# History of Sleep Medicine and Polysomnography

## Overview

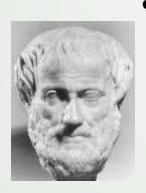
- "More has been learned about sleep in the past 60 years than in the preceding 6,000." –
   J. Allan Hobson, Sleep (1989)
- Sleep research, emphasizing all-night sleep recordings, burgeoned in the 1960s – precursor of sleep medicine and polysomnography

## Timeline

 400 B.C. – Hippocrates wrote, "I have known many persons in sleep groaning and crying out, some in a state of suffocation, some jumping up and fleeing out of doors, and deprived of their reason until they awaken, and afterward becoming well and rational as before, although they be pale and weak; and this will

happen not once but frequently"

- 360 B.C. Dionysius, the tyrant of Heraclea
  - Historical documents describe him as immensely obese and record that he died "choking on his own fat"
  - His physicians may have used the first treatment of apnea – sticking needles through the skin to arouse him from sleep



 350 B.C. – Aristotle wrote about sleep and waking, whether they are a function of the body or the soul, and the significance of dreams; Observed that all creatures sleep

### 1603 A.D. – Shakespeare

Describes sleepwalking in *Macbeth*, act 5, scene 1



- "GENTLEWOMAN. Since his Majesty went into the field, have seen her rise from her bed, throw her nightgown upon her, unlock her closet, take forth paper, fold it, write upon't, read it, afterwards seal it, and again return to bed; yet all this while in a most fast sleep"
- Character Falstaff appears in 3 plays
  - Obese, snored, and fell asleep at inappropriate times – all symptoms of sleep apnea

#### 1605 – Miguel de Cervantes Saavedra

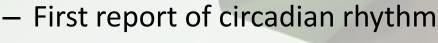
 In The Ingenious Hidalgo Don Quixote of La
 Mancha, described REM behavior disorder in Part 1, Chapter 35

 "And in his right hand he held his unsheathed sword, with which he was slashing about on all sides, uttering exclamations as if he were actually fighting some giant: and the best of it was his eyes were not open, for he was fast asleep, and dreaming that he was doing battle with the giant."



1672 – Sir Thomas Willis (of the circle of Willis) describes the features of restless legs syndrome. The condition does not receive a name until 1945.





- Conducted experiment using mimosa plant that opened leaves at certain time when sunny
- Put plant in box with no exposure to light, and plant's leaves still opened at same time

#### 1816 – William Wadd

- Surgeon Extraordinaire to King of England
- Writes monograph "Cursory Remarks on Corpulence; or Obesity Considered as a Disease" which describes sleepiness in obesity
- One case "became at length so lethargic, that he fell asleep in the act of eating, even in company"



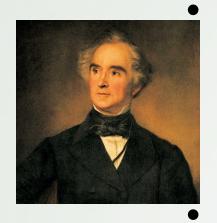


## 1818 – John Cheyne

 Describes breathing pattern named after him in "A Case of Apoplexy in Which the Fleshy Part of the Heart was Converted into Fat"



"For several days his breathing was irregular; it
would cease for a quarter of a minute, then it
would become perceptible, though very low, then
by degrees it became heaving and quick, and then
it would gradually cease again: this revolution in
the state of his breathing occupied about a minute,
during which there were about thirty acts of
respiration."



### 1832 – Justus von Liebig

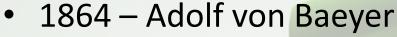
 Discovered chloral hydrate, perhaps the first widely used and abused hypnotic agent

#### 1836 - Charles Dickens



- Published The Posthumous Papers of the Pickwick Club
  - Book describes Joe, the fat boy whose symptoms of snoring and sleepiness form the basis of the first article to describe the Pickwickian syndrome, published in 1956





- Discovers barbituric acid, the parent compound of barbiturates
- 1869 William Hammond



- Publishes Sleep and its Derangements
- Uses phrase "persistent wakefulness" to describe insomnia
- Describes sleep state misperception, blaming the condition on increased blood flow to the brain
- 1875 Richard Caton
  - Demonstrated electrical rhythms in the brains of animals



- 1877 Karl Westphall
  - First to describe sudden bouts of sleeping associated with loss of motor tone



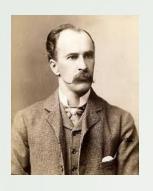
- 1880 Jean Baptiste Edouard Gelineau
  - First to use the term "narcolepsy" to describe a disease with irresistible sleep



- 1895 Nathaniel Kleitman is born
  - First and most famous sleep researcher
- 1898 William Wells makes association of nasal obstruction and daytime sleepiness
- 1902 Leopold Lowenfeld states narcolepsy is associated with cataplexy
- 1902 Emil Fischer and Joseph von Mering



Synthesize barbital, first widely used barbiturate hypnotic



- 1918 William Osler
  - In Principles and Practice of Medicine, describes sleeplessness and mental symptoms including drowsiness in congestive heart failure
- 1929 Hans Berger first to record an electroencephalogram (EEG)
- 1934 W.R. Harrison



- Describes clinical consequences of Cheyne Stokes breathing in heart failure
- Describes sleep onset, sleep maintenance insomnia, paroxysmal nocturnal dyspnea, and shows how periodic breathing pattern improves with treatment of heart failure



- 1935 Alfred Loomis describes the EEG findings of "nonrapid eye movement" (NREM) sleep
- 1937 Annie Spitz describes 3 cases of OSA in patients who have right heart failure, Cheyne Stokes respiration, snoring, and sleepiness



 1939 – Nathaniel Kleitman publishes Sleep and Wakefulness



 1945 – Karl-Axel Ekbom introduces the term restless legs syndrome and describes the condition



• 1949 – Giuseppe Moruzzi and Horace Magoun describe reticular activating system and the neurologic basis for wakefulness and arousal



- 1953 Nathaniel Kleitman
  - At the University of Chicago, assigned grad student Eugene Aserinski to use eye muscle movements as a measure of depth of sleep and documented REM sleep
- 1956 Sydney Burwell and others describe Pickwickian Syndrome



 1957 – William C. Dement, coined the Father of Sleep Medicine, describes REM sleep in the cat

- 1959 Michel Jouvet describes REM sleep atonia in cats
- 1960 Allan Rechtschaffen explores psychophysiology of dreams
- 1960 Gerry Vogel describes sleep-onset REMs in narcolepsy
- 1961 Precursor of the Sleep Research Society is formed
  - Later becomes the Association for the Psychophysiological Study of Sleep (APSS)
  - Ultimately became Sleep Research Society

- 1963 Richard Wurtman's group reports that melatonin synthesis in the pineal gland is controlled by light
- 1964 Association for the Psychophysiological Study of Sleep is founded
  - Became precursor for Association of Sleep
     Disorders Centers (1975), American Sleep
     Disorders Association (1987) and ultimately
     American Academy of Sleep Medicine (1999)
- 1967 Lawrence Monroe reports physiologic findings between good and poor sleepers, precursor to hyperarousal state

- 1969 Allan Rechtschaffen and Anthony Kales produce a sleep scoring manual, which formed the basis of sleep staging for most research for the next 40 years
- 1970 William Dement founds world's first sleep disorders center at Stanford University
- 1972 The Rimini Symposium on Hypersomnia and Periodic breathing, held in Italy, is first major conference in which sleep apnea is main focus. Term "sleep apnea" had not been coined yet.

- 1974 Hewlett-Packard introduces first fiberoptic-based ear oximeter.
- 1974 Dr. Jerome Holland of the Stanford group gives the name "polysomnography" to sleep disorder diagnostic test
- 1976 American Sleep Disorders Association is established. Name later changed to American Academy of Sleep Medicine.
- 1976 Fred Turek and Michael Menaker establish that treatment with melatonin can alter the circadian clock of sparrows, laying foundation for use of melatonin and melatonin agonists as chronobiotic drugs today



- 1976 Mary Carskadon and colleagues report large difference between subjective and objective measures of sleep in insomniacs and showed that arousals increase with age
- 1976 *Sleep Apnea Syndrome* is published
  - Contains papers presented at first
     American symposium on the disorder
  - Term sleep apnea first introduced by Christian Guilleminault's team in 1975
- 1976 Charles Czeisler describes 24hour cortisol secretory patterns in humans

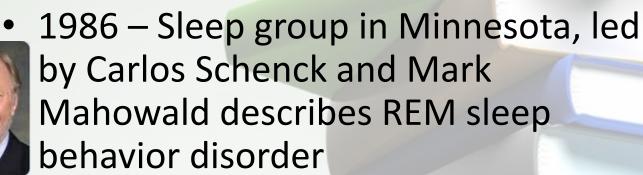


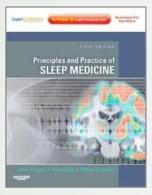


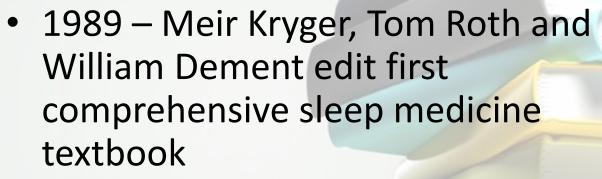
- 1978 First issue of journal Sleep is published, with Christian Guilleminault as editor
- 1981 Colin Sullivan describes use of nasal CPAP in article in *The Lancet*
  - CPAP revolutionized treatment of sleep apnea, which up to that time was treated surgically, usually with tracheostomy
  - Other advancements made by David Rapoport and Mark Sanders
- 1982 National Institutes of Health holds the first consensus symposium conference on insomnia













- 1990 Michael Thorpy spearheads creation of first International Classification of Sleep Disorders
- 1990 National Sleep Foundation is established in the US

- 1993 US legislation establishes Center for Sleep Disorders Research at National Institutes of Health
- 1993 Terry Young shows for first time that sleep apnea found to be extremely common among males. For first time high prevalence among females was shown. Up to that time, thought that sleep apnea was rare among females.
- 1995 Nathaniel Kleitman gives a lecture at age 100 at the APSS



- 1996 AASM granted a seat in House of Delegates of American Medical Association
- 1997 Modafinil shown to be effective as a stimulant. Ultimately found to be an efficacious treatment in narcolepsy and later in excessive sleepiness.
- 1997 Takahashi, Pinto, and Turek discover and clone first mammalian circadian clock gene, called *Clock*. Soon many circadian genes were found to be similar in flies, mice, and humans.

• 1999 – Eve Van Cauter demonstrates that sleep restriction can induce, in otherwise healthy people, physiologic and endocrine changes indicative of early sign of insulin resistance. This led to many studies to investigate relationship between chronic partial sleep loss and obesity, diabetes, and cardiovascular disease.

 1999 – Following 10-year search, Emmanuel Mignot's group found that familial narcolepsy in dogs was due to mutations in the hypocretin receptor-2. Shortly after, a discovery at Stanford University found that most cases of human narcolepsy, with cataplexy and HLA-DQB1\*0602 positive, are associated with a loss of hypocretin peptide in the cerebrospinal fluid and the brain

# Sleep Seen Through the Eyes of History

- Sleep has been expressed in art and literature throughout history
- Something that has been difficult for mankind to understand the functionality of
- Many misconceptions of what happens during sleep
- Only in the last 50 years has sleep's function begun to be understood

# Sleep in Art and Literature

- Depicted through mythology
- Depicted through religion
- Depicted as a state of rest
- Depictions of dreams, danger and death
- Viewers familiar with the stories behind the characters

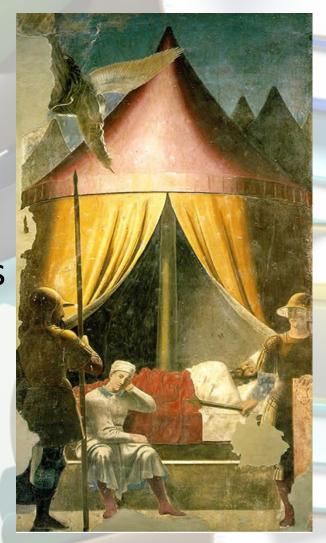
# Sandro Botticelli – Mars and Venus



- Sleeping Mars depicted as incapacitated, exposed, and vulnerable
- Likens sleep to weakness and humiliation

# Piero della Francesca - Dream of Constantine

 Sleep depicted as state in which divine communicates with humans and the sleeper can realize higher states of consciousness



# Jan Steen - The Dissolute Household



- The eye is immediately drawn to woman at the table in restful sleep amongst chaos
- Sleep seen as rest

## Jean-Francois Millet - Noonday Rest



 Sleep seen as wondrous healing state of escape, rest and comfort – common artistic theme

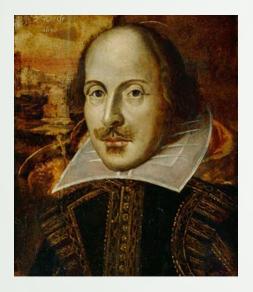
# Berthe Morisot – The Cradle



 Contrasts sleep and awake, unaware and alert, empowered and weakened

# William Shakespeare

- Equated sleep with death
- Hamlet, act 3, scene 1:



To be, or not to be: that is the question. Whether 'tis nobler in the mind to suffer The slings and arrows of outrageous fortune, Or to take arms against a sea of troubles. And by opposing end them? To die: to sleep: No more, and by a sleep to say we end The heartache and the thousand natural shocks That flesh is heir to, --- 'tis a consummation Devoutly to be wish'd. To die, to sleep, To sleep: perchance to dream: ay, there's the rub: For in that sleep of death what dreams may come?

## John Keats

Equated sleep with rest

WHAT is more gentle than a wind in summer? What is more soothing than the pretty hummer That stays one moment in an open flower, And buzzes cheerily from bower to bower? What is more tranquil than a musk-rose blowing In a green island, far from all men's knowing? More healthful than the leafiness of dales? More secret than a nest of nightingales? More serene than Cordelia's countenance? More full of visions than a high romance? What, but thee Sleep? Soft closer of our eyes!

## John Keats

Low murmur of tender lullabies!

Light hoverer around our happy pillows!

Wreather of poppy buds, and weeping willows!

Silent entangler of a beauty's tresses!

Most happy listener! When the morning blesses

Thee for enlivening all the cheerful eyes

That glance so brightly at the new sun-rise.

- From Sleep and Poetry