

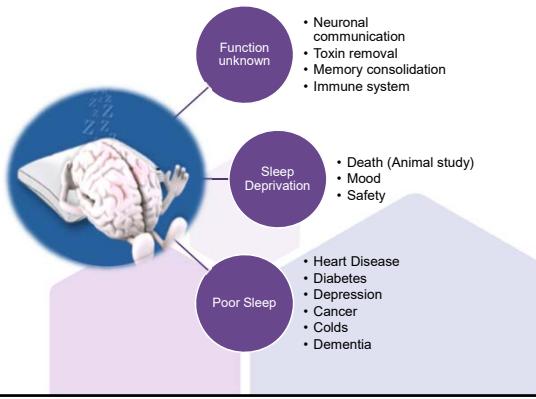
Detox the Brain with Deep Sleep

Shelley A. Tischkau, PhD

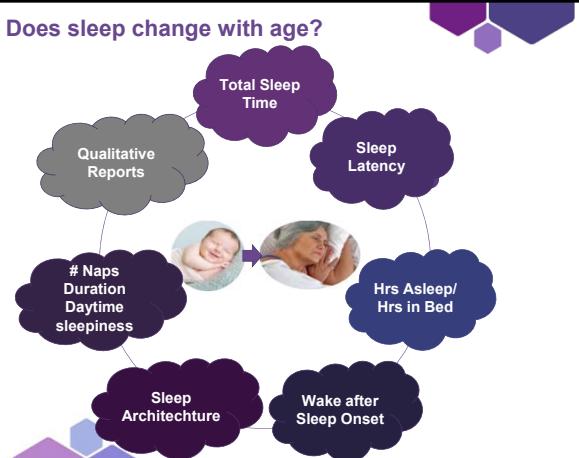
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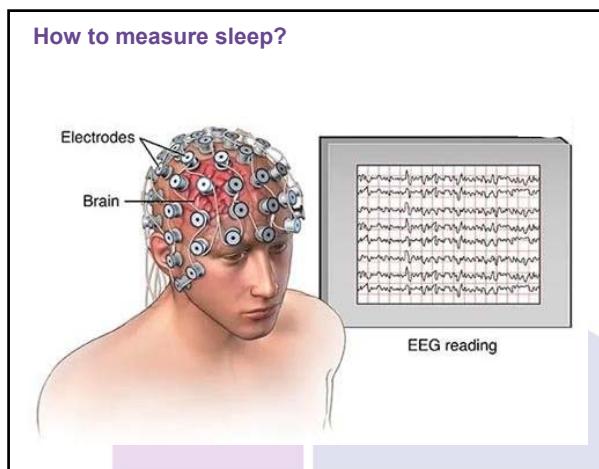


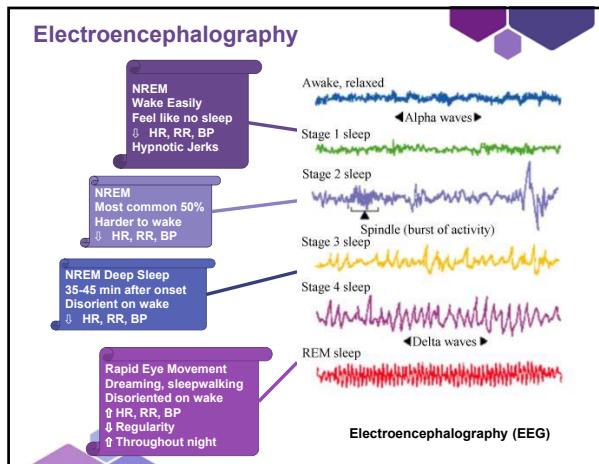
Why Sleep?

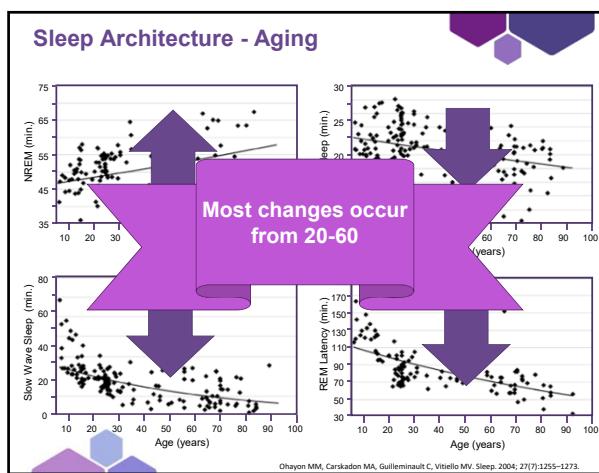


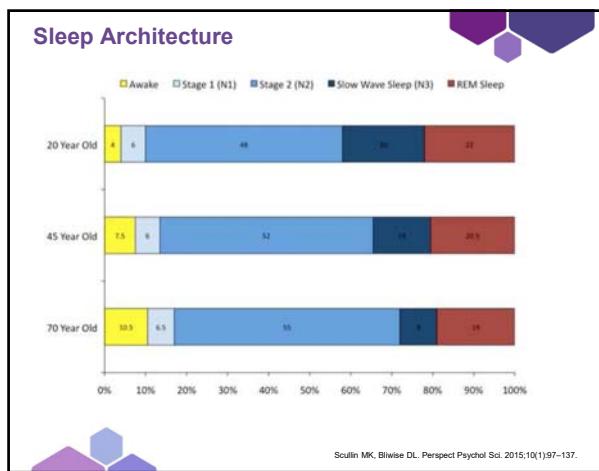
Does sleep change with age?

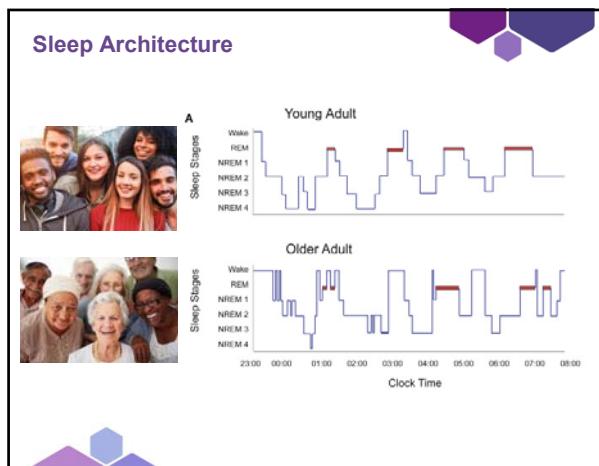


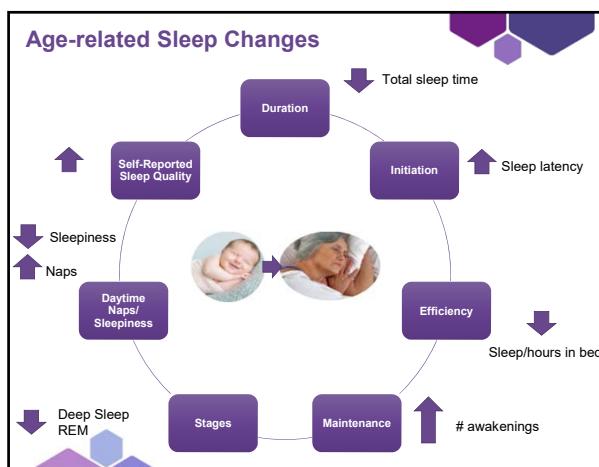


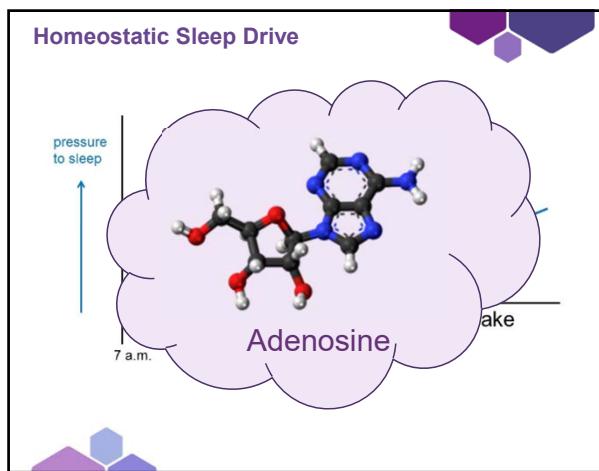


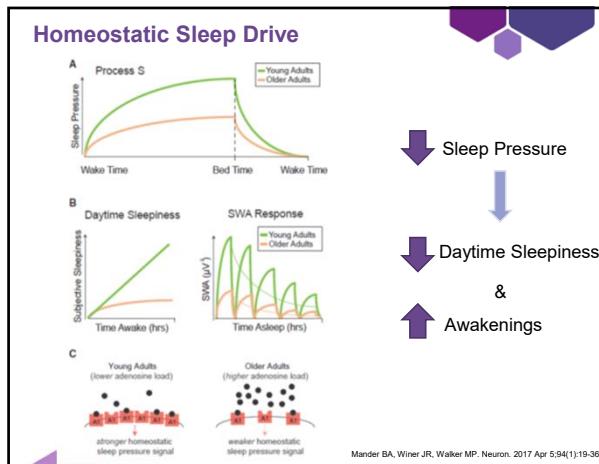


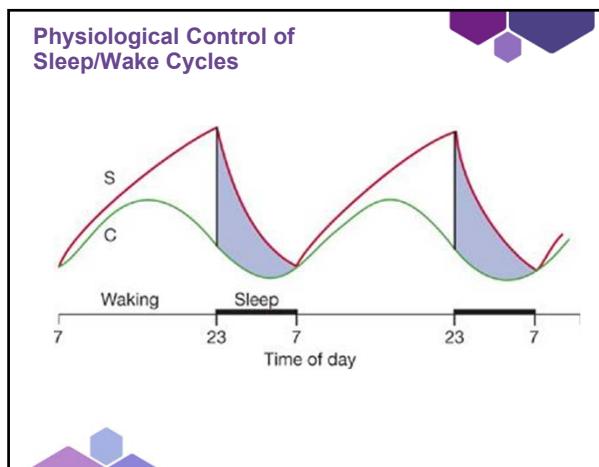


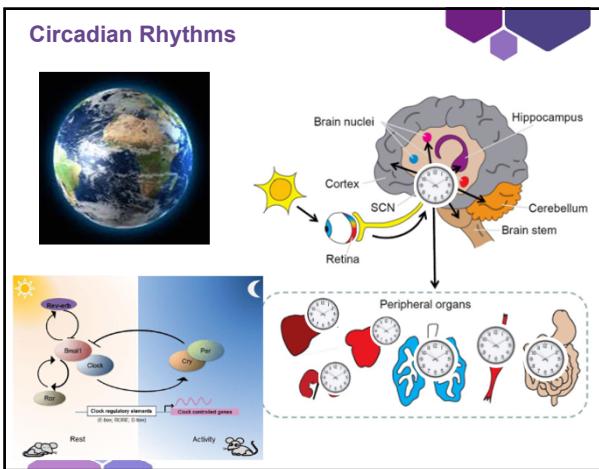


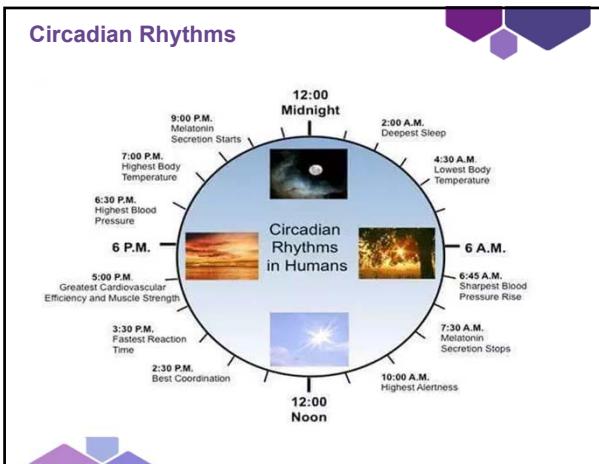


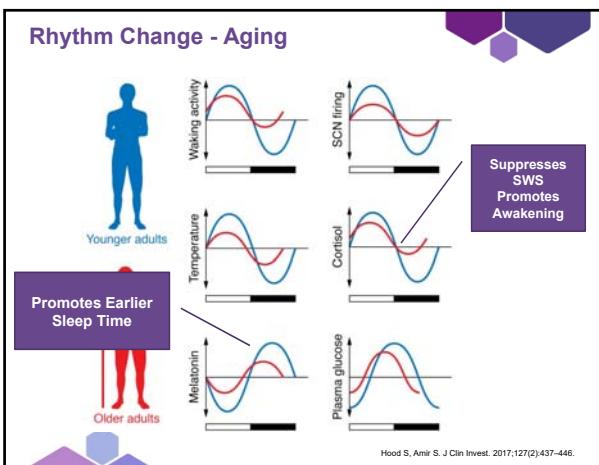


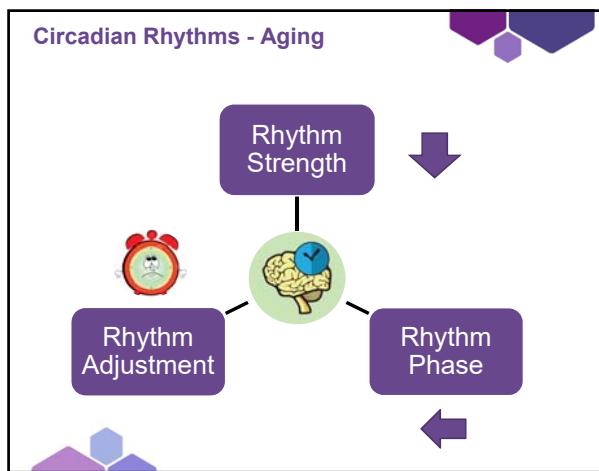


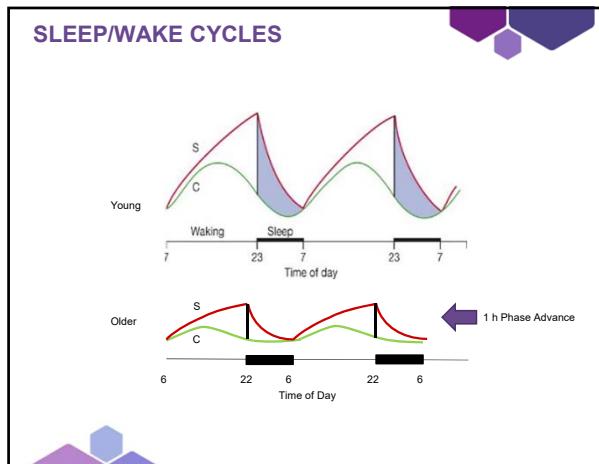




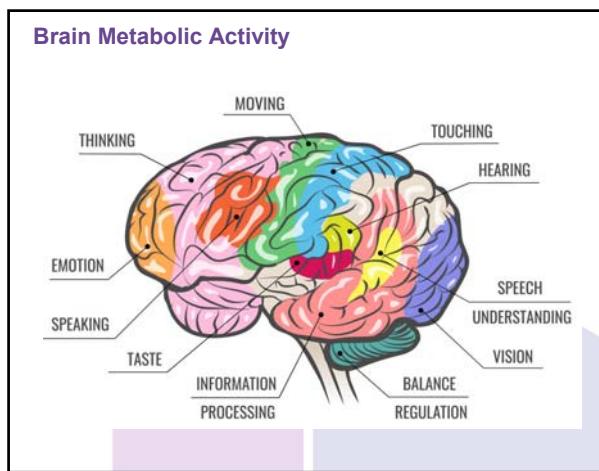




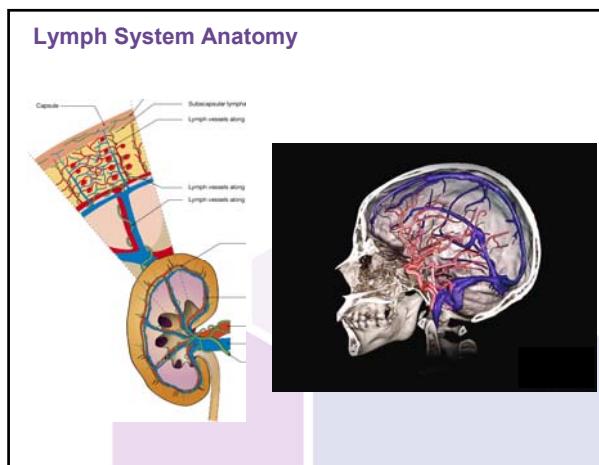




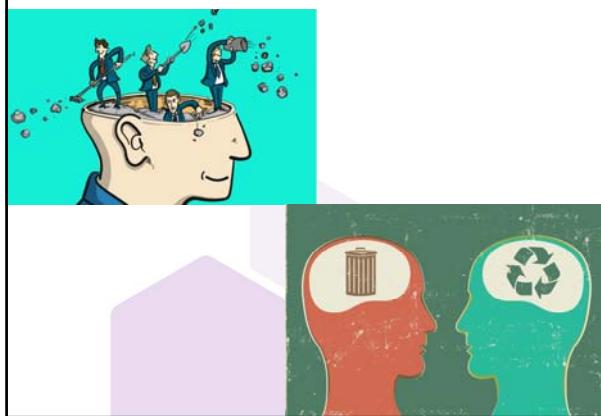




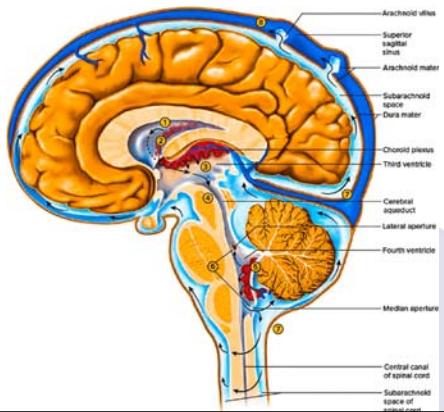




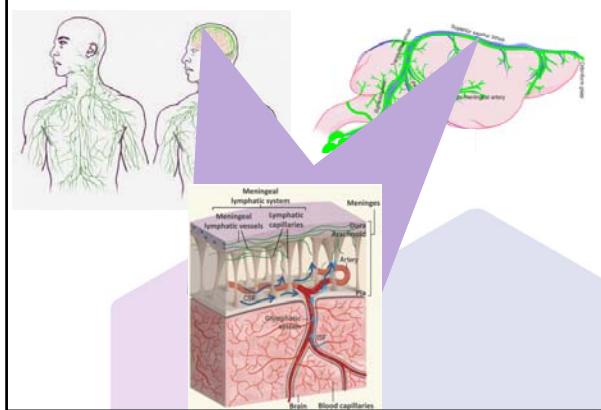
How does the brain handle waste?



CSF – the brain's lymphatic system?

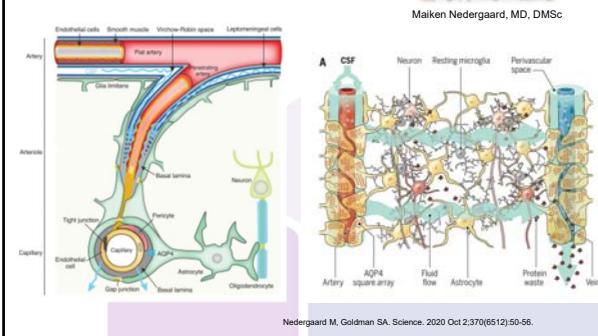


Meningeal Lymphatics

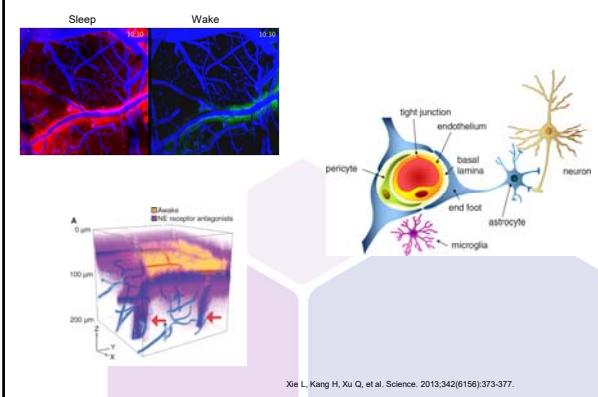


The Glymphatic System

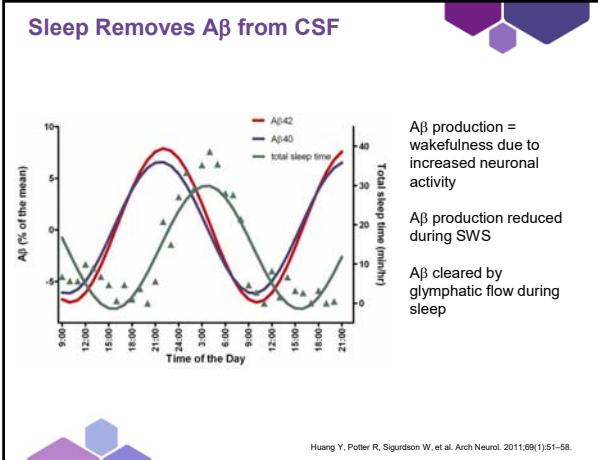
Glial-derived lymphatic-like function

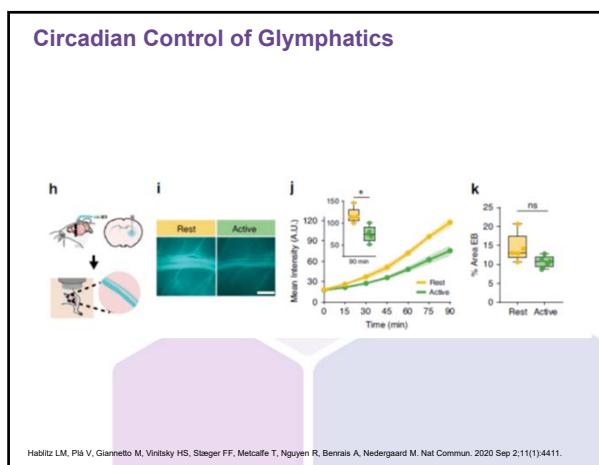
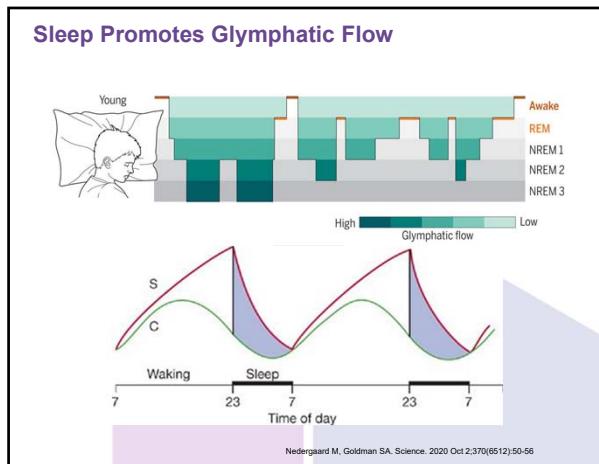
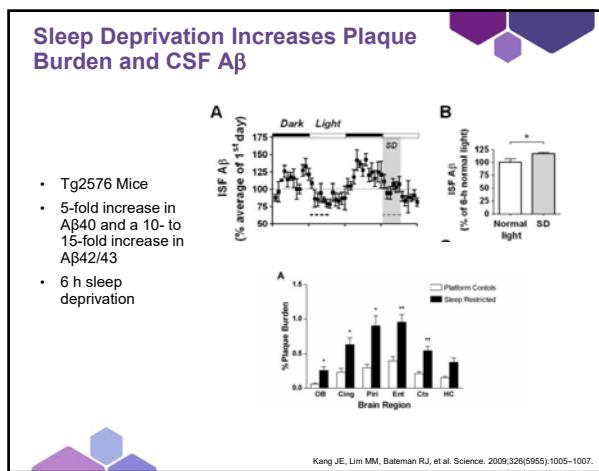


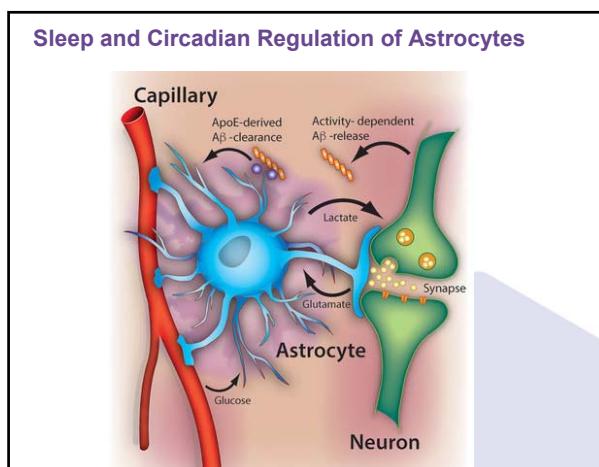
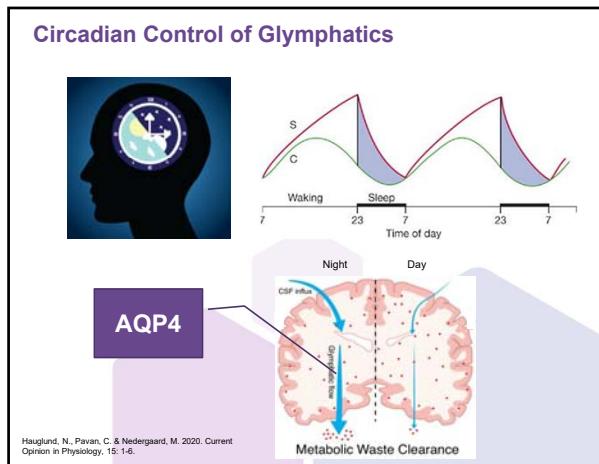
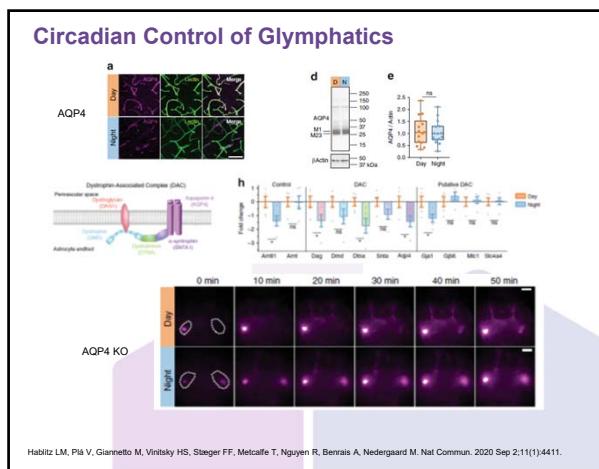
Glymphatic Flow during Sleep

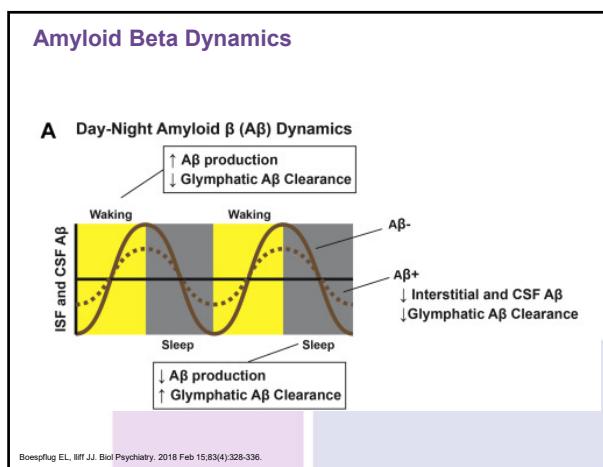
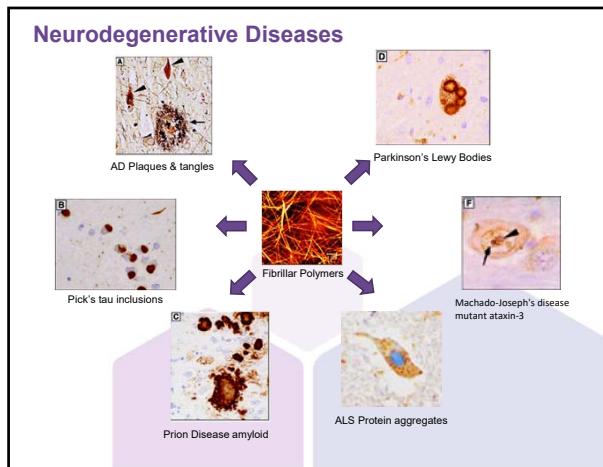
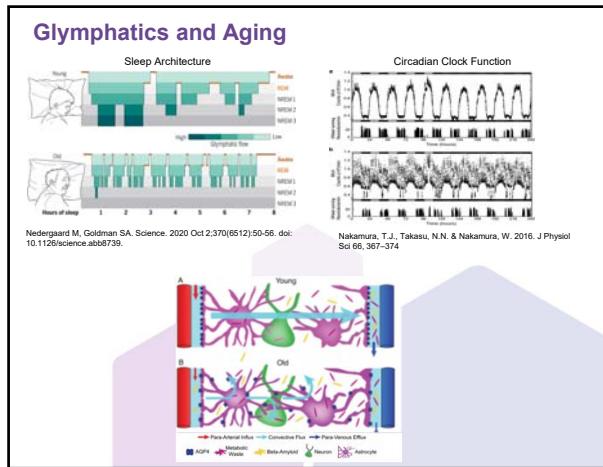


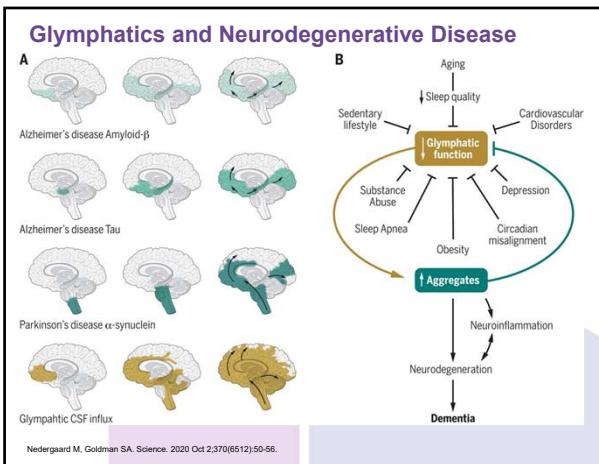
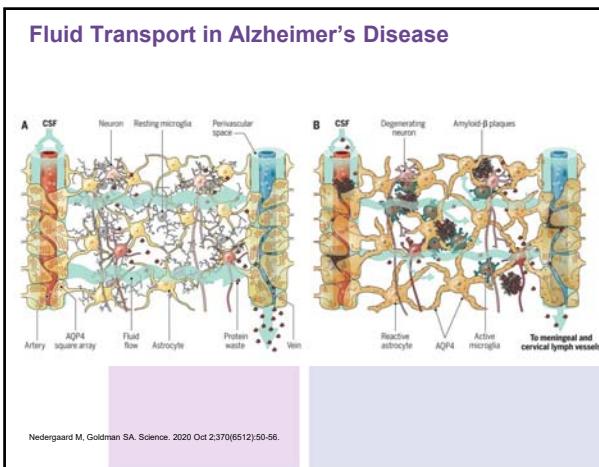
Sleep Removes A β from CSF











References

- Ohayon MM, Carskadon MA, Guilleminault C, Vitello MV. Meta-analysis of quantitative sleep parameters from childhood to old age in healthy individuals: developing normative sleep values across the human lifespan. *Sleep*. 2004 Nov;27(7):1255-73. doi: 10.1093/sleep/27.7.1255. PMID: 15386779.
- Sculley MK, Blawie DL. Sleep, cognition, and normal aging: integrating a half century of multidisciplinary research. *Perspect Psychol Sci*. 2015 Jan;10(1):97-137. doi: 10.1177/1745961614558680. PMID: 25620997. PMCID: PMC4302756.
- Maeder BL, Virelizier JL, Walker MP. Sleep and Human Aging. *Neuron*. 2017 Apr 5;94(1):119-36. doi: 10.1016/j.neuron.2017.02.004. PMID: 28390074. PMCID: PMC5372178.
- Head S, Amie S. The aging clock: circadian rhythms and later life. *J Clin Invest*. 2017 Feb 1;127(2):437-446. doi: 10.1172/JCI90328. Epub 2017 Feb 1. PMID: 28142933. PMCID: PMC5272178.
- Nedergaard M, Goldman SA. Glymphatic failure as a final common pathway to dementia. *Science*. 2020 Oct 2;370(6512):50-56. doi: 10.1126/scienceabb8739. PMID: 33004510. PMCID: PMC8186542.
- Xie L, Kang H, Xu Q, et al. Sleep drives metabolic clearance from the adult brain. *Science*. 2013;342(6156):373-377. doi:10.1126/science.1241224. PMID: 24136970.
- Huang Y, Potter R, Stuurman W, Santacruz A, Shih S, Ju YE, Kastner T, Morris JC, Mintun M, Dunley S, Bateman RJ. Effects of age and amyloid deposition on A_β dynamics in the human central nervous system. *Arch Neurol*. 2012 Jan;69(1):51-8. doi: 10.1007/s00115-011-2355. Epub 2011 Sep 12. PMID: 21911665. PMCID: PMC3254706.
- Kang J, Lin Y, Barnes CA, Ugen JJ, Smith LP, Cirrito JR, Fujiki N, Nishino S, Holtzman DM. Amyloid-beta dynamics are regulated by sleep and the circadian cycle. *Science*. 2009 Nov 13;326(5955):1005-7. doi: 10.1126/science.1183982. Epub 2009 Sep 24. PMID: 19779148. PMCID: PMC2789838.
- Hablitz LM, Pila V, Giannici M, Vrňáky HS, Steger FF, Metcalfe T, Nguyen R, Berndsen A, Nedergaard M. Circadian control of brain glymphatic and lymphatic fluid flow. *Nat Commun*. 2020 Sep 2;11(1):4411. doi: 10.1038/s41467-020-18115-2. PMID: 32879313. PMCID: PMC7371162.
- Haugland A, Pavan C, & Nedergaard, M. 2020. Cleaning sleeping brain – the potential restorative function of the glymphatic system. *Curr Opin in Physiology*, 15, 1-6.
- Nakamura T-J, Takesu N-N & Nakamura W. The suprachiasmatic nucleus: age-related decline in biological rhythms. *J Physiol Sci* 66, 367-374 (2016). <https://doi.org/10.1007/s12576-016-0439-2>
- Boespflug EL, Iff J. The Emerging Relationship Between Interstitial Fluid-Cerebrospinal Fluid Exchange, Amyloid-β, and Sleep. *Biol Psychiatry*. 2018 Feb 15;83(4):328-336. doi: 10.1016/j.biopsych.2017.11.031. Epub 2017 Dec 7. PMID: 29275202. PMCID: PMC5767516.
- Nedergaard EuroPhysiology conference talk: <https://www.youtube.com/watch?v=vtseBfMmMbyA>