Why is this taking so long: Understanding persisting symptoms after pediatric concussion

Julie Wilson, MD, FAAP, CAQSM

Co-Director, Concussion Program
Sports Medicine Center, Children's Hospital Colorado, Aurora, CO
Departments of Orthopedics and Pediatrics, University of Colorado School of Medicine





Disclosures

Grant funding for concussion research





DEPARTMENT OF THE AIR FORCE SOTH MEDICAL WING (AETC) JOINT BASE SAN ANTONIO - LACKLAND TEXAS



Team physician stipend







Objectives

1

Define persisting symptoms after concussion

2

Understand contributing factors to persisting symptoms

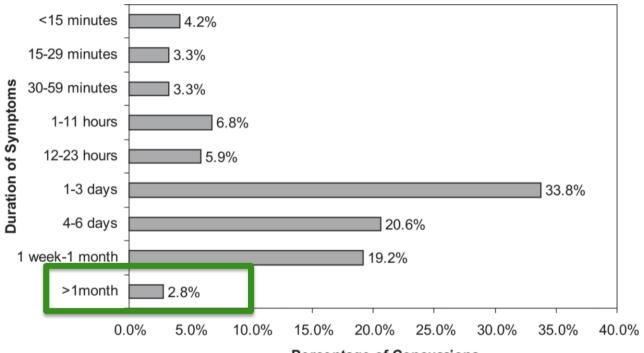
3

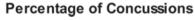
Highlight role of multi-disciplinary team in managing persisting symptoms









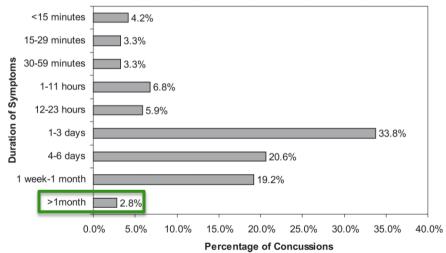






Persisting Symptoms

Used for symptoms that persist >4 weeks



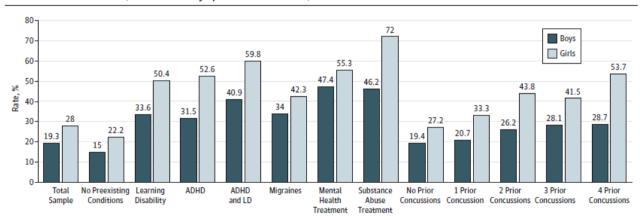




Persisting symptoms

May be pre-existing, concussion-related, or both

Figure. Rates of International Classification of Diseases, 10th Revision (ICD-10), Postconcussional Syndrome Classification in High School Athletes With No Recent Concussion (Mild or Greater Symptoms in Each Domain)



Original Investigation

Factors Associated With Concussion-like Symptom Reporting in High School Athletes

Grant L. Iverson, PhD; Noah D. Silverberg, PhD; Rebekah Mannix, MD, MPH; Bruce A. Maxwell, PhD; Joseph E. Atkins, PhD; Ross Zafonte, DO; Paul D, Berkner, DO



Clinical Risk Score for Persistent Postconcussion Symptoms Among Children With Acute Concussion in the ED

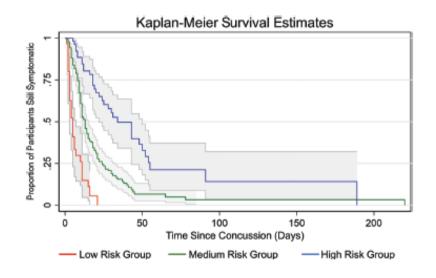
Zemek R, et al. JAMA 2016.

Category		Risk Points
Age group (yrs)	5-7 8-12 13-<18	0 1 2
Sex	Male Female	0 2
Prior concussion / symptom duration	None/Sx duration <1 wk Sx duration ≥1 wk	0 1
Diagnosed migraine history	No Yes	0 1
Answering questions slowly	No Yes	0 1
BESS tandem stance errors	0-3 ≥4 or unable to perform	0 1
Headache	No Yes	0 1
Sensitivity to noise	No Yes	0 1
Fatigue	No Yes	0 2

Utility of a Clinical Prediction Tool for Persisting Postconcussive Symptoms in a Multicenter Sample of Youth Athletes With Concussion

The Sport Concussion Outcomes in Pediatrics (SCOPE) Study

Miller SM, et al. AJSM 2023.







Factors Affecting Recovery Trajectories in Pediatric Female Concussion

Natasha Desai, MD, CAQSM,* Douglas J. Wiebe, PhD,† Daniel J. Corwin, MD,‡ Julia E. Lockyer, MS,§ Matthew F. Grady, MD, CAQSM,§¶ and Christina L. Master, MD, CAQSM§¶

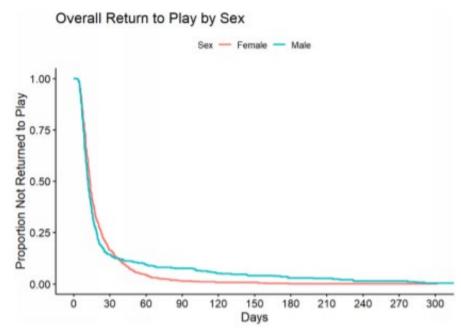
- Female patients with concussion presented later to specialty care evaluation than male patients (15 vs 9 days, p=0.018) and took longer to recover.
- Sex-based recovery differences <u>disappeared</u> when controlling for time to presentation to specialty care





Differences in sport-related concussion for female and male athletes in comparable collegiate sports: a study from the NCAA-DoD Concussion Assessment, Research and Education (CARE) Consortium

Christina L Master ¹ ,^{1,2,3} Barry P Katz, ⁴ Kristy B Arbogast, ^{2,3} Michael A McCrea, ⁵ Thomas W McAllister, ⁶ Paul F Pasquina, ⁷ Michelle Lapradd, ⁴ Wenxian Zhou, ⁴ Steven P Broglio ¹ , ⁸ CARE Consortium Investigators







Paediatric post-concussive symptoms: symptom clusters and clinical phenotypes

Todd W Lyons , ^{1,2} Rebekah Mannix, ^{1,2} Ken Tang, ³ Keith Owen Yeates, ^{4,5} Gurinder Sangha, ^{6,7} Emma CM Burns, ^{8,9} Darcy Beer, ¹⁰ Alexander S Dubrovsky, ^{11,12} Isabelle Gagnon, ¹³ Jocelyn Gravel, ¹⁴ Stephen B Freedman, ¹⁵ William Craig, ¹⁶ Kathy Boutis, ¹⁷ Martin H Osmond, ^{3,18} Gerard Gioia, ¹⁹ Roger Zemek , ^{3,18} The Pediatric Emergency Research Canada (PERC) 5P Concussion Team²⁰

- Fatigue #1 persisting symptom
- Headache #2 persisting symptom
- Physical symptoms occurred in two distinct clusters: vestibular-ocular and headache.
- <u>Emotional and cognitive symptoms occurred together</u> more frequently and with higher symptom severity than physical symptoms.
- <u>Fatigue</u> was more strongly associated with <u>cognitive and emotional symptoms</u> than physical symptoms.







Headache





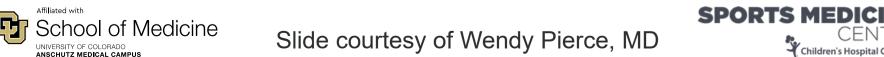
Post traumatic headache* causes

*Defined as HA onset ≤7 days of injury

- Axonal injury
- Altered cerebral metabolism or hemodynamics
- Neuroinflammation
- Underlying genetic predisposition
- Medication overuse

- Patient's expectations of developing headache after head injury
- Sleep disturbances
- Mood disturbances
- Psychosocial and other stressors

Cephalagia 38(1) 2018





Common headache phenotypes after concussion











Headache Treatment – What's the Evidence?

Current evidence for pharmacological management of pediatric concussion: a systematic review

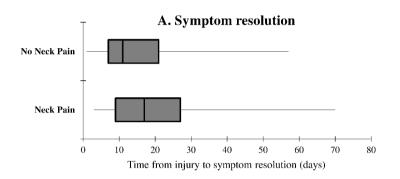
Damla Hanalioglu^{1,2} · Sahin Hanalioglu^{3,4} · Jorge I. Arango¹ · P. David Adelson⁵

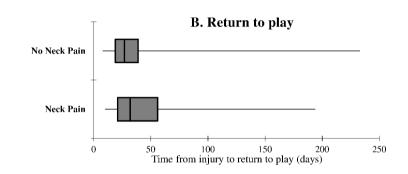
Drug	#	Risk of bias in studies	Symptom (effective for)
Acetaminophen	1	<u> </u>	Acute headache
Amantadine	1	0	Cognitive symptoms
Amitriptyline	4	••••	Persistent headache
Flunarizine	1		Inconclusive
Gabapentin	1		Persistent headache
Hypertonic saline	1		Acute headache
Ibuprofen	1	<u> </u>	Acute headache
Ketorolac	1		Acute headache
Lysdexamphetamine	1		Attention deficit
Magnesium	1		Acute symptom burden
Melatonin	3		Only for sleep and depression
Metoclopramide	2	<u> </u>	Likely no benefit for headache
Nortriptilline	1		Persistent headache (short-term)
Prochlorperazine	1		Inconclusive
Ondansetron	3		Likely no benefit for PCS
Topiramate	2		Likely no benefit for PTH

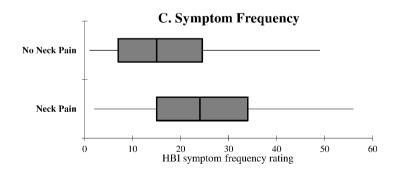




Influence of Cervical Involvement







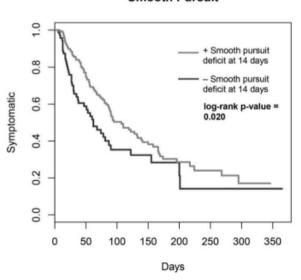




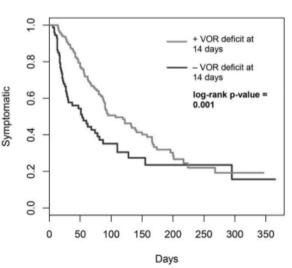
Visual and Vestibular Function and Concussion Recovery



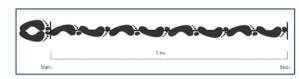
Smooth Pursuit



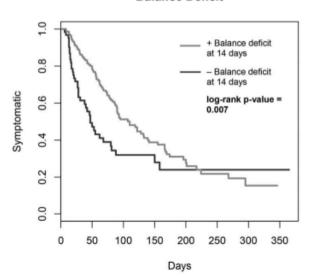
VOR



Master CL, et al. CJSM, 2018.



Balance Deficit



Cervicovestibular rehabilitation in sport-related concussion: a randomised controlled trial

Kathryn J Schneider, ^{1,2} Willem H Meeuwisse, ^{1,3} Alberto Nettel-Aguirre. ^{2,3,4} Karen Barlow, ² Lara Boyd, ⁵ Jian Kang, ¹ Carolyn A Emery ^{1,2,3}

Treatment group 3.91x (95% CI 1.34, 11.34) more likely to be medically cleared by 8 wks

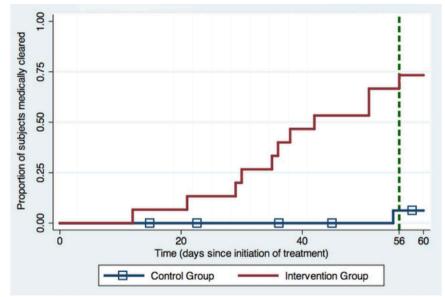


Figure 1 Proportion of patients medically cleared over time.





Fatigue





Sleep outcomes in pediatric mild traumatic brain injury: a systematic review and meta-analysis of prevalence and contributing factors

Suzana Djukic, Natalie Lynette Phillips (i), and Suncica Lah (i)

BRAIN INJURY 2022, VOL. 36, NOS. 12–14, 1289–1322

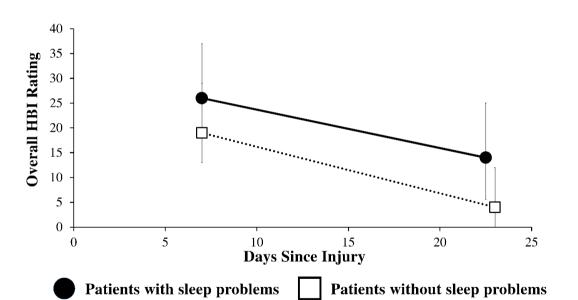
- Post-injury sleep disturbance is common
 - 51% within 1 week
 - 40% 1-4 weeks
 - 9% 1-3 months
 - 21% > 3 months

- Sleeping more than usual most common within 1 week of injury
- Sleeping <u>less than usual</u> common 1-4 weeks post-injury





Sleep Problems after Concussion



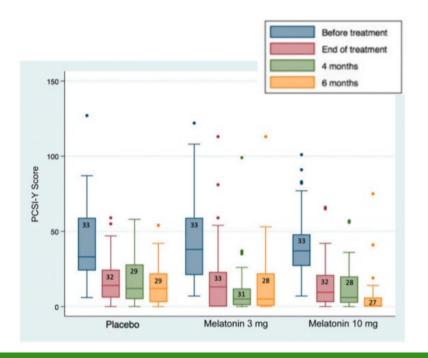
Patients with sleep problems took 7 days longer to recover





Efficacy of Melatonin in Children With Postconcussive Symptoms: A Randomized Clinical Trial

Karen M. Barlow, MBChB, *** Brian L. Brooks, PhD, *** Michael J. Esser, MD, *** Adam Kirton, MD, *** Angelo Mikrogianakis, MD, **
Roger L. Zemek, MD, ** Frank P. MacMaster, PhD, *** Alberto Nettel-Aguirre, PhD, *** Keith Owen Yeates, PhD, *** Valerie Kirk, MD, **
James S. Hutchison, MD, ** Susan Crawford, MSc, ** Brenda Turley, BA, ** Candice Gameron, BA, ** Michael D. Hill, MD, **
Tina Samuel, MBChB, ** Jeffrey Buchhalter, MD, ** Lawrence Richer, MD, ** Robert Platt, PhD, ** Roslyn Boyd, PhD, **
Deborah Dewey, PhD**



Journal of Neurotrauma, VOL. 38, NO. 8 | Original Articles



Efficacy of Melatonin for Sleep Disturbance in Children with Persistent Post-Concussion Symptoms: Secondary Analysis of a Randomized Controlled Trial

Karen Maria Barlow 🖂 Valerie Kirk, Brian Brooks, Michael Joachim Esser, Keith Owen Yeates, Roger Zemek, Adam Kirton,
Angelo Mikrogianakis, Frank MacMaster, Alberto Nettel-Aguirre, James Hutchison, Brenda Turley, Candice Cameron, Michael Hill,
Roslyn Boyd, and Deborah Dewey

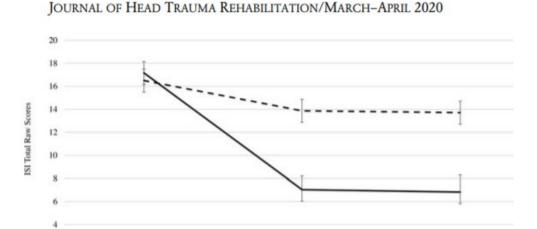
Melatonin

- \$\prec\$ sleep-related problems (3 mg)
- \depressive symptoms (3 mg)
- ↑ sleep duration (3, 10 mg)



A Pilot Randomized Controlled Trial of Cognitive-Behavioral Therapy for Insomnia in Adolescents With Persistent Postconcussion Symptoms

Tomfohr-Madsen, Lianne PhD; Madsen, Joshua W. PhD; Bonneville, Dominique BA; Virani, Shane MSc; Plourc Vickie PhD; Barlow, Karen M. MBChB, MSc, MRCPCH (UK); Yeates, Keith Owen PhD; Brooks, Brian L. PhD



Post-treatment

Baseline

-Cognitive Behavioural Therapy for Insomnia





One-month follow up

- - Treatment as Usual

Cognitive and Emotional Symptoms

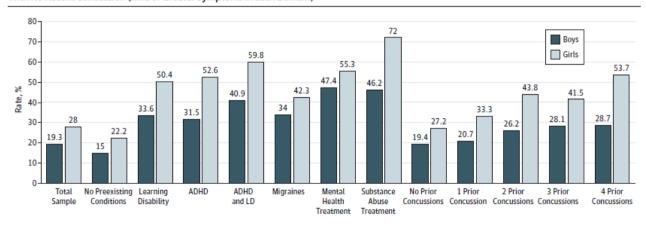




Persisting symptoms

May be pre-existing, concussion-related, or both

Figure. Rates of International Classification of Diseases, 10th Revision (ICD-10), Postconcussional Syndrome Classification in High School Athletes With No Recent Concussion (Mild or Greater Symptoms in Each Domain)

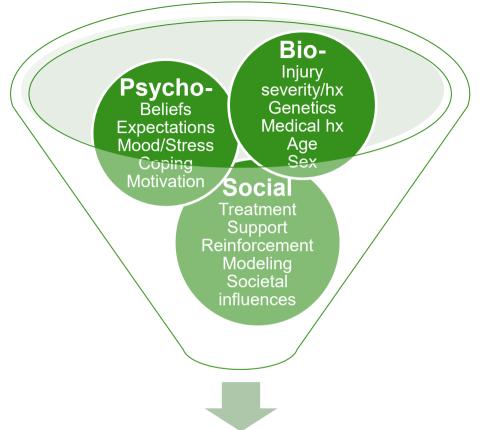


Original Investigation

Factors Associated With Concussion-like Symptom Reporting in High School Athletes

Grant L. Iverson, PhD; Noah D. Silverberg, PhD; Rebekah Mannix, MD, MPH; Bruce A. Maxwell, PhD; Joseph E. Atkins, PhD; Ross Zafonte, DO; Paul D. Berkner, DO







Persisting Symptoms

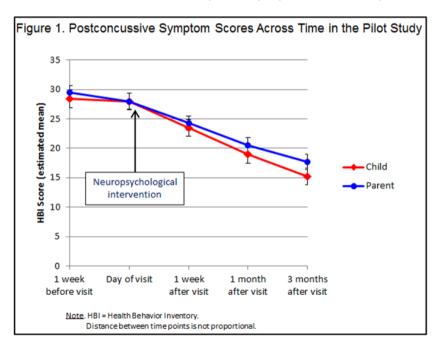
Slide courtesy of Mike Kirkwood, PhD





A Pilot Study Investigating Neuropsychological Consultation as an Intervention for Persistent Postconcussive Symptoms in a Pediatric Sample

Michael W. Kirkwood, PhD1, Robin L. Peterson, PhD1, Amy K. Connery, PsyD1, David A. Baker, PsyD1, and Jeri Forster, PhD1,2





Slide courtesy of Mike Kirkwood, PhD





Brief cognitive behavioral intervention for children and adolescents with persistent post-concussive symptoms: A pilot study

Kelly A. McNally^{a,b}, Kristina E. Patrick^a, Jacob E. LaFleur^c, Jana B. Dykstra^a, Kerry Monahan^a and Kristen R. Hoskinson^{b,c}

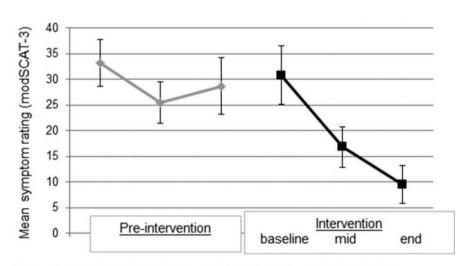


Figure 1. Mean self-reported concussion symptom ratings pre- and post-intervention.

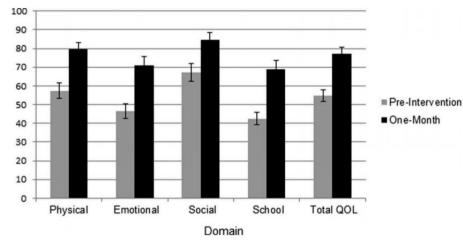


Figure 2. Quality of life mean scores at pre-intervention and one month.





Key Points for Improving Outcomes



Identify and address factors that may impact recovery trajectory (5P risk score)



Optimize
sleep,
introduce
aerobic
exercise and
cervicovestibular
rehab when
appropriate



Consider referral to specialty concussion program





References

- Asken BM, Bauer RM, Guskiewicz KM, McCrea MA, Schmidt JD, Giza CC, Snyder AR, Houck ZM, Kontos AP, McAllister TW, Broglio SP, Clugston JR; CARE Consortium Investigators, Anderson S, Bazarian J, Brooks A, Buckley T, Chrisman S, Collins M, DiFiori J, Duma S, Dykhuizen B, Eckner JT, Feigenbaum L, Hoy A, Kelly L, Langford TD, Lintner L, McGinty G, Mihalik J, Miles C, Ortega J, Port N, Putukian M, Rowson S, Svoboda S. Immediate Removal From Activity After Sport-Related Concussion Is Associated With Shorter Clinical Recovery and Less Severe Symptoms in Collegiate Student-Athletes. Am J Sports Med. 2018 May:46(6):1465-1474.
- Barlow KM, Brooks BL, Esser MJ, Kirton A, Mikrogianakis A, Zemek RL, MacMaster FP, Nettel-Aguirre A, Yeates KO, Kirk V, Hutchison JS, Crawford S, Turley B, Cameron C, Hill MD, Samuel T, Buchhalter J, Richer L, Platt R, Boyd R, Dewey D. Efficacy of Melatonin in Children With Postconcussive Symptoms: A Randomized Clinical Trial. Pediatrics. 2020 Apr;145(4):e20192812.
- Barlow KM, Kirk V, Brooks B, Esser MJ, Yeates KO, Zemek R, Kirton A, Mikrogianakis A, MacMaster F, Nettel-Aguirre A, Hutchison J, Turley B, Cameron C, Hill M, Boyd R, Dewey D. Efficacy of Melatonin for Sleep Disturbance in Children with Persistent Post-Concussion Symptoms: Secondary Analysis of a Randomized Controlled Trial. J Neurotrauma. 2021 Apr 15;38(8):950-959.
- Charek DB, Elbin RJ, Sufrinko A, Schatz P, D'Amico NR, Collins MW, Kontos AP. Preliminary Evidence of a Dose-Response for Continuing to Play on Recovery Time After Concussion. J Head Trauma Rehabil. 2020 Mar/Apr;35(2):85-91.
- Desai N, Wiebe DJ, Corwin DJ, Lockyer JE, Grady MF, Master CL. Factors Affecting Recovery Trajectories in Pediatric Female Concussion. Clin J Sport Med. 2019 Sep;29(5):361-367.
- Djukic S, Phillips NL, Lah S. Sleep outcomes in pediatric mild traumatic brain injury: a systematic review and meta-analysis of prevalence and contributing factors. Brain Inj. 2022 Dec 6;36(12-14):1289-1322. doi: 10.1080/02699052.2022.2140198. Epub 2022 Nov 22. PMID: 36413091.
- Elbin RJ, Sufrinko A, Schatz P, French J, Henry L, Burkhart S, Collins MW, Kontos AP. Removal From Play After Concussion and Recovery Time. Pediatrics. 2016 Sep;138(3):e20160910.
- Hanalioglu D, Hanalioglu S, Arango JI, Adelson PD. Current evidence for pharmacological management of pediatric concussion: a systematic review. Childs Nerv Syst. 2023 Jul;39(7):1831-1849. doi: 10.1007/s00381-023-05960-x. Epub 2023 May 19. PMID: 37208486.
- Headache Classification Committee of the International Headache Society (IHS) The International Classification of Headache Disorders, 3rd edition. Cephalalgia. 2018 Jan;38(1):1-211. doi: 10.1177/0333102417738202. PMID: 29368949.
- Howell DR, O'Brien MJ, Fraser J, Meehan WP 3rd. Continuing Play, Symptom Severity, and Symptom Duration After Concussion in Youth Athletes. Clin J Sport Med. 2020 Mar;30 Suppl 1:S42-S46. doi: 10.1097/JSM.000000000000570. PMID: 32132476.
- Iverson GL, Silverberg ND, Mannix R, Maxwell BA, Atkins JE, Zafonte R, Berkner PD. Factors Associated With Concussion-like Symptom Reporting in High School Athletes. JAMA Pediatr. 2015 Dec;169(12):1132-40. doi: 10.1001/jamapediatrics.2015.2374. PMID: 26457403; PMCID: PMC5333772.
- Kirkwood MW, Peterson RL, Connery AK, Baker DA, Forster J. A Pilot Study Investigating Neuropsychological Consultation as an Intervention for Persistent Postconcussive Symptoms in a Pediatric Sample. J Pediatr. 2016 Feb;169:244-9.e1. doi: 10.1016/j.jpeds.2015.10.014. Epub 2015 Nov 2. PMID: 26541427.





References

- Lyons TW, Mannix R, Tang K, Yeates KO, Sangha G, Burns EC, Beer D, Dubrovsky AS, Gagnon I, Gravel J, Freedman SB, Craig W, Boutis K, Osmond MH, Gioia G, Zemek R; Pediatric Emergency Research Canada (PERC) 5P Concussion Team. Paediatric post-concussive symptoms: symptom clusters and clinical phenotypes. Br J Sports Med. 2022 Jul:56(14):785-791. doi: 10.1136/bjsports-2021-105193. Epub 2022 Mar 10. PMID: 35273038.
- Master CL, Katz BP, Arbogast KB, McCrea MA, McAllister TW, Pasquina PF, Lapradd M, Zhou W, Broglio SP; CARE Consortium Investigators. Differences in sport-related concussion for female and male athletes in comparable collegiate sports: a study from the NCAA-DoD Concussion Assessment, Research and Education (CARE) Consortium. Br J Sports Med. 2021 Dec;55(24):1387-1394. doi: 10.1136/bjsports-2020-103316. Epub 2020 Dec 21. PMID: 33355211.
- Master CL, Master SR, Wiebe DJ, Storey EP, Lockyer JE, Podolak OE, Grady MF. Vision and Vestibular System Dysfunction Predicts Prolonged Concussion Recovery in Children. Clin J Sport Med. 2018 Mar;28(2):139-145. doi: 10.1097/JSM.000000000000507. PMID: 29064869.
- McNally KA, Patrick KE, LaFleur JE, Dykstra JB, Monahan K, Hoskinson KR. Brief cognitive behavioral intervention for children and adolescents with persistent post-concussive symptoms: A pilot study. Child Neuropsychol. 2018 Apr;24(3):396-412. doi: 10.1080/09297049.2017.1280143. Epub 2017 Jan 26. PMID: 28125932.
- Meehan WP 3rd, d'Hemecourt P, Collins CL, Comstock RD. Assessment and management of sport-related concussions in United States high schools. Am J Sports Med. 2011 Nov;39(11):2304-10. doi: 10.1177/0363546511423503. Epub 2011 Oct 3. PMID: 21969181; PMCID: PMC3359792.
- Miller SM, Valovich McLeod TC, Zaslow TL, Wilson JC, Master CL, Snedden TR, Halstead ME, Grady MF, Fazekas ML, Santana JA, Coel RA, Howell DR. Utility of a Clinical Prediction Tool for Persisting Postconcussive Symptoms in a Multicenter Sample of Youth Athletes With Concussion: The Sport Concussion Outcomes in Pediatrics (SCOPE) Study. Am J Sports Med. 2023 Nov;51(13):3546-3553. doi: 10.1177/03635465231201610. Epub 2023 Oct 4. PMID: 37794642.
- Provance AJ, Howell DR, Potter MN, Wilson PE, D'Lauro AM, Wilson JC. Presence of Neck or Shoulder Pain Following Sport-Related Concussion Negatively Influences Recovery. J Child Neurol. 2020 Jun;35(7):456-462. doi: 10.1177/0883073820909046. Epub 2020 Mar 19. PMID: 32192408.
- Schneider KJ, Meeuwisse WH, Nettel-Aguirre A, Barlow K, Boyd L, Kang J, Emery CA. Cervicovestibular rehabilitation in sport-related concussion: a randomised controlled trial. Br J Sports Med. 2014 Sep;48(17):1294-8. doi: 10.1136/bjsports-2013-093267. Epub 2014 May 22. PMID: 24855132.
- Tomfohr-Madsen L, Madsen JW, Bonneville D, Virani S, Plourde V, Barlow KM, Yeates KO, Brooks BL. A Pilot Randomized Controlled Trial of Cognitive-Behavioral Therapy for Insomnia in Adolescents With Persistent Postconcussion Symptoms. J Head Trauma Rehabil. 2020 Mar/Apr;35(2):E103-E112. doi: 10.1097/HTR.0000000000000504. PMID: 31246882.
- Zemek R, Barrowman N, Freedman SB, Gravel J, Gagnon Í, McGahern C, Aglipay M, Sangha G, Boutis K, Beer Ď, Craig W, Burns E, Farion KJ, Mikrogianakis A, Barlow K, Dubrovsky AS, Meeuwisse W, Gioia G, Meehan WP 3rd, Beauchamp MH, Kamil Y, Grool AM, Hoshizaki B, Anderson P, Brooks BL, Yeates KO, Vassilyadi M, Klassen T, Keightley M, Richer L, DeMatteo C, Osmond MH; Pediatric Emergency Research Canada (PERC) Concussion Team. Clinical Risk Score for Persistent Postconcussion Symptoms Among Children With Acute Concussion in the ED. JAMA. 2016 Mar 8;315(10):1014-25. doi: 10.1001/jama.2016.1203. Erratum in: JAMA. 2016 Jun 21;315(23):2624. PMID: 26954410.





Panel Discussion



Ann Lantagne, PhD

Rehab Psychology



Robin Peterson, PhD

Neuropsychology



Nicki Remington, DPT

Physical Therapy



Julie Wilson, MD

Sports Medicine





JN – 17 yo F soccer player

- Hit in the face by soccer ball kicked at close range immediate onset HA, dizziness, nausea, light sensitivity
- Initial visit to CHCO ~3 weeks from injury
 - Feels 65% recovered
 - Continued HA, significant dizziness (motion sensitivity, positional, associated nausea), sleep disturbance
 - Symptoms impacting school tolerance but trying to attend full-time





Prior Concussion History

- 3 prior concussions
 - 2016 and 2017 soccer-related injuries, full recovery took 2-4 wks
 - 2018 basketball-related injury recovery took several months, took 2 years off sports after this injury
 - Played HS soccer in 2021 and 2022 without injury
 - Reports intermittent HA (every 2-3 weeks) since 2018 injury. Maternal relatives with migraines.





Physical Exam

- HR ↑ (58 to 91) with orthostatic VS
- Symptom provocation with all VOMS components (dizziness)
- Positive Romberg (exaggerated loss of balance)
- Slow single (32 sec) and dual-task (54 sec) tandem gait





Initial plan

- Improve hydration and nutrition (Zofran, increase fluid/electrolyte intake)
- Headache management plan
- PT for vestibular rehab
- Sub-symptom threshold exercise
- School support plan



