrics Perelman School of Medicine at The University of Pennsylvania

Micky Collins, PhD

Program Director Sports Medicine Concussion Program University of Pittsburgh Medical Center

Associate Professor Departments of Orthopaedic and Neurological Surgery University of Pittsburgh Medical Center

John DeWitt, PT, DPT, SCS, ATC

Rehab Manager Clinical Development The Ohio State University Sports Medicine

Assistant Clinical Professor Division of Physical Therapy The Ohio State University

Ann-Christine Duhaime, MD

Director Pediatric Neurosurgery Massachusetts General Hospital

Nicholas T. Zervas Professor of Neurosurgery Harvard Medical School

Richard G. Ellenbogen, MD, FACS

Professor and Chairman Department of Neurological Surgery

Director Neuroscience Institute University of Washington Medicine

Director American Board of Neurological Surgery

President American Society of Pediatric Neurosurgeons

Theodore G. Ganiats, MD

Professor Emeritus Department of Family Medicine & Public Health School of Medicine University of California, San Diego

Dr. Ganiats is a Senior Staff Fellow at the Agency for Healthcare Research and Quality (AHRQ). This project was conducted while he was at the University of California San Diego. The views expressed are those of the author; no endorsement by AHRQ or the U.S. Department of Health and Human Services is intended or should be inferred.

Andrew Gregory, MD, FAAP, FACSM

Associate Professor of Orthopedics, Neuorsurgery & Pediatrics Vanderbilt University School of Medicine

Michael Kirkwood, PhD, ABPP/Cn

Associate Clinical Professor Department of Physical Medicine & Rehabilitation University of Colorado School of Medicine Children's Hospital Colorado

Heather Keenan, PhD, MPH, MDCM

Professor of Pediatric Critical Care Department of Pediatrics University of Utah

Karen McAvoy, PsyD

Director Center for Concussion REAP Community-Based Concussion Management Project Rocky Mountain Hospital for Children

Anne Mucha, PT, DPT, MS, NCS

Coordinator Vestibular Rehabilitation Sports Concussion Program & Centers for Rehab Services University of Pittsburgh Medical Center

David Paulk, PA-C, EdD, DFAAPA

Professor & Director Physician Assistant Program Murphy Deming College of Health Sciences Mary Baldwin University

Margot Putukian, MD, FACSM

Director of Athletic Medicine & Head Team Physician Princeton University

Assistant Director of Medical Services Princeton University Associate Clinical Professor Rutgers - Robert Wood Johnson Medical School Past President American Medical Society for Sports Medicine

John Ragheb, MD, FACS, FAAP

Director Neurological Surgery Nicklaus Children's Hospital

Professor of Clinical Neurosurgery and Pediatrics

Affiliated Faculty University of Miami Miller School of Medicine

P.B. Raksin, MD

Director Neurosurgery ICU John H. Stroger Jr Hospital of Cook County (Formerly Cook County Hospital)

Assistant Professor Department of Neurosurgery Rush University Medical Center

Sally Schoessler, MSEd, BSN, RN

Director of Education Allergy & Asthma Network

Former Director of Education National Association of School Nurses

H. Gerry Taylor, PhD

Professor of Pediatrics, Psychiatry, and Psychological Sciences Case Western Reserve University Rainbow Babies & Children's Hospital University Hospitals Case Medical Center

Shari Wade, PhD

Professor Department of Pediatrics Cincinnati Children's Hospital Medical Center and the University of Cincinnati College of Medicine Appendix B: Rationales for Clinical Questions

Appendix B Rationales for Clinical Questions

Selection of Clinical Questions

After presentation of an analytic framework and an introduction to the Patient-Intervention-Comparator or Co-Intervention-Outcome format for questions, Workgroup members independently nominated questions for consideration. Candidate questions were collated and presented to the entire group. Using a modified Delphi process, questions were anonymously ranked on a 9-point ordinal scale of importance over three rounds of voting. Facilitated discussions amongst Workgroup members occurred between voting rounds. Through this process, the Workgroup selected 6 clinical questions. The questions and a brief rationale for their selection are discussed below.

Clinical Questions and Rationale

Although evidence-based clinical decision rules have been recently published to guide the use of head computed tomography (CT) imaging to identify intracranial injury, challenges in the acute diagnosis of mild traumatic brain injury (mTBI) still exist.¹ Advanced imaging techniques such as diffusion tensor imaging, magnetic resonance spectroscopy, perfusion weighted imaging, and functional magnetic resonance imaging (MRI) of the brain have shown changes in patients with mTBI, though their utility in management and feasibility of use has yet to be determined. There is a need for more evidence regarding the reliability and validity of various tools and questionnaires to diagnose mTBI. To address these issues, the Workgroup developed the first research question:

 For children (18 years of age and younger) with suspected mild TBI, do specific tools as compared with a reference standard*, accurately diagnose mild TBI?
 *Acceptable diagnostic reference standards for question 1 will not be pre-specified. We will track the reference standards used in the identified studies during the data extraction process.

A clinical decision rule designed to predict clinically-important brain injuries was recently published based on sign and symptom evaluation of more than 40,000 children with suspected mTBI.¹ Despite the availability of such decision rules, confusion remains regarding how to best distinguish mild versus more significant TBI in acute care settings. Biomarker and imaging diagnostics for mTBI are reported in the literature and continue to be researched. The decision to obtain head CT imaging following suspected mTBI remains a concern due to exposure to ionizing radiation. This led to the development of the next two questions:

¹ Kupperman N, et al. Identification of children at very low risk of clinically-important brain injuries after head trauma: a prospective cohort study. The Lancet, Vol 374, Iss 9696, 1160 - 1170, 3 Oct 2009.

2. For children (18 years of age and younger) presenting to the emergency department (or other acute care setting) with mild TBI, how often does routine head imaging* identify important intracranial injury#?

3. For children (18 years of age and younger) presenting to the emergency department (or other acute care setting) with mild TBI, which features identify patients at risk for important intracranial injury?

*Routine head imaging pertinent to questions 2 and 3 will include skull x-rays, CT and MRI. #Important craniocerebral abnormalities will be those that change acute treatment. For example, those abnormalities which prompt more prolonged emergency room observation, hospitalization or neurosurgical consultation (e.g., intracranial hemorrhage, depressed and basilar skull fractures). Subtle changes of unclear significance (e.g. changes on diffusion tensor MRI images) will not be considered for questions 2 and 3. Such imaging changes will be considered as potential prognostic features in questions 4 and 5.

A significant subset of children with mTBI develop pronounced and prolonged symptoms prompting the need to identify factors that contribute to their course of recovery. A number of factors have previously been identified as potential predictors of recovery following mTBI including: age at injury, individual characteristics including genomics and medical history, social-demographic and family characteristics, the mechanism of the insult, repeat injury, acute medical management and follow-up. The need to identify evidence-based predictors of protracted recovery led to development of the fourth and fifth questions:

4. For children (18 years of age and younger) with mild TBI, what factors* identify patients at increased risk for ongoing impairment, more severe symptoms, or delayed recovery (<1 year post-injury)?

5. For children (18 years of age and younger) with mild TBI, which factors identify patients at increased risk of long-term# (>1 year) sequelae?

*Any feature of potential prognostic importance will be considered relevant to questions 3, 4 and 5. All potential features cannot be exhaustively anticipated before reviewing the literature. Potential features include those already listed in the questions and will also include the mechanism of injury and the timing of the imaging after injury.

#The distinction in timing (> 1 year) of post-injury sequelae between questions 4 and 5 is somewhat arbitrary. The literature relevant to both of these questions will be identified with a single search. The timing of the sequelae described from each identified study will be tracked during the data extraction process. A reasonable distinction between short and long term mTBI consequences will be made based upon the descriptions provided in the literature. At the protocol development stage, 1 year will be working distinction between short and long-term sequelae.

The emergency department and outpatient medical management of children with mTBI is controversial. Expert recommendations for the treatment of mTBI have been published without extensive supporting research. Generally, supportive care, cognitive rest, and a step-wise return to play have been recommended. However, with growing subspecialty clinics, unique application of computerized neurocognitive testing, emerging research on novel therapies, and increasing concerns about potential iatrogenic effects of cognitive rest, further evaluation is required. This led to the development of the sixth and final question:

6. For children (18 years of age and younger) with mild TBI (with ongoing symptoms) which treatments* improve mild TBI-related outcomes?

*Any therapeutic modality will be considered pertinent to question 6.

Appendix C: Literature Search Strategy

mTBI Updated Literature Search: ERIC, CINAHL, PubMed

December 2012-July 2015

ERIC

Strategy	Search Limits	Results
Original Search Strategy	Keyword Identifier tag†; January 1990-November 2012	200
Updated Search _Original Strategy	Keyword Identifier tag [†] ; December 2012-July 2015	16
Search Strategy Not limited by Keyword tag	Search All Fields; January 1990-November 2012	102*
Updated Search_Not limited by Keyword tag	Search All Fields; December 2012-July 2015	15*

*In 2014 ERIC launched a new, more intuitive search algorithm encouraging the use of simple language which the ERIC Thesaurus then uses to produce results, creating a more inclusive search. ERIC discourages large, specific, Boolean searches as too limiting. While speaking with ERIC's Customer Support, we determined the change in search algorithm accounts for the difference in results between searches using the Keyword Identifier and those that do not.

⁺At the time of this search, in the ERIC database the Keyword, or Identifier, tag searched all fields.

ERIC Original Search Strategy

((Keywords:sport* or Keywords:athlete* or Keywords:athletic or Keywords:recreation*) and (Keywords:"head injury" or Keywords:"head injuries" or Keywords:"brain injury" or Keywords:"brain injuries" or Keywords:"head trauma" or Keywords:"brain trauma" or Keywords:"skull injury" or Keywords:"skull injuries" or Keywords:tbi or Keywords:tbis or Keywords:"cerebral trauma" or Keywords:"cerebral injury" or Keywords:"cerebral injuries"))Publication Date:1990-2013

OR

(Keywords:"gcs 13-15" or Keywords:concussion* or Keywords:postconcussion or Keywords:"post concussion" or Keywords:postconcussive or Keywords:"post concussive" or Keywords:"mild tbi" or Keywords:"mild tbis" or Keywords:mtbi or Keywords:mtbis or Keywords:"mild traumatic brain injury" or Keywords:"mild traumatic brain injuries")Publication Date:1990-2013 ((Keywords:"traumatic brain injury" or Keywords:"traumatic brain injuries" or Keywords:tbi or Keywords:tbis or Keywords:"brain contusion" or Keywords:"brain contusions" or Keywords:"brain trauma" or Keywords:"brain laceration" or Keywords:"brain lacerations" or Keywords:"cerebral injury" or Keywords:"cerebral injuries" or Keywords:"head injury" or Keywords:"head injuries" or Keywords:"head trauma" or Keywords:"cerebral trauma" or Keywords:"skull injury" or Keywords:"skull injuries" or Keywords:"craniocerebral trauma") and (Keywords:mild or Keywords:minor))Publication Date:1990-2013

Search Strategy Not Limited by Keyword Tag

((sport* OR athlete* OR athletic OR recreation*) and ("head injury" OR "head injuries" OR "brain injury" OR "brain injuries" OR "head trauma" OR "brain trauma" OR "skull injury" OR "skull injuries" OR tbi OR tbi OR "cerebral trauma" OR "cerebral injury" OR "cerebral injury"))

OR

("gcs 13-15" OR concussion* OR postconcussion OR "post concussion" OR postconcussive OR "post concussive" OR "mild tbi" OR "mild tbis" OR mtbi OR mtbis OR "mild traumatic brain injury" OR "mild traumatic brain injuries")

OR

(("traumatic brain injury" OR "traumatic brain injuries" OR tbi OR tbis OR "brain contusion" OR "brain contusions" OR "brain trauma" OR "brain laceration" OR "brain lacerations" OR "cerebral injury" OR "cerebral injuries" OR "head injury" OR "head injuries" OR "head trauma" OR "cerebral trauma" OR "skull injury" OR "skull injuries" OR "craniocerebral trauma") and (mild OR minor))

CINAHL

Strategy	Search Limits	Results
Original Search	Limiters - Exclude MEDLINE records	551
<u>Strategy</u>	Narrow by SubjectAge6: - infant, newborn: birth-1 month	
	Narrow by SubjectAge5: - infant: 1-23 months	
	Narrow by SubjectAge4: - all infant	
	Narrow by SubjectAge3: - child, preschool: 2-5 years	
	Narrow by SubjectAge2: - adolescent: 13-18 years	
	Narrow by SubjectAge1: - child: 6-12 years	
	Narrow by SubjectAge0: - all child	
	Search modes - Boolean/Phrase	

	January 1990-November 2012	
Updated Search	Limiters - Exclude MEDLINE records	0
	Narrow by SubjectAge6: - infant, newborn: birth-1 month	
	Narrow by SubjectAge5: - infant: 1-23 months	
	Narrow by SubjectAge4: - all infant	
	Narrow by SubjectAge3: - child, preschool: 2-5 years	
	Narrow by SubjectAge2: - adolescent: 13-18 years	
	Narrow by SubjectAge1: - child: 6-12 years	
	Narrow by SubjectAge0: - all child	
	Search modes - Boolean/Phrase	
	December 2012-July 2015	

CINAHL Original Search Strategy

S43	S41	Limiters - Exclude MEDLINE records Narrow by SubjectAge6: - infant, newborn: birth-1 month Narrow by SubjectAge5: - infant: 1-23 months Narrow by SubjectAge4: - all infant Narrow by SubjectAge3: - child, preschool: 2-5 years Narrow by SubjectAge2: - adolescent: 13-18 years Narrow by SubjectAge1: - child: 6-12 years Narrow by SubjectAge0: - all child Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S42	S41	Limiters - Exclude MEDLINE records Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive

S41	S40 AND S15	Search modes -	Interface - EBSCOhost
341	SHO AND STR	Boolean/Phrase	Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
\$40	("NEUROFEEDBACK") OR (S16 OR S17 OR S18 OR S19 OR S20 OR S21 OR S22 OR S23 OR S24 OR S25 OR S26 OR S27 OR S28 OR S29 OR S30 OR S31 OR S32 OR S33 OR S34 OR S35 OR S36 OR S37 OR S38 OR S39)	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S39	"NEUROFEEDBACK"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S38	(MH "Patient Positioning+") OR "POSITIONING"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S37	"RANGE OF MOTION EXERCISE"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S36	(MH "Sensory Deprivation")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S35	(MH "Sensory Stimulation+") OR (MH "Physical Stimulation+") OR (MH "Transcutaneous Electric Nerve Stimulation") OR (MH "Acoustic Stimulation") OR (MH "Electric Stimulation+") OR (MH "Electrical Stimulation, Functional") OR (MH "Electrical Stimulation, Neuromuscular") OR	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive

	(MH "Deep Brain Stimulation") OR "STIMULATION"		
S34	(MH "Massage+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S33	(MH "Dietary Supplements+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S32	(MH "Functional Food")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S31	"COGNITIVE REST"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied <u>Health</u> Collection: Comprehensive
S30	(MH "Stress Management") OR "STRESS REDUCTION"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S29	"PHYSICAL REST"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S28	(MH "Bed Rest")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S27	(MH "Hypothermia, Induced")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;

			Nursing & Allied Health Collection: Comprehensive
S26	(MH "Cognitive Therapy") OR "COGNITIVE THERAPY" OR (MH "Recreational Therapy")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S25	(MH "Physical Therapy+") OR "PHYSICAL THERAPY" OR (MH "Pediatric Physical Therapy")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S24	(MH "Therapeutic Exercise+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S23	(MH "Hyperbaric Oxygenation")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S22	(MH "Biofeedback") OR "BIOFEEDBACK"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S21	(MH "Rehabilitation, Athletic") OR (MH "Rehabilitation") OR "REHABILITATION" OR (MH "Rehabilitation, Psychosocial") OR (MH "Rehabilitation, Speech and Language") OR (MH "Rehabilitation, Pediatric")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S20	(MH "Diet Therapy") OR (MH "Apitherapy") OR (MH "Pediatric Physical Therapy") OR (MH "Behavior Therapy") OR (MH "Pediatric Occupational Therapy") OR (MH "Motion Therapy,	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive

	Continuous Passive") OR (MH "Magnet Therapy") OR (MH "Color Therapy") OR (MH "Electroconvulsive Therapy")		
S19	(MH "Pharmacological and Biological Treatments") OR (MH "Treatment Outcomes") OR "TREATMENT"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S18	(MH "Drug Therapy") OR (MH "Drug Therapy, Combination") OR (MH "Fluid Therapy") OR (MH "Intravenous Therapy") OR (MH "Prescriptions, Drug") OR (MH "Manual Therapy") OR (MH "Manual Therapy") OR (MH "Rediotherapy") OR (MH "Rehabilitation") OR (MH "Respiratory Therapy") OR (MH "Stress Management") OR (MH "Therapy, Computer Assisted") OR (MH "Vestibular Stimulation")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S17	TX THERAPEUTIC	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S16	"THERAPY"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S15	S13 NOT S14	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S14	TI IRAQ OR IRAQI OR AFGHANISTAN OR "GULF WAR" OR "GULF WARS" OR AFGHAN OR "OPERATION ENDURING	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;

	FREEDOM" OR MILITARY OR BATTLEFIELD OR ARMY OR "ARMED FORCES" OR MARINES OR TROOPS OR SERVICEMEN OR "COMBAT RELATED" OR "AIR FORCE" OR SOLDIERS OR "OIF/OEF" OR VETERANS OR "ACTIVE DUTY" OR "SERVICE MEMBERS"		Nursing & Allied Health Collection: Comprehensive
S13	S7 OR S9 OR S10 OR S11	Limiters - Published Date from: 19900101-20121231 Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S12	S7 OR S9 OR S10 OR S11	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S11	TX "MINIMAL TRAUMATIC BRAIN INJURY" OR "MINIMAL TRAUMATIC BRAIN INJURIES"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S10	TX (TBI OR TBIS OR "BRAIN CONTUSION" OR "BRAIN CONTUSIONS" OR "BRAIN TRAUMA" OR "BRAIN LACERATION" OR "BRAIN LACERATIONS" OR "CEREBRAL INJURY" OR "CEREBRAL INJURIES" OR "HEAD INJURY" OR "HEAD INJURIES" OR "HEAD TRAUMA" OR "CEREBRAL TRAUMA" OR "SKULL INJURY" OR "SKULL INJURIES" OR "CRANIOCEREBRAL TRAUMA" OR "BRAIN INJURY" OR "BRAIN INJURIES") AND TX (MILD OR MINOR)	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S 9	"GCS 13-15" OR MTBI OR MTBIS OR "MILD TRAUMATIC BRAIN	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;

	INJURY" OR "MILD TRAUMATIC BRAIN INJURIES"		Nursing & Allied Health Collection: Comprehensive
S8	(MH "Brain Concussion+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S7	S6 AND (S2 OR S1)	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S6	S5 OR S4 OR S3	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S5	(MH "Recreation+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S4	(MH "Athletes+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S 3	(MH "Sports+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S2	(MH "Head Injuries+") OR TX (tbi or tbis)	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text; Nursing & Allied Health Collection: Comprehensive
S1	(MH "Head Injuries+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;

	Nursing & Allied Health	
	Collection: Comprehensive	

PubMed

Strategy	Strategy	Search Limits	Results
Question 1	<u>Original Search</u> <u>Strategy</u>	Human; January 1990-November 2012	380
Question 1	Updated Search _Original Strategy	Human; December 2012-July 2015	78
Question 2 & 3	<u>Original Search</u> <u>Strategy</u>	Human; January 1990-November 2012	956
Question 2 & 3	Updated Search _Original Strategy	Human; December 2012-July 2015	168
Question 4 & 5	<u>Original Search</u> <u>Strategy</u>	Human; January 1990-November 2012	3405
Question 4 & 5	Updated Search _Original Strategy	Human; December 2012-July 2015	790
Question 6	<u>Original Search</u> <u>Strategy</u>	Human; January 1990-November 2012	74
Question 6	Updated Search _Original Strategy	Human; December 2012-July 2015	16

PubMed Question 1_Original Search Strategy

ASSESSMENT* [TIAB] OR DIAGNOSTIC TECHNIQUES, NEUROLOGICAL[MH:NOEXP] OR NEUROLOGIC EXAMINATION[MH] OR DIAGNOSIS, DIFFERENTIAL[MH] OR DIAGNOSIS[MH:NOEXP] OR DIAGNOSIS[SH] OR DIAGNOSTIC ERRORS[MH] OR DIAGNOS*[TIAB] OR DIAGNOSTIC*[TW] OR MEASURES[TW] OR SCALE[TW] OR SCALES[TW] OR SCORE[TW] OR SCORES[TW] OR INDEX[TW] OR INDICES[TW] OR INSTRUMENT[TW] OR INSTRUMENTS[TW] OR MEASURE[TW] OR SENSORY MODALITY ASSESSMENT[TIAB] OR SENSORY STIMULATION

DIAGNOSIS:

(IRAQ WAR, 2003 [MAJR] OR IRAQ WAR [TI] OR GULF WAR [TI] OR OPERATION IRAQI FREEDOM [TI] OR GULF WARS [TI] OR AFGHAN CAMPAIGN 2001[MAJR] OR OPERATION ENDURING FREEDOM [TI] OR AFGHAN WAR [TI] OR WAR [MAJR] OR MILITARY [TI] OR BATTLEFIELD [TI] OR ARMY [TI] OR ARMED FORCES [TI] OR MARINES [TI] OR TROOPS [TI] OR SERVICEMEN [TI] OR COMBAT RELATED [TI] OR MILITARY PERSONNEL [MAJR] OR AIR FORCE [TI] OR SOLDIERS [TI] OR MILITARY MEDICINE [MAJR] OR OIF/OEF [TI] OR COMBAT VETERAN* [TI] OR COMBAT DISORDERS [MAJR] OR VETERANS [MAJR] OR ACTIVE DUTY [TI] OR SERVICE MEMBERS [TI]) NOT (MILITARY [TW] AND (CIVILIAN* [TIAB] OR SPORTS [TIAB] OR ATHLETE* [TIAB] OR SPORT RELATED [TI] OR SPORTS RELATED [TI]))

NOT

(ADULT[MH] NOT (ADULT[MH] AND (CHILD[MH] OR ADOLESCENT*[MH] OR INFANT*[MH]))

NOT

(ANIMALS[MH] NOT (ANIMALS[MH] AND HUMAN*[MH]))

NOT

MINIMAL TRAUMATIC BRAIN INJUR* [TIAB] OR MINIMAL TBI [TIAB]

OR

OR

OR

(TRAUMATIC BRAIN INJUR*[TIAB] OR TBI[TIAB] OR TBIS[TIAB] OR BRAIN CONTUSION*[TW] OR BRAIN TRAUMA*[TW] OR BRAIN LACERATION*[TW] OR CEREBRAL INJUR*[TW] OR HEAD INJUR*[TW] OR HEAD TRAUMA*[TW] OR CEREBRAL TRAUMA*[TIAB] OR SKULL INJUR*[TW] OR CRANIOCEREBRAL TRAUMA[MH:NOEXP] OR BRAIN INJUR* [TW]) AND (MILD[TIAB] OR MINOR[TIAB])

GCS 13-15[TIAB] OR BRAIN CONCUSSION[MH] OR CONCUSSION*[TW] OR CONCUSSIVE[TIAB] OR COMMOTIO CEREBRI[TW] OR POST CONCUSSIVE[TW] OR POSTCONCUSSIVE[TW] OR MILD TBI[TIAB] OR MILD TBIS[TIAB] OR MTBI[TIAB] OR MTBIS[TIAB] OR MILD TRAUMATIC BRAIN INJUR*[TIAB]

SPORT*[TW] OR ATHLETE[TW] OR ATHLETES[TW] OR ATHLETIC*[TW] OR RECREATION*[TIAB]) AND (HEAD INJUR*[TW] OR BRAIN INJUR*[TW] OR HEAD TRAUMA*[TIAB] OR BRAIN TRAUMA*[TIAB] OR SKULL INJUR*[TIAB] OR TBI[TIAB] OR TBIS[TIAB] OR CEREBRAL TRAUMA*[TW] OR CEREBRAL INJUR*[TW]

POPULATION AND DIAGNOSIS

POPULATION:

ASSESSMENT[TIAB] OR WESSEX HEAD INJURY MATRIX[TIAB] OR WESTERN NEURO SENSORY STIMULATION PROFILE[TIAB] OR SENSITIVITY AND SPECIFICITY[MH] OR SPECIFICITY[TIAB] OR PREDICTIVE VALUE OF TESTS[MH] OR CONFIDENCE INTERVAL[MH] OR DETECT*[TIAB] OR CHARACTERISTIC*[TIAB] OR CATEGORIZ*[TIAB] OR CATEGORIS*[TIAB] OR DISTINGUISH*[TIAB] OR DIFFERENTIAT*[TIAB] OR BEHAVIOURAL ASSESSMENT*[TIAB] OR BEHAVIORAL ASSESSMENT*[TIAB] OR BEHAVIOR ASSESSMENT*[TIAB] OR BEHAVIOUR ASSESSMENT*[TIAB] OR NEUROPSYCHOLOGICAL TESTS[MH:NOEXP] OR TRAUMA SEVERITY INDICES [MH] OR DETERMINANT* [TIAB] OR KING DEVICK TEST [TIAB] OR MILITARY ACUTE CONCUSSION EVALUATION [TIAB] OR REACTION TIME [MH] OR REACTION TIME* [TIAB] OR RESPONSE LATENCY [TW] OR SCREENING TEST* [TIAB] OR SCREENING TOOL* [TIAB] OR BALANCE TEST* [TW] OR POSTURAL BALANCE [MH] OR POSTURAL EQUILIBRIUM [TW] OR SENSATION DISORDERS [MH] OR BALANCE ASSESSMENT [TIAB] OR MEDICAL SYMPTOM VALIDITY TEST [TIAB] OR EVOKED POTENTIALS[MH] OR EVOKED POTENTIAL*[TIAB]OR CLASSIFICATION [MH] OR CLASSIFICATION [SH]OR NEUROCOGNITIVE TEST* [TIAB]OR PSYCHOMOTOR PERFORMANCE [MH] OR MEMORY [MH]OR IMMEDIATE POST-CONCUSSION ASSESSMENT AND COGNITIVE TEST* [TIAB] OR ELECTROENCEPHALOGRAPHY[MH] OR ELECTROENCEPHALOGRAPHY[TIAB] OR ELECTROENCEPHALOGRAM*[TIAB] OR ECHOENCEPHALOGRAPHY[MH] OR EEG[TIAB] OR EEGS[TIAB] OR QEEG[TIAB] OR QEEGS[TIAB] OR ELECTROMYOGRAPH*[TW] OR EMG[TIAB] OR EMGS[TIAB] OR ELECTROPHYSIOLOGIC TECHNIQUE*[TIAB]

OR

PREDICTIVE[TIAB] AND VALUE*[TIAB]

PubMed Question 2 & 3_Original Search Strategy

(POPULATION AND ACUTE CARE SETTING) AND (IMAGING OR INTRACRANIAL INJURY)

POPULATION:

GCS 13-15[TIAB] OR BRAIN CONCUSSION[MH] OR CONCUSSION*[TW] OR COMMOTIO CEREBRI[TW] OR POST CONCUSSIVE[TW] OR POSTCONCUSSIVE[TW] OR MTBI[TIAB] OR MTBIS[TIAB] OR BRAIN INJURIES[MH] OR BRAIN INJUR*[TW] OR TBI[TIAB] OR TBIS[TIAB] OR BRAIN CONTUSION*[TW] OR BRAIN TRAUMA*[TW] OR BRAIN LACERATION*[TW] OR HEAD TRAUMA*[TW] OR CEREBRAL TRAUMA*[TW] OR CEREBRAL INJUR*[TW] OR SKULL INJUR*[TW] OR SKULL TRAUMA[TW] OR CRANIOCEREBRAL TRAUMA[MH:noexp] OR HEAD INJUR*[TW]

NOT

(IRAQ WAR, 2003 [MAJR] OR IRAQ WAR [TI] OR GULF WAR [TI] OR OPERATION IRAQI FREEDOM [TI] OR GULF WARS [TI] OR AFGHAN CAMPAIGN 2001[MAJR] OR OPERATION ENDURING FREEDOM [TI] OR AFGHAN WAR [TI] OR WAR [MAJR] OR MILITARY [TI] OR BATTLEFIELD [TI] OR ARMY [TI] OR ARMED FORCES [TI] OR MARINES [TI] OR TROOPS [TI] OR SERVICEMEN [TI] OR COMBAT RELATED [TI] OR MILITARY PERSONNEL [MAJR] OR AIR FORCE [TI] OR SOLDIERS [TI] OR MILITARY MEDICINE [MAJR] OR OIF/OEF [TI] OR COMBAT VETERAN* [TI] OR COMBAT

(ADULT[MH] NOT (ADULT[MH] AND (CHILD[MH] OR ADOLESCENT*[MH] OR INFANT*[MH]))

NOT

NOT

(ANIMALS[MH] NOT (ANIMALS[MH] AND HUMAN*[MH]))

DISORDERS [MAJR] OR VETERANS [MAJR] OR ACTIVE DUTY [TI] OR SERVICE MEMBERS [TI]) NOT (MILITARY [TW] AND (CIVILIAN* [TIAB] OR SPORTS [TIAB] OR ATHLETE* [TIAB] OR SPORT RELATED [TI] OR SPORTS RELATED [TI]))

ACUTE CARE SETTING:

MEDICAL CENTRE*[TIAB] OR TRAUMA CENTRE*[TIAB] OR NON TRAUMA CENTRE* [TIAB] OR NONTRAUMA CENTRE* [TIAB] OR INPATIENT*[TW] OR MEDICAL CENTER*[TW] OR ACADEMIC MEDICAL CENTERS[MH] OR HEALTH FACILITIES[MH] OR ACUTE CARE[TW] OR HOSPITALS[MH] OR HOSPITAL*[TW] OR NON TRAUMA CENTER*[TW] OR NONTRAUMA CENTER*[TW] OR TRAUMA CENTERS[MH] OR TRAUMA CENTER*[TW] OR TRAUMA UNIT*[TW] OR EMERGENCY SERVICE, HOSPITAL[MH] OR EMERGENCY[TW] OR ACUTE SETTING* [TIAB] OR TERTIARY CARE [TIAB]

IMAGING:

MAGNETIC RESONANCE IMAGING [MH] OR ZEUGMATOGRAPHY [TW] OR TOMOGRAPHY [TW] OR MRI SCAN* [TW] OR FMRI [TW] OR FMRIS [TW] OR MRI [TW] OR MRIS [TW] OR IMAGING* [TW] OR NEUROIMAGING [MH:NOEXP] OR NEUROIMAGING* [TIAB] OR NEURO IMAGING* [TIAB] OR TOMOGRAPHY, X-RAY COMPUTED [MH] OR CAT SCAN* [TW] OR CT SCAN* [TW] OR CT X RAY* [TW] OR TOMODENSITOMETRY [TW] OR CINE-CT [TW] OR X-RAYS [MH] OR X RAY* [TIAB] OR RADIOGRAPHY [SH]

INTRACRANIAL INJURY:

INTRACRANIAL INJUR* [TIAB] OR INTRACRANIAL HEMORRHAGES[MH] OR BRAIN HEMORRHAGE, TRAUMATIC[MH] OR HEMATOMA, SUBDURAL[MH] OR HEMORRHAGE*[TW] OR HAEMORRHAGE*[TIAB] OR BRAIN BLEED*[TIAB] OR HAEMATOMA*[TIAB] OR HEMATOMA*[TIAB] OR PITUITARY APOPLEXY[TW] OR EDEMA[TW] OR DIFFUSE AXONAL INJURY[MH] OR AXONAL SWELLING[TW] OR RETRACTION BALLS[TIAB] OR CEREBRAL ATROPHY[TIAB] OR INTRACEREBRAL INJUR*[TIAB] OR OEDEMA[TIAB] OR BRAIN CONTUSION*[TIAB] OR CEREBRAL CONTUSION*[TIAB] OR AXONAL INJUR*[TW] OR PNEUMOCEPHALUS[MH] OR PNEUMOCEPHALUS[TIAB] OR CRANIAL PNEUMOCYST*[TW] OR INTRACRANIAL GAS[TW] OR MICROBLEED*[TIAB] OR MICRO BLEED* [TIAB] OR WHITE MATTER INJUR*[TIAB] OR SKULL FRACTURES[MH] OR SKULL FRACTURE*[TIAB] OR CRANIAL INJUR*[TW] OR LESION*[TW] OR NEUROPATHOLOG*[TW] OR BLEEDING[TIAB] OR MICROHEMORRHAGE*[TIAB] OR MICROHAEMORRHAGE*[TIAB] OR MICRO HEMORRHAGE*[TIAB] OR MICRO HAEMORRHAGE*[TIAB] OR INTRACRANIAL SEQUELAE[TIAB]

PubMed Question 4 & 5_Original Search Strategy

POPULATION AND (PROGNOSIS OR RISK FACTORS FOR SEQUELAE)

POPULATION:

SPORT*[TW] OR ATHLETE[TW] OR ATHLETES[TW] OR ATHLETIC*[TW] OR RECREATION*[TIAB]) AND (HEAD INJUR*[TW] OR BRAIN INJUR*[TW] OR HEAD TRAUMA*[TIAB] OR BRAIN TRAUMA*[TIAB] OR SKULL INJUR*[TIAB] OR TBI[TIAB] OR TBIS[TIAB] OR CEREBRAL TRAUMA*[TW] OR CEREBRAL INJUR*[TW]

OR

GCS 13-15[TIAB] OR BRAIN CONCUSSION[MH] OR CONCUSSION*[TW] OR CONCUSSIVE[TIAB] OR COMMOTIO CEREBRI[TW] OR POST CONCUSSIVE[TW] OR POSTCONCUSSIVE[TW] OR MILD TBI[TIAB] OR MILD TBIS[TIAB] OR MTBI[TIAB] OR MTBIS[TIAB] OR MILD TRAUMATIC BRAIN INJUR*[TIAB] (TRAUMATIC BRAIN INJUR*[TIAB] OR TBI[TIAB] OR TBIS[TIAB] OR BRAIN CONTUSION*[TW] OR BRAIN TRAUMA*[TW] OR BRAIN LACERATION*[TW] OR CEREBRAL INJUR*[TW] OR HEAD INJUR*[TW] OR HEAD TRAUMA*[TW] OR CEREBRAL TRAUMA*[TIAB] OR SKULL INJUR*[TW] OR CRANIOCEREBRAL TRAUMA[MH:NOEXP] OR BRAIN INJUR* [TW]) AND (MILD[TIAB] OR MINOR[TIAB])

OR

OR

NOT

MINIMAL TRAUMATIC BRAIN INJUR*[TIAB] OR MINIMAL TBI[TIAB]

(ADULT[MH] NOT (ADULT[MH] AND (CHILD[MH] OR ADOLESCENT*[MH] OR INFANT*[MH]))

(IRAQ WAR, 2003 [MAJR] OR IRAQ WAR [TI] OR GULF WAR [TI] OR OPERATION IRAQI FREEDOM [TI] OR GULF WARS [TI] OR AFGHAN CAMPAIGN 2001[MAJR] OR OPERATION ENDURING FREEDOM [TI] OR AFGHAN WAR [TI] OR WAR [MAJR] OR MILITARY [TI] OR BATTLEFIELD [TI] OR ARMY [TI] OR ARMED FORCES [TI] OR MARINES [TI] OR TROOPS [TI] OR SERVICEMEN [TI] OR COMBAT RELATED [TI] OR MILITARY PERSONNEL [MAJR] OR AIR FORCE [TI] OR SOLDIERS [TI] OR MILITARY MEDICINE [MAJR] OR OIF/OEF [TI] OR COMBAT VETERAN* [TI] OR COMBAT DISORDERS [MAJR] OR VETERANS [MAJR] OR ACTIVE DUTY [TI] OR SERVICE MEMBERS [TI]) NOT (MILITARY [TW] AND (CIVILIAN* [TIAB] OR SPORTS [TIAB] OR ATHLETE* [TIAB] OR SPORT RELATED [TI] OR SPORTS RELATED [TI]))

EXTENDED [TIAB] OR LASTING [TIAB] OR LONGTERM [TIAB] OR LONG TERM [TIAB] OR "LONGER THAN" [TIAB] OR "MORE THAN" [TIAB] OR "GREATER THAN" [TIAB] OR DELAYED [TIAB] OR FAILURE* [TW] OR PROLONG* [TIAB] OR PROTRACTED [TIAB] OR COMPLICATED [TIAB] OR ENDURING [TIAB] OR CONTINUING [TIAB] OR CONTINUED [TIAB]OR UNRESOLVED [TIAB] OR UNREMITTING [TIAB] OR LINGERING [TIAB] OR PROBLEMATIC [TIAB]) AND

CONVALESCE[TIAB] OR CONVALESCED[TIAB] OR CONVALESCES[TIAB] OR CONVALESCING[TIAB] OR RECOVER[TIAB] OR RECOVERS[TIAB] OR RECOVERY[TIAB] OR RECOVERING[TIAB] OR RECOVERIES[TIAB] OR IMPROVE* [TW] OR IMPROVEMENT* [TIAB] OR IMPROVING[TW] OR RESTORATION[TW] OR RESTORE*[TW] OR RECOVERY OF

FUNCTION[TW] OR RESOLUTION* [TW] OR PROGNOSIS [MH] OR DISEASE FREE SURVIVAL [MH OR PROGNOS*[TW] OR PREDICT* [TIAB] OR NOMOGRAM* [TW] OR MEDICAL FUTILITY [MH] OR FUTIL* [TW] OR TREATMENT OUTCOME [MH:NOEXP] OR TREATMENT FAILURE [MH] OR PREDICTIVE VALUE OF TESTS [MH] OR FORECASTING [MH] OR

(PATHOPHYSIOLOG* [TW] OR DYSFUNCTION [TIAB] OR SYMPTOMATOLOGY [TIAB] OR SUBSYMPTOM* [TIAB] OR SUB SYMPTOM* [TIAB] OR SYMPTOMS [TW] OR SYMPTOMATIC [TIAB] OR DISEASE-FREE SURVIVAL[MH] OR TREATMENT FAILURE[MH] OR DISEASE PROGRESSION[MH] OR DISEASE PROGRESSION [TIAB] OR SEQUELAE [TW] OR ENDURING [TW] OR DELAYED [TIAB] OR PROLONGED [TIAB] OR ONGOING [TIAB] OR PERSISTENT [TIAB] OR

(FUNCTIONAL LEVEL [TIAB] OR PROGRESS*[TW] OR RECUPERAT*[TIAB] OR CONVALESCENCE[TW] OR

NOT

NOT

PROGNOSIS:

FORECAST* [TIAB] OR PROJECTION* [TW]

RISK FACTORS FOR SEQUELAE:

(ANIMALS[MH] NOT (ANIMALS[MH] AND HUMAN*[MH]))

NOT

MINIMAL TRAUMATIC BRAIN INJUR*[TIAB] OR MINIMAL TBI[TIAB]

OR

(TRAUMATIC BRAIN INJUR*[TIAB] OR TBI[TIAB] OR TBIS[TIAB] OR BRAIN CONTUSION*[TW] OR BRAIN TRAUMA*[TW] OR BRAIN LACERATION*[TW] OR CEREBRAL INJUR*[TW] OR HEAD INJUR*[TW] OR HEAD TRAUMA*[TW] OR CEREBRAL TRAUMA*[TIAB] OR SKULL INJUR*[TW] OR CRANIOCEREBRAL TRAUMA[MH:NOEXP] OR BRAIN INJUR* [TW]) AND (MILD[TIAB] OR MINOR[TIAB])

OR

GCS 13-15[TIAB] OR BRAIN CONCUSSION[MH] OR CONCUSSION*[TW] OR CONCUSSIVE[TIAB] OR COMMOTIO CEREBRI[TW] OR POST CONCUSSIVE[TW] OR POSTCONCUSSIVE[TW] OR MILD TBI[TIAB] OR MILD TBIS[TIAB] OR MTBI[TIAB] OR MTBIS[TIAB] OR MILD TRAUMATIC BRAIN INJUR*[TIAB]

OR

SPORT*[TW] OR ATHLETE[TW] OR ATHLETES[TW] OR ATHLETIC*[TW] OR RECREATION*[TIAB]) AND (HEAD INJUR*[TW] OR BRAIN INJUR*[TW] OR HEAD TRAUMA*[TIAB] OR BRAIN TRAUMA*[TIAB] OR SKULL INJUR*[TIAB] OR TBI[TIAB] OR TBIS[TIAB] OR CEREBRAL TRAUMA*[TW] OR CEREBRAL INJUR*[TW]

POPULATION:

POPULATION AND ONGOING SYMPTOMS AND TREATMENTS

PubMed Question 6_Original Search Strategy

(RISK [MH:NOEXP]OR RISK ASSESSMENT [MH] OR RISK FACTORS [MH] OR RISK* [TIAB] OR RISK ADJUSTMENT [MH] OR PRECIPITATING FACTORS [MH] OR EPIDEMIOLOGIC FACTORS [MH] OR CAUSALITY [MH] OR CAUSALITY [TW] OR CAUSATION* [TW] OR ETIOLOGY [SH:NOEXP] OR EPIDEMIOLOGY [SH] OR FEATURE* [TIAB] OR FACTOR [TIAB] OR FACTORS [TIAB] OR IDENTIFY [TIAB] OR IDENTIFICATION [TIAB] OR IDENTIFIES [TIAB] OR INDENTIFIED [TIAB] OR AGE FACTORS [MH] OR SEX [TW] OR SEXES [TW] OR GENDER [TW] OR TIME SINCE INJUR* [TIAB] OR TIME FACTORS [MH] OR BETWEEN GROUP DIFFERENCE* [TIAB] OR MECHANISM OF INJURY [TIAB] OR MECHANISMS OR INJURY [TIAB] OR INDICATOR* [TIAB] OR CHARACTERISTIC* [TIAB] OR ANALYSIS OF VARIANCE [MH])

AND

PERSISTENCE [TIAB] OR PERSISTING [TIAB] OR LASTING [TIAB] OR LONGTERM [TIAB] OR LONG TERM [TIAB] OR POST CONCUSSION SYNDROME* [TIAB] OR POST-CONCUSSION SYNDROME [MH] OR POST-CONCUSSIVE SYNDROME [TW] OR POSTCONCUSSION SYNDROME* [TIAB] OR POSTCONCUSSIVE SYNDROME* [TIAB] OR UNRESOLVED [TIAB] OR CONTINUING [TIAB] OR CONTINUE* [TIAB] OR UNREMITTING [TIAB] OR LINGERING [TIAB] OR PROTRACTED [TIAB] OR POSTTRAUMATIC [TW] OR POST TRAUMATIC [TIAB] OR COMPLICATIONS [SH] OR COMPLICATION* [TIAB] OR COMPLICATED [TIAB] OR EXTENDED [TIAB] OR DISABILITY [TW] OR DISABILITIES [TW] OR DISABLED [TW] OR IMPAIRMENT [TW] OR IMPAIRED [TIAB] OR FAILURE [TW] OR REFRACTORY [TW] OR MORE SEVERE* [TIAB])

(ANIMALS[MH] NOT (ANIMALS[MH] AND HUMAN*[MH]))

NOT

(ADULT[MH] NOT (ADULT[MH] AND (CHILD[MH] OR ADOLESCENT*[MH] OR INFANT*[MH]))

NOT

(IRAQ WAR, 2003 [MAJR] OR IRAQ WAR [TI] OR GULF WAR [TI] OR OPERATION IRAQI FREEDOM [TI] OR GULF WARS [TI] OR AFGHAN CAMPAIGN 2001[MAJR] OR OPERATION ENDURING FREEDOM [TI] OR AFGHAN WAR [TI] OR WAR [MAJR] OR MILITARY [TI] OR BATTLEFIELD [TI] OR ARMY [TI] OR ARMED FORCES [TI] OR MARINES [TI] OR TROOPS [TI] OR SERVICEMEN [TI] OR COMBAT RELATED [TI] OR MILITARY PERSONNEL [MAJR] OR AIR FORCE [TI] OR SOLDIERS [TI] OR MILITARY MEDICINE [MAJR] OR OIF/OEF [TI] OR COMBAT VETERAN* [TI] OR COMBAT DISORDERS [MAJR] OR VETERANS [MAJR] OR ACTIVE DUTY [TI] OR SERVICE MEMBERS [TI]) NOT (MILITARY [TW] AND (CIVILIAN* [TIAB] OR SPORTS [TIAB] OR ATHLETE* [TIAB] OR SPORT RELATED [TI] OR SPORTS RELATED [TI]))

ONGOING SYMPTOMS:

LONGTERM [TIAB] OR SYMPTOMS [TW] OR SYMPTOMATIC [TIAB] OR SEQUELAE [TW] OR ENDURING [TW] OR DELAYED [TIAB] OR PROLONG* [TIAB] OR ONGOING [TIAB] OR PERSISTENT [TIAB] OR PERSISTENCE [TIAB] OR PERSISTING [TIAB] OR LASTING [TIAB] OR LONG TERM [TIAB] OR POST-CONCUSSION SYNDROME [MH] OR POST-CONCUSSIVE SYNDROME [TW] OR POSTCONCUSSION SYNDROME* [TIAB] OR POSTCONCUSSIVE SYNDROME* [TIAB] OR SYMPTOMATOLOGY [TIAB] OR SUBSYMPTOM* [TIAB] OR UNRESOLVED [TIAB] OR CONTINUING [TIAB] OR CONTINUED [TIAB] OR UNREMITTING [TIAB] OR LINGERING [TIAB] OR PROTRACTED [TIAB] OR "LONGER THAN" [TIAB] OR "MORE THAN" [TIAB] OR "GREATER THAN" [TIAB]OR POST CONCUSSION SYNDROME [MH] OR POST CONCUSSION SYNDROME* [TIAB] OR POSTCONCUSSION SYNDROME* [TIAB] OR POSTTRAUMATIC [TW] OR POST TRAUMATIC [TIAB] OR COMPLICATIONS [SH] OR COMPLICATION* [TIAB] OR COMPLICATED [TIAB] OR EXTENDED [TIAB] OR RECOVERY OR RECOVER [TIAB] OR RECOVERED [TIAB] OR COMMUNITY DWELLING [TIAB] OR DISABILITY [TW] OR DISABILITIES [TW] OR DISABLED [TW] OR IMPAIRMENT [TW] OR IMPAIRED [TIAB] OR REHABILITATION [MH] OR REINTEGRATION [TW] OR REHABILITATION [TIAB] OR RE-INTEGRATION [TIAB] OR FAILURE [TW] OR REMEDIATION [TW] OR REMEDIATE [TW] OR REFRACTORY [TW] OR EXTENDED [TIAB] OR "LONGER THAN" [TIAB] OR "MORE THAN" [TIAB] OR "GREATER THAN" [TIAB] OR PROBLEMATIC [TIAB]

TREATMENTS:

THERAPY [SH] OR THERAPEUTIC USE [SH] OR DRUG THERAPY[MH]OR PHARMACOTHERAP* [TW] OR PHARMACOLOGIC* [TW] OR DRUG THERAP* [TIAB]OR THERAPY [TW] OR THERAPIES [TW] OR THERAPEUTIC* [TW] OR TREATMENT* [TW] OR TREATMENT OUTCOME [MH] OR DIET THERAPY [MH] OR DIET THERAPY [SH] OR NUTRITION THERAPY [MH] OR REHABILITATION OUTCOME* [TW] OR REHABILITATION [SH] OR REHABILITAT* [TIAB] OR TREATED [TIAB] OR MANAGEMENT [TIAB] OR INTERVENTION* [TIAB] OR BIOFEEDBACK, PSYCHOLOGY [MH] OR BIOFEEDBACK [TIAB] OR HYPERBARIC OXYGENATION [MH] OR EXERCISE THERAPY [MH] OR PHYSICAL THERAPY MODALITIES [MH] OR PHYSIOTHERAPY [TW] OR COGNITIVE THERAPY [MH] OR HYPOTHERMIA, INDUCED [MH:NOEXP] OR HYPOTHERMIA [TIAB] OR REST [MH] OR NEUROREHABILIT* [TIAB] OR STRESS REDUCTION [TIAB] OR PHYSICAL REST [TIAB] OR COGNITIVE REST [TIAB] OR NEUROPROTECTIVE AGENTS [MH] OR SALINE SOLUTION, HYPERTONIC [MH] OR NUTRACEUTICAL* [TIAB] OR DIETARY SUPPLEMENTS [MH] OR DIETARY SUPPLEMENT* [TIAB] OR NEUTRACEUTICAL* [TIAB] OR NEUROFEEDBACK [TIAB] OR NEURO FEEDBACK [TIAB] OR BIO FEEDBACK [TIAB] OR MASSAGE [MH] OR MASSAGE [TIAB] OR COGNITIVE RETRAINING [TIAB] OR STIMULATION [TW] OR SENSORY [TW] OR DEPRIVATION[TW] OR RANGE OF MOTION EXERCISE* [TW] OR POSITIONING [TIAB] OR POSITIONAL CHANGE* [TW] OR POSTURAL CHANGE* [TIAB] OR ACETAMINOPHEN [TW] OR ASPIRIN [TW] OR IBUPROFEN [TW] NAPROXEN [TW] OR DICLOFENAC [TW] OR ANALGESICS [TW] OR NSAIDS [TW] OR ANTIDEPRESSANT* [TW] OR AMITRIPTYLINE [TW] OR NORTRIPTYLINE [TW] OR ANTICONVULSANT* [TW] OR VALPROIC ACID [TW] OR TOPIRAMATE [TW] OR GABAPENTIN [TW] OR PROPRANOLOL [TW] OR B-ADRENERGIC ANTAGONIST* [TW] OR METOPROLOL [TW] OR DIHYDROERGOTAMINE [TW] OR ERGOT [TW] OR SUMATRIPTAN [TW] OR TRIPTANS [TW] OR ZOLMITRIPTAN [TW] OR RIZATRIPTAN [TW] OR VESTIBULAR SUPPRESSANT* [TW] OR MECLIZINE [TW] OR SCOPOLAMINE [TW] OR DIMEHYDRINATE [TW] OR BENZODIAZEPINES [TW] OR LORAZEPAM [TW] OR CLONAZEPAM [TW] OR DIAZEPAM [TW] OR NEUROSTIMULANT* [TW] OR METHYLPHENIDATE [TW] OR DEXTROAMPHETAMINE [TW] OR MODAFANIL [TW] OR AMANTADINE [TW] OR ATOMOXETINE [TW] OR ANTIEMETIC* [TW] OR ONDANSETRON [TW] OR PHENERGAN [TW] OR SEDATIVE-HYPNOTIC* [TW] OR ZOLPIDEM [TW] OR SEROTONIN MODULATOR* [TW] OR TRAZODONE [TW] OR A-ADRENERGIC ANTAGONIST* [TW] OR PRAZOSIN [TW] OR MELATONIN [TW] OR SERTRALINE [TW] OR CITALOPRAM [TW] OR ESCITALOPRAM [TW] OR PAROXETINE [TW] OR FLUOXETINE [TW] OR BUPROPION [TW] OR ACETYLCHOLINESTERASE INHIBITOR* [TW] OR SELECTIVE SEROTONIN REUPTAKE INHIBITOR* [TW] OR DONEPEZIL [TW] OR RIVASTIGMINE [TW] OR GALANTAMINE [TW] OR CYTIDINE DIPHOSPHATE CHOLINE [TW]

EMBASE SEARCH STRATEGIES—Updated Search

December 2012-July 2015

Question 1:

<u>2,208</u> #2.3 AND #4.31

<u>3,075,946</u>

#2.3 #2.1 OR #2.2

696,823

#2.2

'evoked response'/exp OR 'response time'/exp OR 'body equilibrium'/exp OR 'psychomotor performance'/de OR 'task performance'/exp OR 'memory'/exp OR 'sensory dysfunction'/exp

2,515,609

#2.1

'diagnostic procedure'/de OR 'clinical assessment tool'/exp OR 'clinical observation'/exp OR 'computer assisted diagnosis'/de OR 'delayed diagnosis'/exp OR 'diagnostic accuracy'/exp OR 'diagnostic error'/exp OR 'diagnostic reasoning'/exp OR 'diagnostic test'/de OR 'diagnostic test accuracy study'/exp OR 'diagnostic value'/exp OR 'differential diagnosis'/exp OR 'early diagnosis'/exp OR 'electrodiagnosis'/de OR 'laboratory diagnosis'/exp OR 'molecular diagnosis'/exp OR 'physical examination'/de OR 'qualitative diagnosis'/exp OR 'quantitative diagnosis'/exp OR 'diagnosis'/de OR 'diagnostic approach route'/exp OR 'evaluation or follow up' OR 'clinical assessment'/exp OR 'clinical evaluation'/exp OR 'clinical global impression scale'/exp OR 'course evaluation'/exp OR 'expanded disability status scale'/exp OR 'functional assessment inventory'/exp OR 'gross motor function classification system'/exp OR 'gross motor function measure'/exp OR 'international classification of functioning, disability or health' OR 'oars multidimensional functional assessment questionnaire'/exp OR 'patient assessment'/exp OR 'sensory evaluation'/exp OR 'examination'/de OR 'contact examination'/exp OR 'medical examination'/de OR 'clinical examination'/exp OR 'functional assessment'/exp OR 'measurement'/de OR 'named inventories, questionnaires or rating scales' OR 'neurologic examination'/de OR 'dubowitz neurological assessment'/exp OR 'electroencephalography'/exp OR 'neuropsychological test'/exp OR 'stabilography'/exp OR 'tilt table test'/exp OR 'non invasive measurement/exp OR 'rating scale'/exp OR 'scoring system'/exp OR 'screening'/de OR 'screening test'/exp OR 'sensory system examination'/de OR 'vestibular test'/exp OR 'summated rating scale'/exp

<u>5,412</u>

#4.31 #4.27 OR #4.30

<u>163</u> #4.30 #4.29 NOT #4.19

<u>163</u>

#4.29 #4.28 NOT #4.24

<u>163</u>

#4.28

('glasgow coma scale'/exp OR 'glasgow outcome scale'/exp) AND '13-15':ab,ti

<u>5,295</u> #4.27 #4.23 NOT #4.26

<u>11,241</u>

#4.26 #4.24 OR #4.25

<u>30</u> #4.25 'oif/oef':ti AND [1990-2013]/py

<u>11,235</u>

#4.24

'iraq war':ti OR 'gulf war':ti OR 'gulf wars':ti OR 'operation iraqi freedom':ti OR 'afghan campaign':ti OR 'operation enduring freedom':ti OR military:ti OR battlefield:ti OR army:ti OR armed AND forces:ti OR marines:ti OR troops:ti OR servicemen:ti OR 'combat related':ti OR 'military personnel':ti OR 'air force':ti OR soldiers:ti OR 'combat veteran':ti OR 'combat disorder':ti OR 'combat disorders':ti OR veterans:ti OR 'active duty':ti OR 'service members':ti AND [1990-2013]/py

<u>5,335</u>

#4.23

#4.22 AND ('case report'/de OR 'clinical article'/de OR 'clinical protocol'/de OR 'clinical trial'/de OR 'cohort analysis'/de OR 'comparative study'/de OR 'control group'/de OR 'controlled clinical trial'/de OR 'controlled study'/de OR 'human'/de OR 'human tissue'/de OR 'interview'/de OR 'major clinical study'/de OR 'medical record review'/de OR 'methodology'/de OR 'model'/de OR 'nonhuman'/de OR 'normal human'/de OR 'outcomes research'/de OR 'practice guideline'/de OR 'prospective study'/de OR 'questionnaire'/de OR 'retrospective study'/de OR 'systematic review'/de) AND ('article'/it OR 'article in press'/it OR 'conference abstract'/it OR 'conference paper'/it OR 'letter'/it OR 'review'/it OR 'short survey'/it)

<u>6,018</u> #4.22 #4.20 NOT #4.21

452,516

#4.21 'editorial'/exp

<u>6,140</u>

#4.20 #4.16 NOT #4.19

<u>30,949</u>

#4.19 #4.17 NOT #4.18

<u>1,241</u>

#4.18

('military phenomena'/exp/mj OR 'veteran'/exp/mj) AND (civilian*:ab,ti OR sport* OR athlete*) AND [1990-2013]/py

<u>32,190</u>

#4.17 'military phenomena'/exp/mj OR 'veteran'/exp/mj

<u>6,202</u>

#4.16 #4.12 NOT #4.15

8,001,374

#4.15 #4.13 NOT #4.14

<u>10,227,651</u>

#4.14 'animal'/exp AND 'human'/exp AND [1990-2013]/py

<u>18,229,025</u>

#4.13 'animal'/exp

<u>7,286</u>

#4.12 #4.8 NOT #4.11

4,719,417

#4.11 #4.9 NOT #4.10

665,955

#4.10

'adult'/exp OR 'aged'/exp OR 'middle aged'/exp OR 'embryo'/exp OR 'fetus'/exp AND ('newborn'/exp OR 'child'/exp OR 'adolescent'/exp) AND [1990-2013]/py

<u>5,385,372</u>

#4.9

'adult'/exp OR 'aged'/exp OR 'middle aged'/exp OR 'embryo'/exp OR 'fetus'/exp

<u>10,092</u>

#4.8

#4.4 OR #4.5 OR #4.6 OR #4.7 AND [embase]/lim AND [1990-2013]/py

<u>974</u>

#4.7

mtbi:ab,ti OR mtbis:ab,ti OR 'minimal traumatic brain injury':ab,ti OR 'minimal traumatic brain injuries':ab,ti OR 'minimal tbi':ab,ti OR 'minimal tbis':ab,ti

<u>9,904</u> #4.6 #4.2 AND (mild:ab,ti OR minor:ab,ti)

<u>3,409</u> #4.5 'brain concussion'/exp

<u>2,878</u> #4.4 #4.2 AND #4.3

<u>103,017</u> #4.3 'sport'/exp OR 'athlete'/exp

<u>156,672</u> #4.2 'brain injury'/exp OR 'head injury'/de OR 'skull injury'/exp

<u>3,578,947</u>

#4.1

'therapy'/de OR 'acupuncture'/exp OR 'animal assisted therapy'/exp OR 'apitherapy'/exp OR 'biological therapy'/exp OR 'computer assisted therapy'/exp OR 'conservative treatment'/exp OR 'counseling'/exp OR 'device therapy'/exp OR 'drug therapy'/exp OR 'early intervention'/exp OR 'experimental therapy'/exp OR 'fluid therapy'/exp OR 'homeopathy'/exp OR 'intensive care'/de OR 'neuromonitoring'/exp OR 'blood pressure monitoring'/exp OR 'body temperature monitoring'/exp OR 'capnography'/exp OR 'life sustaining treatment'/exp OR 'maintenance therapy'/de OR 'patient positioning'/exp OR 'rehabilitation care'/exp OR 'radiotherapy'/de OR 'cognitive rehabilitation'/exp OR 'functional training'/exp OR 'muscle training'/exp OR 'occupational therapy'/exp OR 'rejuvenation'/exp OR 'sensorimotor integration'/exp OR 'speech rehabilitation'/exp OR 'sleep therapy'/exp OR 'supplementation'/exp OR 'systemic therapy'/exp OR 'therapy effect'/exp OR 'treatment response'/exp

Questions 2 & 3

<u>3,539</u>

2,194,721 #7.19 #7.17 OR #7.18

<u>897,748</u>

#7.18

'diffuse axonal injury'/exp OR 'nerve fiber degeneration'/exp OR 'axonal swelling':ab,ti OR 'retraction balls':ab,ti OR 'brain atrophy'/exp OR 'intracerebral injury':ab,ti OR 'intracerebral injuries':ab,ti OR 'edema'/de OR 'brain edema'/exp OR oedema:ab,ti OR 'axonal injury'/exp OR 'pneumocephalus'/exp OR 'cranial pneumocyst':ab,ti OR 'cranial pneumocysts':ab,ti OR 'intracranial gas':ab,ti OR 'white matter injury':ab,ti OR 'white matter injuries':ab,ti OR 'white matter injury'/exp OR 'skull fracture'/exp OR 'cranial injury':ab,ti OR 'cranial injuries':ab,ti OR lesion*:ab,ti OR 'neuropathology'/exp OR microhemorrhage*:ab,ti OR 'micro hemorrhage':ab,ti OR 'micro hemorrhages':ab,ti OR 'intracranial sequelae':ab,ti

<u>1,504,683</u>

#7.17 #7.13 OR #7.16

<u>347,602</u>

#7.16 #7.14 OR #7.15

347,060

#7.15

'bleeding'/de OR 'brain hemorrhage'/exp OR 'hematoma'/de OR 'brain hematoma'/exp OR 'traumatic hematoma'/exp OR 'meningeal hemorrhage'/exp OR 'brain bleed':ab,ti OR 'brain bleeding':ab,ti OR microbleed*:ab,ti OR 'micro-bleed':ab,ti OR 'micro-bleed':ab,ti OR 'micro-bleed':ab,ti OR hemorrhage*:ab,ti OR haematoma*:ab,ti

<u>782</u>

#7.14

'intracranial injury':ab,ti OR 'intracranial injuries':ab,ti

<u>1,213,947</u>

#7.13

'nuclear magnetic resonance imaging'/exp OR zeugmatography:ab,ti OR 'tomography'/de OR 'brain tomography'/exp OR 'computer assisted tomography'/exp OR 'emission tomography'/exp OR 'optical coherence tomography'/exp OR tomography:ab,ti OR 'mri scan':ab,ti OR 'mri scans':ab,ti OR fmri:ab,ti OR fmris:ab,ti OR mri:ab,ti OR mris:ab,ti OR 'neuroimaging'/exp OR 'cat scan':ab,ti OR 'cat scans':ab,ti OR 'ct scan':ab,ti OR 'ct scans':ab,ti OR 'radiodensitometry'/exp OR 'x ray'/exp OR 'x-ray':ab,ti OR 'x ray':ab,ti OR 'x rays':ab,ti OR 'xrays':ab,ti OR xray*:ab,ti

<u>8,388</u>

#7.12 #7.10 AND #7.11

<u>1,466,199</u>

#7.11

'health care facility'/de OR 'hospital'/exp OR 'emergency health service'/exp OR 'hospital service'/exp OR 'emergency department':ab,ti OR 'emergency departments':ab,ti OR 'hospital patient'/de OR 'hospitalized adolescent'/exp OR 'hospitalized child'/exp OR 'hospitalized infant'/exp OR 'emergency patient'/exp OR 'critically ill patient'/exp OR 'high risk patient'/exp OR 'medical center':ab,ti OR 'medical centers':ab,ti OR 'acute care':ab,ti OR hospital*:ab,ti OR 'trauma center':ab,ti OR 'trauma centers':ab,ti OR 'trauma unit':ab,ti OR 'trauma units':ab,ti OR 'emergency service':ab,ti OR 'nontrauma center':ab,ti OR 'non trauma center':ab,ti OR 'nontrauma centers':ab,ti OR 'non trauma centers':ab,ti

54,306

#7.10 #7.8 NOT #7.9

<u>32,259</u>

#7.9 'military phenomena'/exp/mj OR 'veteran'/exp/mj OR 'veterans health'/exp/mj

<u>54,493</u>

#7.8

#7.4 NOT #7.7

4,153,342

#7.7 #7.5 NOT (#7.5 AND #7.6)

<u>2,632,147</u>

#7.6 'child'/exp OR 'adolescent'/exp OR 'newborn'/exp

5,090,845

#7.5 'adult'/exp OR 'aged'/exp OR 'middle aged'/exp

<u>82,997</u>

#7.4

#7.3 AND (1990:py OR 1991:py OR 1992:py OR 1993:py OR 1994:py OR 1995:py OR 1996:py OR 1997:py OR 1998:py OR 1999:py OR 2000:py OR 2001:py OR 2002:py OR 2003:py OR 2004:py OR 2005:py OR 2006:py OR 2007:py OR 2008:py OR 2009:py OR 2010:py OR 2011:py OR 2012:py OR 2013:py) AND ('article'/it OR 'article in press'/it OR 'conference paper'/it OR 'letter'/it OR 'review'/it)

<u>137,019</u>

#7.3 #7.1 NOT #7.2

<u>4,236,760</u>

#7.2 'animal'/exp NOT ('animal'/exp AND 'human'/exp)

<u>159,318</u>

#7.1

'brain concussion'/exp OR 'brain injury'/exp OR mtbi:ab,ti OR mtbis:ab,ti OR 'head injury'/de OR 'skull injury'/de OR 'second impact syndrome'/exp OR tbi:ab,ti OR tbis:ab,ti OR concussion*:ab,ti OR 'brain laceration':ab,ti OR 'brain lacerations':ab,ti OR 'cerebral injury':ab,ti OR 'cerebral injuries':ab,ti OR 'brain contusion':ab,ti OR 'brain contusions':ab,ti OR 'brain trauma':ab,ti OR 'brain traumas':ab,ti OR 'brain injury':ab,ti OR 'brain injuries':ab,ti OR 'skull fracture'/de OR 'skull base fracture'/de

Questions 4 & 5

<u>3,229</u> #3.39

#3.36 NOT #3.38

<u>241</u>

#3.38 #3.36 AND #3.37

<u>1,895,899</u>

#3.37 'case report'/exp

<u>3,470</u>

#3.36 #3.35 NOT #3.32

<u>3,541</u>

#3.35 #3.31 AND #3.34

4,251,395

#3.34 #3.1 OR #3.33

<u>1,628,084</u>

#3.33

'risk factor'/exp OR 'risk'/de OR 'high risk patient'/exp OR 'high risk infant'/exp OR 'risk assessment'/exp OR 'precipitating factor':ab,ti OR 'precipitating factors':ab,ti OR 'age factor':ab,ti OR 'age factors':ab,ti OR 'time factor':ab,ti OR 'time factors':ab,ti OR 'analysis of variance'/exp OR 'sex difference'/exp OR 'mechanism of injury':ab,ti OR 'mechanisms of injury':ab,ti OR 'time since injury':ab,ti OR 'between group difference':ab,ti OR 'between group differences':ab,ti OR epidemiology:de,ab,ti OR 'age distribution':de OR 'seasonal variation':de

<u>1,470,038</u>

#3.32

rat:ti OR rats:ti OR rodent*:ti OR mice:ti OR mouse:ti OR geriatric:ti OR elderly:ti OR 'animal model':ti OR alzheimer*:ti OR 'oif/oef':ti OR 'oef/oif':ti OR military:ti OR iraq:ti OR afghanistan:ti OR 'combat related':ti OR army:ti OR soldiers:ti OR veterans:ti OR 'air force':ti OR 'gulf war':ti OR 'gulf wars':ti OR 'operation enduring freedom':ti OR 'operation iraqi freedom':ti OR 'afghan war':ti OR 'afghan campaign':ti OR marines:ti OR troops:ti OR servicemen:ti OR 'combat disorders':ti OR 'combat disorder':ti OR 'active duty':ti

<u>5,412</u>

#3.31 #3.27 OR #3.30

<u>163</u> #3.30 #3.29 NOT #3.19

<u>163</u>

#3.29 #3.28 NOT #3.24

<u>163</u>

#3.28

'glasgow coma scale'/exp OR 'glasgow outcome scale'/exp AND '13-15':ab,ti

<u>5,295</u> #3.27 #3.23 NOT #3.26

<u>11,241</u>

#3.26 #3.24 OR #3.25

<u>30</u> #3.25 'oif/oef':ti AND [1990-2013]/py

<u>11,235</u>

#3.24

'iraq war':ti OR 'gulf war':ti OR 'gulf wars':ti OR 'operation iraqi freedom':ti OR 'afghan campaign':ti OR 'operation enduring freedom':ti OR military:ti OR battlefield:ti OR army:ti OR armed AND forces:ti OR marines:ti OR troops:ti OR servicemen:ti OR 'combat related':ti OR 'military personnel':ti OR 'air force':ti OR soldiers:ti OR 'combat veteran':ti OR 'combat disorder':ti OR 'combat disorders':ti OR veterans:ti OR 'active duty':ti OR 'service members':ti AND [1990-2013]/py

<u>5,335</u>

#3.23

#3.22 AND ('case report'/de OR 'clinical article'/de OR 'clinical protocol'/de OR 'clinical trial'/de OR 'cohort analysis'/de OR 'comparative study'/de OR 'control group'/de OR 'controlled clinical trial'/de OR 'controlled study'/de OR 'human'/de OR 'human tissue'/de OR 'interview'/de OR 'major clinical study'/de OR 'medical record review'/de OR 'methodology'/de OR 'model'/de OR 'nonhuman'/de OR 'normal human'/de OR 'outcomes research'/de OR 'practice guideline'/de OR 'prospective study'/de OR 'questionnaire'/de OR 'retrospective study'/de OR 'systematic review'/de) AND ('article'/it OR 'article in press'/it OR 'conference abstract'/it OR 'conference paper'/it OR 'letter'/it OR 'review'/it OR 'short survey'/it)

<u>6,018</u> #3.22 #3.20 NOT #3.21

452,516

#3.21 'editorial'/exp

<u>6,140</u>

#3.20 #3.16 NOT #3.19

<u>30,949</u>

#3.19 #3.17 NOT #3.18

<u>1,241</u>

#3.18

('military phenomena'/exp/mj OR 'veteran'/exp/mj) AND (civilian*:ab,ti OR sport* OR athlete*) AND [1990-2013]/py

<u>32,190</u>

#3.17 'military phenomena'/exp/mj OR 'veteran'/exp/mj

<u>6,202</u>

#3.16 #3.12 NOT #3.15

8,001,374

#3.15 #3.13 NOT #3.14

<u>10,227,651</u>

#3.14 'animal'/exp AND 'human'/exp AND [1990-2013]/py

<u>18,229,025</u>

#3.13 'animal'/exp

<u>7,286</u>

#3.12 #3.8 NOT #3.11

<u>4,719,417</u> #3.11

#3.9 NOT #3.10

<u>665,955</u>

#3.10

('adult'/exp OR 'aged'/exp OR 'middle aged'/exp OR 'embryo'/exp OR 'fetus'/exp) AND ('newborn'/exp OR 'child'/exp OR 'adolescent'/exp) AND [1990-2013]/py

<u>5,385,372</u>

#3.9

'adult'/exp OR 'aged'/exp OR 'middle aged'/exp OR 'embryo'/exp OR 'fetus'/exp

<u>10,092</u>

#3.8

#3.4 OR #3.5 OR #3.6 OR #3.7 AND [embase]/lim AND [1990-2013]/py

<u>974</u>

#3.7

mtbi:ab,ti OR mtbis:ab,ti OR 'minimal traumatic brain injury':ab,ti OR 'minimal traumatic brain injuries':ab,ti OR 'minimal tbi':ab,ti OR 'minimal tbis':ab,ti

<u>9,904</u> #3.6 #3.2 AND (mild:ab,ti OR minor:ab,ti)

<u>3,409</u> #3.5 'brain concussion'/exp

<u>2,878</u> #3.4 #3.2 AND #3.3

<u>103,017</u> #3.3 'sport'/exp OR 'athlete'/exp

156,672
#3.2
'brain injury'/exp OR 'head injury'/de OR 'skull injury'/exp

<u>3,030,325</u>

#3.1

'complication'/de OR 'disease exacerbation'/exp OR 'neurological complication'/exp OR 'posttraumatic complication'/exp OR 'posttraumatic pain'/exp OR 'prognosis'/de OR 'disease course'/de OR 'chronic disease'/exp OR 'illness trajectory'/exp OR 'recurrent disease'/exp OR 'relapse'/exp OR 'terminal disease'/exp OR 'acute disease'/exp OR 'adverse outcome'/exp OR 'bothersomeness'/exp OR 'deterioration'/exp OR 'disability severity'/exp OR 'injury severity'/exp OR 'nomogram'/exp OR 'treatment failure'/exp OR 'forecasting'/exp OR 'prediction'/exp OR 'predictive value'/exp OR 'death'/de OR 'lethality'/exp OR 'disability'/exp OR 'disease severity'/de OR impairment*:ab,ti OR sequelae:ab,ti OR impairment*:de OR 'dysfunction:ab,ti OR refractory:ab,ti OR 'more severe':ab,ti OR 'persisting symptoms':ab,ti OR 'prolonged:ab,ti OR 'long term symptoms':ab,ti OR 'longterm symptoms':ab,ti OR 'symptomatology'/exp

Question 6

<u>976</u>

#4.44 #4.41 NOT #4.43 <u>126</u> #4.43 #4.41 AND #4.42

1,894,200

#4.42 'case report'/exp

<u>1,102</u>

#4.41 #4.39 NOT #4.40

1,385,362

#4.40 rat:ti OR rats:ti OR mouse:ti OR mice:ti OR rodent*:ti OR geriatric:ti OR elderly:ti OR 'animal model':ti OR 'oif/oef':ti OR 'oef/oif':ti

<u>1,111</u>

#4.39 #4.37 AND #4.38

<u>5,588,005</u>

#4.38

longterm:ab,ti OR symptom*:de,ab,ti OR symptomatic:ab,ti OR sequelae:ab,ti OR enduring:ab,ti OR delayed:ab,ti OR prolong*:ab,ti OR ongoing:ab,ti OR persistent:ab,ti OR persistence:ab,ti OR

persisting:ab,ti OR lasting:ab,ti OR 'long term':ab,ti OR 'long-term':ab,ti OR 'post-concussion syndrome':de,ab,ti OR 'postconcussive syndrome':ab,ti OR 'post-concussive syndrome':ab,ti OR 'postconcussion syndrome':ab,ti OR symptomatology:de,ab,ti OR subsymptom*:ab,ti OR unresolved:ab,ti OR continuing:ab,ti OR continued:ab,ti OR unremitting:ab,ti OR lingering:ab,ti OR protracted:ab,ti OR 'greater than':ab,ti OR posttraumatic:ab,ti OR 'post-traumatic':ab,ti OR complication*:de,ab,ti OR complicated:ab,ti OR recovery:de,ab,ti OR recover:de,ab,ti OR recovered:ab,ti OR disability:de,ab,ti OR disabilities:de,ab,ti OR disabled:de,ab,ti OR impairment*:de,ab,ti OR impaired:ab,ti OR rehabilitation:de,ab,ti OR reintegration:ab,ti OR 'reintegration':ab,ti OR failure:de,ab,ti OR remediation:ab,ti OR remediate:ab,ti OR refractory:ab,ti OR extended:ab,ti OR 'longer than':ab,ti OR 'more than':ab,ti OR problematic:ab,ti

<u>1,546</u>

#4.37 #4.31 AND #4.36

<u>6,141,110</u>

#4.36 #4.1 OR #4.32 OR #4.33 OR #4.34 OR #4.35

<u>1,610,488</u>

#4.35

'paracetamol'/exp OR acetaminophen:de,ab,ti OR 'acetylsalicylic acid'/exp OR 'ibuprofen'/exp OR 'naproxen'/exp OR 'diclofenac'/exp OR 'analgesic agent'/exp OR 'nonsteroid antiinflammatory agent'/exp OR nsaid*:ab,ti OR 'antidepressant agent'/exp OR 'amitriptyline'/exp OR 'nortriptyline'/exp OR 'anticonvulsive agent'/exp OR 'valproic acid'/exp OR 'topiramate'/exp OR 'gabapentin'/exp OR 'propranolol'/exp OR 'beta adrenergic receptor blocking agent'/exp OR 'metoprolol'/exp OR 'dihydroergotamine'/exp OR ergot*:de,ab,ti OR 'sumatriptan'/exp OR 'antimigraine agent'/exp OR 'vestibular suppressant':ab,ti OR 'vestibular suppressants':ab,ti OR 'meclozine'/exp OR 'scopolamine'/exp OR 'benzodiazepine derivative'/exp OR neurostimulant*:ab,ti OR 'methylphenidate'/exp OR 'dexamphetamine'/exp OR modafanil:ab,ti OR 'amantadine'/exp OR 'atomoxetine'/exp OR 'antiemetic agent'/exp OR 'promethazine'/exp OR 'hypnotic sedative agent'/exp OR 'zolpidem'/exp OR 'serotonin modulator':ab,ti OR 'serotonin modulators':ab,ti OR 'trazodone'/exp OR 'alpha adrenergic receptor blocking agent'/exp OR 'melatonin'/exp OR 'sertraline'/exp OR 'citalopram'/exp OR 'escitalopram'/exp OR 'paroxetine'/exp OR 'fluoxetine'/exp OR 'amfebutamone'/exp OR 'cholinesterase inhibitor'/exp OR 'serotonin uptake inhibitor'/exp OR 'donepezil'/exp OR 'rivastigmine'/exp OR 'galantamine'/exp OR 'cytidine diphosphate choline':ab,ti

569,867

#4.34

'neurofeedback'/exp OR biofeedback:de,ab,ti OR 'hyperbaric oxygen'/exp OR 'kinesiotherapy'/exp OR 'physiotherapy'/exp OR 'cognitive therapy'/exp OR 'induced hypothermia'/exp OR 'rest'/exp OR neurorehabilitation:ab,ti OR 'stress reduction':ab,ti OR 'cognitive rest':ab,ti OR 'neuroprotective agent'/exp OR 'sodium chloride'/exp OR 'hypertonic saline solution':ab,ti OR 'nutraceutical'/exp OR 'diet supplementation'/exp OR 'massage'/exp OR 'cognitive retraining':ab,ti OR 'sensory stimulation'/exp OR 'exercise'/exp OR 'positional change':ab,ti OR 'postural change':ab,ti

144,728

#4.33 intervention*:de,ti

<u>2,010,750</u>

#4.32 treatment*:de,ti

<u>5,412</u>

#4.31 #4.27 OR #4.30

<u>163</u> #4.30 #4.29 NOT #4.19

<u>163</u>

#4.29 #4.28 NOT #4.24

<u>163</u>

#4.28 ('glasgow coma scale'/exp OR 'glasgow outcome scale'/exp) AND '13-15':ab,ti

<u>5,295</u>

#4.27 #4.23 NOT #4.26

<u>11,241</u>

#4.26 #4.24 OR #4.25

<u>30</u>

#4.25 'oif/oef':ti AND [1990-2013]/py

11,235

#4.24

'iraq war':ti OR 'gulf war':ti OR 'gulf wars':ti OR 'operation iraqi freedom':ti OR 'afghan campaign':ti OR 'operation enduring freedom':ti OR military:ti OR battlefield:ti OR army:ti OR armed AND forces:ti OR marines:ti OR troops:ti OR servicemen:ti OR 'combat related':ti OR 'military personnel':ti OR 'air force':ti OR soldiers:ti OR 'combat veteran':ti OR 'combat disorder':ti OR 'combat disorders':ti OR veterans:ti OR 'active duty':ti OR 'service members':ti AND [1990-2013]/py

<u>5,335</u>

#4.23

#4.22 AND ('case report'/de OR 'clinical article'/de OR 'clinical protocol'/de OR 'clinical trial'/de OR 'cohort analysis'/de OR 'comparative study'/de OR 'control group'/de OR 'controlled clinical trial'/de OR 'controlled study'/de OR 'human'/de OR 'human tissue'/de OR 'interview'/de OR 'major clinical study'/de OR 'medical record review'/de OR 'methodology'/de OR 'model'/de OR 'nonhuman'/de OR 'normal human'/de OR 'outcomes research'/de OR 'practice guideline'/de OR 'prospective study'/de OR 'questionnaire'/de OR 'retrospective study'/de OR 'systematic review'/de) AND ('article'/it OR 'article in press'/it OR 'conference abstract'/it OR 'conference paper'/it OR 'letter'/it OR 'review'/it OR 'short survey'/it)

6,018

#4.22 #4.20 NOT #4.21

<u>452,516</u>

#4.21 'editorial'/exp

<u>6,140</u>

#4.20 #4.16 NOT #4.19

30,949

#4.19 #4.17 NOT #4.18

<u>1,241</u>

#4.18

'military phenomena'/exp/mj OR 'veteran'/exp/mj AND (civilian*:ab,ti OR sport* OR athlete*) AND [1990-2013]/py

<u>32,190</u>

#4.17

'military phenomena'/exp/mj OR 'veteran'/exp/mj

<u>6,202</u> #4.16 #4.12 NOT #4.15

<u>8,001,374</u> #4.15 #4.13 NOT #4.14

10,227,651

#4.14 'animal'/exp AND 'human'/exp AND [1990-2013]/py

<u>18,229,025</u> #4.13 'animal'/exp

<u>7,286</u> #4.12 #4.8 NOT #4.11

<u>4,719,417</u>

#4.11 #4.9 NOT #4.10

665,955

#4.10

'adult'/exp OR 'aged'/exp OR 'middle aged'/exp OR 'embryo'/exp OR 'fetus'/exp AND ('newborn'/exp OR 'child'/exp OR 'adolescent'/exp) AND [1990-2013]/py

<u>5,385,372</u>

#4.9

'adult'/exp OR 'aged'/exp OR 'middle aged'/exp OR 'embryo'/exp OR 'fetus'/exp

<u>10,092</u>

#4.8

#4.4 OR #4.5 OR #4.6 OR #4.7 AND [embase]/lim AND [1990-2013]/py

<u>974</u>

#4.7

mtbi:ab,ti OR mtbis:ab,ti OR 'minimal traumatic brain injury':ab,ti OR 'minimal traumatic brain injuries':ab,ti OR 'minimal tbi':ab,ti OR 'minimal tbis':ab,ti

<u>9,904</u>

#4.6 #4.2 AND (mild:ab,ti OR minor:ab,ti)

<u>3,409</u>

#4.5 'brain concussion'/exp

<u>2,878</u> #4.4 #4.2 AND #4.3

<u>103,017</u>

#4.3 'sport'/exp OR 'athlete'/exp

156,672

#4.2 'brain injury'/exp OR 'head injury'/de OR 'skull injury'/exp

3,578,947

#4.1

'therapy'/de OR 'acupuncture'/exp OR 'animal assisted therapy'/exp OR 'apitherapy'/exp OR 'biological therapy'/exp OR 'computer assisted therapy'/exp OR 'conservative treatment'/exp OR 'counseling'/exp OR 'device therapy'/exp OR 'drug therapy'/exp OR 'early intervention'/exp OR 'experimental therapy'/exp OR 'fluid therapy'/exp OR 'homeopathy'/exp OR 'intensive care'/de OR 'neuromonitoring'/exp OR 'blood pressure monitoring'/exp OR 'body temperature monitoring'/exp OR 'capnography'/exp OR 'life sustaining treatment'/exp OR 'maintenance therapy'/de OR 'patient positioning'/exp OR 'rehabilitation care'/exp OR 'radiotherapy'/de OR 'cognitive rehabilitation'/exp OR 'functional training'/exp OR 'muscle training'/exp OR 'occupational therapy'/exp OR 'rejuvenation'/exp OR 'sensorimotor integration'/exp OR 'speech rehabilitation'/exp OR 'sleep therapy'/exp OR 'supplementation'/exp OR 'systemic therapy'/exp OR 'therapy effect'/exp OR 'treatment response'/exp

SportDiscus: Updated Search

Run: August 3, 2015

Contact: Alix Hayden, PhD, University of Calgary ahayden@ucalgary.ca

	Query	Limiters/Expanders	Results
S6	TX (youth or child or children or childhood or toddler* or infant* or teen or teens or teenager* or "teen ager" or "teen agers" or adolescent* or adolescence or pediatric* or paediatric*) AND S4	Limiters - Published Date: 19900101-20151231; Language: English Search modes - Boolean/Phrase	1,441
S5	TX (youth or child or children or childhood or toddler* or infant* or teen or teens or teenager* or "teen ager" or "teen agers" or adolescent* or adolescence or pediatric* or paediatric*) AND S4	Search modes - Boolean/Phrase	1,482
S4	S1 OR S2 OR S3	Search modes - Find all my search terms	4,851
S 3	("head injury" or "head injuries" or "brain injury" or "brain injuries" or "head trauma" or "brain trauma" or "skull injury" or "skull injuries" or tbi or tbis or "cerebral trauma" or "cerebral injury" or "cerebral injuries") AND (sport* or athlete* or athletic or recreation)	Search modes - Boolean/Phrase	1,616
S2	("gcs 13-15" or concussion* or concussive or postconcussion or "post concussion" or postconcussive or "post concussive" or "mild tbi" or "mild tbis" or mtbi or mtbis or "mild traumatic brain injury" or "mild traumatic brain injuries")	Search modes - Boolean/Phrase	Display
S1	("traumatic brain injury" or "traumatic brain injuries" or tbi or tbis or "brain contusion" or "brain contusions" or "brain trauma" or "brain laceration" or "brain lacerations" or "cerebral injury" or "cerebral injuries" or "head injury" or "head injuries" or "head trauma" or "cerebral trauma" or "skull injury" or "skull injuries" or "craniocerebral trauma" or "brain injury" or "brain injuries") AND TX (mild or minor)	Search modes - Boolean/Phrase	1,549

mTBI – SEARCH STRATEGIES—Initial Search

January 1, 1990 to December 31, 2013

All searches were limited by date from January 1, 1990 to December 31, 2013. No languages were excluded.

PUBMED SEARCH STRATEGIES

Publication types excluded from PubMed searches were: comments, editorials, patient education handouts, newspaper articles, biographies, autobiographies, and festschrifts.

Question 1: Last run 11/25/12

This search included a POPULATION component and a DIAGNOSIS component. In *Boolean* terms, these components were combined as follows:

POPULATION <u>AND</u> DIAGNOSIS

POPULATION:

SPORT*[TW] OR ATHLETE[TW] OR ATHLETES[TW] OR ATHLETIC*[TW] OR RECREATION*[TIAB]) AND (HEAD INJUR*[TW] OR BRAIN INJUR*[TW] OR HEAD TRAUMA*[TIAB] OR BRAIN TRAUMA*[TIAB] OR SKULL INJUR*[TIAB] OR TBI[TIAB] OR TBIS[TIAB] OR CEREBRAL TRAUMA*[TW] OR CEREBRAL INJUR*[TW]

OR

GCS 13-15[TIAB] OR BRAIN CONCUSSION[MH] OR CONCUSSION*[TW] OR CONCUSSIVE[TIAB] OR COMMOTIO CEREBRI[TW] OR POST CONCUSSIVE[TW] OR POSTCONCUSSIVE[TW] OR MILD TBI[TIAB] OR MILD TBIS[TIAB] OR MTBI[TIAB] OR MTBIS[TIAB] OR MILD TRAUMATIC BRAIN INJUR*[TIAB]

OR

(TRAUMATIC BRAIN INJUR*[TIAB] OR TBI[TIAB] OR TBIS[TIAB] OR BRAIN CONTUSION*[TW] OR BRAIN TRAUMA*[TW] OR BRAIN LACERATION*[TW] OR CEREBRAL INJUR*[TW] OR HEAD INJUR*[TW] OR HEAD TRAUMA*[TW] OR CEREBRAL TRAUMA*[TIAB] OR SKULL INJUR*[TW] OR CRANIOCEREBRAL TRAUMA[MH:NOEXP] OR BRAIN INJUR* [TW]) AND (MILD[TIAB] OR MINOR[TIAB]) OR

MINIMAL TRAUMATIC BRAIN INJUR*[TIAB] OR MINIMAL TBI[TIAB]

NOT

(ANIMALS[MH] NOT (ANIMALS[MH] AND HUMAN*[MH]))

NOT

(ADULT[MH] NOT (ADULT[MH] AND (CHILD[MH] OR ADOLESCENT*[MH] OR INFANT*[MH]))

NOT

(IRAQ WAR, 2003 [MAJR] OR IRAQ WAR [TI] OR GULF WAR [TI] OR OPERATION IRAQI FREEDOM [TI] OR GULF WARS [TI] OR AFGHAN CAMPAIGN 2001[MAJR] OR OPERATION ENDURING FREEDOM [TI] OR AFGHAN WAR [TI] OR WAR [MAJR] OR MILITARY [TI] OR BATTLEFIELD [TI] OR ARMY [TI] OR ARMED FORCES [TI] OR MARINES [TI] OR TROOPS [TI] OR SERVICEMEN [TI] OR COMBAT RELATED [TI] OR MILITARY PERSONNEL [MAJR] OR AIR FORCE [TI] OR SOLDIERS [TI] OR MILITARY MEDICINE [MAJR] OR OIF/OEF [TI] OR COMBAT VETERAN* [TI] OR COMBAT DISORDERS [MAJR] OR VETERANS [MAJR] OR ACTIVE DUTY [TI] OR SERVICE MEMBERS [TI]) NOT (MILITARY [TW] AND (CIVILIAN* [TIAB] OR SPORTS [TIAB] OR ATHLETE* [TIAB] OR SPORT RELATED [TI] OR SPORTS RELATED [TI]))

DIAGNOSIS:

ASSESSMENT* [TIAB] OR DIAGNOSTIC TECHNIQUES, NEUROLOGICAL[MH:NOEXP] OR NEUROLOGIC EXAMINATION[MH] OR DIAGNOSIS, DIFFERENTIAL[MH] OR DIAGNOSIS[MH:NOEXP] OR DIAGNOSIS[SH] OR DIAGNOSTIC ERRORS[MH] OR DIAGNOS*[TIAB] OR DIAGNOSTIC*[TW] OR MEASURES[TW] OR SCALE[TW] OR SCALES[TW] OR SCORE[TW] OR SCORES[TW] OR INDEX[TW] OR INDICES[TW] OR INSTRUMENT[TW] OR INSTRUMENTS[TW] OR MEASURE[TW] OR SENSORY MODALITY ASSESSMENT[TIAB] OR SENSORY STIMULATION ASSESSMENT [TIAB] OR WESSEX HEAD INJURY MATRIX [TIAB] OR WESTERN NEURO SENSORY STIMULATION PROFILE[TIAB] OR SENSITIVITY AND SPECIFICITY[MH] OR SPECIFICITY[TIAB] OR PREDICTIVE VALUE OF TESTS[MH] OR CONFIDENCE INTERVAL[MH] OR DETECT*[TIAB] OR CHARACTERISTIC*[TIAB] OR CATEGORIZ*[TIAB] OR CATEGORIS*[TIAB] OR DISTINGUISH*[TIAB] OR DIFFERENTIAT*[TIAB] OR BEHAVIOURAL ASSESSMENT*[TIAB] OR BEHAVIORAL ASSESSMENT*[TIAB] OR BEHAVIOR ASSESSMENT*[TIAB] OR BEHAVIOUR ASSESSMENT*[TIAB] OR NEUROPSYCHOLOGICAL TESTS[MH:NOEXP] OR TRAUMA SEVERITY INDICES [MH] OR DETERMINANT* [TIAB] OR KING DEVICK TEST [TIAB]

OR MILITARY ACUTE CONCUSSION EVALUATION [TIAB] OR REACTION TIME [MH] OR REACTION TIME* [TIAB] OR RESPONSE LATENCY [TW] OR SCREENING TEST* [TIAB] OR SCREENING TOOL* [TIAB] OR BALANCE TEST* [TW] OR POSTURAL BALANCE [MH] OR POSTURAL EQUILIBRIUM [TW] OR SENSATION DISORDERS [MH] OR BALANCE ASSESSMENT [TIAB] OR MEDICAL SYMPTOM VALIDITY TEST [TIAB] OR EVOKED POTENTIALS[MH] OR EVOKED POTENTIAL*[TIAB]OR CLASSIFICATION [MH] OR CLASSIFICATION [SH]OR NEUROCOGNITIVE TEST* [TIAB]OR PSYCHOMOTOR PERFORMANCE [MH] OR MEMORY [MH]OR IMMEDIATE POST-CONCUSSION ASSESSMENT AND COGNITIVE TEST* [TIAB] OR ELECTROENCEPHALOGRAPHY[MH] OR ELECTROENCEPHALOGRAPHY[TIAB] OR ELECTROENCEPHALOGRAM*[TIAB] OR ECHOENCEPHALOGRAPHY[MH] OR EEG[TIAB] OR EEGS[TIAB] OR QEEG[TIAB] OR QEEGS[TIAB] OR ELECTROMYOGRAPH*[TW] OR EMG[TIAB] OR EMGS[TIAB] OR ELECTROPHYSIOLOGIC TECHNIQUE*[TIAB]

OR

PREDICTIVE[TIAB] AND VALUE*[TIAB]

Questions 2 and 3: Last run 11/25/12

This search included a larger (more results) POPULATION component than Question 1. It also included a component to identify emergency or other acute care settings, and components for IMAGING and for INTRACRANIAL INJURY. In <u>Boolean</u> terms, the components were combined as follows:

(POPULATION AND ACUTE CARE SETTING) AND (IMAGING OR INTRACRANIAL INJURY)

POPULATION:

GCS 13-15[TIAB] OR BRAIN CONCUSSION[MH] OR CONCUSSION*[TW] OR COMMOTIO CEREBRI[TW] OR POST CONCUSSIVE[TW] OR POSTCONCUSSIVE[TW] OR MTBI[TIAB] OR MTBIS[TIAB] OR BRAIN INJURIES[MH] OR BRAIN INJUR*[TW] OR TBI[TIAB] OR TBIS[TIAB] OR BRAIN CONTUSION*[TW] OR BRAIN TRAUMA*[TW] OR BRAIN LACERATION*[TW] OR HEAD TRAUMA*[TW] OR CEREBRAL TRAUMA*[TW] OR CEREBRAL INJUR*[TW] OR SKULL INJUR*[TW] OR SKULL TRAUMA[TW] OR CRANIOCEREBRAL TRAUMA[MH:noexp] OR HEAD INJUR*[TW] NOT

(ANIMALS[MH] NOT (ANIMALS[MH] AND HUMAN*[MH]))

NOT

(ADULT[MH] NOT (ADULT[MH] AND (CHILD[MH] OR ADOLESCENT*[MH] OR INFANT*[MH]))

NOT

(IRAQ WAR, 2003 [MAJR] OR IRAQ WAR [TI] OR GULF WAR [TI] OR OPERATION IRAQI FREEDOM [TI] OR GULF WARS [TI] OR AFGHAN CAMPAIGN 2001[MAJR] OR OPERATION ENDURING FREEDOM [TI] OR AFGHAN WAR [TI] OR WAR [MAJR] OR MILITARY [TI] OR BATTLEFIELD [TI] OR ARMY [TI] OR ARMED FORCES [TI] OR MARINES [TI] OR TROOPS [TI] OR SERVICEMEN [TI] OR COMBAT RELATED [TI] OR MILITARY PERSONNEL [MAJR] OR AIR FORCE [TI] OR SOLDIERS [TI] OR MILITARY MEDICINE [MAJR] OR OIF/OEF [TI] OR COMBAT VETERAN* [TI] OR COMBAT DISORDERS [MAJR] OR VETERANS [MAJR] OR ACTIVE DUTY [TI] OR SERVICE MEMBERS [TI]) NOT (MILITARY [TW] AND (CIVILIAN* [TIAB] OR SPORTS [TIAB] OR ATHLETE* [TIAB] OR SPORT RELATED [TI] OR SPORTS RELATED [TI]))

ACUTE CARE SETTING:

MEDICAL CENTRE*[TIAB] OR TRAUMA CENTRE*[TIAB] OR NON TRAUMA CENTRE* [TIAB] OR NONTRAUMA CENTRE* [TIAB] OR INPATIENT*[TW] OR MEDICAL CENTER*[TW] OR ACADEMIC MEDICAL CENTERS[MH] OR HEALTH FACILITIES[MH] OR ACUTE CARE[TW] OR HOSPITALS[MH] OR HOSPITAL*[TW] OR NON TRAUMA CENTER*[TW] OR NONTRAUMA CENTER*[TW] OR TRAUMA CENTERS[MH] OR TRAUMA CENTER*[TW] OR TRAUMA UNIT*[TW] OR EMERGENCY SERVICE, HOSPITAL[MH] OR EMERGENCY[TW] OR ACUTE SETTING* [TIAB] OR TERTIARY CARE [TIAB]

IMAGING:

MAGNETIC RESONANCE IMAGING [MH] OR ZEUGMATOGRAPHY [TW] OR TOMOGRAPHY [TW] OR MRI SCAN* [TW] OR FMRI [TW] OR FMRIS [TW] OR MRI [TW] OR MRIS [TW] OR IMAGING* [TW] OR NEUROIMAGING [MH:NOEXP] OR NEUROIMAGING* [TIAB] OR NEURO IMAGING* [TIAB] OR TOMOGRAPHY, X-RAY COMPUTED [MH] OR CAT SCAN* [TW] OR CT SCAN* [TW] OR CT X RAY* [TW] OR TOMODENSITOMETRY [TW] OR CINE-CT [TW] OR X-RAYS [MH] OR X RAY* [TIAB] OR RADIOGRAPHY [SH]

INTRACRANIAL INJURY:

INTRACRANIAL INJUR* [TIAB] OR INTRACRANIAL HEMORRHAGES[MH] OR BRAIN HEMORRHAGE, TRAUMATIC[MH] OR HEMATOMA, SUBDURAL[MH] OR HEMORRHAGE*[TW] OR HAEMORRHAGE*[TIAB] OR BRAIN BLEED*[TIAB] OR HAEMATOMA*[TIAB] OR HEMATOMA*[TIAB] OR PITUITARY APOPLEXY[TW] OR EDEMA[TW] OR DIFFUSE AXONAL INJURY[MH] OR AXONAL SWELLING[TW] OR RETRACTION BALLS[TIAB] OR CEREBRAL ATROPHY[TIAB] OR INTRACEREBRAL INJUR*[TIAB] OR OEDEMA[TIAB] OR BRAIN CONTUSION*[TIAB] OR CEREBRAL CONTUSION*[TIAB] OR AXONAL INJUR*[TW] OR PNEUMOCEPHALUS[MH] OR PNEUMOCEPHALUS[TIAB] OR CRANIAL PNEUMOCYST*[TW] OR INTRACRANIAL GAS[TW] OR MICROBLEED*[TIAB] OR MICRO BLEED* [TIAB] OR WHITE MATTER INJUR*[TIAB] OR SKULL FRACTURES[MH] OR SKULL FRACTURE*[TIAB] OR CRANIAL INJUR*[TW] OR LESION*[TW] OR NEUROPATHOLOG*[TW] OR BLEEDING[TIAB] OR MICROHEMORRHAGE*[TIAB] OR MICROHAEMORRHAGE*[TIAB] OR MICRO HEMORRHAGE*[TIAB] OR MICRO HAEMORRHAGE*[TIAB] OR INTRACRANIAL SEQUELAE[TIAB]

Questions 4 and 5: Last run 11/25/12

This search included the same POPULATION component as Question 1. In addition it included components for PROGNOSIS and for RISK FACTORS FOR SEQUELAE. In <u>Boolean</u> terms, they were combined as follows:

POPULATION AND (PROGNOSIS OR RISK FACTORS FOR SEQUELAE)

POPULATION:

SPORT*[TW] OR ATHLETE[TW] OR ATHLETES[TW] OR ATHLETIC*[TW] OR RECREATION*[TIAB]) AND (HEAD INJUR*[TW] OR BRAIN INJUR*[TW] OR HEAD TRAUMA*[TIAB] OR BRAIN TRAUMA*[TIAB] OR SKULL INJUR*[TIAB] OR TBI[TIAB] OR TBIS[TIAB] OR CEREBRAL TRAUMA*[TW] OR CEREBRAL INJUR*[TW]

OR

GCS 13-15[TIAB] OR BRAIN CONCUSSION[MH] OR CONCUSSION*[TW] OR CONCUSSIVE[TIAB] OR COMMOTIO CEREBRI[TW] OR POST CONCUSSIVE[TW] OR POSTCONCUSSIVE[TW] OR MILD TBI[TIAB] OR MILD TBIS[TIAB] OR MTBI[TIAB] OR MTBIS[TIAB] OR MILD TRAUMATIC BRAIN INJUR*[TIAB]

OR

(TRAUMATIC BRAIN INJUR*[TIAB] OR TBI[TIAB] OR TBIS[TIAB] OR BRAIN CONTUSION*[TW] OR BRAIN TRAUMA*[TW] OR BRAIN LACERATION*[TW] OR CEREBRAL INJUR*[TW] OR HEAD INJUR*[TW] OR HEAD TRAUMA*[TW] OR CEREBRAL TRAUMA*[TIAB] OR SKULL INJUR*[TW] OR CRANIOCEREBRAL TRAUMA[MH:NOEXP] OR BRAIN INJUR* [TW]) AND (MILD[TIAB] OR MINOR[TIAB]) OR

MINIMAL TRAUMATIC BRAIN INJUR*[TIAB] OR MINIMAL TBI[TIAB]

NOT

(ANIMALS[MH] NOT (ANIMALS[MH] AND HUMAN*[MH]))

NOT

(ADULT[MH] NOT (ADULT[MH] AND (CHILD[MH] OR ADOLESCENT*[MH] OR INFANT*[MH]))

NOT

(IRAQ WAR, 2003 [MAJR] OR IRAQ WAR [TI] OR GULF WAR [TI] OR OPERATION IRAQI FREEDOM [TI] OR GULF WARS [TI] OR AFGHAN CAMPAIGN 2001[MAJR] OR OPERATION ENDURING FREEDOM [TI] OR AFGHAN WAR [TI] OR WAR [MAJR] OR MILITARY [TI] OR BATTLEFIELD [TI] OR ARMY [TI] OR ARMED FORCES [TI] OR MARINES [TI] OR TROOPS [TI] OR SERVICEMEN [TI] OR COMBAT RELATED [TI] OR MILITARY PERSONNEL [MAJR] OR AIR FORCE [TI] OR SOLDIERS [TI] OR MILITARY MEDICINE [MAJR] OR OIF/OEF [TI] OR COMBAT VETERAN* [TI] OR COMBAT DISORDERS [MAJR] OR VETERANS [MAJR] OR ACTIVE DUTY [TI] OR SERVICE MEMBERS [TI]) NOT (MILITARY [TW] AND (CIVILIAN* [TIAB] OR SPORTS [TIAB] OR ATHLETE* [TIAB] OR SPORT RELATED [TI] OR SPORTS RELATED [TI]))

PROGNOSIS:

EXTENDED [TIAB] OR LASTING [TIAB] OR LONGTERM [TIAB] OR LONG TERM [TIAB] OR "LONGER THAN" [TIAB] OR "MORE THAN" [TIAB] OR "GREATER THAN" [TIAB] OR DELAYED [TIAB] OR FAILURE* [TW] OR PROLONG* [TIAB] OR PROTRACTED [TIAB] OR COMPLICATED [TIAB] OR ENDURING [TIAB] OR CONTINUING [TIAB] OR CONTINUED [TIAB]OR UNRESOLVED [TIAB] OR UNREMITTING [TIAB] OR LINGERING [TIAB] OR PROBLEMATIC [TIAB]) AND (FUNCTIONAL LEVEL [TIAB] OR PROGRESS*[TW] OR RECUPERAT*[TIAB] OR CONVALESCENCE[TW] OR CONVALESCE[TIAB] OR CONVALESCED[TIAB] OR CONVALESCES[TIAB] OR CONVALESCING[TIAB] OR RECOVER[TIAB] OR RECOVERS[TIAB] OR RECOVERY[TIAB] OR RECOVERING[TIAB] OR RECOVERIES[TIAB] OR IMPROVE* [TW] OR IMPROVEMENT* [TIAB] OR IMPROVING[TW] OR RESTORATION[TW] OR RESTORE*[TW] OR RECOVERY OF FUNCTION[TW] OR RESOLUTION* [TW] OR PROGNOSIS [MH] OR DISEASE FREE SURVIVAL [MH OR PROGNOS*[TW] OR PREDICT*[TIAB] OR NOMOGRAM*[TW] OR MEDICAL FUTILITY [MH] OR FUTIL* [TW] OR TREATMENT OUTCOME [MH:NOEXP] OR TREATMENT FAILURE [MH] OR PREDICTIVE VALUE OF TESTS [MH] OR FORECASTING [MH] OR FORECAST* [TIAB] OR PROJECTION* [TW]

RISK FACTORS FOR SEQUELAE:

(PATHOPHYSIOLOG* [TW] OR DYSFUNCTION [TIAB] OR SYMPTOMATOLOGY [TIAB] OR SUBSYMPTOM* [TIAB] OR SUB SYMPTOM* [TIAB] OR SYMPTOMS [TW] OR SYMPTOMATIC [TIAB] OR DISEASE-FREE SURVIVAL[MH] OR TREATMENT FAILURE[MH] OR DISEASE PROGRESSION[MH] OR DISEASE PROGRESSION [TIAB] OR SEQUELAE [TW] OR ENDURING [TW] OR DELAYED [TIAB] OR PROLONGED [TIAB] OR ONGOING [TIAB] OR PERSISTENT [TIAB] OR PERSISTENCE [TIAB] OR PERSISTING [TIAB] OR LASTING [TIAB] OR LONGTERM [TIAB] OR LONG TERM [TIAB] OR POST CONCUSSION SYNDROME* [TIAB] OR POST-CONCUSSION SYNDROME [MH] OR POST-CONCUSSIVE SYNDROME [TW] OR POSTCONCUSSION SYNDROME* [TIAB] OR POSTCONCUSSIVE SYNDROME* [TIAB] OR UNRESOLVED [TIAB] OR CONTINUING [TIAB] OR CONTINUE* [TIAB] OR UNREMITTING [TIAB] OR LINGERING [TIAB] OR PROTRACTED [TIAB] OR POSTTRAUMATIC [TW] OR POST TRAUMATIC [TIAB] OR COMPLICATIONS [SH] OR COMPLICATION* [TIAB] OR COMPLICATED [TIAB] OR EXTENDED [TIAB] OR DISABILITY [TW] OR DISABILITIES [TW] OR DISABLED [TW] OR IMPAIRMENT [TW] OR IMPAIRED [TIAB] OR FAILURE [TW] OR REFRACTORY [TW] OR MORE SEVERE* [TIAB])

AND

(RISK [MH:NOEXP]OR RISK ASSESSMENT [MH] OR RISK FACTORS [MH] OR RISK* [TIAB] OR RISK ADJUSTMENT [MH] OR PRECIPITATING FACTORS [MH] OR EPIDEMIOLOGIC FACTORS [MH] OR CAUSALITY [MH] OR CAUSALITY [TW] OR CAUSATION* [TW] OR ETIOLOGY [SH:NOEXP] OR EPIDEMIOLOGY [SH] OR FEATURE* [TIAB] OR FACTOR [TIAB] OR FACTORS [TIAB] OR IDENTIFY [TIAB] OR IDENTIFICATION [TIAB] OR IDENTIFIES [TIAB] OR INDENTIFIED [TIAB] OR AGE FACTORS [MH] OR SEX [TW] OR SEXES [TW] OR GENDER [TW] OR TIME SINCE INJUR* [TIAB] OR TIME FACTORS [MH] OR BETWEEN GROUP DIFFERENCE* [TIAB] OR MECHANISM OF INJURY [TIAB] OR MECHANISMS OR INJURY [TIAB] OR INDICATOR* [TIAB] OR CHARACTERISTIC* [TIAB] OR ANALYSIS OF VARIANCE [MH])

Question 6: Last run 11/25/12

This search included the same POPULATION component as Question 1. It also included a TREATMENT component and a component for the concept of ONGOING SYMPTOMS. In *Boolean* terms, the components were combined as follows:

POPULATION AND ONGOING SYMPTOMS AND TREATMENTS

SPORT*[TW] OR ATHLETE[TW] OR ATHLETES[TW] OR ATHLETIC*[TW] OR RECREATION*[TIAB]) AND (HEAD INJUR*[TW] OR BRAIN INJUR*[TW] OR HEAD TRAUMA*[TIAB] OR BRAIN TRAUMA*[TIAB] OR SKULL INJUR*[TIAB] OR TBI[TIAB] OR TBIS[TIAB] OR CEREBRAL TRAUMA*[TW] OR CEREBRAL INJUR*[TW]

OR

GCS 13-15[TIAB] OR BRAIN CONCUSSION[MH] OR CONCUSSION*[TW] OR CONCUSSIVE[TIAB] OR COMMOTIO CEREBRI[TW] OR POST CONCUSSIVE[TW] OR POSTCONCUSSIVE[TW] OR MILD TBI[TIAB] OR MILD TBIS[TIAB] OR MTBI[TIAB] OR MTBIS[TIAB] OR MILD TRAUMATIC BRAIN INJUR*[TIAB]

OR

(TRAUMATIC BRAIN INJUR*[TIAB] OR TBI[TIAB] OR TBIS[TIAB] OR BRAIN CONTUSION*[TW] OR BRAIN TRAUMA*[TW] OR BRAIN LACERATION*[TW] OR CEREBRAL INJUR*[TW] OR HEAD INJUR*[TW] OR HEAD TRAUMA*[TW] OR CEREBRAL TRAUMA*[TIAB] OR SKULL INJUR*[TW] OR CRANIOCEREBRAL TRAUMA[MH:NOEXP] OR BRAIN INJUR* [TW]) AND (MILD[TIAB] OR MINOR[TIAB])

OR

MINIMAL TRAUMATIC BRAIN INJUR*[TIAB] OR MINIMAL TBI[TIAB]

NOT

(ANIMALS[MH] NOT (ANIMALS[MH] AND HUMAN*[MH]))

NOT

(ADULT[MH] NOT (ADULT[MH] AND (CHILD[MH] OR ADOLESCENT*[MH] OR INFANT*[MH]))

NOT

(IRAQ WAR, 2003 [MAJR] OR IRAQ WAR [TI] OR GULF WAR [TI] OR OPERATION IRAQI FREEDOM [TI] OR GULF WARS [TI] OR AFGHAN CAMPAIGN 2001[MAJR] OR OPERATION ENDURING FREEDOM [TI] OR AFGHAN WAR [TI] OR WAR [MAJR] OR MILITARY [TI] OR BATTLEFIELD [TI] OR ARMY [TI] OR ARMED FORCES [TI] OR MARINES [TI] OR TROOPS [TI] OR SERVICEMEN [TI] OR COMBAT RELATED [TI] OR MILITARY PERSONNEL [MAJR] OR AIR FORCE [TI] OR SOLDIERS [TI] OR MILITARY MEDICINE [MAJR] OR OIF/OEF [TI] OR COMBAT VETERAN* [TI] OR COMBAT DISORDERS [MAJR] OR VETERANS [MAJR] OR ACTIVE DUTY [TI] OR SERVICE MEMBERS [TI]) NOT (MILITARY [TW] AND (CIVILIAN* [TIAB] OR SPORTS [TIAB] OR ATHLETE* [TIAB] OR SPORT RELATED [TI] OR SPORTS RELATED [TI]))

ONGOING SYMPTOMS:

LONGTERM [TIAB] OR SYMPTOMS [TW] OR SYMPTOMATIC [TIAB] OR SEQUELAE [TW] OR ENDURING [TW] OR DELAYED [TIAB] OR PROLONG* [TIAB] OR ONGOING [TIAB] OR PERSISTENT [TIAB] OR PERSISTENCE [TIAB] OR PERSISTING [TIAB] OR LASTING [TIAB] OR LONG TERM [TIAB] OR POST-CONCUSSION SYNDROME [MH] OR POST-CONCUSSIVE SYNDROME [TW] OR POSTCONCUSSION SYNDROME* [TIAB] OR POSTCONCUSSIVE SYNDROME* [TIAB] OR SYMPTOMATOLOGY [TIAB] OR SUBSYMPTOM* [TIAB] OR UNRESOLVED [TIAB] OR CONTINUING [TIAB] OR CONTINUED [TIAB] OR UNREMITTING [TIAB] OR LINGERING [TIAB] OR PROTRACTED [TIAB] OR "LONGER THAN" [TIAB] OR "MORE THAN" [TIAB] OR "GREATER THAN" [TIAB] OR POST CONCUSSION SYNDROME [MH] OR POST CONCUSSION SYNDROME* [TIAB] OR POSTCONCUSSION SYNDROME* [TIAB]OR POSTTRAUMATIC [TW] OR POST TRAUMATIC [TIAB] OR COMPLICATIONS [SH] OR COMPLICATION* [TIAB] OR COMPLICATED [TIAB] OR EXTENDED [TIAB] OR RECOVERY OR RECOVER [TIAB] OR RECOVERED [TIAB] OR COMMUNITY DWELLING [TIAB] OR DISABILITY [TW] OR DISABILITIES [TW] OR DISABLED [TW] OR IMPAIRMENT [TW] OR IMPAIRED [TIAB] OR REHABILITATION [MH] OR REINTEGRATION [TW] OR REHABILITATION [TIAB] OR RE-INTEGRATION [TIAB] OR FAILURE [TW] OR REMEDIATION [TW] OR REMEDIATE [TW] OR REFRACTORY [TW] OR EXTENDED [TIAB] OR "LONGER THAN" [TIAB] OR "MORE THAN" [TIAB] OR "GREATER THAN" [TIAB] OR PROBLEMATIC [TIAB]

TREATMENTS:

THERAPY [SH] OR THERAPEUTIC USE [SH] OR DRUG THERAPY[MH]OR PHARMACOTHERAP* [TW] OR PHARMACOLOGIC* [TW] OR DRUG THERAP* [TIAB]OR THERAPY [TW] OR THERAPIES [TW] OR THERAPEUTIC* [TW] OR TREATMENT* [TW] OR TREATMENT OUTCOME [MH] OR DIET THERAPY [MH] OR DIET THERAPY [SH] OR NUTRITION THERAPY [MH] OR REHABILITATION OUTCOME* [TW] OR REHABILITATION [SH] OR REHABILITAT* [TIAB] OR TREATED [TIAB] OR MANAGEMENT [TIAB] OR INTERVENTION* [TIAB] OR BIOFEEDBACK, PSYCHOLOGY [MH] OR BIOFEEDBACK [TIAB] OR HYPERBARIC OXYGENATION [MH] OR EXERCISE THERAPY [MH] OR PHYSICAL THERAPY MODALITIES [MH] OR PHYSIOTHERAPY [TW] OR COGNITIVE THERAPY [MH] OR HYPOTHERMIA, INDUCED [MH:NOEXP] OR HYPOTHERMIA [TIAB] OR REST [MH] OR NEUROREHABILIT* [TIAB] OR STRESS REDUCTION [TIAB] OR PHYSICAL REST [TIAB] OR COGNITIVE REST [TIAB] OR NEUROPROTECTIVE AGENTS [MH] OR

SALINE SOLUTION, HYPERTONIC [MH] OR NUTRACEUTICAL* [TIAB] OR DIETARY SUPPLEMENTS [MH] OR DIETARY SUPPLEMENT* [TIAB] OR NEUTRACEUTICAL* [TIAB] OR NEUROFEEDBACK [TIAB] OR NEURO FEEDBACK [TIAB] OR BIO FEEDBACK [TIAB] OR MASSAGE [MH] OR MASSAGE [TIAB] OR COGNITIVE RETRAINING [TIAB] OR STIMULATION [TW] OR SENSORY [TW] OR DEPRIVATION TW] OR RANGE OF MOTION EXERCISE* TW] OR POSITIONING [TIAB] OR POSITIONAL CHANGE* [TW] OR POSTURAL CHANGE* [TIAB] OR ACETAMINOPHEN [TW] OR ASPIRIN [TW] OR IBUPROFEN [TW] NAPROXEN [TW] OR DICLOFENAC [TW] OR ANALGESICS [TW] OR NSAIDS [TW] OR ANTIDEPRESSANT* [TW] OR AMITRIPTYLINE [TW] OR NORTRIPTYLINE [TW] OR ANTICONVULSANT* [TW] OR VALPROIC ACID [TW] OR TOPIRAMATE [TW] OR GABAPENTIN [TW] OR PROPRANOLOL [TW] OR B-ADRENERGIC ANTAGONIST* [TW] OR METOPROLOL [TW] OR DIHYDROERGOTAMINE [TW] OR ERGOT [TW] OR SUMATRIPTAN [TW] OR TRIPTANS [TW] OR ZOLMITRIPTAN [TW] OR RIZATRIPTAN [TW] OR VESTIBULAR SUPPRESSANT* [TW] OR MECLIZINE [TW] OR SCOPOLAMINE [TW] OR DIMEHYDRINATE [TW] OR BENZODIAZEPINES [TW] OR LORAZEPAM [TW] OR CLONAZEPAM [TW] OR DIAZEPAM [TW] OR NEUROSTIMULANT* [TW] OR METHYLPHENIDATE [TW] OR DEXTROAMPHETAMINE [TW] OR MODAFANIL [TW] OR AMANTADINE [TW] OR ATOMOXETINE [TW] OR ANTIEMETIC* [TW] OR ONDANSETRON [TW] OR PHENERGAN [TW] OR SEDATIVE-HYPNOTIC* [TW] OR ZOLPIDEM [TW] OR SEROTONIN MODULATOR* [TW] OR TRAZODONE [TW] OR A-ADRENERGIC ANTAGONIST* [TW] OR PRAZOSIN [TW] OR MELATONIN [TW] OR SERTRALINE [TW] OR CITALOPRAM [TW] OR ESCITALOPRAM [TW] OR PAROXETINE [TW] OR FLUOXETINE [TW] OR BUPROPION [TW] OR ACETYLCHOLINESTERASE INHIBITOR* [TW] OR SELECTIVE SEROTONIN REUPTAKE INHIBITOR* [TW] OR DONEPEZIL [TW] OR RIVASTIGMINE [TW] OR GALANTAMINE [TW] OR CYTIDINE **DIPHOSPHATE CHOLINE [TW]**

EMBASE SEARCH STRATEGIES

Question 1: Last run Nov 29, 2012

<u>2,208</u> #2.3 AND #4.31

3,075,946

#2.3 #2.1 OR #2.2

696,823

#2.2

'evoked response'/exp OR 'response time'/exp OR 'body equilibrium'/exp OR 'psychomotor performance'/de OR 'task performance'/exp OR 'memory'/exp OR 'sensory dysfunction'/exp

<u>2,515,609</u>

#2.1

'diagnostic procedure'/de OR 'clinical assessment tool'/exp OR 'clinical observation'/exp OR 'computer assisted diagnosis'/de OR 'delayed diagnosis'/exp OR 'diagnostic accuracy'/exp OR 'diagnostic error'/exp OR 'diagnostic reasoning'/exp OR 'diagnostic test'/de OR 'diagnostic test accuracy study'/exp OR 'diagnostic value'/exp OR 'differential diagnosis'/exp OR 'early diagnosis'/exp OR 'electrodiagnosis'/de OR 'laboratory diagnosis'/exp OR 'molecular diagnosis'/exp OR 'physical examination'/de OR 'qualitative diagnosis'/exp OR 'quantitative diagnosis'/exp OR 'diagnosis'/de OR 'diagnostic approach route'/exp OR 'evaluation or follow up' OR 'clinical assessment'/exp OR 'clinical evaluation'/exp OR 'clinical global impression scale'/exp OR 'course evaluation'/exp OR 'expanded disability status scale'/exp OR 'functional assessment inventory'/exp OR 'gross motor function classification system'/exp OR 'gross motor function measure'/exp OR 'international classification of functioning, disability or health' OR 'oars multidimensional functional assessment questionnaire'/exp OR 'patient assessment'/exp OR 'sensory evaluation'/exp OR 'examination'/de OR 'contact examination'/exp OR 'medical examination'/de OR 'clinical examination'/exp OR 'functional assessment'/exp OR 'measurement'/de OR 'named inventories, questionnaires or rating scales' OR 'neurologic examination'/de OR 'dubowitz neurological assessment'/exp OR 'electroencephalography'/exp OR 'neuropsychological test'/exp OR 'stabilography'/exp OR 'tilt table test'/exp OR 'non invasive measurement/exp OR 'rating scale'/exp OR 'scoring system'/exp OR 'screening'/de OR 'screening test'/exp OR 'sensory system examination'/de OR 'vestibular test'/exp OR 'summated rating scale'/exp

<u>5,412</u>

#4.31 #4.27 OR #4.30

<u>163</u>

#4.30 #4.29 NOT #4.19

<u>163</u>

#4.29 #4.28 NOT #4.24

<u>163</u>

#4.28

('glasgow coma scale'/exp OR 'glasgow outcome scale'/exp) AND '13-15':ab,ti

<u>5,295</u> #4.27 #4.23 NOT #4.26

<u>11,241</u>

#4.26 #4.24 OR #4.25

<u>30</u> #4.25 'oif/oef':ti AND [1990-2013]/py

<u>11,235</u>

#4.24

'iraq war':ti OR 'gulf war':ti OR 'gulf wars':ti OR 'operation iraqi freedom':ti OR 'afghan campaign':ti OR 'operation enduring freedom':ti OR military:ti OR battlefield:ti OR army:ti OR armed AND forces:ti OR marines:ti OR troops:ti OR servicemen:ti OR 'combat related':ti OR 'military personnel':ti OR 'air force':ti OR soldiers:ti OR 'combat veteran':ti OR 'combat disorder':ti OR 'combat disorders':ti OR veterans:ti OR 'active duty':ti OR 'service members':ti AND [1990-2013]/py

<u>5,335</u>

#4.23

#4.22 AND ('case report'/de OR 'clinical article'/de OR 'clinical protocol'/de OR 'clinical trial'/de OR 'cohort analysis'/de OR 'comparative study'/de OR 'control group'/de OR 'controlled clinical trial'/de OR 'controlled study'/de OR 'human'/de OR 'human tissue'/de OR 'interview'/de OR 'major clinical study'/de OR 'medical record review'/de OR 'methodology'/de OR 'model'/de OR 'nonhuman'/de OR 'normal human'/de OR 'outcomes research'/de OR 'practice guideline'/de OR 'prospective study'/de OR 'questionnaire'/de OR 'retrospective study'/de OR 'systematic review'/de) AND ('article'/it OR 'article in press'/it OR 'conference abstract'/it OR 'conference paper'/it OR 'letter'/it OR 'review'/it OR 'short survey'/it)

<u>6,018</u>

#4.22 #4.20 NOT #4.21

452,516

#4.21 'editorial'/exp

<u>6,140</u>

#4.20 #4.16 NOT #4.19

<u>30,949</u>

#4.19 #4.17 NOT #4.18

<u>1,241</u>

#4.18 ('military phenomena'/exp/mj OR 'veteran'/exp/mj) AND (civilian*:ab,ti OR sport* OR athlete*) AND [1990-2013]/py

<u>32,190</u>

#4.17 'military phenomena'/exp/mj OR 'veteran'/exp/mj

<u>6,202</u>

#4.16 #4.12 NOT #4.15

8,001,374

#4.15 #4.13 NOT #4.14

<u>10,227,651</u>

#4.14 'animal'/exp AND 'human'/exp AND [1990-2013]/py

<u>18,229,025</u>

#4.13 'animal'/exp

<u>7,286</u>

#4.12 #4.8 NOT #4.11

<u>4,719,417</u>

#4.11 #4.9 NOT #4.10

<u>665,955</u>

#4.10 'adult'/exp OR 'aged'/exp OR 'middle aged'/exp OR 'embryo'/exp OR 'fetus'/exp AND ('newborn'/exp OR 'child'/exp OR 'adolescent'/exp) AND [1990-2013]/py

<u>5,385,372</u>

#4.9

'adult'/exp OR 'aged'/exp OR 'middle aged'/exp OR 'embryo'/exp OR 'fetus'/exp

<u>10,092</u>

#4.8 #4.4 OR #4.5 OR #4.6 OR #4.7 AND [embase]/lim AND [1990-2013]/py

<u>974</u>

#4.7

mtbi:ab,ti OR mtbis:ab,ti OR 'minimal traumatic brain injury':ab,ti OR 'minimal traumatic brain injuries':ab,ti OR 'minimal tbi':ab,ti OR 'minimal tbis':ab,ti

<u>9,904</u> #4.6 #4.2 AND (mild:ab,ti OR minor:ab,ti)

<u>3,409</u>

#4.5 'brain concussion'/exp

<u>2,878</u>

#4.4 #4.2 AND #4.3

103,017

#4.3 'sport'/exp OR 'athlete'/exp

156,672 #4.2 'brain injury'/exp OR 'head injury'/de OR 'skull injury'/exp

<u>3,578,947</u>

#4.1

'therapy'/de OR 'acupuncture'/exp OR 'animal assisted therapy'/exp OR 'apitherapy'/exp OR 'biological therapy'/exp OR 'computer assisted therapy'/exp OR 'conservative treatment'/exp OR 'counseling'/exp OR 'device therapy'/exp OR 'drug therapy'/exp OR 'early intervention'/exp OR 'experimental therapy'/exp OR 'fluid therapy'/exp OR 'homeopathy'/exp OR 'intensive care'/de OR 'neuromonitoring'/exp OR 'blood pressure monitoring'/exp OR 'body temperature monitoring'/exp OR 'capnography'/exp OR 'life sustaining treatment'/exp OR 'maintenance therapy'/de OR 'patient positioning'/exp OR 'rehabilitation care'/exp OR 'radiotherapy'/de OR 'cognitive rehabilitation'/exp OR 'functional training'/exp OR 'muscle training'/exp OR 'occupational therapy'/exp OR 'rejuvenation'/exp OR 'sensorimotor integration'/exp OR 'speech rehabilitation'/exp OR 'sleep therapy'/exp OR 'supplementation'/exp OR 'systemic therapy'/exp OR 'therapy effect'/exp OR 'treatment response'/exp

Questions 2 & 3 – Last run on Dec. 1, 2012

<u>3,539</u>

<u>2,194,721</u>

#7.19 #7.17 OR #7.18

<u>897,748</u>

#7.18

'diffuse axonal injury'/exp OR 'nerve fiber degeneration'/exp OR 'axonal swelling':ab,ti OR 'retraction balls':ab,ti OR 'brain atrophy'/exp OR 'intracerebral injury':ab,ti OR 'intracerebral injuries':ab,ti OR 'edema'/de OR 'brain edema'/exp OR oedema:ab,ti OR 'axonal injury'/exp OR 'pneumocephalus'/exp OR 'cranial pneumocyst':ab,ti OR 'cranial pneumocysts':ab,ti OR 'intracranial gas':ab,ti OR 'white matter injury':ab,ti OR 'white matter injuries':ab,ti OR 'white matter injury'/exp OR 'skull fracture'/exp OR 'cranial injury':ab,ti OR 'cranial injuries':ab,ti OR lesion*:ab,ti OR 'neuropathology'/exp OR microhemorrhage*:ab,ti OR 'micro hemorrhage':ab,ti OR 'micro hemorrhages':ab,ti OR 'intracranial sequelae':ab,ti

1,504,683

#7.17 #7.13 OR #7.16

347,602

#7.16 #7.14 OR #7.15

<u>347,060</u>

#7.15

'bleeding'/de OR 'brain hemorrhage'/exp OR 'hematoma'/de OR 'brain hematoma'/exp OR 'traumatic hematoma'/exp OR 'meningeal hemorrhage'/exp OR 'brain bleed':ab,ti OR 'brain bleed':ab,ti OR 'brain bleeding':ab,ti OR microbleed*:ab,ti OR 'micro-bleed':ab,ti OR 'micro-bleed':ab,ti OR 'micro-bleed':ab,ti OR hemorrhage*:ab,ti OR haematoma*:ab,ti

<u>782</u>

#7.14

'intracranial injury':ab,ti OR 'intracranial injuries':ab,ti

<u>1,213,947</u>

#7.13

'nuclear magnetic resonance imaging'/exp OR zeugmatography:ab,ti OR 'tomography'/de OR 'brain tomography'/exp OR 'computer assisted tomography'/exp OR 'emission tomography'/exp OR 'optical coherence tomography'/exp OR tomography:ab,ti OR 'mri scan':ab,ti OR 'mri scans':ab,ti OR fmri:ab,ti OR fmris:ab,ti OR mri:ab,ti OR mris:ab,ti OR 'neuroimaging'/exp OR 'cat scan':ab,ti OR 'cat scans':ab,ti OR 'ct scan':ab,ti OR 'ct scans':ab,ti OR 'radiodensitometry'/exp OR 'x ray'/exp OR 'x-ray':ab,ti OR 'x ray':ab,ti OR 'x rays':ab,ti OR 'xrays':ab,ti OR xray*:ab,ti

<u>8,388</u>

#7.12 #7.10 AND #7.11

<u>1,466,199</u>

#7.11

'health care facility'/de OR 'hospital'/exp OR 'emergency health service'/exp OR 'hospital service'/exp OR 'emergency department':ab,ti OR 'emergency departments':ab,ti OR 'hospital patient'/de OR 'hospitalized adolescent'/exp OR 'hospitalized child'/exp OR 'hospitalized infant'/exp OR 'emergency patient'/exp OR 'critically ill patient'/exp OR 'high risk patient'/exp OR 'medical center':ab,ti OR 'medical centers':ab,ti OR 'acute care':ab,ti OR hospital*:ab,ti OR 'trauma center':ab,ti OR 'trauma centers':ab,ti OR 'trauma unit':ab,ti OR 'trauma units':ab,ti OR 'emergency service':ab,ti OR 'nontrauma center':ab,ti OR 'non trauma center':ab,ti OR 'nontrauma centers':ab,ti OR 'non trauma centers':ab,ti

<u>54,306</u>

#7.10 #7.8 NOT #7.9

<u>32,259</u>

#7.9 'military phenomena'/exp/mj OR 'veteran'/exp/mj OR 'veterans health'/exp/mj

<u>54,493</u>

#7.8

#7.4 NOT #7.7

4,153,342

#7.7 #7.5 NOT (#7.5 AND #7.6)

<u>2,632,147</u>

#7.6 'child'/exp OR 'adolescent'/exp OR 'newborn'/exp

5,090,845

#7.5 'adult'/exp OR 'aged'/exp OR 'middle aged'/exp

<u>82,997</u>

#7.4

#7.3 AND (1990:py OR 1991:py OR 1992:py OR 1993:py OR 1994:py OR 1995:py OR 1996:py OR 1997:py OR 1998:py OR 1999:py OR 2000:py OR 2001:py OR 2002:py OR 2003:py OR 2004:py OR 2005:py OR 2006:py OR 2007:py OR 2008:py OR 2009:py OR 2010:py OR 2011:py OR 2012:py OR 2013:py) AND ('article'/it OR 'article in press'/it OR 'conference paper'/it OR 'letter'/it OR 'review'/it)

<u>137,019</u>

#7.3 #7.1 NOT #7.2

4,236,760

#7.2 'animal'/exp NOT ('animal'/exp AND 'human'/exp)

<u>159,318</u>

#7.1

'brain concussion'/exp OR 'brain injury'/exp OR mtbi:ab,ti OR mtbis:ab,ti OR 'head injury'/de OR 'skull injury'/de OR 'second impact syndrome'/exp OR tbi:ab,ti OR tbis:ab,ti OR concussion*:ab,ti OR 'brain laceration':ab,ti OR 'brain lacerations':ab,ti OR 'cerebral injury':ab,ti OR 'cerebral injuries':ab,ti OR 'brain contusion':ab,ti OR 'brain contusions':ab,ti OR 'brain trauma':ab,ti OR 'brain traumas':ab,ti OR 'brain injury':ab,ti OR 'brain injuries':ab,ti OR 'skull fracture'/de OR 'skull base fracture'/de

Questions 4 & 5 – Last run on Dec. 1, 2012

<u>3,229</u> #3.39

#3.36 NOT #3.38

<u>241</u>

#3.38 #3.36 AND #3.37

<u>1,895,899</u>

#3.37 'case report'/exp

<u>3,470</u>

#3.36 #3.35 NOT #3.32

<u>3,541</u>

#3.35 #3.31 AND #3.34

4,251,395

#3.34 #3.1 OR #3.33

<u>1,628,084</u>

#3.33

'risk factor'/exp OR 'risk'/de OR 'high risk patient'/exp OR 'high risk infant'/exp OR 'risk assessment'/exp OR 'precipitating factor':ab,ti OR 'precipitating factors':ab,ti OR 'age factor':ab,ti OR 'age factors':ab,ti OR 'time factor':ab,ti OR 'time factors':ab,ti OR 'analysis of variance'/exp OR 'sex difference'/exp OR 'mechanism of injury':ab,ti OR 'mechanisms of injury':ab,ti OR 'time since injury':ab,ti OR 'between group difference':ab,ti OR 'between group differences':ab,ti OR epidemiology:de,ab,ti OR 'age distribution':de OR 'seasonal variation':de

<u>1,470,038</u>

#3.32

rat:ti OR rats:ti OR rodent*:ti OR mice:ti OR mouse:ti OR geriatric:ti OR elderly:ti OR 'animal model':ti OR alzheimer*:ti OR 'oif/oef':ti OR 'oef/oif':ti OR military:ti OR iraq:ti OR afghanistan:ti OR 'combat related':ti OR army:ti OR soldiers:ti OR veterans:ti OR 'air force':ti OR 'gulf war':ti OR 'gulf wars':ti OR 'operation enduring freedom':ti OR 'operation iraqi freedom':ti OR 'afghan war':ti OR 'afghan campaign':ti OR marines:ti OR troops:ti OR servicemen:ti OR 'combat disorders':ti OR 'combat disorder':ti OR 'active duty':ti

<u>5,412</u>

#3.31 #3.27 OR #3.30

<u>163</u>

#3.30 #3.29 NOT #3.19

<u>163</u>

#3.29 #3.28 NOT #3.24

<u>163</u>

#3.28 'glasgow coma scale'/exp OR 'glasgow outcome scale'/exp AND '13-15':ab,ti

<u>5,295</u> #3.27 #3.23 NOT #3.26

<u>11,241</u>

#3.26 #3.24 OR #3.25

<u>30</u> #3.25 'oif/oef':ti AND [1990-2013]/py

<u>11,235</u>

#3.24

'iraq war':ti OR 'gulf war':ti OR 'gulf wars':ti OR 'operation iraqi freedom':ti OR 'afghan campaign':ti OR 'operation enduring freedom':ti OR military:ti OR battlefield:ti OR army:ti OR armed AND forces:ti OR marines:ti OR troops:ti OR servicemen:ti OR 'combat related':ti OR 'military personnel':ti OR 'air force':ti OR soldiers:ti OR 'combat veteran':ti OR 'combat disorder':ti OR 'combat disorders':ti OR veterans:ti OR 'active duty':ti OR 'service members':ti AND [1990-2013]/py

<u>5,335</u>

#3.23

#3.22 AND ('case report'/de OR 'clinical article'/de OR 'clinical protocol'/de OR 'clinical trial'/de OR 'cohort analysis'/de OR 'comparative study'/de OR 'control group'/de OR 'controlled clinical trial'/de OR 'controlled study'/de OR 'human'/de OR 'human tissue'/de OR 'interview'/de OR 'major clinical study'/de OR 'medical record review'/de OR 'methodology'/de OR 'model'/de OR 'nonhuman'/de OR 'normal human'/de OR 'outcomes research'/de OR 'practice guideline'/de OR 'prospective study'/de OR 'questionnaire'/de OR 'retrospective study'/de OR 'systematic review'/de) AND ('article'/it OR 'article in press'/it OR 'conference abstract'/it OR 'conference paper'/it OR 'letter'/it OR 'review'/it OR 'short survey'/it)

<u>6,018</u>

#3.22 #3.20 NOT #3.21

<u>452,516</u>

#3.21 'editorial'/exp

<u>6,140</u>

#3.20 #3.16 NOT #3.19

<u>30,949</u>

#3.19 #3.17 NOT #3.18

<u>1,241</u>

#3.18

('military phenomena'/exp/mj OR 'veteran'/exp/mj) AND (civilian*:ab,ti OR sport* OR athlete*) AND [1990-2013]/py

<u>32,190</u>

#3.17 'military phenomena'/exp/mj OR 'veteran'/exp/mj

<u>6,202</u>

#3.16 #3.12 NOT #3.15

8,001,374

#3.15 #3.13 NOT #3.14

<u>10,227,651</u>

#3.14 'animal'/exp AND 'human'/exp AND [1990-2013]/py

<u>18,229,025</u>

#3.13 'animal'/exp

<u>7,286</u>

#3.12 #3.8 NOT #3.11

4,719,417

#3.11 #3.9 NOT #3.10

<u>665,955</u>

#3.10

('adult'/exp OR 'aged'/exp OR 'middle aged'/exp OR 'embryo'/exp OR 'fetus'/exp) AND ('newborn'/exp OR 'child'/exp OR 'adolescent'/exp) AND [1990-2013]/py

<u>5,385,372</u>

#3.9

'adult'/exp OR 'aged'/exp OR 'middle aged'/exp OR 'embryo'/exp OR 'fetus'/exp

<u>10,092</u>

#3.8 #3.4 OR #3.5 OR #3.6 OR #3.7 AND [embase]/lim AND [1990-2013]/py

<u>974</u>

#3.7

mtbi:ab,ti OR mtbis:ab,ti OR 'minimal traumatic brain injury':ab,ti OR 'minimal traumatic brain injuries':ab,ti OR 'minimal tbi':ab,ti OR 'minimal tbis':ab,ti

<u>9,904</u> #3.6 #3.2 AND (mild:ab,ti OR minor:ab,ti)

<u>3,409</u> #3.5 'brain concussion'/exp

<u>2,878</u> #3.4 #3.2 AND #3.3

<u>103,017</u>

#3.3 'sport'/exp OR 'athlete'/exp

<u>156,672</u> #3.2 'brain injury'/exp OR 'head injury'/de OR 'skull injury'/exp

<u>3,030,325</u>

#3.1

'complication'/de OR 'disease exacerbation'/exp OR 'neurological complication'/exp OR 'posttraumatic complication'/exp OR 'posttraumatic pain'/exp OR 'prognosis'/de OR 'disease course'/de OR 'chronic disease'/exp OR 'illness trajectory'/exp OR 'recurrent disease'/exp OR 'relapse'/exp OR 'terminal disease'/exp OR 'acute disease'/exp OR 'adverse outcome'/exp OR 'bothersomeness'/exp OR 'deterioration'/exp OR 'disability severity'/exp OR 'injury severity'/exp OR 'nomogram'/exp OR 'treatment failure'/exp OR 'forecasting'/exp OR 'prediction'/exp OR 'predictive value'/exp OR 'death'/de OR 'lethality'/exp OR 'disability'/exp OR 'disease severity'/de OR impairment*:ab,ti OR sequelae:ab,ti OR impairment*:de OR 'dysfunction:ab,ti OR refractory:ab,ti OR 'more severe':ab,ti OR 'persisting symptoms':ab,ti OR 'prolonged:ab,ti OR 'long term symptoms':ab,ti OR 'longterm symptoms':ab,ti OR 'symptomatology'/exp

Question 6 – Last run on Dec. 1, 2012

<u>976</u>

#4.44 #4.41 NOT #4.43 <u>126</u> #4.43 #4.41 AND #4.42

1,894,200

#4.42 'case report'/exp

<u>1,102</u>

#4.41 #4.39 NOT #4.40

1,385,362

#4.40 rat:ti OR rats:ti OR mouse:ti OR mice:ti OR rodent*:ti OR geriatric:ti OR elderly:ti OR 'animal model':ti OR 'oif/oef':ti OR 'oef/oif':ti

<u>1,111</u>

#4.39 #4.37 AND #4.38

<u>5,588,005</u>

#4.38

longterm:ab,ti OR symptom*:de,ab,ti OR symptomatic:ab,ti OR sequelae:ab,ti OR enduring:ab,ti OR delayed:ab,ti OR prolong*:ab,ti OR ongoing:ab,ti OR persistent:ab,ti OR persistence:ab,ti OR

persisting:ab,ti OR lasting:ab,ti OR 'long term':ab,ti OR 'long-term':ab,ti OR 'post-concussion syndrome':de,ab,ti OR 'postconcussive syndrome':ab,ti OR 'post-concussive syndrome':ab,ti OR 'postconcussion syndrome':ab,ti OR symptomatology:de,ab,ti OR subsymptom*:ab,ti OR unresolved:ab,ti OR continuing:ab,ti OR continued:ab,ti OR unremitting:ab,ti OR lingering:ab,ti OR protracted:ab,ti OR 'greater than':ab,ti OR posttraumatic:ab,ti OR 'post-traumatic':ab,ti OR complication*:de,ab,ti OR complicated:ab,ti OR recovery:de,ab,ti OR recover:de,ab,ti OR recovered:ab,ti OR disability:de,ab,ti OR disabilities:de,ab,ti OR disabled:de,ab,ti OR impairment*:de,ab,ti OR impaired:ab,ti OR rehabilitation:de,ab,ti OR reintegration:ab,ti OR 'reintegration':ab,ti OR failure:de,ab,ti OR remediation:ab,ti OR remediate:ab,ti OR refractory:ab,ti OR extended:ab,ti OR 'longer than':ab,ti OR 'more than':ab,ti OR problematic:ab,ti

<u>1,546</u>

#4.37 #4.31 AND #4.36

<u>6,141,110</u>

#4.36 #4.1 OR #4.32 OR #4.33 OR #4.34 OR #4.35

<u>1,610,488</u>

#4.35

'paracetamol'/exp OR acetaminophen:de,ab,ti OR 'acetylsalicylic acid'/exp OR 'ibuprofen'/exp OR 'naproxen'/exp OR 'diclofenac'/exp OR 'analgesic agent'/exp OR 'nonsteroid antiinflammatory agent'/exp OR nsaid*:ab,ti OR 'antidepressant agent'/exp OR 'amitriptyline'/exp OR 'nortriptyline'/exp OR 'anticonvulsive agent'/exp OR 'valproic acid'/exp OR 'topiramate'/exp OR 'gabapentin'/exp OR 'propranolol'/exp OR 'beta adrenergic receptor blocking agent'/exp OR 'metoprolol'/exp OR 'dihydroergotamine'/exp OR ergot*:de,ab,ti OR 'sumatriptan'/exp OR 'antimigraine agent'/exp OR 'vestibular suppressant':ab,ti OR 'vestibular suppressants':ab,ti OR 'meclozine'/exp OR 'scopolamine'/exp OR 'benzodiazepine derivative'/exp OR neurostimulant*:ab,ti OR 'methylphenidate'/exp OR 'dexamphetamine'/exp OR modafanil:ab,ti OR 'amantadine'/exp OR 'atomoxetine'/exp OR 'antiemetic agent'/exp OR 'promethazine'/exp OR 'hypnotic sedative agent'/exp OR 'zolpidem'/exp OR 'serotonin modulator':ab,ti OR 'serotonin modulators':ab,ti OR 'trazodone'/exp OR 'alpha adrenergic receptor blocking agent'/exp OR 'melatonin'/exp OR 'sertraline'/exp OR 'citalopram'/exp OR 'escitalopram'/exp OR 'paroxetine'/exp OR 'fluoxetine'/exp OR 'amfebutamone'/exp OR 'cholinesterase inhibitor'/exp OR 'serotonin uptake inhibitor'/exp OR 'donepezil'/exp OR 'rivastigmine'/exp OR 'galantamine'/exp OR 'cytidine diphosphate choline':ab,ti

569,867

#4.34

'neurofeedback'/exp OR biofeedback:de,ab,ti OR 'hyperbaric oxygen'/exp OR 'kinesiotherapy'/exp OR 'physiotherapy'/exp OR 'cognitive therapy'/exp OR 'induced hypothermia'/exp OR 'rest'/exp OR neurorehabilitation:ab,ti OR 'stress reduction':ab,ti OR 'cognitive rest':ab,ti OR 'neuroprotective agent'/exp OR 'sodium chloride'/exp OR 'hypertonic saline solution':ab,ti OR 'nutraceutical'/exp OR 'diet supplementation'/exp OR 'massage'/exp OR 'cognitive retraining':ab,ti OR 'sensory stimulation'/exp OR 'exercise'/exp OR 'positional change':ab,ti OR 'postural change':ab,ti

144,728

#4.33 intervention*:de,ti

<u>2,010,750</u>

#4.32 treatment*:de,ti

<u>5,412</u>

#4.31 #4.27 OR #4.30

<u>163</u>

#4.30 #4.29 NOT #4.19

<u>163</u>

#4.29 #4.28 NOT #4.24

<u>163</u>

#4.28 ('glasgow coma scale'/exp OR 'glasgow outcome scale'/exp) AND '13-15':ab,ti

<u>5,295</u>

#4.27 #4.23 NOT #4.26

<u>11,241</u>

#4.26 #4.24 OR #4.25

<u>30</u>

<u>11,235</u>

#4.24

'iraq war':ti OR 'gulf war':ti OR 'gulf wars':ti OR 'operation iraqi freedom':ti OR 'afghan campaign':ti OR 'operation enduring freedom':ti OR military:ti OR battlefield:ti OR army:ti OR armed AND forces:ti OR marines:ti OR troops:ti OR servicemen:ti OR 'combat related':ti OR 'military personnel':ti OR 'air force':ti OR soldiers:ti OR 'combat veteran':ti OR 'combat disorder':ti OR 'combat disorders':ti OR veterans:ti OR 'active duty':ti OR 'service members':ti AND [1990-2013]/py

<u>5,335</u>

#4.23

#4.22 AND ('case report'/de OR 'clinical article'/de OR 'clinical protocol'/de OR 'clinical trial'/de OR 'cohort analysis'/de OR 'comparative study'/de OR 'control group'/de OR 'controlled clinical trial'/de OR 'controlled study'/de OR 'human'/de OR 'human tissue'/de OR 'interview'/de OR 'major clinical study'/de OR 'medical record review'/de OR 'methodology'/de OR 'model'/de OR 'nonhuman'/de OR 'normal human'/de OR 'outcomes research'/de OR 'practice guideline'/de OR 'prospective study'/de OR 'questionnaire'/de OR 'retrospective study'/de OR 'systematic review'/de) AND ('article'/it OR 'article in press'/it OR 'conference abstract'/it OR 'conference paper'/it OR 'letter'/it OR 'review'/it OR 'short survey'/it)

<u>6,018</u>

#4.22 #4.20 NOT #4.21

<u>452,516</u>

#4.21 'editorial'/exp

<u>6,140</u>

#4.20 #4.16 NOT #4.19

<u>30,949</u>

#4.19 #4.17 NOT #4.18

<u>1,241</u>

#4.18

'military phenomena'/exp/mj OR 'veteran'/exp/mj AND (civilian*:ab,ti OR sport* OR athlete*) AND [1990-2013]/py

<u>32,190</u>

#4.17

'military phenomena'/exp/mj OR 'veteran'/exp/mj

<u>6,202</u>

#4.16 #4.12 NOT #4.15

8,001,374

#4.15 #4.13 NOT #4.14

<u>10,227,651</u>

#4.14 'animal'/exp AND 'human'/exp AND [1990-2013]/py

18,229,025

#4.13 'animal'/exp

<u>7,286</u>

#4.12 #4.8 NOT #4.11

<u>4,719,417</u>

#4.11 #4.9 NOT #4.10

<u>665,955</u>

#4.10

'adult'/exp OR 'aged'/exp OR 'middle aged'/exp OR 'embryo'/exp OR 'fetus'/exp AND ('newborn'/exp OR 'child'/exp OR 'adolescent'/exp) AND [1990-2013]/py

<u>5,385,372</u>

#4.9

'adult'/exp OR 'aged'/exp OR 'middle aged'/exp OR 'embryo'/exp OR 'fetus'/exp

<u>10,092</u>

#4.8 #4.4 OR #4.5 OR #4.6 OR #4.7 AND [embase]/lim AND [1990-2013]/py

<u>974</u>

#4.7

mtbi:ab,ti OR mtbis:ab,ti OR 'minimal traumatic brain injury':ab,ti OR 'minimal traumatic brain injuries':ab,ti OR 'minimal tbi':ab,ti OR 'minimal tbis':ab,ti

<u>9,904</u>

#4.6 #4.2 AND (mild:ab,ti OR minor:ab,ti)

<u>3,409</u>

#4.5 'brain concussion'/exp

<u>2,878</u> #4.4 #4.2 AND #4.3

103,017

#4.3 'sport'/exp OR 'athlete'/exp

<u>156,672</u>

#4.2 'brain injury'/exp OR 'head injury'/de OR 'skull injury'/exp

<u>3,578,947</u>

#4.1

'therapy'/de OR 'acupuncture'/exp OR 'animal assisted therapy'/exp OR 'apitherapy'/exp OR 'biological therapy'/exp OR 'computer assisted therapy'/exp OR 'conservative treatment'/exp OR 'counseling'/exp OR 'device therapy'/exp OR 'drug therapy'/exp OR 'early intervention'/exp OR 'experimental therapy'/exp OR 'fluid therapy'/exp OR 'homeopathy'/exp OR 'intensive care'/de OR 'neuromonitoring'/exp OR 'blood pressure monitoring'/exp OR 'body temperature monitoring'/exp OR 'capnography'/exp OR 'life sustaining treatment'/exp OR 'maintenance therapy'/de OR 'patient positioning'/exp OR 'rehabilitation care'/exp OR 'radiotherapy'/de OR 'cognitive rehabilitation'/exp OR 'functional training'/exp OR 'muscle training'/exp OR 'occupational therapy'/exp OR 'rejuvenation'/exp OR 'sensorimotor integration'/exp OR 'speech rehabilitation'/exp OR 'sleep therapy'/exp OR 'supplementation'/exp OR 'systemic therapy'/exp OR 'therapy effect'/exp OR 'treatment response'/exp

ERIC SEARCH STRATEGY

Due to the subject content of the ERIC database (education), only one search strategy was used. The rationale is that there will be so few citations relative to the major biomedical databases, such as PubMed and Embase, that searching only for the *population* will not generate enough results to necessitate narrowing down by the focus of each individual question.

((Keywords:sport* or Keywords:athlete* or Keywords:athletic or Keywords:recreation*) and (Keywords:"head injury" or Keywords:"head injuries" or Keywords:"brain injury" or Keywords:"brain injuries" or Keywords:"head trauma" or Keywords:"brain trauma" or Keywords:"skull injury" or Keywords:"skull injuries" or Keywords:tbi or Keywords:tbis or **Keywords:**"cerebral trauma" or **Keywords:**"cerebral injury" or **Keywords:**"cerebral injuries"))**Publication Date:**1990-2013

OR

(Keywords:"gcs 13-15" or Keywords:concussion* or Keywords:postconcussion or Keywords:"post concussion" or Keywords:postconcussive or Keywords:"post concussive" or Keywords:"mild tbi" or Keywords:"mild tbis" or Keywords:mtbi or Keywords:mtbis or Keywords:"mild traumatic brain injury" or Keywords:"mild traumatic brain injuries")Publication Date:1990-2013

Or

((Keywords:"traumatic brain injury" or Keywords:"traumatic brain injuries" or Keywords:tbi or Keywords:tbis or Keywords:"brain contusion" or Keywords:"brain contusions" or Keywords:"brain trauma" or Keywords:"brain laceration" or Keywords:"brain lacerations" or Keywords:"cerebral injury" or Keywords:"cerebral injuries" or Keywords:"head injury" or Keywords:"head injuries" or Keywords:"head trauma" or Keywords:"cerebral trauma" or Keywords:"skull injury" or Keywords:"skull injuries" or Keywords:"craniocerebral trauma") and (Keywords:mild or Keywords:minor))Publication Date:1990-2013

SPORTDISCUS SEARCH STRATEGY

Due to the subject content of the SportDiscus database (sports/recreation), only one search strategy was used. The rationale is that there will be so few citations relative to the major biomedical databases, such as PubMed and Embase, that searching only for the *population* will not generate enough results to necessitate narrowing down by the focus of each individual question.

S6	TX (youth or child or children or childhood or toddler* or infant* or teen or teens or teenager* or "teen ager" or "teen agers" or adolescent* or adolescence or pediatric* of paediatric*) AND S5	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - SPORTDiscus with Full Text	171
S5	KW s4	Limiters - Publication Type: Journal Article,	Interface - EBSCOhost Search Screen - Advanced	471

54	KW S3 or S2 or S1	Thesis or dissertation Search modes - Boolean/Phrase Limiters - Published Date: 19900101-20121231; Publication Type: Journal Article, Thesis or dissertation Search modes - Boolean/Phrase	Search Database - SPORTDiscus with Full Text Interface - EBSCOhost Search Screen - Advanced Search Database - SPORTDiscus with Full Text	471
53	KW ("traumatic brain injury" or "traumatic brain injuries" or tbi or tbis or "brain contusion" or "brain contusions" or "brain laceration" or "brain lacerations" or "cerebral injury" or "cerebral injuries" or "head injury" or "head injuries" or "head trauma" or "cerebral trauma" or "skull injury" or "skull injuries" or "craniocerebral trauma" or "brain injury" or "brain injuries") AND TX (mild or minor)	Limiters - Published Date: 19900101-20121231; Publication Type: Journal Article, Thesis or dissertation Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - SPORTDiscus with Full Text	322
S2	KW "gcs 13-15" or concussion* or concussive or postconcussion or "post concussion" or postconcussive or "post	Limiters - Published Date: 19900101-20121231; Publication Type: Journal Article, Thesis or dissertation	Interface - EBSCOhost Search Screen - Advanced Search Database - SPORTDiscus with Full Text	259

	concussive" or "mild tbi" or "mild tbis" or mtbi or mtbis or "mild traumatic brain injury" or "mild traumatic brain injuries"	Search modes - Boolean/Phrase		
S1	KW ("head injury" or "head injuries" or "brain injury" or "brain injuries" or "head trauma" or "brain trauma" or "skull injury" or "skull injuries" or tbi or tbis or "cerebral trauma" or "cerebral injury" or "cerebral injuries") AND KW (sport* or athlete* or athletic or recreation)	Limiters - Published Date: 19900101-20121231; Publication Type: Journal Article, Thesis or dissertation <u>Search</u> modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - SPORTDiscus with Full Text	43

CINAHL SEARCH STRATEGY

CINAHL was searched only for Question 6, dealing with treatment. The rationale is that CINAHL could have unique studies dealing with treatments provided by allied health professionals. The treatments searched did not include specific medications since it is unlikely that drug studies would appear in CINAHL, but not PubMed or Embase.

S43	S41	Limiters - Exclude MEDLINE records Narrow by SubjectAge6: - infant, newborn: birth-1 month Narrow by SubjectAge5: - infant: 1-23 months Narrow by SubjectAge4: - all infant Narrow by SubjectAge3: -	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	369
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		child, preschool: 2-5 years Narrow by SubjectAge2: - adolescent: 13-18 years Narrow by SubjectAge1: - child: 6-12 years Narrow by SubjectAge0: - all child Search modes - Boolean/Phrase		
S42	S41	Limiters - Exclude MEDLINE records Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	6,650
S41	S40 AND S15	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	9,413
S40	("NEUROFEEDBACK") OR (S16 OR S17 OR S18 OR S19 OR S20 OR S21 OR S22 OR S23 OR S24 OR S25 OR S26 OR S27 OR S28 OR S29 OR S30 OR S31 OR S32 OR S33 OR S34 OR S35 OR S36 OR S37 OR S38 OR S39)	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	1,064,298

S39	"NEUROFEEDBACK"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	168
S38	(MH "Patient Positioning+") OR "POSITIONING"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	10,049
S37	"RANGE OF MOTION EXERCISE"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	34
S36	(MH "Sensory Deprivation")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	218
S35	(MH "Sensory Stimulation+") OR (MH "Physical Stimulation+") OR (MH "Transcutaneous Electric Nerve Stimulation") OR (MH "Acoustic Stimulation") OR (MH "Electric	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL	24,968

	Stimulation+") OR (MH "Electrical Stimulation, Functional") OR (MH "Electrical Stimulation, Neuromuscular") OR (MH "Deep Brain Stimulation") OR "STIMULATION"		with Full Text;Nursing & Allied Health Collection: Comprehensive	
S34	(MH "Massage+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	7,089
S33	(MH "Dietary Supplements+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	9,906
S32	(MH "Functional Food")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	565
S31	"COGNITIVE REST"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	4

S30	(MH "Stress Management") OR "STRESS REDUCTION"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	5,530
S29	"PHYSICAL REST"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	13
S28	(MH "Bed Rest")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	773
S27	(MH "Hypothermia, Induced")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	1,405
S26	(MH "Cognitive Therapy") OR "COGNITIVE THERAPY" OR (MH "Recreational Therapy")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL	8,545

			with Full Text;Nursing & Allied Health Collection: Comprehensive	
S25	(MH "Physical Therapy+") OR "PHYSICAL THERAPY" OR (MH "Pediatric Physical Therapy")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	81,215
S24	(MH "Therapeutic Exercise+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	24,929
S23	(MH "Hyperbaric Oxygenation")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	1,245
S22	(MH "Biofeedback") OR "BIOFEEDBACK"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	2,578

S21	(MH "Rehabilitation, Athletic") OR (MH "Rehabilitation") OR "REHABILITATION" OR (MH "Rehabilitation, Psychosocial") OR (MH "Rehabilitation, Speech and Language") OR (MH "Rehabilitation, Pediatric")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	114,697
S20	(MH "Diet Therapy") OR (MH "Apitherapy") OR (MH "Pediatric Physical Therapy") OR (MH "Behavior Therapy") OR (MH "Pediatric Occupational Therapy") OR (MH "Motion Therapy, Continuous Passive") OR (MH "Magnet Therapy") OR (MH "Color Therapy") OR (MH "Electroconvulsive Therapy")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	11,143
S19	(MH "Pharmacological and Biological Treatments") OR (MH "Treatment Outcomes") OR "TREATMENT"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	431,742
S18	(MH "Drug Therapy") OR (MH "Drug Therapy, Combination") OR (MH "Fluid Therapy") OR (MH "Intravenous Therapy") OR (MH "Prescriptions, Drug") OR (MH "Manual Therapy") OR (MH "Phototherapy") OR (MH "Radiotherapy") OR (MH "Rehabilitation") OR (MH "Respiratory Therapy") OR (MH "Stress Management") OR (MH "Therapy, Computer Assisted") OR (MH "Vestibular Stimulation")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	63,672
S17	TX THERAPEUTIC	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL	392,389

			with Full Text;Nursing & Allied Health Collection: Comprehensive	
S16	"THERAPY"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	603,914
S15	S13 NOT S14	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	16,007
S14	TI IRAQ OR IRAQI OR AFGHANISTAN OR "GULF WAR" OR "GULF WARS" OR AFGHAN OR "OPERATION ENDURING FREEDOM" OR MILITARY OR BATTLEFIELD OR ARMY OR "ARMED FORCES" OR MARINES OR TROOPS OR SERVICEMEN OR "COMBAT RELATED" OR "AIR FORCE" OR SOLDIERS OR "OIF/OEF" OR VETERANS OR "ACTIVE DUTY" OR "SERVICE MEMBERS"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	14,619
S13	S7 OR S9 OR S10 OR S11	Limiters - Published Date from: 19900101- 20121231 Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	16,356

S12	S7 OR S9 OR S10 OR S11	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	16,948
S11	TX "MINIMAL TRAUMATIC BRAIN INJURY" OR "MINIMAL TRAUMATIC BRAIN INJURIES"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	1
S10	TX (TBI OR TBIS OR "BRAIN CONTUSION" OR "BRAIN CONTUSIONS" OR "BRAIN TRAUMA" OR "BRAIN LACERATION" OR "BRAIN LACERATIONS" OR "CEREBRAL INJURY" OR "CEREBRAL INJURIES" OR "HEAD INJURY" OR "HEAD INJURIES" OR "HEAD TRAUMA" OR "CEREBRAL TRAUMA" OR "SKULL INJURY" OR "SKULL INJURIES" OR "CRANIOCEREBRAL TRAUMA" OR "BRAIN INJURY" OR "BRAIN INJURIES") AND TX (MILD OR MINOR)	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	15,832
S9	"GCS 13-15" OR MTBI OR MTBIS OR "MILD TRAUMATIC BRAIN INJURY" OR "MILD TRAUMATIC BRAIN INJURIES"	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	791
S8	(MH "Brain Concussion+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL	1,395

			with Full Text;Nursing & Allied Health Collection: Comprehensive	
S7	S6 AND (S2 OR S1)	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	1,314
S6	S5 OR S4 OR S3	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	51,836
S5	(MH "Recreation+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	11,629
S4	(MH "Athletes+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	12,216

\$3	(MH "Sports+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	36,674
52	(MH "Head Injuries+") OR TX (tbi or tbis)	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	37,418
S1	(MH "Head Injuries+")	Search modes - Boolean/Phrase	Interface - EBSCOhost Search Screen - Advanced Search Database - CINAHL with Full Text;Nursing & Allied Health Collection: Comprehensive	18,968

Appendix D: Classification of Evidence Scheme

Appendix D Classification of Evidence Scheme

American Academy of Neurology Classification of Evidence Schemes

Criteria for Rating Therapeutic Studies

Class I Criteria

- Randomized controlled clinical trial (RCT) in a representative population

- Masked or objective outcome assessment

- Relevant baseline characteristics are presented and substantially equivalent between treatment groups, or there is appropriate statistical adjustment for differences

- Additional Class I criteria:

a. Concealed allocation

b. No more than two primary outcomes specified

c. Exclusion and inclusion criteria clearly defined

d. Adequate accounting of dropouts (with at least 80 percent of participants completing the study) and crossovers

e. For noninferiority or equivalence trials claiming to prove efficacy for one or both drugs, the following are also required^{*}:

i. The authors explicitly state the clinically meaningful difference to be excluded by defining the threshold for equivalence or noninferiority

ii. The standard treatment used in the study is substantially similar to that used in previous studies establishing efficacy of the standard treatment (e.g., for a drug, the mode of administration, dose, and dosage adjustments are similar to those previously shown to be effective)

iii. The inclusion and exclusion criteria for participant selection and the outcomes of participants on the standard treatment are comparable with those of previous studies establishing efficacy of the standard treatment

iv. The interpretation of the study results is based on a per-protocol analysis that accounts for dropouts or crossovers

f. For crossover trials, both period and carryover effects are examined and statistical adjustments performed, if appropriate

Class II Criteria

- RCT that lacks one or two Class I criteria a-e (see above)

- Cohort study meeting Class I criteria b–e (see above)
- Randomized crossover trial missing one of the following two criteria:
 - a. Period and carryover effects described
 - b. Baseline characteristics of treatment order groups presented
- All relevant baseline characteristics are presented and substantially equivalent across treatment groups, or there is appropriate statistical adjustment for differences
- Masked or objective outcome assessment

Class III Criteria

- Controlled studies (including studies with external controls such as well-defined natural history controls)

- Crossover trial missing both of the following two criteria:

- a. Period and carryover effects
- b. Presentation of baseline characteristics

- A description of major confounding differences between treatment groups that could affect outcome^{**}

- Outcome assessment masked, objective,** or performed by someone who is not a member of the treatment team

Class IV Criteria

If any therapeutic study meets any one of the following criteria, the study must be classified as Class IV.

- Patients with the disease are not included
- Patients receiving different interventions are not included
- Interventions or outcome measures are undefined or unaccepted
- Measures of effectiveness or statistical precision are either not presented or not calculable

*Numbers i–iii in Class Ie are required for Class II in equivalence trials. If any one of the three is missing, the class is automatically downgraded to Class III

**Objective outcome measurement: an outcome measure that is unlikely to be affected by an observer's (patient, treating physician, investigator) expectation or bias (e.g., blood tests, administrative outcome data)

Criteria for Rating Causation Studies

Class I Criteria

- Cohort survey with prospective data collection

- All relevant confounding characteristics are presented and substantially equivalent between comparison groups, or there is appropriate statistical adjustment for differences

- Outcome measurement is objective or determined without knowledge of risk factor status
- Additional Class I criteria:
 - a. No more than two primary outcomes specified
 - b. Exclusion and inclusion criteria clearly defined

c. Accounting of dropouts indicated (with at least 80 percent of participants completing the study)

Class II Criteria

- Cohort study with retrospective data collection or case-control study. Study meets Class I criteria a–c (see above)

- All relevant confounding characteristics are presented and substantially equivalent across comparison groups, or there is appropriate statistical adjustment for differences

- Masked or objective outcome assessment

Class III Criteria

- Cohort or case-control study designs

- A description of major confounding differences between risk groups that could affect outcome^{**}

- Outcome assessment masked, objective,** or performed by someone other than the

investigator who measured the risk factor

Class IV Criteria

If any study meets any one of the following criteria, the study must be classified as Class IV Persons at risk for the disease are not included

- Persons at risk for the disease are not included
- Both patients with and patients without the risk factor are not included
- Measures of risk factor or outcomes are undefined or unaccepted
- Measures of association or statistical precision are either not presented or not calculable

**Objective outcome measurement: an outcome measure that is unlikely to be affected by an observer's (patient, treating physician, investigator) expectation or bias (e.g., blood tests, administrative outcome data)

Criteria for Rating Prognostic Accuracy Studies Class I Criteria

- Cohort survey with prospective data collection
- Inclusion of a broad spectrum of persons at risk for developing the outcome
- Outcome measurement is objective or determined without knowledge of risk factor status
- Additional Class I criteria:
 - a. Exclusion/inclusion criteria clearly defined
 - b. Both the risk factor and the outcome measured in at least 80 percent of participants

Class II Criteria

- Cohort study with retrospective data collection or case-control study. Study meets Class I criteria a and b (see above)

- Inclusion of a broad spectrum of persons with and persons without both the risk factor and the outcome

- Presence of the risk factor and outcome are determined objectively or without the investigator's knowledge of both in each study participant

Class III Criteria

- Cohort or case-control study
- Narrow spectrum of persons with or without the disease
- Presence of the risk factor and outcome are determined objectively, without the investigator's knowledge of both in each study participant, or by different investigators

Class IV Criteria

If any study meets any one of the following criteria, the study must be classified as Class IV

- Persons at risk for the outcome are not included

- Patients with and patients without the risk factor are not included
- Measures of risk factor or outcomes are undefined or unaccepted

- Measures of association or statistical precision are either not presented or not calculable

Criteria for Rating Diagnostic Accuracy Studies Class I Criteria

- Cohort survey with prospective data collection

- Inclusion of a broad spectrum of persons suspected of having the disease

- Disease status determination is objective or made without knowledge of diagnostic test result

- Additional Class I criteria:

a. Exclusion/inclusion criteria clearly defined

b. Both the diagnostic test and disease status measured in at least 80 percent of participants

Class II Criteria

- Cohort study with retrospective data collection or case-control study. Study meets criteria a and b (see Class I)

- Includes a broad spectrum of persons with and persons without the disease

- The diagnostic test result and disease status are determined objectively or without knowledge of one another

Class III Criteria

- Cohort or case-control study

- Narrow spectrum of persons with or without the disease

- The diagnostic test result and disease status are determined objectively, without the investigator's knowledge of both in each participant, or by different investigators

Class IV Criteria

If any study meets any one of the following criteria, the study must be classified as Class IV

- Persons suspected of having the disease are not included
- Patients with and patients without the disease are not included

- Independent reference standard is undefined or unaccepted

- Measures of diagnostic accuracy or statistical precision are either not presented or not calculable

Criteria for Rating Population Screening Studies

Class I Criteria

- Study of a cohort of patients at risk for the outcome from a defined geographic area (i.e., population based)

- Outcome is objective
- Additional Class I criteria:
 - a. Exclusions and inclusion criteria clearly defined
 - b. At least 80 percent of patients undergo the screening of interest

Class II Criteria

- Non–population-based nonclinical cohort (e.g., mailing list, volunteer panel) or a general medical/neurology clinic/center without a specialized interest in the outcome. Study meets Class I criteria a and b (see above)
- Outcome is objective

Class III Criteria

- Referral cohort from a center with a potential specialized interest in the outcome

Class IV Criteria

If any study meets any one of the following criteria, the study must be classified as Class IV - Persons at risk for the outcome are not included

- Patients were not statistically sampled, or patients were specifically selected for inclusion by outcome

- Screening procedure or outcome measure is undefined or unaccepted

- Measures of frequency or statistical precision are either not presented or not calculable

Appendix E: Evidence Tables

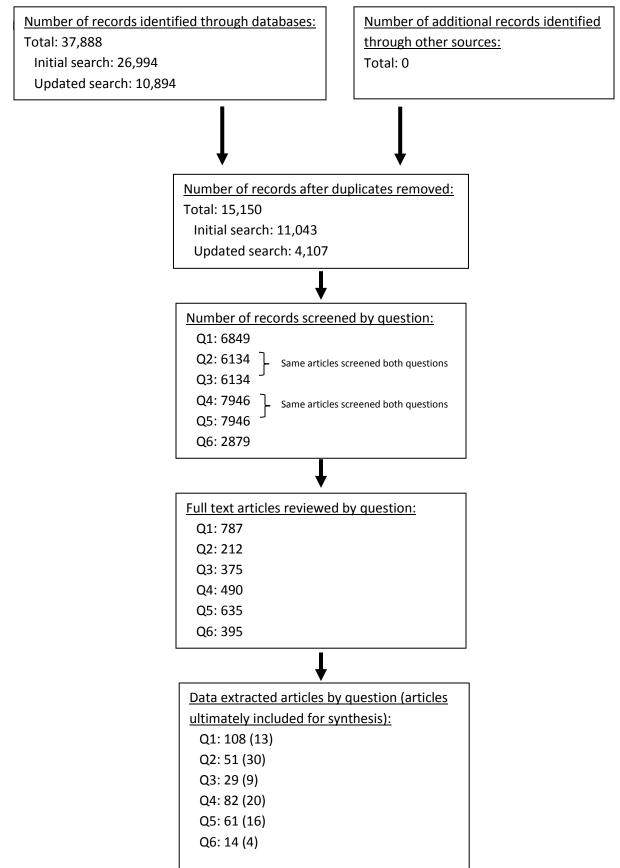
Appendix E Evidence Tables

Please locate Appendix E in a separate Excel file.

Appendix F: PRISMA Diagram

Appendix F PRISMA Diagram

Appendix. Flow of article selection process



Appendix G: Recommendation Process Methodology

Appendix G Recommendation Process Methodology

Constructing the recommendation and its rationale

Rationale for recommendation summarized in the Clinical Context includes three categories of premises

- □ Evidence-based conclusions based on the systematic review
- □ Stipulated axiomatic principles of care
- □ Strong related evidence not systematically reviewed
- \Box Inferences from the other premises

Actionable recommendations include the following mandatory elements

- \Box The patient population that is the subject of the recommendation
- $\hfill\square$ The person performing the action of the recommendation statement
- \Box The specific action to be performed
- \Box The expected outcome to be attained

Assigning a level of obligation

Modal modifiers used to indicate the final level of obligation (LOO)

- □ Level A: "Must"
- □ Level B: "Should"
- □ Level C: "May"
- □ Level U: No recommendation supported

□ Level R: The intervention should not be used outside of a research setting

LOO assigned by eliciting panel members' judgments regarding multiple domains, using a modified Delphi process. Goal is to attain consensus after a maximum of four rounds of voting. Consensus is defined by:

 $\square > 80\%$ agreement on dichotomous judgments

 $\square > 80\%$ agreement, within one point for ordinal judgments

□ If consensus is obtained, LOO assigned at the median. If not obtained, LOO assigned at the 10th percentile

Three steps used to assign final LOO

1. Initial LOO determined by the cogency of the deductive inference supporting the recommendation on the basis of ratings within four domains. Initial LOO anchored to lowest

LOO supported by any domain.

□ Confidence in evidence. LOO anchored to confidence in evidence determined by modified form of the Grading of Recommendations Assessment, Development and Evaluation process

 \Box Level A: High confidence

- \square Level B: Moderate confidence
- \Box Level C: Low confidence
- \Box Level U: Very low confidence

 \Box Soundness of inference assuming all premises are true. LOO anchored to proportion of panel members convinced of soundness of the inference

 \Box Level A: 100%

 \Box Level B: > 80% to < 100%

 \Box Level C: > 50% to < 80%

 \Box Level U or R: < 50%

□ Acceptance of axiomatic principles: LOO anchored to proportion of panel members who accept principles

□ Level A: 100%

 \Box Level B: > 80% to < 100%

 \Box Level C: > 50% to < 80%

 \Box Level U or R: < 50%

□ Belief that evidence cited from related conditions is strong: LOO anchored to proportion of panel members who believe the related evidence is strong

 \Box Level B: > 80% to 100% (recommendations dependent on inferences from non-systematically reviewed evidence cannot be anchored to a Level A LOO)

 \Box Level C: > 50% to < 80%

 \Box Level U or R: < 50%

2. LOO is modified mandatorily on the basis of the judged magnitude of benefit relative to harm expected to be derived from complying with the recommendation

 $\hfill\square$ Magnitude relative to harm rated on 4-point ordinal scale

□ Large benefit relative to harm: benefit judged large, harm judged none

□ Moderate benefit relative to harm: benefit judged large, harm judged minimal; or benefit judged moderate, harm judged none

□ Small benefit relative to harm: benefit judged large, harm judged moderate; or benefit judged moderate, harm judged minimal; or benefit judged small, harm judged none

□ Benefit to harm judged too close to call: benefit and harm judged to be substantially similar

 $\hfill\square$ Regardless of cogency of the recommendation the LOO can be no higher than that supported

by the rating of the magnitude of benefit relative to harm

 \Box Level A: large benefit relative to harm

□ Level B: moderate benefit relative to harm

 $\hfill\square$ Level C: small benefit relative to harm

 \Box Level U: too close to call

 \Box LOO can be increased by one grade if LOO corresponding to benefit relative to harm greater than LOO corresponding to the cogency of the recommendation

3. LOO optionally downgraded on the basis of the following domains

□ Importance of the outcome: critical, important, mildly important, not important

□ Expected variation in patient preferences: none, minimal, moderate, large

 $\hfill\square$ Financial burden relative to benefit expected: none, minimal, moderate, large

□ Availability of intervention: universal, usually, sometimes, limited

Appendix H: Clinical Contextual Profiles

Appendix H Clinical Contextual Profiles

Diagnosing Mild Traumatic Brain Injury Following Head Injury in Children (as compared to absence of brain injury or more severe injuries)

A. Risk Factor Identification and Imaging

• Risk Factors for Intracranial Injury and CT

Rationale: Up to 7.5% of children with mTBI will have intracranial injury (ICI).^{C1-C15} Identification of risk factors for ICI in children presenting with possible mTBI in the acute setting is important to the diagnosis of more severe forms of TBI, further directing observation and the possible need for emergent head CT. ICI further directs the prognosis of patients with mTBI (see Prognosis Recommendations). There is moderate evidence that several risk factors identify those patients with increased risk of ICI.^{C3,C5,C16-C18} These risk factors include age younger than 2 years, vomiting, loss of consciousness, severe mechanism of injury, severe or worsening headache, amnesia, non-frontal scalp hematoma, Glasgow Coma Score less than 15, and clinical suspicion for skull fracture. Notably, upon literature review, there is insufficient evidence to report seizures as a risk factor for ICI at this time. There is strong clinical evidence that use of clinical decision rules are effective in identifying children at low risk for ICI.^{C3,C5,C16,C18} The use of clinical decision rules may minimize the risk of failure to identify important ICI while avoiding unnecessary radiation exposure from head CT. Head CT is the preferred diagnostic tool in acute care settings to rapidly identify ICI. However, higher doses of radiation attributable to this type of imaging in children have been associated with an increase in the lifetime cancer mortality risk.^{C19-C22} Further, certain pediatric populations will require sedation in order to obtain adequate neuroimaging, increasing the overall risk related to imaging processes.^{C23} Families require clinical counseling regarding these risks in order to understand best practices for the clinical care of their child.

Pediatric head CT following mTBI in the acute care setting may possibly identify ICI in 7.5% (95% CI, 6.0-9.1%) of patients in the acute setting based on 16 Class III studies.^{C1-C15} However, this is likely an overestimate of the rate of findings due to bias in the selection of children for imaging. ICI included combinations of epidural hematoma, subdural hematoma, intracranial hemorrhage, subarachnoid hemorrhage, intraventricular hemorrhage, cerebral edema, and depressed skull fractures. Simple skull fractures were not included as ICI unless they occurred concomitantly with another intracranial finding.

In addition, head CT performed on children diagnosed initially with mTBI presenting to an acute care setting may possibly identify abnormalities resulting in clinically important outcomes in 1.9% (95% CI, 1.3-2.5) of patients based on 16 class III studies.^{C1-C3,C8-C10,C12,C14,C16,C24-C30} Clinically important outcomes included, but were not limited to, death, neurosurgical intervention, intubation for more than 24 hours, hospital admission of more than two nights for TBI, placement of intracranial pressure monitor, or other neurosurgical procedures. This is probably an overestimation of the rate of findings due to bias in the selection of children for imaging. Routine head CT in the acute care setting is possibly associated with neurosurgical intervention in 0.9% (95% CI, 0.5-1.2%) of patients based on 14 Class III studies.^{C1-C3,C8-C10,C12,C14,C16,C24-C30} Neurosurgical intervention was defined differently by the different authors, but included all craniotomies and occasionally intracranial pressure (ICP) monitors.

Following seemingly minor head injuries and mTBI, ICI resulting in the above stated clinically important outcomes is rare.^{C1-C3,C8-C10,C12,C16,C24-C31} Clinical evaluation of the child with possible mTBI includes balancing the likelihood of potentially devastating complications of a more severe injury against the risks associated with head CT (as well as possible concomitant sedation for imaging).

1. Healthcare providers *should not* routinely obtain head CT for diagnostic purposes in children with mTBI. (Level B)

Domain	Rating				Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 2	Benefit >> Harm 4	Benefit >>> Harm 11	Yes
Importance of outcomes	Not Important or 0	Mildly 0	Very 8	Critically Important 9	Yes
Variation in preferences	Large	Moderate	Modest 12	Minimal 3	Yes
Feasible	Rarely 0	Occasionally 0	Usually 3	Always 14	Yes
Cost relative to net benefit	Very Large 0	Large	Moderate 2	Small 14	Yes
Strength of recommendation	R/U	С	В	Α	

2. Healthcare providers *should* use validated clinical decision rules to identify children at low risk for intracranial injury, in whom head CT is not indicated, as well as children who may be at higher risk for clinically important ICI, and therefore may warrant head CT. Existing decision rules combine a variety of risk factors, including the following:

- Age < 2 years old
- Vomiting
- Loss of consciousness
- Severe mechanism of injury
- Severe or worsening headache
- Amnesia
- Nonfrontal scalp hematoma
- Glasgow Coma Score < 15
- Clinical suspicion for skull fracture (Level B)

Domain	Rating			•	Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 2	Benefit > Harm 4	Benefit >> Harm 6	Benefit >>> Harm 7	No
Importance of outcomes	Not Important or 1	Mildly 1	Very 8	Critically Important 9	Yes
Variation in preferences	Large	Moderate 5	Modest 6	Minimal 7	No
Feasible	Rarely 1	Occasionally 0	Usually 5	Always 13	Yes
Cost relative to net benefit	Very Large 0	Large 3	Moderate 6	Small 10	Yes
Strength of recommendation	R/U	С	В	Α	

3. For children diagnosed with mTBI, healthcare providers *should* discuss the risks of pediatric head CT in the context of risk factors for ICI with the patient and his/her family. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 3	Benefit >>> Harm 15	Yes
Importance of outcomes	Not Important or 0	Mildly 1	Very 7	Critically Important 10	Yes
Variation in preferences	Large O	Moderate 0	Modest 5	Minimal 13	Yes
Feasible	Rarely O	Occasionally 0	Usually 6	Always 12	Yes
Cost relative to net benefit	Very Large 0	Large O	Moderate 3	Small 15	Yes
Strength of recommendation	R/U	C	В	Α	

• Brain Magnetic Resonance Imaging (MRI)

Rationale: This review did not find any studies that met inclusion criteria addressing the use of brain MRI in the diagnosis of mTBI in children. MRI is more sensitive in identifying structural abnormalities than CT,^{C32,C33} and MRI avoids the use of ionizing radiation associated with CT. However, MRI more often requires sedation due to longer imaging acquisition times, and is more expensive than CT, though recently rapid, non-sedated MRI has been successfully employed in children with suspected acute TBI.^{C34} While current standard of care does not support the use of brain MRI in the diagnosis of pediatric mTBI for these reasons, specialized MRI imaging sequences such as functional MRI (fMRI), diffusion tensor imaging (DTI), Gradient recalled echo imaging (GREI), as well as others, have shown promise in research settings.^{C35} Studies on mixed groups, including children and adults, have found that early brain MRI can stratify outcome in mTBI.^{C36}

4. There is currently insufficient evidence to recommend the use of brain MRI in the diagnosis of mTBI in children. Healthcare providers *should not* routinely use MRI in the acute evaluation of suspected or diagnosed mTBI. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 2	Benefit > Harm 1	Benefit >> Harm 9	Benefit >>> Harm 7	Yes
Importance of outcomes	Not Important or 2	Mildly 1	Very 11	Critically Important 5	Yes
Variation in preferences	Large O	Moderate 1	Modest 7	Minimal 11	Yes
Feasible	Rarely O	Occasionally 2	Usually 7	Always 10	Yes
Cost relative to net benefit	Very Large 2	Large 1	Moderate 4	Small 12	Yes
Strength of recommendation	R/U	С	В	А	

• Single Photon Emission Computed Tomography (SPECT)

Rationale: SPECT may demonstrate brain hypoperfusion or hypometabolism following mTBI, but the qualitative nature of the images results in significant variability.^{C37-C39} A single Class I pediatric study looked at the presence of medial temporal hypometabolism on SPECT within 3 days of mTBI and reported a significant risk difference (RD: 0.732 [0.487–0.976]) for children with medial temporal hypometabolism having postconcussion syndrome.^{C37} However, confidence in this evidence was low due to significant issues with the generalizability of the data and this study did not use SPECT to diagnose mTBI specifically. SPECT is not commonly used in the clinical setting of TBI in children, may require patient sedation employing additional risks, requires intravenous access in the child with the injection of a radiopharmeceutical, and may be more expensive than head CT alone as it is often employed in conjunction with CT. This review did not find any studies that met our inclusion criteria addressing the use of SPECT in the diagnosis of mTBI in children.

5. Insufficient evidence currently exists to recommend the use of SPECT in the diagnosis of mTBI in children. Healthcare providers *should not* use SPECT in the acute evaluation of cases of suspected or diagnosed mTBI. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 1	Benefit > Harm 2	Benefit >> Harm 3	Benefit >>> Harm 12	Yes
Importance of outcomes	Not Important or 4	Mildly 3	Very 8	Critically Important 3	No
Variation in preferences	Large 1	Moderate 0	Modest 0	Minimal 17	Yes
Feasible	Rarely O	Occasionally 5	Usually 2	Always 11	No
Cost relative to net benefit	Very Large	Large 1	Moderate 2	Small 14	Yes
Strength of recommendation	R/U	С	В	Α	

• Skull X-ray

Rationale: This review identified two Class III studies evaluating the use of skull X-rays in children following minor head injury. The evidence identified a possible skull fracture in 7.14% (95% CI, 4.0-10.3%) of patients.^{C2} Because related literature reports that skull X-ray has a 63% sensitivity for diagnosing a single skull fracture in children, X-ray cannot detect intracranial injuries such as hemorrhage, shift from midline, or edema, and because X-ray employs radiation for imaging, it is not the best test to diagnose skull fracture with ICI following mTBI.^{C40} Clinical suspicion for skull fracture is a risk factor for other ICI following mTBI in children.^{C5,C16-C18} Neuroimaging modalities, such as head CT, better detect intracranial injuries, including skull fractures, making it the more appropriate diagnostic imaging choice when imaging is clinically indicated to assess for acute TBI.^{C40} In the instances where CT is not available, validated clinical decision rules are better than skull X-rays when screening patients with increased risk for ICI prior to determining the need for transfer to a facility with neuroimaging capabilities.^{C3, C5,C16,C18,C41}

6. Skull X-rays *should not* be used in the diagnosis of pediatric mTBI. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate	High 10	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 2	Benefit >> Harm 3	Benefit >>> Harm 13	Yes
Importance of outcomes	Not Important or 0	Mildly 2	Very 9	Critically Important 7	Yes
Variation in preferences	Large O	Moderate 0	Modest 3	Minimal 15	Yes
Feasible	Rarely 0	Occasionally 0	Usually 4	Always 14	Yes
Cost relative to net benefit	Very Large 0	Large O	Moderate 4	Small 14	Yes
Strength of recommendation	R/U	С	В	Α	

7. Skull X-rays *should not* be used in the screening for ICI. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate	High 10	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 5	Benefit >>> Harm 13	Yes
Importance of outcomes	Not Important or 0	Mildly 1	Very 8	Critically Important 9	Yes
Variation in preferences	Large O	Moderate 0	Modest 3	Minimal 15	Yes
Feasible	Rarely 0	Occasionally 0	Usually 3	Always 15	Yes
Cost relative to net benefit	Very Large 0	Large 1	Moderate 3	Small 14	Yes
Strength of recommendation	R/U	С	В	А	

B. Neuropsychological Tools:

• Symptom Scales

Rationale:

This review demonstrated with moderate confidence that the Graded Symptom Checklist (GSC) is useful in distinguishing children ages 6 years and older with mTBI from those without TBI within the first 2 days after injury.^{C42} This review demonstrated with moderate confidence that the Postconcussion Symptom Scale used in the ImPACT neurocognitive testing battery distinguishes high school athletes with mTBI from those without TBI within the first 4 days after injury.^{C43,C44} There are several other validated symptom scales that are reliable in the diagnosis of mTBI and have demonstrated validity at ages younger than high school.^{C45} The consequences of missing a diagnosis of mTBI include failure to recommend appropriate treatment and management that may contribute to prolongation of symptoms and increased risk of re-injury. Symptom inventories can be applied quickly and inexpensively.

8. Healthcare providers *should* use an age-appropriate, validated symptom rating scale as a component of the diagnostic evaluation in children presenting with acute mTBI. (Level B)

Domain		Rating	S		Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit	Benefit > Harm	Benefit >> Harm	Benefit >>> Harm 17	Yes
Importance of outcomes	Not Important or	Mildly	Very 7	Critically Important 11	Yes
Variation in preferences	Large	Moderate	Modest 4	Minimal 11	Yes
Feasible	Rarely C	Occasionally C	Usually 3	Always 15	Yes
Cost relative to net benefit	Very Large	Large	Moderate 3	Small 14	Yes
Strength of recommendation	R/U	С	В	А	

• Computerized Cognitive Testing

Rationale: This review identified two Class II studies meeting inclusion criteria evaluating computerized cognitive testing and the diagnosis of mTBI in children. ^{C43,C44} These studies only evaluated ImPACT cognitive testing and demonstrated that ImPACT cognitive testing probably distinguishes high school athletes with and without mTBI in the first 4 days post-injury and may add sensitivity to use of a symptom rating scale alone. ^{C43,C44} While these two studies only reviewed ImPACT testing, related evidence demonstrates that other validated computerized cognitive tests are also accurate in the diagnosis of mTBI in adults and children. ^{C46,C47} There is insufficient evidence to determine whether baseline testing in children better identifies mTBI post-injury as compared to post-injury scores alone, though evidence in adults currently suggests that baseline testing may be unnecessary in most cases. ^{C48,C49} The consequences of missing a diagnosis of mTBI include failure to recommend appropriate treatment and management that may contribute to prolongation of symptoms and increased risk of re-injury.

9. Healthcare providers *may* use validated, age-appropriate computerized cognitive testing in the acute period of injury as a component of the diagnosis of mTBI. (Level C)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm ≥ Benefit 1	Benefit > Harm 2	Benefit >> Harm 9	Benefit >>> Harm 6	Yes
Importance of outcomes	Not Important or 0	Mildly 3	Very 12	Critically Important 3	Yes
Variation in preferences	Large O	Moderate 5	Modest 7	Minimal 6	No
Feasible	Rarely O	Occasionally 12	Usually 6	Always O	Yes
Cost relative to net benefit	Very Large	Large 7	Moderate 8	Small 3	Yes
Strength of recommendation	R/U	С	В	Α	

• Standardized Assessment of Concussion

Rationale: This review demonstrated that cognitive screening using the Standardized Assessment of Concussion (SAC) was not accurate in distinguishing those children with mTBI from those without mTBI due to lack of statistical significance from a single Class III study.^{C42} Mixed literature in high school and collegiate athletes suggests that the SAC may detect effects of acute mTBI; however, these data were unable to be applied specifically to children in general or children specifically outside of the sports setting.^{C50} The consequences of missing a diagnosis of mTBI include failure to recommend appropriate treatment and management and may contribute to prolongation of symptoms and increased risk of re-injury.

10. There is insufficient evidence to support the use of the SAC in the diagnosis of children with mTBI, and this test *should not* be exclusively used to identify mTBI in children 6-18 years of age. (Level B)

Domain		Rating		•	Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 1	Benefit > Harm 1	Benefit >> Harm 5	Benefit >>> Harm 10	Yes
Importance of outcomes	Not Important or 0	Mildly 1	Very 14	Critically Important 2	Yes
Variation in preferences	Large O	Moderate 0	Modest 4	Minimal 13	Yes
Feasible	Rarely 0	Occasionally 3	Usually 3	Always 11	Yes
Cost relative to net benefit	Very Large 0	Large O	Moderate 3	Small 14	Yes
Strength of recommendation	R/U	С	B	Α	

C. Serum Markers

Rationale: Blood serum markers that may aid in the diagnosis of pediatric mTBI would be beneficial due to their low-risk profile. In two Class II studies, S100B was shown to be associated with a low sensitivity but high specificity in severe TBI patients, with no discrimination in mild to moderate TBI.^{C51,C52} Low sensitivity limits the usefulness of biomarkers, including S100B, for identifying or categorizing mTBI in children. In a Class II study, Tau was significantly different between pediatric mTBI patients with normal head CT, abnormal CT, and with non-TBI control subjects.^{C53} The effect size was small, and the control group was poorly defined, thus limiting the application of the findings. Serum potassium, sodium, glucose, and white blood cell count were examined in a single Class II study.^{C54} There were significant differences between these tests in children with mTBI versus the control groups.^{C54} Because the effect sizes were small and the groups were ill-defined, conclusions from this study are limited. A single Class II study explored the use of autoantibodies against glutamate receptors and oxide metabolites as a marker to discriminate between severe and mild pediatric TBI.^{C55} There was good discrimination between the two groups, but there was no uninjured control group, and further data will be needed before the test's value in pediatric mTBI can be determined. A single Class III study examined multiplex bead array biomarkers in a small number of infants with TBI compared to controls and found significant differences in a number of biomarkers.^{C56} The size of the study and the specialized population limit the applicability of the results. Related studies have demonstrated associations between neuronal ubiquitin C-terminal hydrolase-L1 and glial fibrillary acidic protein biomarker levels and ICI in adults following mild to moderate TBI.^{C57-C60} This review evaluated a single Class II study of 23 children and determined that there is insufficient evidence to determine whether ubiquitin C-terminal hydrolase-L1 and glial fibrillary acidic protein biomarker levels are useful tools in distinguishing children with or without mTBI.^{C61} Biomarker studies are not generally obtained in the clinical setting of mTBI and are costly. The time to report serum biomarker results represents a limitation in their clinical use in the acute setting. The risk of utilizing unvalidated biomarkers to clinically diagnose pediatric mTBI is false identification or under-identification of the injury.

11. There is insufficient evidence to currently recommend any of the studied biomarkers for the diagnosis of mTBI in children. Healthcare providers *should not* perform these tests outside of a research setting at this time for the diagnosis of children with mTBI. (Level R)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate	High 10	
Benefit relative to Harm	Harm <u>></u> Benefit 1	Benefit > Harm 0	Benefit >> Harm 7	Benefit >>> Harm 9	Yes
Importance of outcomes	Not Important or 2	Mildly 2	Very 9	Critically Important 4	No
Variation in preferences	Large O	Moderate 1	Modest	Minimal 15	Yes
Feasible	Rarely 1	Occasionally 5	Usually O	Always 11	No
Cost relative to net benefit	Very Large 0	Large	Moderate 2	Small 14	Yes
Strength of recommendation	R/U	С	В	А	

Prognosis of mTBI in Children

A. General HealthCare Provider Counseling of Prognosis

Rationale: Recovery from pediatric mTBI is variable,^{C62-C64} and no factors can individually predict recovery of symptoms or outcome.^{C65} Therefore, much of the variance in outcomes remains unaccounted for, even when multiple factors are considered. Evidence also suggests that the symptoms experienced by the majority of children with mTBI resolve within 1-3 months post injury.^{C62} A single Class III study reported that providing informational booklets to families that counseled on symptoms and coping strategies for children with mTBI resulted in improved patient outcomes at 3 months.^{C66} Although some effect sizes were statistically significant, there is insufficient evidence to determine the clinical significance of this specific intervention. However, related studies in children and adults with mTBI report direct patient benefits of counseling by healthcare providers.^{C67,C68} Public health campaigns have emphasized the importance of parent and family education in mTBI because health outcomes in general are optimized through patient health literacy and the resulting behavior modifications.^{C69-C71} Important aspects of healthcare provider counseling include education regarding warning signs of more serious injury (Centers of Disease Control and Prevention [CDC] Heads Up "12 Danger Signs"); review of expected symptoms, monitoring of postconcussive symptoms, prevention of further injury, cognitive and physical activity/rest, instructions regarding return to play/recreation and school (www.cdc.gov/HEADSUP), and clear clinician follow-up instructions.

12. Healthcare providers *should* counsel patients and families that the large majority (70-80%) of children with mTBI do not show significant difficulties that last more than 1-3 months post injury. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 5	Benefit >>> Harm 11	Yes
Importance of outcomes	Not Important or 0	Mildly O	Very 11	Critically Important 5	Yes
Variation in preferences	Large O	Moderate 0	Modest 2	Minimal 14	Yes
Feasible	Rarely 0	Occasionally 0	Usually 2	Always 14	Yes
Cost relative to net benefit	Very Large 0	Large 1	Moderate 1	Small 14	Yes
Strength of recommendation	R/U	С	В	А	

13. Healthcare providers *should* counsel patients and families that although some factors predict an increased or decreased risk for prolonged symptoms, each child's recovery from mTBI is unique and will follow its own trajectory. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 2	Benefit >>> Harm 14	Yes
Importance of outcomes	Not Important or 0	Mildly 1	Very 10	Critically Important 5	Yes
Variation in preferences	Large O	Moderate 0	Modest 1	Minimal 15	Yes
Feasible	Rarely 0	Occasionally 1	Usually 4	Always 11	Yes
Cost relative to net benefit	Very Large 0	Large 0	Moderate 3	Small 13	Yes
Strength of recommendation	R/U	С	В	А	

B. Prognosis Related to Premorbid Conditions

Rationale: Weak to strong evidence indicates that there is an increased risk of delayed recovery or prolonged symptoms associated with the following premorbid conditions in children with mTBI: premorbid history of concussion, ^{C72,C73} lower cognitive ability in pediatric mTBI with intracranial lesion, ^{C74} neurological or psychiatric disorder, ^{C75,C76} learning difficulties, ^{C63} increased preinjury symptoms, ^{C75-C77} and family and social stressors. ^{C77,C78} The assessment of premorbid history is likely to be most accurate when completed prior to injury (eg, as part of pre-participation athletic examinations) or as soon as possible post injury to avoid biases or inaccuracies in recall. ^{C79} Healthcare providers can more effectively counsel patients with mTBI when they have assessed the risks of premorbid conditions for prognosis.

14. Healthcare providers *should* assess the premorbid history of children either prior to injury as a part of pre-participation athletic examinations, or as soon as possible post injury in children with mTBI, to assist in determining prognosis. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 3	Benefit >>> Harm 13	Yes
Importance of outcomes	Not Important or 0	Mildly O	Very 11	Critically Important 5	Yes
Variation in preferences	Large O	Moderate 1	Modest 4	Minimal 11	Yes
Feasible	Rarely O	Occasionally 0	Usually 5	Always 11	Yes
Cost relative to net benefit	Very Large 0	Large 2	Moderate 2	Small 12	Yes
Strength of recommendation	R/U	С	В	Α	

15. Healthcare providers *should* counsel children and families completing pre-participation athletic examinations and children with mTBI as well as their families that recovery from mTBI might be delayed in those with: (Level B)

- Premorbid histories of mTBI
- Lower cognitive ability (for children with an intracranial lesion)
- Neurological or psychiatric disorder
- Learning difficulties
- Increased pre-injury symptoms (i.e., similar to those commonly referred to as "postconcussive")
- Family and social stressors

Domain		Rating				
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes	
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes	
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes	
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes	
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes	
Confidence in Inference	Very low	Low	Moderate 10	High		
Benefit relative to Harm	Harm <u>></u> Benefit 1	Benefit > Harm 0	Benefit >> Harm 5	Benefit >>> Harm 9	Yes	
Importance of outcomes	Not Important or 1	Mildly 0	Very 10	Critically Important 4	Yes	
Variation in preferences	Large O	Moderate 2	Modest 2	Minimal 11	Yes	
Feasible	Rarely O	Occasionally 0	Usually 5	Always 10	Yes	
Cost relative to net benefit	Very Large 0	Large 3	Moderate	Small 11	Yes	
Strength of recommendation	R/U	С	В	А		

C. Assessment of Cumulative Risk Factors and Prognosis

Rationale:

The outcomes of pediatric mTBI are known to be heterogeneous.^{C62-C64} Weak to strong evidence indicates that a variety of demographic and injury-related factors predict outcomes in pediatric mTBI, including age, gender, ethnicity, severity of injury, presence of ICI, and acute postconcussive symptoms. More specifically, symptoms may last longer for older children/adolescents, ^{C62,C80,C81} for children of Hispanic ethnicity as compared with White ethnicity, ^{C81} for children from lower socioeconomic status, ^{C78,C81} for children with more severe presentations of mTBI^{C64,C82,C83} (including those associated with intracranial injury), ^{C82,C84} and for children who report higher level of acute postconcussion symptoms. ^{C63,C73,C85} Additionally, headaches persist longer in girls.^{C80} However, no single factor is strongly predictive of outcome.^{C65}

A 2016 prospective multicenter cohort study of 3,063 children with mTBI seen in emergency department (ED) settings demonstrated that an empirically derived and cross-validated prediction rule combining multiple risk factors stratified the risk of persistent postconcussion symptoms at 28 days.^{C86} Healthcare providers can more effectively counsel patients with mTBI when they have assessed cumulative risk factors for prognosis.

16. Healthcare providers *should* screen for known risk factors for persistent symptoms in children with mTBI. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate	High 10	
Benefit relative to Harm	Harm <u>></u> Benefit 1	Benefit > Harm 0	Benefit >> Harm 6	Benefit >>> Harm 9	Yes
Importance of outcomes	Not Important or 1	Mildly 2	Very 11	Critically Important 2	Yes
Variation in preferences	Large	Moderate 1	Modest 4	Minimal 10	Yes
Feasible	Rarely 1	Occasionally 1	Usually 8	Always 6	Yes
Cost relative to net benefit	Very Large 0	Large O	Moderate 6	Small 10	Yes
Strength of recommendation	R/U	С	В	А	

17. Healthcare providers *may* use validated prediction rules, which combine information about multiple risk factors for persistent symptoms, to provide prognostic counseling to children with mTBI evaluated in ED settings. (Level C)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	No
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	No
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low 10	Moderate	High	
Benefit relative to Harm	Harm <u>></u> Benefit 1	Benefit > Harm 1	Benefit >> Harm 7	Benefit >>> Harm 7	Yes
Importance of outcomes	Not Important or 0	Mildly O	Very 15	Critically Important	Yes
Variation in preferences	Large 0	Moderate 1	Modest 4	Minimal 11	Yes
Feasible	Rarely 1	Occasionally 2	Usually 7	Always 6	Yes
Cost relative to net benefit	Very Large 0	Large 0	Moderate 7	Small 9	Yes
Strength of recommendation	R/U	C	В	А	

D. Assessment Tools and Prognosis

Rationale: No single assessment tool is strongly predictive of outcome in children with mTBI.^{C65} However, multiple tools have shown utility in the assessment of individual patients and their recovery from mTBI.^{C87-C89} Multiple tools are likely to be necessary because recovery trajectories can differ across specific domains of assessment, including symptom report, cognitive test performance, and balance.^{C90,C91} Symptom scales and cognitive testing (including measures of reaction time) have the strongest evidence in terms of their contribution to diagnosis (see Diagnosis Recommendations, as well as predicting and assessing recovery^{C92}). There is less evidence supporting balance testing as a predictor for prognosis in children, but it has shown utility in older adolescent athletes.^{C93} Healthcare providers can more effectively counsel patients with mTBI when they have assessed risk factors for outcome and recovery.

18. Healthcare providers *should* use a combination of tools to assess recovery in children with mTBI. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 5	Benefit >>> Harm 11	Yes
Importance of outcomes	Not Important or 0	Mildly 1	Very 12	Critically Important 3	Yes
Variation in preferences	Large 1	Moderate 0	Modest 4	Minimal 11	Yes
Feasible	Rarely 0	Occasionally 4	Usually 9	Always 3	Yes
Cost relative to net benefit	Very Large	Large 1	Moderate 10	Small 5	Yes
Strength of recommendation	R/U	С	В	А	

19. Healthcare providers *should* use validated symptom scales to assess recovery in children with mTBI. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 4	Benefit >>> Harm 13	Yes
Importance of outcomes	Not Important or 0	Mildly 0	Very 12	Critically Important 5	Yes
Variation in preferences	Large O	Moderate 0	Modest 3	Minimal 14	Yes
Feasible	Rarely O	Occasionally 1	Usually 10	Always 6	Yes
Cost relative to net benefit	Very Large 0	Large 0	Moderate 8	Small 9	Yes
Strength of recommendation	R/U	С	В	А	

20. Healthcare providers *may* use validated cognitive testing (including measures of reaction time) to assess recovery in children with mTBI. (Level C)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 1	Benefit >> Harm 7	Benefit >>> Harm 8	Yes
Importance of outcomes	Not Important or 1	Mildly 2	Very 12	Critically Important	Yes
Variation in preferences	Large 1	Moderate 2	Modest 6	Minimal 7	Yes
Feasible	Rarely 0	Occasionally 8	Usually 8	Always 0	Yes
Cost relative to net benefit	Very Large 0	Large 4	Moderate 11	Small 1	Yes
Strength of recommendation	R/U	С	В	А	

21. Healthcare providers *may* use balance testing to assess recovery in adolescent athletes with mTBI. (Level C)

Domain		Rating				
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes	
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes	
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes	
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes	
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes	
Confidence in Inference	Very low	Low	Moderate 10	High		
Benefit relative to Harm	Harm <u>></u> Benefit 1	Benefit > Harm 3	Benefit >> Harm 6	Benefit >>> Harm 6	No	
Importance of outcomes	Not Important or 2	Mildly 2	Very 11	Critically Important	Yes	
Variation in preferences	Large 1	Moderate 1	Modest 7	Minimal 7	Yes	
Feasible	Rarely O	Occasionally 8	Usually 5	Always 3	Yes	
Cost relative to net benefit	Very Large	Large 3	Moderate 9	Small 3	No	
Strength of recommendation	R/U	С	В	Α		

E. Interventions for mTBI With Poor Prognosis

Rationale: The symptoms experienced by the majority of children with mTBI resolve within 1-3 months post injury,^{C62} but some children are at risk for persistent symptoms and delayed recovery (ie, those who demonstrate certain premorbid characteristics and other risk factors; see rationales for recommendation items 5: Prognosis Related to Premorbid Conditions and 6: Assessment of Cumulative Risk Factors and Prognosis. Children with mTBI who are at high risk for persistent symptoms or delayed recovery are more likely to require intervention than children at low risk. Healthcare providers can more effectively counsel patients with mTBI when they have assessed risks for prognosis.

22. Healthcare providers should monitor children with mTBI who are determined to be at high risk for persistent symptoms based on premorbid history, demographics, and/or injury characteristics.

Domain		Rating		-	Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	No
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low 10	Moderate	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 3	Benefit >>> Harm 13	Yes
Importance of outcomes	Not Important or 0	Mildly 1	Very 9	Critically Important 6	Yes
Variation in preferences	Large O	Moderate 2	Modest 7	Minimal 7	Yes
Feasible	Rarely 0	Occasionally 0	Usually 10	Always 6	Yes
Cost relative to net benefit	Very Large 1	Large 1	Moderate 3	Small 11	Yes
Strength of recommendation	R/U	C	В	А	

23. For children with mTBI whose symptoms do not resolve as expected with standard care (i.e., within 4-6 weeks), healthcare providers *should* provide or refer for appropriate assessments and/or interventions (see Recommendations for Treatment and Management). (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 4	Benefit >>> Harm 12	Yes
Importance of outcomes	Not Important or 0	Mildly 1	Very 9	Critically Important 6	Yes
Variation in preferences	Large O	Moderate 2	Modest 9	Minimal 5	Yes
Feasible	Rarely 0	Occasionally 5	Usually 10	Always 1	Yes
Cost relative to net benefit	Very Large 0	Large 2	Moderate 11	Small 3	Yes
Strength of recommendation	R/U	С	В	Α	

Treatment and Management of mTBI in Children

A. Patient/Family Education and Reassurance

Rationale: There is no definitive evidence to support specific methods of patient and family education and reassurance following pediatric mTBI that are associated with clear improvements in patient health outcomes. Regardless, public health campaigns have emphasized the importance of parent and family education in mTBI because health outcomes in general are optimized through patient health literacy and the resulting behavior modifications.^{C69-C71}

Patient and family education and reassurance are key components of mTBI care initiatives and ED discharge instructions.^{C66-C68,C94} Standardized processes of evaluation and discharge instruction provide significant benefit to pediatric mTBI patient outcomes.^{C67} Important aspects of healthcare provider counseling include education regarding warning signs of more serious injury (CDC Heads Up "12 Danger Signs"), review of expected symptoms, monitoring of postconcussive symptoms, prevention of further injury, cognitive and physical activity/rest, instructions regarding return to play/recreation and school (<u>www.cdc.gov/HEADSUP</u>), and clear clinician follow-up instructions. 24. In providing education and reassurance to the family, the healthcare provider should include the following information:

- a. Warning signs of more serious injury
- b. Description of injury and expected course of symptoms and recovery
- c. Instructions on how to monitor postconcussive symptoms
- d. Prevention of further injury
- e. Management of cognitive and physical activity/rest
- f. Instructions regarding return to play/recreation and school
- g. Clear clinician follow-up instructions

(Level B)

Domain		Rating					
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes		
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes		
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes		
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes		
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes		
Confidence in Inference	Very low	Low	Moderate	High 10			
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 1	Benefit >> Harm 3	Benefit >>> Harm 15	Yes		
Importance of outcomes	Not Important or 0	Mildly 0	Very 7	Critically Important 12	Yes		
Variation in preferences	Large O	Moderate 0	Modest 4	Minimal 15	Yes		
Feasible	Rarely 0	Occasionally 0	Usually 8	Always 11	Yes		
Cost relative to net benefit	Very Large 0	Large O	Moderate 6	Small 13	Yes		
Strength of recommendation	R/U	С	B	А			

B. Cognitive/Physical Rest and Aerobic Activity

Rationale:

Historically, "rest" has been a foundation in the treatment of acute mTBI.^{C95,C96} However, scientific evidence supporting its timing. duration, and efficacy is limited.^{C97} A clear definition of rest is not provided in the literature and interpretations range from full bedrest to a reduced level of activity.^{C97} Related evidence suggests that rest or reduction in cognitive/physical activity is beneficial immediately following mTBI and, for those who are slow to recover, may help accelerate recovery.^{C98-C100} The rationale for rest is based on the attempted reduction of neurometabolic demand in the context of post-injury symptoms.^{C101} Exertional, early post-injury activity increases the metabolic demand of impaired neural cells, and may result in increased symptom manifestation. ^{C101} Animal literature suggests that too much physical activity early post-injury may be counterproductive to recovery, but that later physical activity may accelerate recovery.^{C102} The post-injury period is a posited temporal window of vulnerability for re-injury,^{C103,C104} because the re-injury threshold is lower during recovery and the symptom burden may be greater.^{C105,C107,C126} Re-injury during this window of vulnerability has been associated with catastrophic injury in rare pediatric cases via unclear mechanisms.^{C108-C110} Studies in children and adults with prolonged symptoms beyond 4 weeks demonstrate that physical exercise managed below symptom exacerbation reduced postconcussive symptoms in active rehabilitation models.^{C111-C114} Animal studies demonstrate that physical exercise facilitates key neurobiological factors (eg, increased brain derived neurotrophic factor, positive changes in neurotransmitters), which may support recovery from brain injury.^{C102} However, these studies found that physical activity that was initiated early post injury had worse outcomes and may compromise the positive effects of exercise. ^{C102} Related evidence demonstrates the deleterious effects of significant inactivity as well as significant health benefits of a regular program of exercise in other medical conditions in humans.^{C97,C115-C119}

The optimal timing to initiate an aerobic program following pediatric mTBI has not been established and only limited studies have applied this treatment to patients with symptoms persisting past 4 weeks.^{C111-C113} No evidence exists to provide guidance on the exact timing of activity onset, dosing (how much), and the progression of activity post injury for a given symptom profile.^{C97} Related evidence suggests that early rest within the first 3 days of the injury may be beneficial,^{C95,C120} but that inactivity beyond this time period for most children may worsen their symptom report.^{C121} As a result, the gradual resumption of noncontact activity that does not exacerbate symptoms has replaced the prescription of full rest until asymptomatic.

25. Healthcare providers *should* counsel patients to observe more restrictive physical and cognitive activity during the first several days following mTBI in children.

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 1	Benefit > Harm 2	Benefit >> Harm 3	Benefit >>> Harm 12	Yes
Importance of outcomes	Not Important or 0	Mildly O	Very 10	Critically Important 8	Yes
Variation in preferences	Large O	Moderate 2	Modest 10	Minimal 6	Yes
Feasible	Rarely 1	Occasionally 0	Usually 5	Always 12	Yes
Cost relative to net benefit	Very Large	Large O	Moderate 6	Small 12	Yes
Strength of recommendation	R/U	С	В	А	

26. Following these first several days, healthcare providers *should* counsel patients and families to resume a gradual schedule of activity that does not exacerbate symptoms, with close monitoring of symptom expression (number, severity). (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 1	Benefit > Harm 1	Benefit >> Harm 5	Benefit >>> Harm 11	Yes
Importance of outcomes	Not Important or 0	Mildly O	Very 16	Critically Important 2	Yes
Variation in preferences	Large O	Moderate 1	Modest 10	Minimal 7	Yes
Feasible	Rarely O	Occasionally 0	Usually 5	Always 13	Yes
Cost relative to net benefit	Very Large 0	Large O	Moderate 7	Small 11	Yes
Strength of recommendation	R/U	С	В	А	

27. Following the successful resumption of a gradual schedule of activity (see 26), healthcare providers *should* offer an active rehabilitation program of progressive reintroduction of noncontact aerobic activity that does not exacerbate symptoms, with close monitoring of symptom expression (number, severity). (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate	High 10	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 3	Benefit >>> Harm 13	Yes
Importance of outcomes	Not Important or 0	Mildly 0	Very 11	Critically Important 5	Yes
Variation in preferences	Large O	Moderate	Modest 9	Minimal 6	Yes
Feasible	Rarely 0	Occasionally 0	Usually 11	Always 5	Yes
Cost relative to net benefit	Very Large 0	Large 0	Moderate 7	Small 9	Yes
Strength of recommendation	R/U	С	В	Α	

28. Healthcare providers *should* counsel patients to return to full activity when they return to premorbid performance if they have remained symptom free at rest and with increasing levels of physical exertion (see 25-27). (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 5	Benefit >>> Harm 11	Yes
Importance of outcomes	Not Important or 0	Mildly 0	Very 10	Critically Important 6	Yes
Variation in preferences	Large O	Moderate 0	Modest 4	Minimal 12	Yes
Feasible	Rarely 0	Occasionally 0	Usually 7	Always 9	Yes
Cost relative to net benefit	Very Large 0	Large 3	Moderate 5	Small 8	Yes
Strength of recommendation	R/U	С	В	А	

C. Psychosocial/Emotional Support

Rationale:

Social support exerts a powerful influence on a variety of health issues, including chronic diseases.^{C122,C123} Social support is positively associated with healthy behaviors and adherence, improved overall quality of life, and reduced deleterious effects of stress on health.^{C122-C124} Conversely, lack of social support (perceived or actual) increases morbidity and a greater likelihood of hospital admissions or re-hospitalizations.^{C124} Social isolation has been identified as an independent risk factor for all-cause mortality.^{C125} It is reasonable to assume that the role of social support in any human interaction is beneficial.

Social support takes many forms, including emotional, with the provision of empathy, love, trust, and caring; instrumental, involving the provision of tangible aid/services directly assisting persons in need; informational, with its provision of suggestions, advice, or information used to address problems; and appraisal, which provides information useful for self-evaluation such as constructive feedback and positive affirmations. Social support has proven useful in the recovery of persons with TBI, particularly those with cognitive deficits.^{C126,C127} Limited research with those who have suffered an mTBI demonstrates similar benefits.^{C128,C129} Direct, ancillary, and extrapolated evidence is strongly suggestive of the utility of social support in the management of mTBI.

29. Healthcare providers *may* assess the extent and types of social support (ie, emotional, informational, instrumental, appraisal) in children with mTBI and emphasize social support as a key element in the education of caregivers and educators. (Level C)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 9	Benefit >>> Harm 9	Yes
Importance of outcomes	Not Important or 1	Mildly 6	Very 9	Critically Important 2	Yes
Variation in preferences	Large 1	Moderate 2	Modest 8	Minimal 7	Yes
Feasible	Rarely O	Occasionally 3	Usually 8	Always 7	Yes
Cost relative to net benefit	Very Large 0	Large O	Moderate 11	Small 7	Yes
Strength of recommendation	R/U	С	В	А	

D. Return to School

Rationale:

Return to school following mTBI must be carefully planned given the adverse effects (eg, headaches and fatigue interfering with learning, greater problems concentrating on schoolwork, difficulty taking notes) that can affect learning and performance.^{C130,C131} Limited evidence exists to guide the timing or progression of return to activity in relation to academic activities.^{C121} A subset of children with mTBI is at higher risk for more severe or prolonged postconcussive symptoms (see Prognosis Recommendations), which may interfere substantially with resumption of academic activities.^{C62,C65,C73,C78,C80-C85,C132}

Consensus-based recommendations for returning to school after mTBI attempt to minimize cognitive and physical overexertion.^{C96} Return to school protocols affirm the need for continued collaboration among medical, school, and family systems to gradually adjust interventions and return the child to full participation without significant worsening of symptoms. ^{C96,C130,C133-C136} The protocols target the student's symptoms as the focus of intervention, linking specific accommodations in efforts to limit symptom expression. Because postconcussive symptoms resolve at different rates in different children after mTBI, individualization of return to school programming is necessary.

To protect their legal right for an appropriate education, children with mTBI who have greater symptom burden and prolonged recoveries may require formal educational planning incorporating protections under federal statutes.^{C137,C138} These protections are provided to qualifying students under Section 504 of the Rehabilitation Act and the Traumatic Brain Injury guidelines under the Individuals with Disabilities Education Act.^{C137}

30. To assist children returning to school following mTBI, medical and school-based teams *should* counsel the student and family regarding the process of gradually increasing the duration and intensity of academic activities as tolerated, with the goal of increasing participation without significantly exacerbating symptoms. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 1	Benefit > Harm 1	Benefit >> Harm 1	Benefit >>> Harm 13	Yes
Importance of outcomes	Not Important or 0	Mildly 0	Very 6	Critically Important 10	Yes
Variation in preferences	Large O	Moderate 1	Modest 5	Minimal 10	Yes
Feasible	Rarely O	Occasionally 0	Usually 7	Always 9	Yes
Cost relative to net benefit	Very Large	Large 3	Moderate 4	Small 9	Yes
Strength of recommendation	R/U	С	В	А	

31. Return to school protocols *should* be customized based on the severity of postconcussion symptoms in children with mTBI as determined jointly by medical and school-based teams. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 3	Benefit >>> Harm 12	Yes
Importance of outcomes	Not Important or 0	Mildly 0	Very 4	Critically Important 11	Yes
Variation in preferences	Large O	Moderate 2	Modest 6	Minimal 7	Yes
Feasible	Rarely O	Occasionally 0	Usually 7	Always 8	Yes
Cost relative to net benefit	Very Large 0	Large 2	Moderate 4	Small 9	Yes
Strength of recommendation	R/U	С	В	А	

32. For any student with prolonged symptoms that interfere with academic performance, school-based teams *should* assess the educational needs of that student and determine the student's need for additional educational supports, including those described under pertinent federal statutes (e.g., Section 504, IDEA).^{C137} (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate	High 10	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 6	Benefit >>> Harm 10	Yes
Importance of outcomes	Not Important or 0	Mildly O	Very 11	Critically Important 5	Yes
Variation in preferences	Large O	Moderate 4	Modest 7	Minimal 5	No
Feasible	Rarely O	Occasionally 2	Usually 13	Always 1	Yes
Cost relative to net benefit	Very Large 0	Large O	Moderate 16	Small 0	Yes
Strength of recommendation	R/U	С	В	А	

33. Postconcussion symptoms and academic progress in school *should* be monitored collaboratively by the student, family, healthcare provider, and school teams, who jointly determine what modifications or accommodations are needed to maintain an academic workload without significantly exacerbating symptoms. (Level B)

Domain		Rating		•	Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate	High 10	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 7	Benefit >>> Harm 11	Yes
Importance of outcomes	Not Important or 0	Mildly 0	Very 11	Critically Important 7	Yes
Variation in preferences	Large 1	Moderate 3	Modest 8	Minimal 6	No
Feasible	Rarely O	Occasionally 4	Usually 11	Always 3	Yes
Cost relative to net benefit	Very Large 0	Large 0	Moderate 17	Small 1	Yes
Strength of recommendation	R/U	С	B	А	

34. The provision of educational supports *should* be monitored and adjusted on an ongoing basis by the school-based team until the student's academic performance has returned to preinjury levels. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 3	Benefit >>> Harm 7	Yes
Importance of outcomes	Not Important or 0	Mildly 0	Very 6	Critically Important 4	Yes
Variation in preferences	Large O	Moderate 2	Modest 5	Minimal 3	Yes
Feasible	Rarely 0	Occasionally 0	Usually 9	Always 1	Yes
Cost relative to net benefit	Very Large 0	Large 2	Moderate 5	Small 3	Yes
Strength of recommendation	R/U	С	В	А	

35. For students who demonstrate prolonged symptoms and academic difficulties despite an active treatment approach, healthcare providers *should* refer the child for a formal evaluation by a specialist in pediatric mTBI. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 1	Benefit >> Harm 4	Benefit >>> Harm 13	Yes
Importance of outcomes	Not Important or 0	Mildly 1	Very 8	Critically Important 9	Yes
Variation in preferences	Large 2	Moderate 3	Modest 9	Minimal 4	No
Feasible	Rarely O	Occasionally 7	Usually 11	Always O	Yes
Cost relative to net benefit	Very Large 0	Large 1	Moderate 17	Small 0	Yes
Strength of recommendation	R/U	С	В	А	

Symptom/problem-specific treatment/management:

A. Post-traumatic Headache Management

Rationale:

Headache is the most common symptom of pediatric mTBI in the acute setting. Children presenting with a headache, including worsening or severe headache, following mTBI are probably at moderate risk for ICI reflected by risk difference of 1.86% (CI 0.12%-3.59%) from three Class I studies and one Class II study.^{C3,C5,C16,C18} This evidence supports that the risk of not identifying more severe forms of TBI presenting with a progressive, severe headache in a child with or without other risk factors outweighs the risk of ionizing radiation.

There is no evidence supporting a relationship between headache severity on postconcussion symptom assessment in the ED and neurocognitive function during the acute period of recovery. Additionally, insufficient data exists to determine a relationship between early postconcussion symptoms, including headache, and later neurocognitive outcomes or behavioral function among children with mTBI.^{C132,C139} There is no evidence to support a relationship between age and headache following mTBI. However, among children presenting to an ED following mTBI, those injured above the age

of 6 years are probably at a 5-10% increased risk of remaining symptomatic (including headache) for 12 months or longer as compared to children 6 years of age or younger.^{C62} There was a relationship between gender and headache with girls reporting recurrent and persisting headache after 3 months compared with boys.^{C80,C140}

Painful headache in children requires intervention. Non-narcotic analgesics such as ibuprofen and acetaminophen are often effective in treating headaches in children and opioids are not generally recommended as therapy for headaches.^{C141,C142,C143} While common clinical practice supports use of non-narcotic analgesics and avoidance of exertional activities for the treatment of headache secondary to pediatric mTBI, there is no evidence to support the success of such interventions in the acute setting or their impact on headache recurrence in the subacute or chronic setting.^{C144} Non-narcotic analgesic overuse carries important risks of toxicity and rebound headache.^{C145}

A single Class I study evaluated 3% hypertonic saline as a treatment for headache in children following acute mTBI presenting to an ED.^{C146} In children with mTBI and headache, hypertonic saline possibly decreases pain with headache immediately following the intervention, with a Z score of 1.53 (CI 0.85-2.200), though the study failed to show an effect at 3 days post intervention. ^{C146} However, there are limitations in the study's sample size and inability to show sustained effect on pain improvement, as well as lack of related evidence and concerns for risks versus benefits of intravenous medication administration in children that preclude further recommendation at this time.

36. Healthcare providers in the ED *should* clinically observe and consider obtaining a head CT in children presenting with severe and worsening headache following mTBI to evaluate for ICI requiring further management in accordance with validated clinical decision making rules. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate	High 10	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 5	Benefit >>> Harm 11	Yes
Importance of outcomes	Not Important or 0	Mildly 0	Very 5	Critically Important 11	Yes
Variation in preferences	Large O	Moderate 0	Modest 5	Minimal 11	Yes
Feasible	Rarely 0	Occasionally 0	Usually 6	Always 10	Yes
Cost relative to net benefit	Very Large 0	Large O	Moderate 12	Small 4	Yes
Strength of recommendation	R/U	С	В	Α	

37. Children undergoing observation periods for headache with acutely worsening symptoms *should* undergo emergent neuroimaging. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate	High 10	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 9	Benefit >>> Harm 10	Yes
Importance of outcomes	Not Important or 0	Mildly 1	Very 3	Critically Important 15	Yes
Variation in preferences	Large O	Moderate 1	Modest 3	Minimal 15	Yes
Feasible	Rarely 0	Occasionally 1	Usually 11	Always 7	Yes
Cost relative to net benefit	Very Large 0	Large 0	Moderate 16	Small 3	Yes
Strength of recommendation	R/U	С	В	А	

38. Healthcare providers and caregivers *should* offer non-narcotic analgesia (i.e., ibuprofen or acetaminophen) to children with painful headache following acute mTBI, but also provide counseling to the family regarding the risks of analgesic overuse, including rebound headache. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 2	Benefit >> Harm 5	Benefit >>> Harm 9	Yes
Importance of outcomes	Not Important or 0	Mildly 1	Very 13	Critically Important 2	Yes
Variation in preferences	Large O	Moderate 0	Modest 6	Minimal 10	Yes
Feasible	Rarely O	Occasionally 0	Usually 8	Always 8	Yes
Cost relative to net benefit	Very Large 0	Large 0	Moderate 8	Small 8	Yes
Strength of recommendation	R/U	С	В	Α	

39. There is insufficient evidence to currently recommend the administration of 3% hypertonic saline as a treatment for acute headache following mTBI in children. Healthcare providers should not administer this medication to children with mTBI for treatment of symptoms outside of a research setting at this time. (Level R)

Domain		Rating		•	Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 1	Benefit > Harm 2	Benefit >> Harm 8	Benefit >>> Harm 6	Yes
Importance of outcomes	Not Important or 1	Mildly 5	Very 9	Critically Important 2	Yes
Variation in preferences	Large O	Moderate 2	Modest	Minimal 14	Yes
Feasible	Rarely O	Occasionally 1	Usually 4	Always 12	Yes
Cost relative to net benefit	Very Large 0	Large O	Moderate 6	Small 11	Yes
Strength of recommendation	R/U	С	В	Α	

40. Chronic headache following mTBI is likely to be multifactorial, and, therefore, healthcare providers *should* refer children with chronic headache after mTBI for multidisciplinary evaluation and treatment, with consideration of analgesic overuse as a contributory factor. (Level B)

Domain		Rating		•	Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate	High 10	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 6	Benefit >>> Harm 11	Yes
Importance of outcomes	Not Important or 0	Mildly 0	Very 16	Critically Important	Yes
Variation in preferences	Large O	Moderate 2	Modest 10	Minimal 5	Yes
Feasible	Rarely O	Occasionally 7	Usually 8	Always 2	Yes
Cost relative to net benefit	Very Large 0	Large 2	Moderate 14	Small 1	Yes
Strength of recommendation	R/U	С	В	А	

B. Vestibulo-ocular Motor Dysfunction

Rationale:

Dizziness is a pervasive and debilitating symptom reported following mTBI in children. A single Class II study reported that vestibular and oculomotor dysfunction may contribute to the diagnosis of mTBI and longer symptom duration.^{C147} Gaining interest as an area for screening, as well as treatment, limited evidence suggests that early vestibular physical therapy may be of benefit for patients presenting with subjective complaints (symptom of dizziness) or objective physical examination findings.^{C148-C151} The optimal time to initiate vestibular physical therapy, the specific order and intensity of exercises, and longitudinal outcomes have yet to be studied.

41. Healthcare providers *may* refer children with subjective or objective evidence of persistent vestibulo-ocular motor dysfunction following mTBI to a program of vestibular rehabilitation. (Level C)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 1	Benefit > Harm 0	Benefit >> Harm 10	Benefit >>> Harm 6	Yes
Importance of outcomes	Not Important or 2	Mildly 2	Very 12	Critically Important	Yes
Variation in preferences	Large 1	Moderate 4	Modest 7	Minimal 5	No
Feasible	Rarely 1	Occasionally 12	Usually 4	Always 0	Yes
Cost relative to net benefit	Very Large 0	Large 4	Moderate 13	Small 0	Yes
Strength of recommendation	R/U	C	В	А	

C. Sleep

Rationale:

Sleep disturbance is a common problem following TBI, and may lead to ongoing disability.^{C152-C157} Sleep disturbance may impede the recovery process given the critical need for the availability of appropriate energy to support neurobiological recovery and daily functioning, and worsen symptoms. Related evidence in adolescents with mTBI reported poorer sleep quality and demonstrated significantly shorter actigraphic-measured sleep duration, poorer sleep efficiency, and more wake time after onset of sleep, compared with healthy adolescents (all, p<0.05).^{C158} Receiving adequate sleep has been shown to facilitate health,^{C159} and when not appropriate, adversely affects medical conditions, including TBI.^{C154,C160,C161} Practices that promote healthy sleep include (1) age-appropriate and consistent bedtimes and wake times, (2) establishing bedtime routines, (3) maintaining appropriate lighting and sound control in the bedroom or before bed, (6) positivity, (7) independence when falling asleep, and (8) meeting the child's emotional needs during the day.^{C162-C164} Related evidence from studies in adults with TBI discuss potential treatments including cognitive behavior therapy supporting lifestyle modifications, pharmacologic treatments with modafinil and melatonin, and light therapy.^{C152,C165,C166} While limited evidence supports a recommendation for sleep hygiene specifically in children with mTBI, related evidence in adults with TBI indicates benefits, suggesting that the maintenance of appropriate sleep and management of disrupted sleep may be a critical target of treatment for the child with mTBI.

42. Healthcare providers *should* provide guidance on proper sleep hygiene methods to facilitate recovery from pediatric mTBI. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 1	Benefit >>> Harm 16	Yes
Importance of outcomes	Not Important or 0	Mildly O	Very 14	Critically Important 3	Yes
Variation in preferences	Large O	Moderate 0	Modest 7	Minimal 10	Yes
Feasible	Rarely 0	Occasionally 0	Usually 6	Always 11	Yes
Cost relative to net benefit	Very Large 0	Large O	Moderate 2	Small 15	Yes
Strength of recommendation	R/U	С	В	А	

43. If sleep problems emerge or continue despite appropriate sleep hygiene measures, healthcare providers *may* refer children with mTBI to a sleep disorder specialist for further assessment. (Level C)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 1	Benefit >> Harm 8	Benefit >>> Harm 8	Yes
Importance of outcomes	Not Important or 0	Mildly 2	Very 14	Critically Important	Yes
Variation in preferences	Large 1	Moderate 4	Modest 7	Minimal 5	No
Feasible	Rarely O	Occasionally 10	Usually 7	Always O	Yes
Cost relative to net benefit	Very Large 0	Large 2	Moderate 15	Small 0	Yes
Strength of recommendation	R/U	C	В	А	

D. Cognitive Impairment

Rationale:

Cognitive impairment occurs following mTBI and includes the following areas: attention, memory and learning, response speed, and aspects of executive functions.^{C43,C44,C167,C168} Dysfunctional attention or memory may result in significant problems with learning in school or social interactions.^{C131,C134,Error!} Reference source not found. Current literature is insufficient to determine whether cognitive impairment is directly related to the pathology of the brain injury (i.e., impaired neurotransmission) or secondary effects of the plethora of other symptoms (e.g., ongoing headache pain, fatigue/low energy, low frustration tolerance), which, as a result of their distracting effects, may produce a disruption in cognitive processing. Understanding the etiology of the cognitive dysfunction is important to direct treatment/management appropriately. For example, primary cognitive impairment suggests the need to apply direct therapeutic interventions to the affected cognitive process (e.g., teaching memory or attentional strategies). In contrast, if the cognitive dysfunction is secondary to another symptom (e.g., headache pain), then the primary therapeutic intervention would be directed toward the reduction of the headaches. Neuropsychological evaluations can assist in determining etiology of cognitive impairment and directing treatment.²

44. Healthcare providers *should* attempt to determine the etiology of cognitive dysfunction, within the context of other mTBI symptoms. (Level B)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate 10	High	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 1	Benefit >> Harm 4	Benefit >>> Harm 11	Yes
Importance of outcomes	Not Important or 0	Mildly 1	Very 10	Critically Important 5	Yes
Variation in preferences	Large 1	Moderate 2	Modest 5	Minimal 8	Yes
Feasible	Rarely 1	Occasionally 3	Usually 9	Always 3	No
Cost relative to net benefit	Very Large 0	Large 3	Moderate 8	Small 5	Yes
Strength of recommendation	R/U	С	В	А	

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate	High 10	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 0	Benefit >> Harm 4	Benefit >>> Harm 13	Yes
Importance of outcomes	Not Important or 0	Mildly 0	Very 13	Critically Important 4	Yes
Variation in preferences	Large 0	Moderate 3	Modest 11	Minimal 3	Yes
Feasible	Rarely 0	Occasionally 4	Usually 12	Always 1	Yes
Cost relative to net benefit	Very Large 0	Large O	Moderate 14	Small 3	Yes
Strength of recommendation	R/U	С	В	А	

45. Healthcare providers *should* recommend treatment for cognitive dysfunction that reflects its presumed etiology. (Level B)

46. Healthcare providers *may* refer children with persisting complaints related to cognitive function for a formal neuropsychological evaluation to assist in determining etiology and recommending targeted treatment. (Level C)

Domain		Rating			Consensus
Rationale is logical	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Evidence statements accurate	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Axioms true	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Related evidence strong & applicable	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Internal inferences logically follow	< 50%	50% to < 80%	80% to < 100%	100%	Yes
Confidence in Inference	Very low	Low	Moderate	High 10	
Benefit relative to Harm	Harm <u>></u> Benefit 0	Benefit > Harm 1	Benefit >> Harm 6	Benefit >>> Harm 10	Yes
Importance of outcomes	Not Important or 0	Mildly O	Very 11	Critically Important 6	Yes
Variation in preferences	Large O	Moderate 4	Modest 10	Minimal 3	Yes
Feasible	Rarely 0	Occasionally 7	Usually 10	Always O	Yes
Cost relative to net benefit	Very Large 0	Large 1	Moderate 16	Small 0	Yes
Strength of recommendation	R/U	С	В	А	

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