

Neurobiology of

Sleep*

Sreekanth

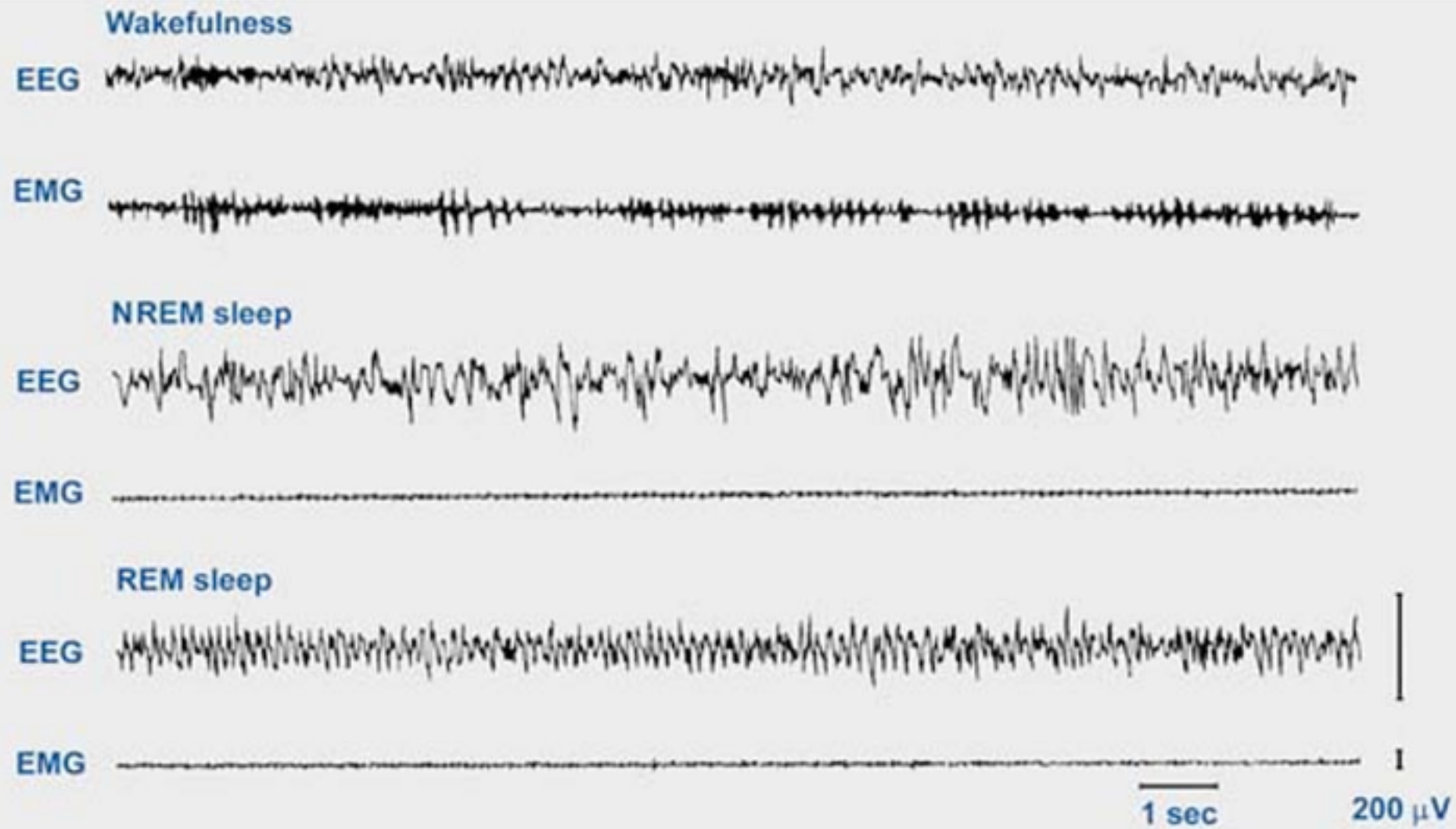
* As I understand it and from my readings, See References

What is Sleep?

- ❖ Natural (all animals have it)
- ❖ Low Motor Activity
- ❖ Suspended sensory activity
- ❖ Reversible

What is Wakefulness?

- ❖ Complementary to sleep
- ❖ Receive and respond to stimulus
- ❖ Active Cerebral cortex
- ❖ Low voltage high frequency EEG and muscle tone.



EEG/EMG patterns

http://www.scholarpedia.org/article/File:Sch_Figure1.jpg

Sleep and Wakefulness

The Differences from the perspective of a scientist

Why Sleep?

- ❖ Intellectual function is impaired.
- ❖ Brain is not taking a break.
- ❖ Some parts of the brain see more blood flow during sleep than wakefulness esp. during REM

Why Sleep*?

- ❖ Recovery and Restoration
- ❖ Energy conservation(like hibernation)
- ❖ Brain Plasticity Theory

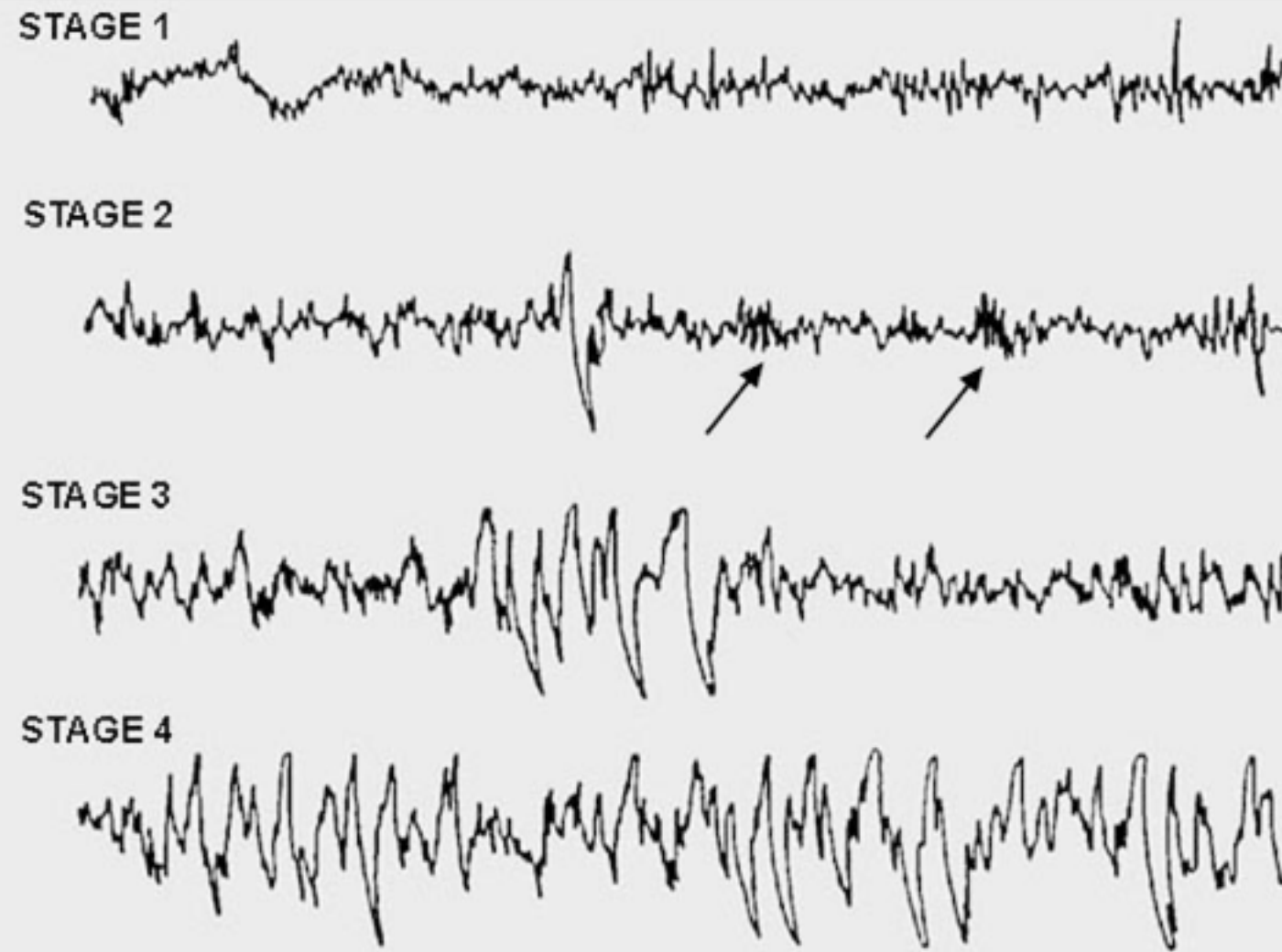
* There are only strong or weak hypotheses

Restoration and Recovery

- ❖ Increased Growth hormones at sleep onset
- ❖ Increased exercise leads to increased NREM sleep
- ❖ REM sleep seems to help rebalance neurotransmitter levels in the brain
- ❖ Improved immune system and wound healing

Energy Conservation

- ❖ Metabolic rate and body temperature drops to conserve energy during sleep
- ❖ Vasodilatation causes temperature to drop(1-2 C)
- ❖ Sleep onset leads to reduction of the body thermostat in the Hypothalamus



From wakefulness to sleep

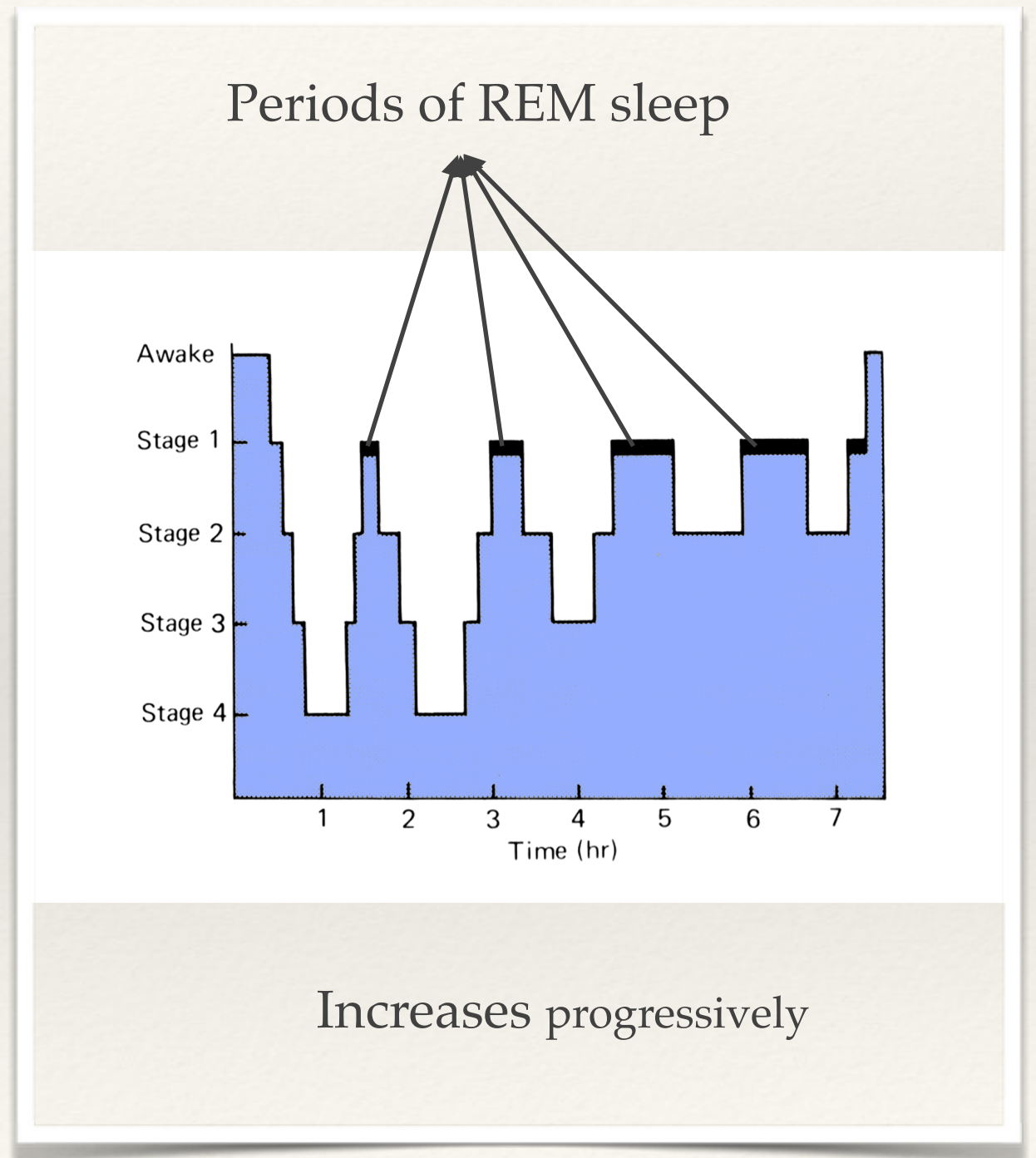
http://www.scholarpedia.org/article/File:Sch_Figure2.jpg

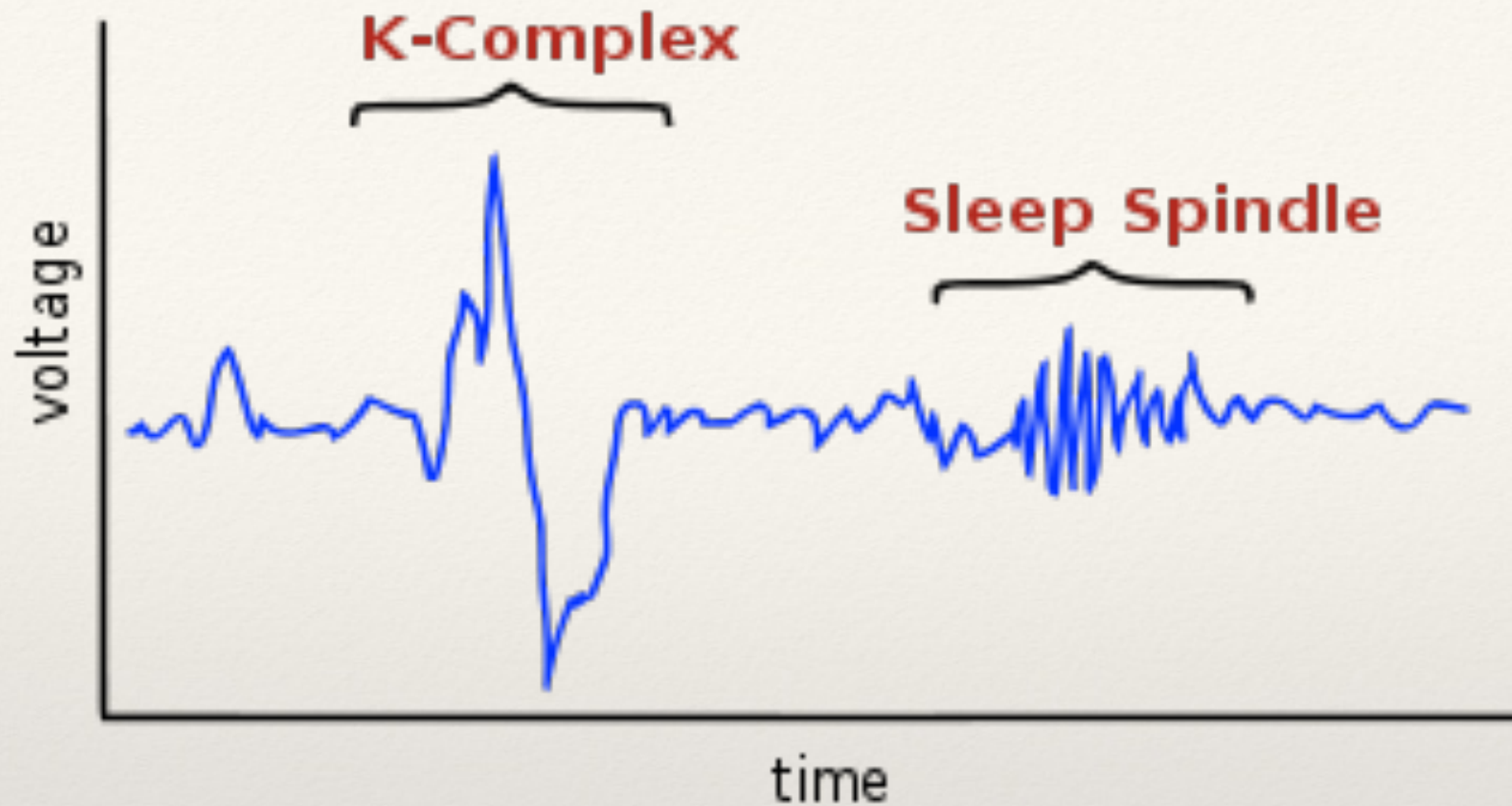
Stages of sleep

Increased voltage differences
Reduced frequency

REM sleep

- ❖ Characterised by Eyes moving rapidly
- ❖ Dreams
- ❖ Linked to consolidation of memories





Stage 2 sleep

Sleep Spindles and K Complexes

Sleep Spindles

- ❖ Typically beginning or end of NREM stage 2 sleep
- ❖ Short burst of high frequency waves
- ❖ Brain is activated and ends with muscle twitching
- ❖ Possibly mapping motor neurons to muscles
- ❖ Maintain tranquil sleep in presence of external sounds

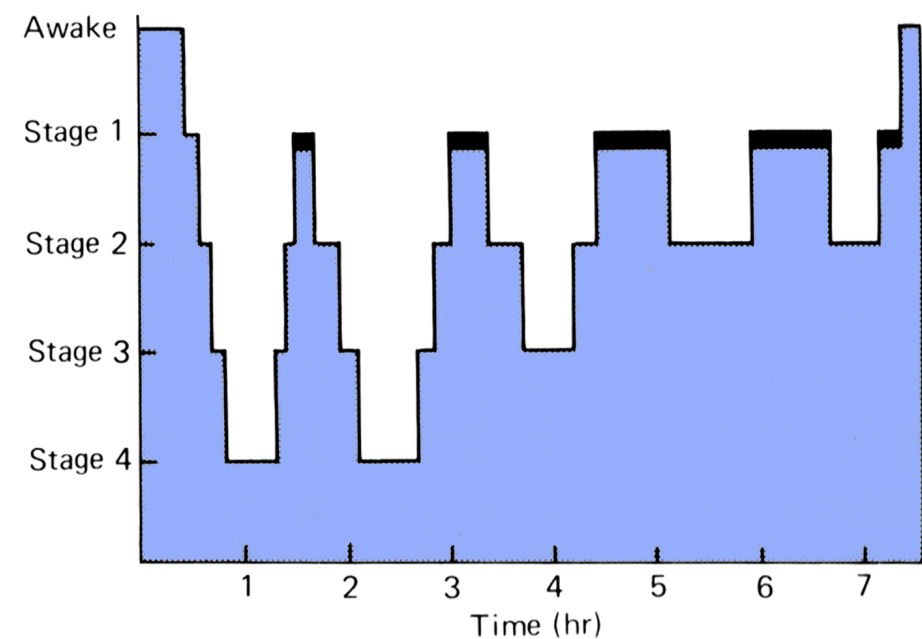
K Complexes

- ❖ Typically beginning or end of NREM stage 2 sleep
- ❖ Single delta wave(high voltage). It is largest 'healthy' event on an EEG.
- ❖ Also occur in response to external stimuli
- ❖ Linked to memory consolidation
- ❖ Cortical "down" state?

NREM Sleep

- ❖ Highly active brain regions
- ❖ Consists of multiple stages of varying patterns and length
- ❖ Probably important for homeostasis
- ❖ Important for Visuomotor and perceptual learning.

Stages of NREM sleep



Reduces progressively
Deep sleep consolidated early

Slow wave sleep

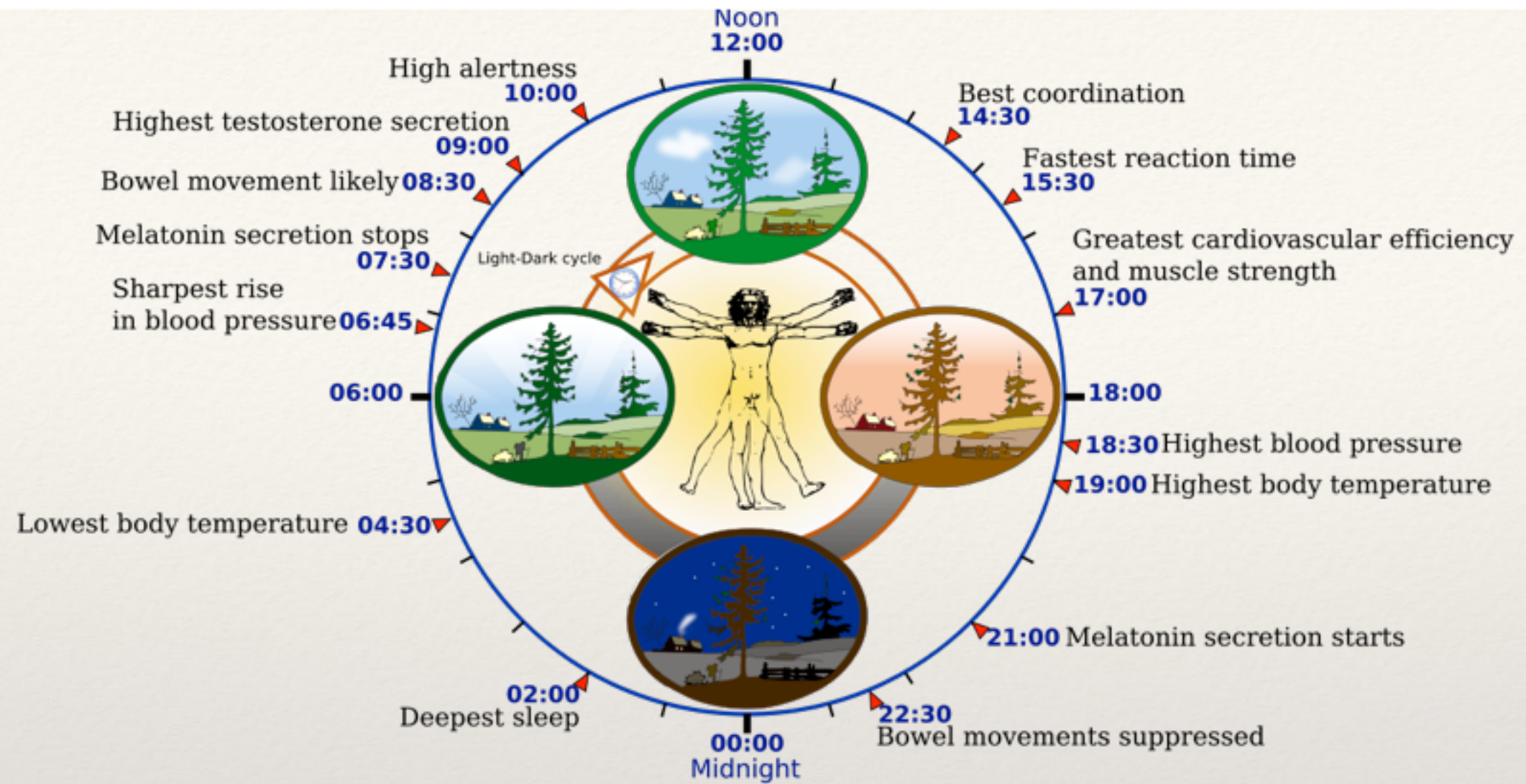
- ❖ Stages 3 and 4
- ❖ Relates to our experience of deep sleep
- ❖ Have "Up" states. Intense firing of cortical neurons
- ❖ And "Down" States. Periods of neuronal silence



<https://flic.kr/p/jZbArT>

What Causes Sleep?

All animals sleep and it's origins are primitive



Humans follow a daily pattern

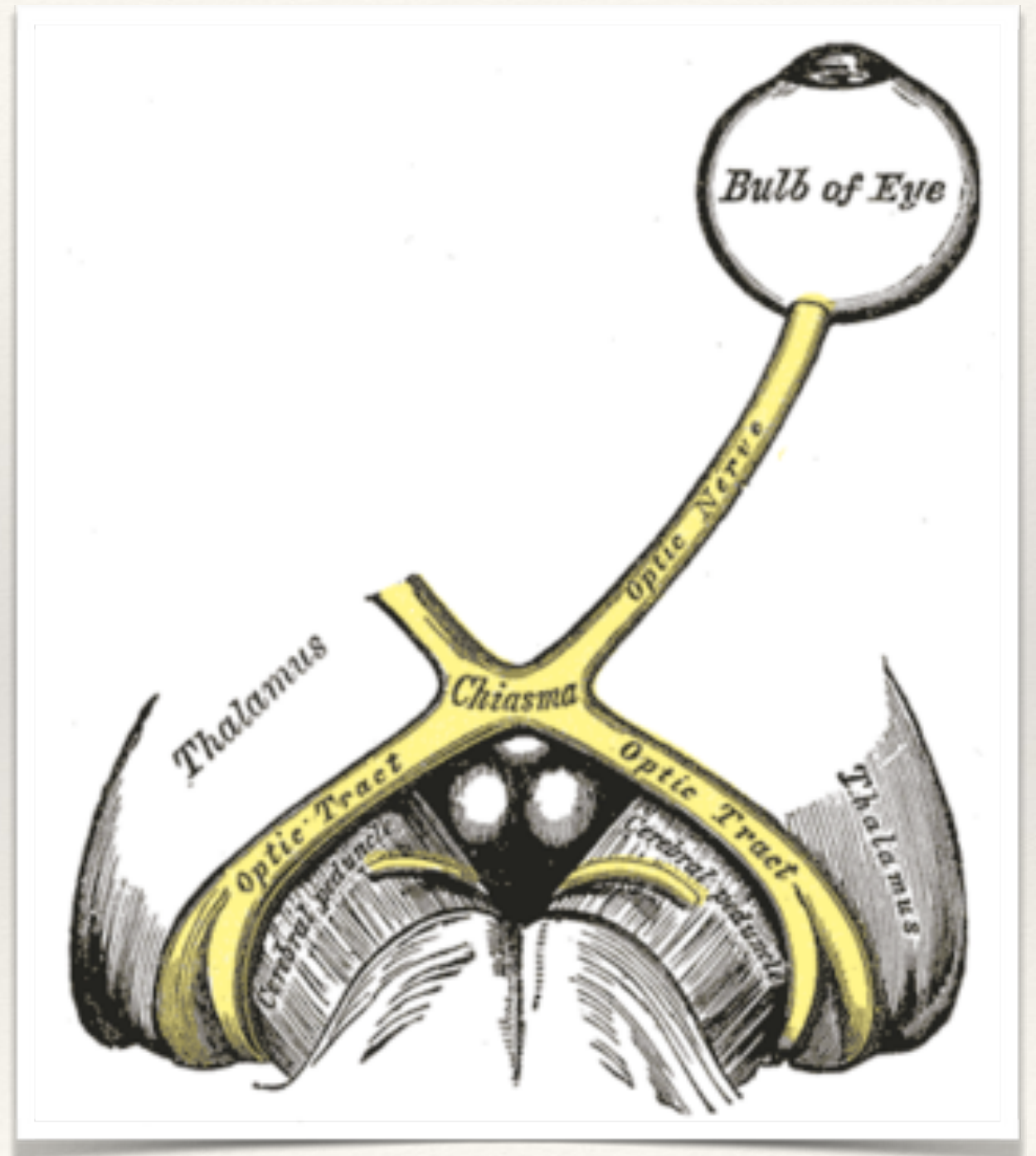
https://en.wikipedia.org/wiki/Sleep#mediaviewer/File:Biological_clock_human.svg

Circadian Rhythm

A lot of biological processes are driven by day / night. These patterns are built in and self sustained

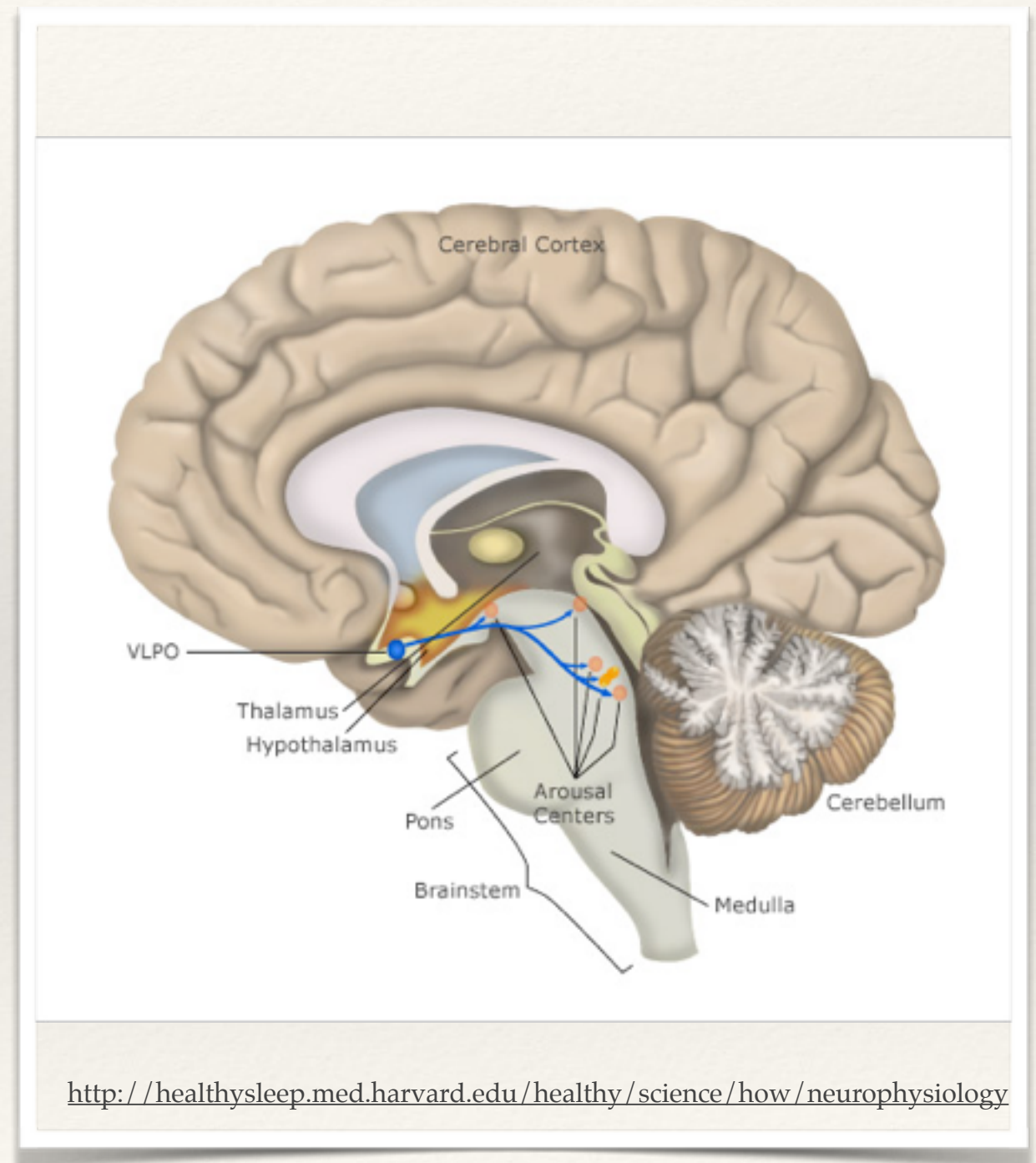
Suprachiasmatic Nucleus(SCN)

- ❖ Situated right above the optic chiasma and gets input from retina
- ❖ Part of Hypothalamus
- ❖ The master clock to control Circadian Rhythms
- ❖ Controls endocrinal activities via Pineal gland control to produce melanin



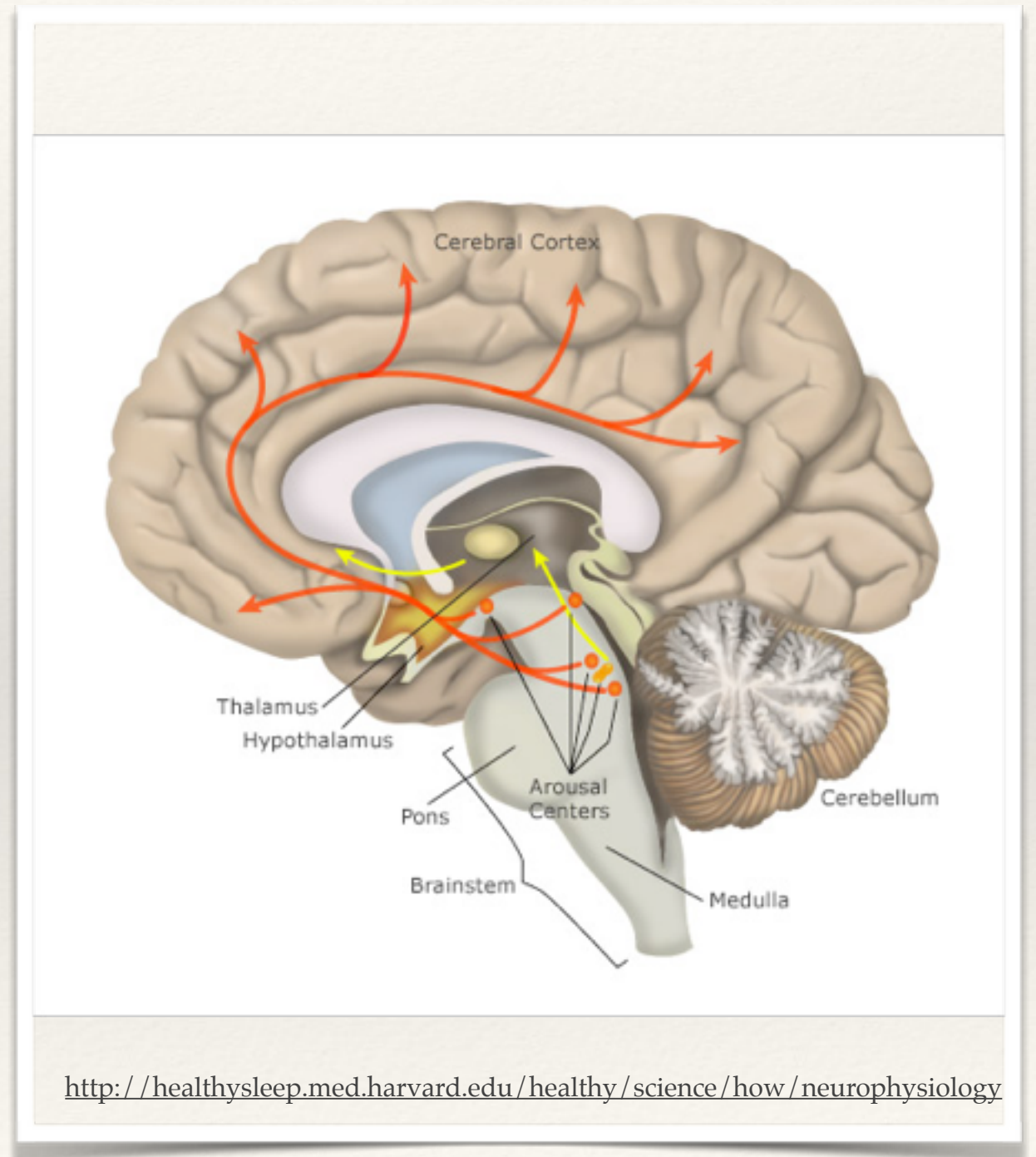
Ventrolateral preoptic nucleus(VLPO)

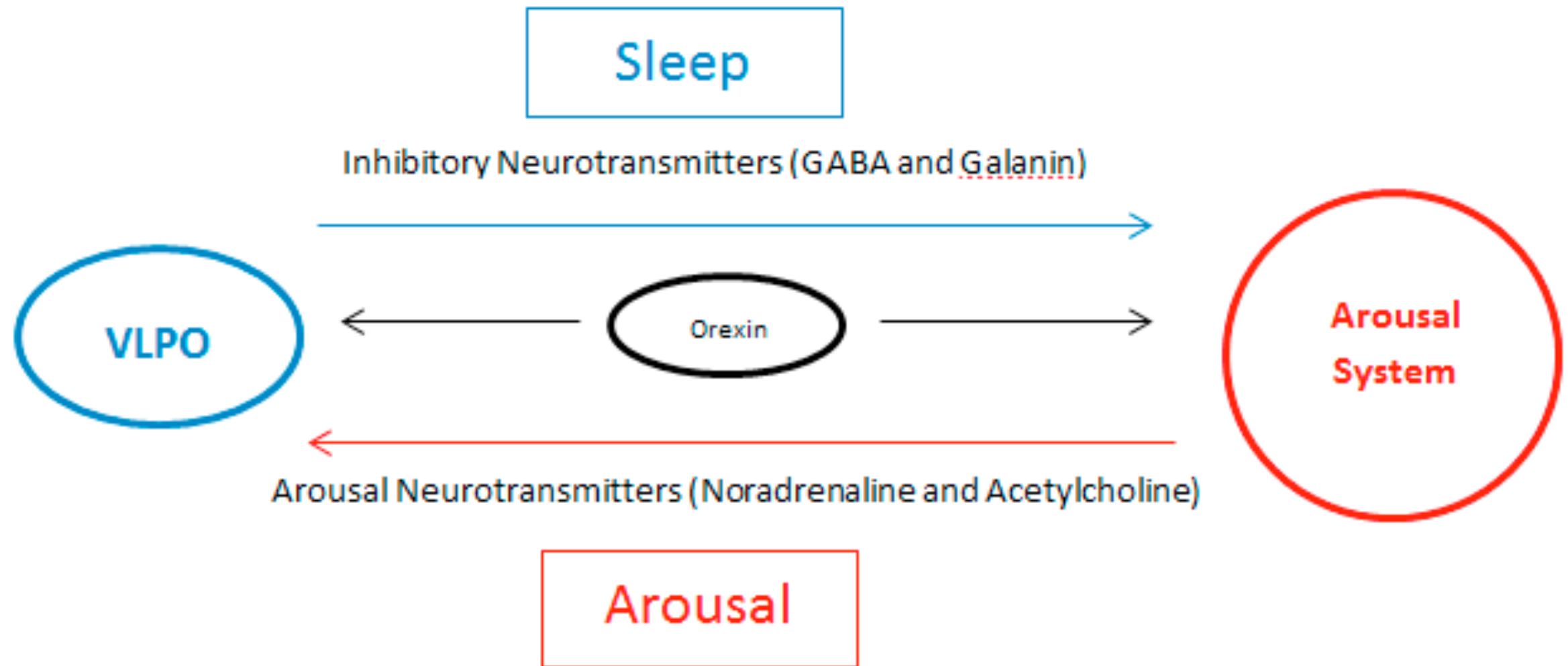
- ❖ Part of anterior Hypothalamus on the side of optic chiasm
- ❖ Inhibits the arousal systems and promotes sleep
- ❖ Inhibited by Orexin neurons during transition to wakefulness



Arousal Centers

- ❖ Produce alertness and consciousness (wakefulness)
- ❖ Start at the brain stem
- ❖ Enable cerebral cortex to be active.
- ❖ Orexin in Hypothalamus also affects cortex and arousal centres





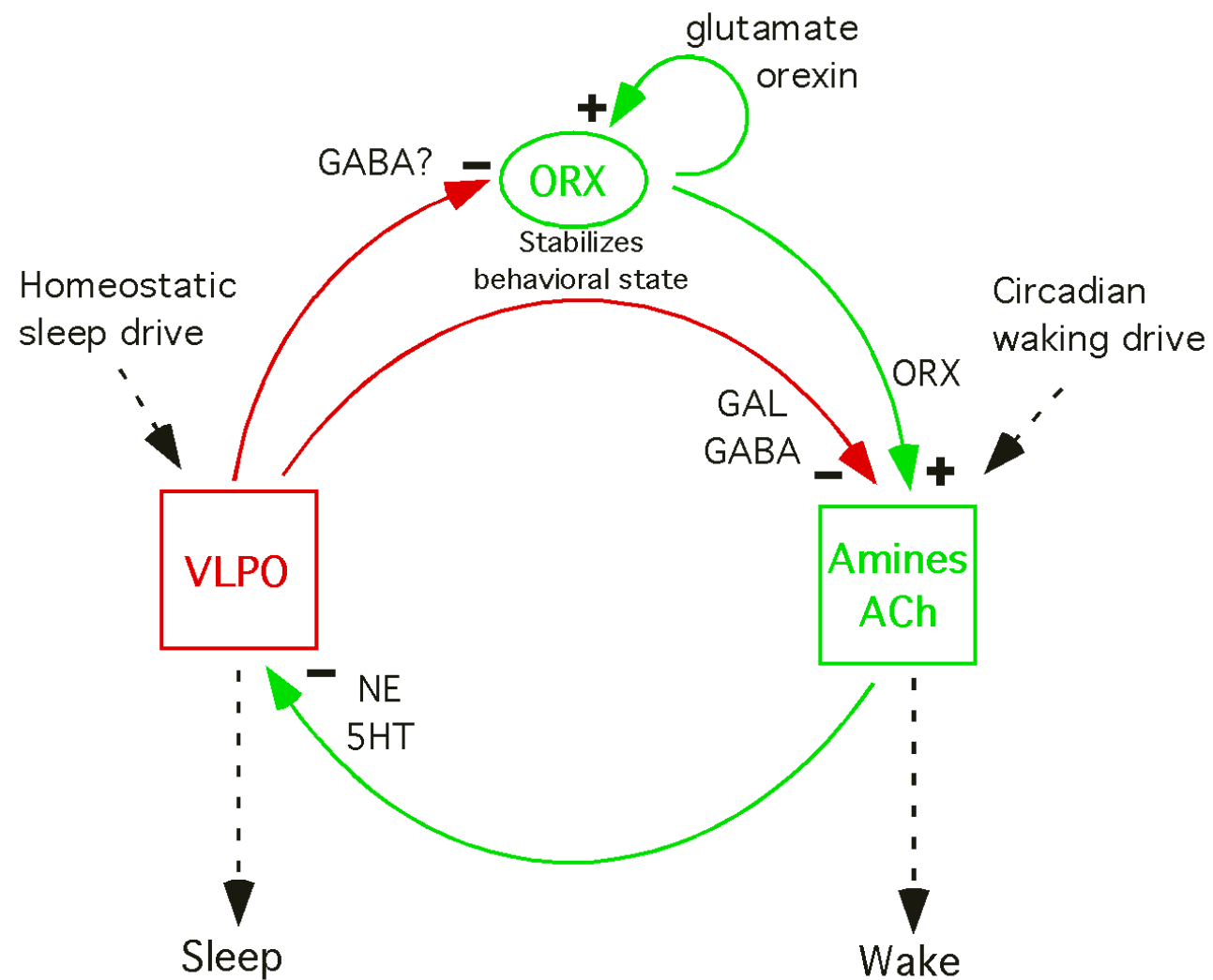
Lorem Ipsum Dolor

Sleep wake Balance

The neurotransmitter Orexin modulates the sleep wake cycle.

Homeostatic sleep need

- ❖ Humans need 1 hr of sleep for 2 waking hours
- ❖ Long sleep compensates for prolonged wakefulness
- ❖ Arousal systems in the brain cause Adenosine accumulation
- ❖ This detected in the brain stem and excites the VLPO inhibitory system to promote sleep.



The balancing act

Sleep and Wake balance

The cast

- ❖ Cortex
- ❖ Brain stem (Pons specifically)
- ❖ Hypothalamus
 - ❖ SCN
 - ❖ VLPO(Anterior Hypothalamus)
 - ❖ Orexin

REM sleep activation

- ❖ Ultradian oscillator residing in mesopontine junction(pontine taguntum)
- ❖ Rem "off" cells : Acetocholine triggered neurons
- ❖ Rem "off" cells: monoamine triggered neurons
- ❖ Rem "on" and "off" neurons form an A-B feedback to generate the cycles of REM sleep

References

- ❖ Circadian Rhythm
- ❖ REM sleep
- ❖ The science of sleep
- ❖ Role of ATP in sleep regulation
- ❖ Sleep
- ❖ Sleep resources