# Physiology of Sleep

Dr Nervana Mostafa





#### **Objectives:**

- 1. Explain the difference between **sleep** and **coma**.
- 2. Define **NREM** and **REM** sleep.
- 3. Describe how NREM and REM sleep **are distributed** during a normal night sleep in the average adult human.
- 4. Describe the **behavioral** and **autonomic** features associated with **NREM** and **REM** sleep.
- 5. Describe **EEG**, as a **physiological tool**, is being used to delineate in which stage of sleep (or wakefulness) a person is.
- 6. Appreciate how the **total sleep duration** and **different sleep stages** vary with different ages in normal humans.
- 7. Describe the **current theories** about the neural basis of sleep.

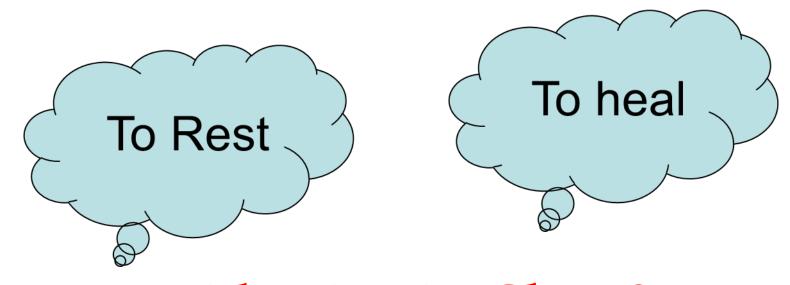
## WHAT IS SLEEP?

Sleep: State of unconsciousness from which a subject can be aroused by appropriate sensory or other stimuli.

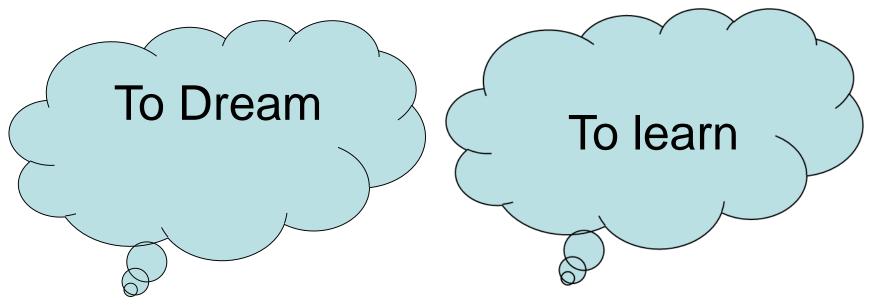
- Sleep may also be defined as a normal, periodic, inhibition of the reticular Activating system.

Awake: State of readiness/alertness & ability to react consciously to various stimuli.

Coma: State of unconsciousness from which a person cannot be aroused by any external stimuli.



## Why Do We Sleep?



# Why do we sleep?

- Sleep Theory #1 To Rest: to gain relief from this hyperactive state.
- Sleep Theory #2 To Heal: Sleep also allows us to heal our bodies.
  the immune system (our ability to fight disease) sleep deprivation
  affects our metabolism (our internal chemical reactions). It may also
  help us save energy for when we most need it.
- Sleep Theory #3 To Learn: Sleep may help the human brain get better organized - by filing away important memories and discarding unwanted information.
- Sleep Theory #4 To Dream : Dreaming appears to be a by-product of REM sleep. So is it possible that the main reason why we sleep is to dream? If so, why do we dream?

## Theories of sleep

#### **Restoration theory:**

Body wears out during the day and sleep is necessary to put it back in shape.

\* This is supported by findings that many of the major restorative functions in the body like muscle growth, tissue repair, protein synthesis, and growth hormone release occur mostly, or in some cases only, during sleep.

#### Preservation & protection theory:

Sleep preserve energy and it provides protection.

<sup>\*</sup> For example, both body temperature and caloric demand decrease during sleep, as compared to wakefulness.

Why Do We Sleep?

## Ultimate goal:

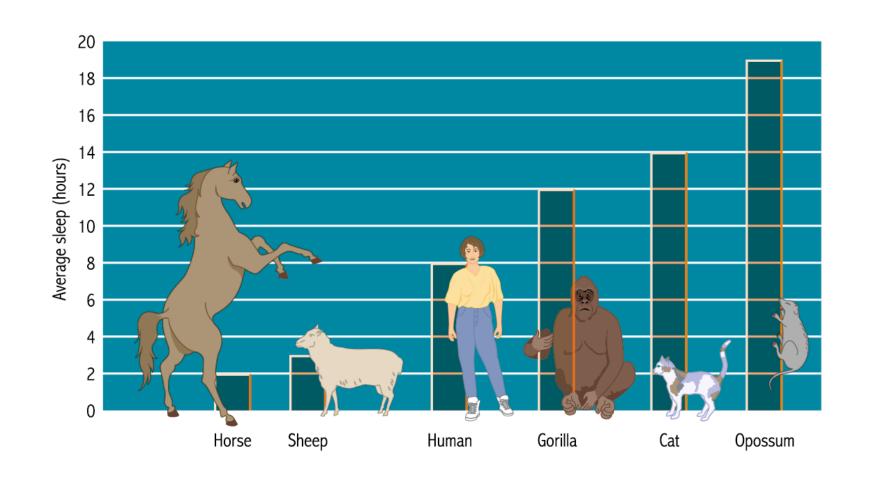
Sustains our ability to reproduce successfully, by maintaining good health.

## **Humans (Sleep Duration)**

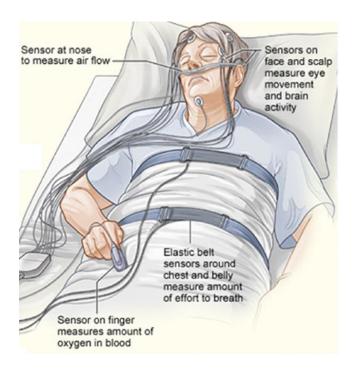
New born 15 - 20 hours. Children 10 -15 hours.

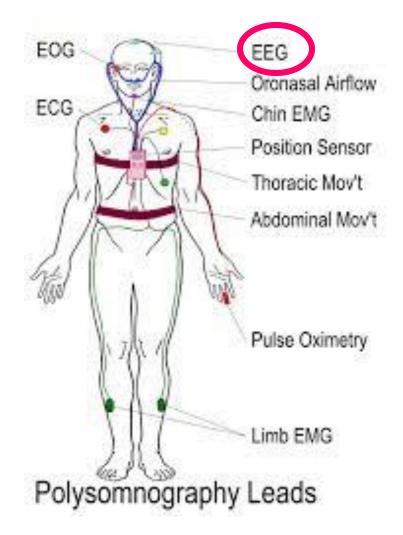
Adults 6 - 9 hours.

Old age 5 - 6 hours.







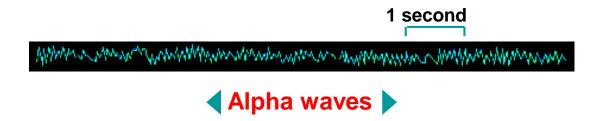


## Stages of Sleep: recorded by EEG

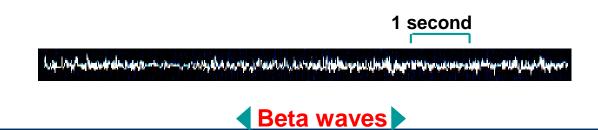


## **EEG WAVES IN WAKKFULNESS**

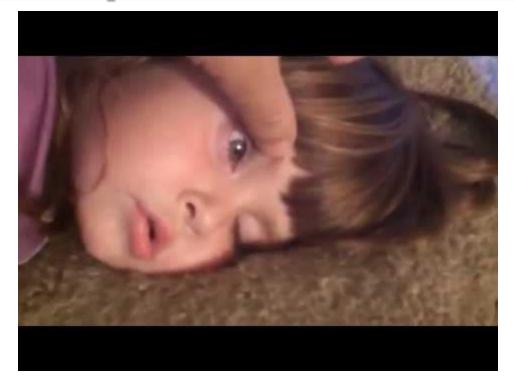
Awake, but non-attentive, large, regular alpha waves



Awake and attentive - low amplitude, fast, irregular beta waves



# **Sleep Classification**



NREM (non-rapid eye movement) - (SWS) slow wave Sleep

**REM** (rapid eye movement) Sleep

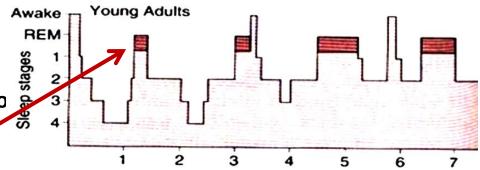
#### # Types of Sleep:

- (1) SWS (Slow-Wave Sleep),
- EEG waves are generally of low frequency.
- Non-Rapid Eye Movement (NREM) sleep is not associated with rapid eye movements.
  - (2) REM sleep (Rapid Eye Movement)
- this type of sleep rapid eye movements occur (Neurons of the pons)

#### **Distribution of Sleep Stages**

#### In a typical night of sleep, a young Adult

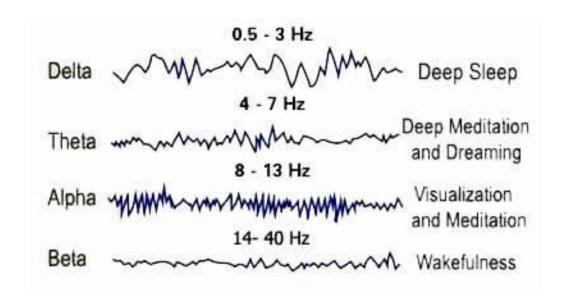
- SWS occupies most of the total night sleep time (≅ 75-80%), it is interrupted by intervening REM sleep periods, ≅ every 90 minutes .
- first enters into NREM sleep, passes
  through stages 1, 2, 3 &4 SWS,
  then, 60-100 min from sleep onset, goes into
  the first REM sleep episode.
- This cycle is repeated at intervals of about 90 min throughout the 8 hours of night sleep.
- There are 4-6 sleep cycles/night (& 4-6 REM periods per night)
- As the night goes on → there is progressive reduction in stages 3 & 4 sleep and a progressive ↑ in REM sleep.

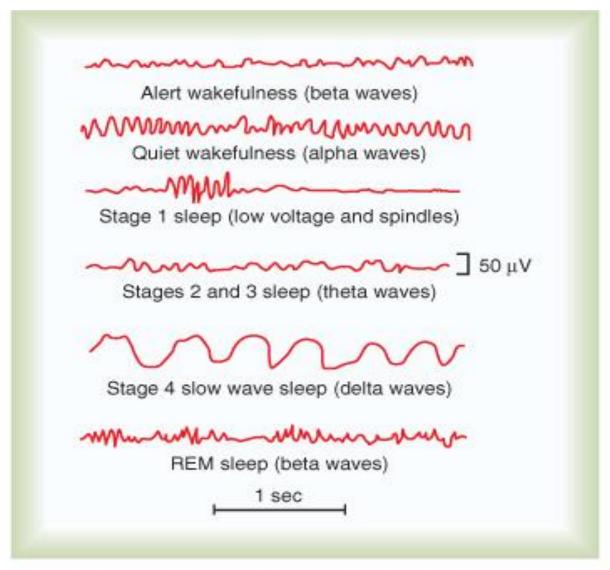


REM sleep periods are shown in red

In young adults SWS (NREM sleep) occupies <u>75-80%</u> of night sleep time & REM sleep occupies <u>20-25%</u> of sleep time.

| EEG waves |                |                                 |  |  |  |  |
|-----------|----------------|---------------------------------|--|--|--|--|
| Waves     | Frequency      | Type of activity                |  |  |  |  |
| Alpha     | 8 – 13 hz      | Smooth electrical activity      |  |  |  |  |
| Beta      | 14– 40 hz      | Irregular electrical activity   |  |  |  |  |
|           | 4– 7 hz        |                                 |  |  |  |  |
| Theta     |                |                                 |  |  |  |  |
|           | Less than 4 hz | Regular, synchronous electrical |  |  |  |  |
| Delta     |                | activity                        |  |  |  |  |



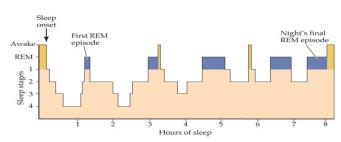


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| EEG waves |   |  |  |  |
|-----------|---|--|--|--|
| Waves     | Associated with / occurs  |  |  |  |
| Alpha     | Associated with a state of relaxation.                                      |  |  |  |
| Beta      | Associated with a state of arousal.   |  |  |  |
| Theta     | Occurs intermittently during early stages of slow wave sleep and REM sleep. |  |  |  |
| Delta     | Occurs during the deepest stages of slow-wave sleep.                        |  |  |  |

## **REM Sleep (Paradoxical Sleep)**

- In a normal night of sleep, episodes of REM sleep lasting 5 to 30 minutes usually appear on the average every 90 minutes.
- REM sleep is not as restful as SWS.
- When the person is extremely sleepy, each episode of REM sleep is short, and it may even be absent.
- Conversely, as the person becomes more rested through the night, the durations of the REM episodes increase.
- <u>Characteristics of REM sleep</u>:
- (1) There are rapid eye movements.



(2) Muscle tone throughout the body (except eye muscles) is exceedingly depressed.

- REM sleep is a type of sleep in which the brain is quite active, but this brain →
- (1) is not aware cut-off the external world.
- (2) its activity is not channeled into purposeful external motor activity.

REM Sleep \_\_\_\_\_cont..

 REM sleep occupies 80% of total sleep time in premature infants & 50% in full-term neonates.

- Thereafter, the proportion of REM sleep falls rapidly and plateaus at about 25% until it falls further in old age.
- Children have more total sleep time & stage 4 sleep than adults.

## PHYSIOLOGICAL CHANGES IN SLEEP

CVS: Pulse Rate, cardiac output, blood pressure & vasomotor tone are decreased but the *blood volume* is increased.

Respiration: Tidal volume & respiratory rate are decreased. BMR is decreased 10-15%.

*Urine volume:* decreased.

Secretions: Salivary/lacrimal secretions are reduced, gastric/sweat secretions are increased.

**TABLE 14.1** Properties of Slow-Wave and REM Sleep

| Property                             | Slow-wave sleep                       | REM sleep                     |
|--------------------------------------|---------------------------------------|-------------------------------|
| AUTONOMIC ACTIVITIES                 |                                       |                               |
| Heart rate                           | Slow decline                          | Variable with high bursts     |
| Respiration                          | Slow decline                          | Variable with high bursts     |
| Thermoregulation                     | Maintained                            | Impaired                      |
| Brain temperature                    | Decreased                             | Increased -                   |
| Cerebral blood flow                  | Reduced                               | High ←                        |
| SKELETAL MUSCULAR SYSTEM             |                                       |                               |
| Postural tension                     | Progressively reduced                 | Eliminated                    |
| Knee jerk reflex                     | Normal                                | Suppressed                    |
| Phasic twitches                      | Reduced                               | Increased -                   |
| Eye movements                        | Infrequent, slow, uncoordinated       | Rapid, coordinated            |
| COGNITIVE STATE                      | Vague thoughts                        | Vivid dreams, well organized  |
| HORMONE SECRETION                    |                                       |                               |
| Growth hormone secretion             | High ←                                | Low                           |
| NEURAL FIRING RATES                  |                                       |                               |
| Cerebral cortex (sustained) activity | Many cells reduced and<br>more phasic | Increased firing rates; tonic |
| EVENT-RELATED POTENTIALS             |                                       |                               |
| Sensory-evoked                       | Large -                               | Reduced                       |

## **Mechanisms of Sleep**

- Although several theories of sleep have been proposed, most current evidence is in favor of the following:
- (1)<u>Serotonin</u>, produced by the **Raphe Nuclei**, induces **SWS** sleep.
- (2) The mechanism that triggers **REM** sleep is located in the **Pontine Reticular Formation** & the **Ponto-Geniculo-Occipital circuit** is instrumental in generation of **REM sleep**.
- (3) <u>Melatonin</u> (released from **Pineal Gland**) plays a role in day-night alternation of sleep.

#### **Role of Serotonin & Melatonin in SWS:**

- Raphe nucleus:
- Stimulation of Raphe Nuclei (in the lower pons & medulla) induces SW.

[Destruction of the Raphe Nuclei renders the animal **sleepless** for several days until it dies]. [Administration of drugs that **block serotonin** formation make the animal sleepless for several days].

- **Transecting the brainstem at the level of the mid pons**, leaves the animal in a state of intense wakefulness for a period of days. [The transection cuts the nerves going from the inhibitory serotonin-secreting Raphe Nuclei to the Bulboreticular Facilitory area of the RAS].

Indication that the serotonin-secreting Raphe fibers normally **inhibit** the Bulboreticular Facilitory area to produce sleep.

**Injections of melatonin** induce sleep.

- Stimulation of the Supra-chiasmal Nucleus (SCN) of hypothalamus by light falling on the retina

inhibits Melatonin release from Pineal gland & produces wakefulness.

## ■Physiological Mechanisms of Sleep & Waking:

Acetylcholine:

One of the most important neurotransmitters involved in arousal.

Two groups of acetylcholinergic neurons are located in <u>pons</u> & basal forebrain.

Muramyl peptide: induces sleep

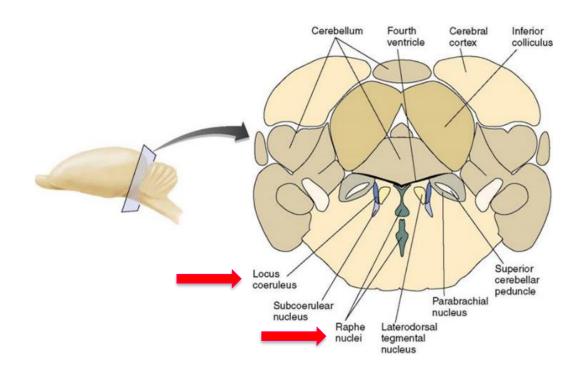
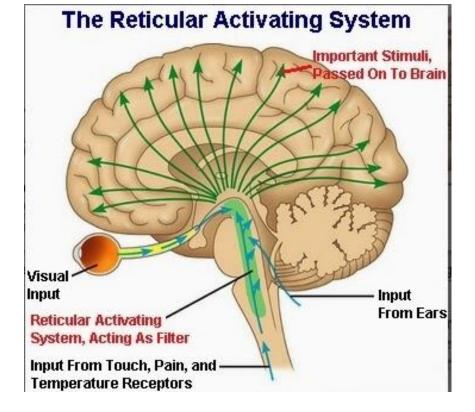
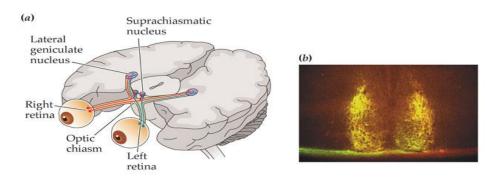


TABLE 14.2 Neural Activity of Neurotransmitter Systems during Sleep and Arousal

|                  |                     | Activity during |     |          |
|------------------|---------------------|-----------------|-----|----------|
| Neurotransmitter | Site of cell bodies | Wakefulness     | sws | REM      |
| Serotonin        | Raphe nuclei        | High            | Low | Very low |
| Norepinephrine   | Locus coeruleus     | High            | Low | Very low |
| Acetylcholine    | Brainstem           | High            | Low | High     |





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RAS: is a diffuse network of nerve pathways in the brainstem connecting the spinal cord, cerebrum, and cerebellum, and mediating the overall level of consciousness.

## SLEEP DISORDERS/BURDEN

70 million people in the US suffer from sleep problems [50% have chronic sleep disorder]

Insomnia = 30 million sleep apnea= 18 million Narcolepsy= 250,000 Americans have Motor Car Accidents= 100,000 traffic fatalities =1500 drowsy driving/annum

Approximately \$16 billion annually the cost of healthcare in USA & result in \$50 billion annually in lost productivity.

## Disorders of Sleep:

- Insomnia (inability to sleep).

- Somnolence: Extreme sleepiness.

## Drug dependency insomnia:

•An insomnia caused by the side effects of ever increasing doses of sleeping medications.

## Sleep apnea.

Cessation of breathing while sleeping.

#### – Narcolepsy:

 A sleep disorder characterized by periods of irresistible sleep, attacks of cataplexy, sleep paralysis, and hypnagogic hallucinations.

#### – Sleep attack:

 A symptom of narcolepsy; an irresistible urge to sleep during the day, after which the person awakes feeling refreshed.

## – Cataplexy:

 A symptom of narcolepsy; complete paralysis that occurs during waking.

#### – Sleep paralysis:

 A symptom of narcolepsy; paralysis occurring just before a person falls asleep.

# Good night ... Sleep tight ...