Physiology of Sleep and Clinical Characteristics

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Mini Review

General Considerations

Sleep is defined as a natural decrease in the perception [1] of the external environment that occurs periodically and reversibly, but retaining a certain degree of reactivity towards the environment and autonomous functions [2]. Sleep is considered an active process of biological, cyclical and, it is essential for survival. Most adults require an amount of sleep between 7-8 hours per day; however, there are individual variations regarding the schedule, duration, and internal structure of the dream [3].

Sleep is associated with the state of physical and mental health. It is documented that during this process of neurobiological nature, functions inherent to the organism are verified [3]. Through sleep, the organism recovers from everyday fatigue [4]. Likewise; sleep plays a relevant role in the consolidation of learning and memory. On the contrary, sleep deprivation impairs psychomotor and cognitive functions including the time of reaction to a stimulus, surveillance capacity, judgment, attention and, information processing. In addition, sleep deprivation is associated with decreased concentration and high frequency of affective disorders [5-7]. It is estimated that sleep disorders and alterations in the circadian rhythm affect 15-20% of adult individuals [7,8].

Numerous studies show that sleep disorders favor the development of various diseases, and can adversely affect their evolution, including cardiovascular disease, glucose intolerance, hypertension (HAS) and obesity. On the other hand, in several investigations it is shown that sleep disorders can adversely affect the state of mental health causing excessive drowsiness, fatigue, cognitive alterations and affective disorders [5,6,7,9].

Neurobiology of Normal Sleep

The characteristics of normal sleep are described by three electrophysiological parameters that include: [10].

a. Brain electrical activity (electroencephalogram, EEG).

b. Eye movements (electro-oculogram, EOG).

c. Muscle activity (electromyogram, EMG).

From a physiological point of view, the continuous recording of these parameters allows distinguishing the waking state of the dream and, within it; different phases are identified [10]. The EEG record of a healthy adult who is in the waking state and with his eyes closed demonstrates basal electrical activity consisting of a series of waves of relatively fast frequencies and low to moderate amplitude. This pattern is designated as beta waves (13-25cycles/s) and alpha waves (8-12cycles/s); In addition, the EMG shows high tonic activity, and the EOG records a series of rapid eye movements and flickering.

When numbness occurs, eye movements slow down, EMG activity decreases and the alpha wave record is replaced by a pattern that is characterized by a series of slow frequency waves (3-7cycles/s) and low amplitude. In the center of the skull there are outbreaks of waves that show greater voltage. This period is designated as N1 Stadium of the dream [10,11]. Next, the EEG registers the spindles that are characteristic of the dream (outbreaks of 12-15 cycles/s), and the K complexes (slow wave outbreaks of 1cycle/s) that constitute markers of the N2 Stage. From this moment, an individual stops responding to simple orders and cannot remember events that have occurred. Additionally, as the depth of sleep increases, the EEG registers a type of electrical activity called slow delta (1-3cycles/s) with high amplitude (Stage N3) [11].

After Stage N3, the transition to the so-called rapid eye movement dream or REM (or REM) dream is evident. REM sleep is characterized by having this type of movements in the eyes that are similar to those registered during the waking state (with closed...
eyes) and with minimal muscle activity; in addition, the EEG records a pattern of intermediate activity between the waking state and the N1 stage, consisting of wave sprouts with triangular morphology of 3-5 cycles/s [12]. The electrophysiological recordings that are evidenced during REM sleep are characteristic and allow the classification of sleep in two stages: REM sleep and non-REM sleep (includes stages N1-N3) [12,13].

During sleep various variations are recorded in some physiological processes including heart rate, body temperature, respiratory rate and hormonal secretion [14]. In general terms, basal cardiac activity is greater in the waking state, and decreases during sleep; however, the heart rate increases at specific times. Systemic blood pressure and heart rate show lower values during Stage N3 of non-REM sleep in relation to baseline; however, during REM sleep, lower values are recorded with respect to the waking state [14].

In the first stages of sleep (Stages N1 and N2), breathing is irregular, with episodes in which the respiratory amplitude increases, but there are also periods in which it decreases. In Stage N3, breathing is regular in both frequency and amplitude; it also decreases ventilation per minute [14]. In the secretion of hypothalamic-pituitary hormones rhythmcity associated with sleep and wakefulness is demonstrated. The plasma levels of growth hormone and prolactin increase during sleep. Cortisol is secreted with periodicity independent of environmental influences. The highest concentration of this hormone is demonstrated in the early hours of the morning, prior to awakening. The concentration of cortisol decreases during the day to reach its lowest levels at the beginning of sleep [14]. Body temperature decreases at the onset of sleep, and its lowest level is recorded during the second half of the night [14-16]. It is postulated that the generation of the process of sleep and awakening occurs along a neuronal axis comprised between the brainstem and the anterior portion of the brain [17].

Cardiac Rhythms

Circadian rhythms are regulated by a “biological clock” located at the level of the suprachiasmatic nucleus of the hypothalamus in the Central Nervous System (CNS) [17]. It is postulated that the light-dark level influences the endogenous biological clock, in virtue of direct neural connections between the suprachiasmatic nucleus and the retina. The biological clock demonstrates an approximate rhythm city of 24 hours (h) with respect to the sleep-wake cycle, various organic functions and in cognition [17].

Periodic repetition of sleep and the waking state every 24 hours depends not only on the exterior (day and night), but is determined by genetic mechanisms, since a similar rhythm persists (approximately 25 hours) after the external influences associated with some schedule [17]. In addition, various biological functions show approximate rhythmcity of 24h, including urinary excretion for potassium, gastric secretion and acid renal secretion. In general terms, these rhythms are called “circadian” (circa = around), by virtue of their duration of approximately one day [18].

Recent evidence shows that at the level of the CNS there are neural feedback loops in which a genetic transcription process that forms specific proteins to integrate the molecular basis of the endogenous biological clock is verified. This molecular basis is demonstrated in some peripheral tissues that indicates the physiological relevance of circadian rhythmcity throughout the organism [17,18].

Classification of Sleep Disorders

Being according to the international classification of sleep disorders (2005):

a. Insomnia.

b. Sleep disorders secondary to respiratory disorders.

c. Hypersomnia of central origin not related to circadian rhythm disorders, sleep disorders due to respiratory disorders, or other cause that interrupts nighttime sleep.

d. Circadian rhythm disorders.

e. Parasomnias.

f. SAW. Sleep disorders due to movement disorders.

g. Isolated symptoms, seemingly normal variations and unresolved issues.

h. Other sleep disorders

Excessive daytime sleepiness It refers that sleepiness consists of a propensity to sleep, and represents the transition from the waking state to the beginning of sleep. Several studies indicate that 4-9% of adult individuals show excessive daytime sleepiness [19]. The main cause of this disorder is Obstructive Sleep Apnea Syndrome (OSAS) [20].

In OSAS, there is a respiratory dysfunction that is associated with intermittent hypoxia during sleep, resulting in sleep fragmentation, sleep deprivation and insomnia [20]. This syndrome occurs frequently in men who are overweight - obese, and in older adults. Estimates indicate that it affects 2 to 5 million adults in the United States of America. OSAS is due to episodic occlusion in the airway that results in a decrease or cessation of breathing lasting between 10-150s, and repeated thirty to several hundred times during the night. OSAS causes excessive daytime sleepiness with decreased alertness, hyperactivity of the Sympathetic Nervous System, oxidative stress, systemic arterial hypertension and cardiovascular complications [20].
Excessive sleepiness usually occurs in situations that require poor physical activity, as well as relative alertness. In addition, excessive sleepiness can be evidenced as an automatic behavior in which routine activities are carried out but without being fully aware of them such as writing or walking, and which register duration from seconds to minutes [21]. The results of electrophysiological records in people with excessive daytime sleepiness evidence shortening in latencies that corresponds to the onset of sleep. These records are consistent with the existence of micro-dreams that interfere with the state of alert, and affect the ability to respond appropriately to specific requirements of cognitive functioning. The above, causes variability in cognitive functions that results in an unstable alert state, and favors errors by omission, or errors by commission. The errors by omission constitute behavioral lapses evidenced by the inability to respond to a stimulus during the appropriate time. On the other hand, commission errors are behavioral type responses without any stimulus, or represent incorrect responses to a specific stimulus [21]. Excessive daytime sleepiness is associated with problems in social relationships and decreased individual performance that leads to job loss, school failure and high risk of work and traffic accidents, among other consequences [22].

Insomnia

In studies carried out in various countries, insomnia is considered to be the most frequent sleep disorder. Estimates indicate that its prevalence varies according to age (higher in adults) and gender (higher in females). Insomnia is rare and is estimated to affect 6% of the adult population [21,23].

Atypical behaviors during sleep

Atypical behaviors during sleep or parasomnias are rare disorders in adults. However, it is pointed out that 15% of children occasionally have an episode of sleepwalking, between 1 - 6% present night terrors, and approximately 20% experience nightmares [24].

Disorders in the sleep cycle – wakefulness

Sleep deprivation caused by an imbalance in circadian rhythmicity results in excessive drowsiness, cognitive dysfunction and insomnia [7].

Some studies report that this disorder may be transient or persistent

The transitory imbalance of the sleep-wake cycle is verified during air travel through areas that have different hours, or by frequent modifications in the work schedule. On the other hand, the persistent circadian maladjustment includes two syndromic pictures [25].

Syndrome with delayed sleep phase

This syndrome consists of being able to sleep until 2:00 or 3:00 a.m. It should be noted that sleep is normal in both quantity and quality, and should not be confused with insomnia.

Syndrome with advanced sleep phase

In this syndrome, a very early awakening is verified.

The role of the imbalance in the circadian rhythm as a trigger for the development of insomnia has yet to be elucidated. Authors postulate that insomnia can start as a mismatch in circadian rhythmicity and later evolve into a chronic type disorder [7].

References


