## **PUBERTY**

**Reproduction Block** Physiology Lecture # 4 (Puberty) Prof. Abdelfattah A. Alhader **Department of Physiology College of Medicine King Saud University** 

#### **PUBERTY**

A stage of human development when sexual maturation and growth are completed and result in ability to reproduce.

Accelerated somatic growth

Maturation of *primary sexual characteristics* (gonads and genitals)

Appearance of secondary sexual characteristics (pubic and axillary hair, female breast development, male voice changes,...)

Menstruation and spermatogenesis begin

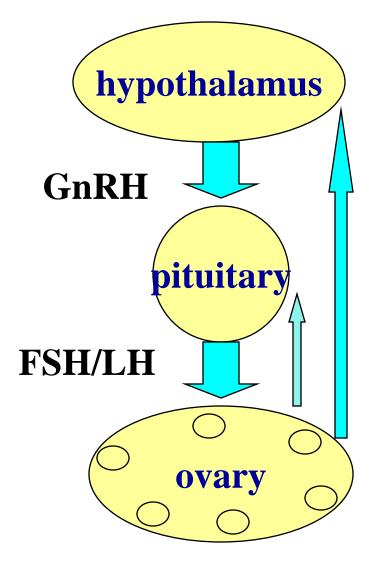
#### Puberty – Terms & Events

- Thelarche: development of breast
- Puberache: development of axillary & pubic hair
- Menarche: the first menstrual period
- Adrenarche: the onset of an increase in the secretion of androgens

Hormonal changes procede physical changes Increased stimulation of hypothalamo-pituitarygonadal axis

- gradual activation of the GnRH (LHRH)
- increases frequency and amplitude of LH pulses.
- gonadotropins stimulate secretion of sexual steroids (estrogenes and androgenes)
- extragonadal hormonal changes (elevation of IGF-I, and adrenal steroids)

#### Hypothalamic-Pituitary-Gonadal Axis



Steroidal & Non-steroidal hormones

Nocturnal GnRH pulsatility (LH secretion) precedes phenotypic changes by several years

First phenotypic changes:

breast development / testicular enlargement

- in young children, LH and FSH levels insufficient to initiate gonadal function
- between 9-12 yrs., blood levels of LH, FSH increase
- amplitude of pulses increases, especially during sleep
- high levels of LH, FSH initiate gonadal development

- GH secretion from pituitary also increases
- TSH (thyroid stimulating hormone) secretion from pituitary increases in both sexes:
  - increases metabolic rate
  - promotes tissue growth

#### Puberty – Female hormonal changes

- surge of LH release initiates 1st ovarian cycle
- usually not sufficient to cause ovulation during 1st cycle
- brain and endocrine systems mature soon thereafter
- estrogen levels in blood increase, due to growing follicles

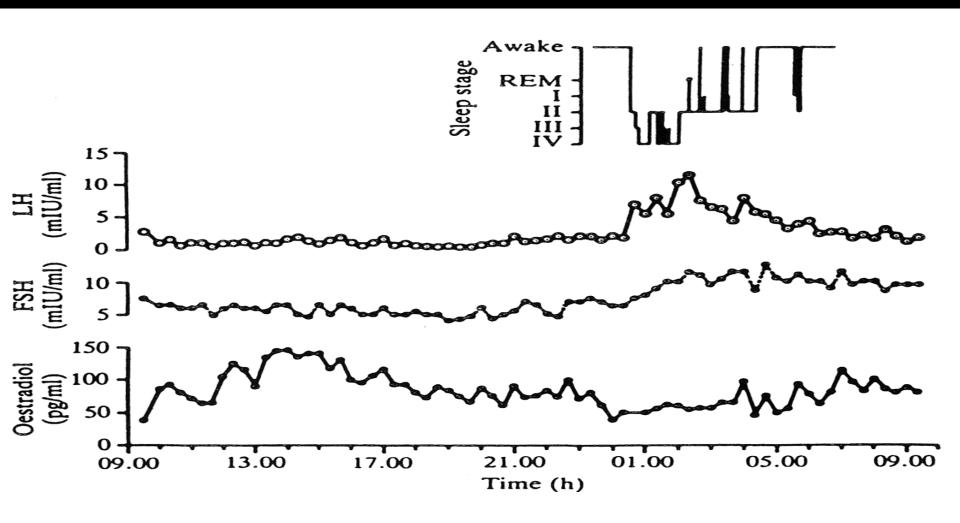
#### Puberty – Female hormonal changes

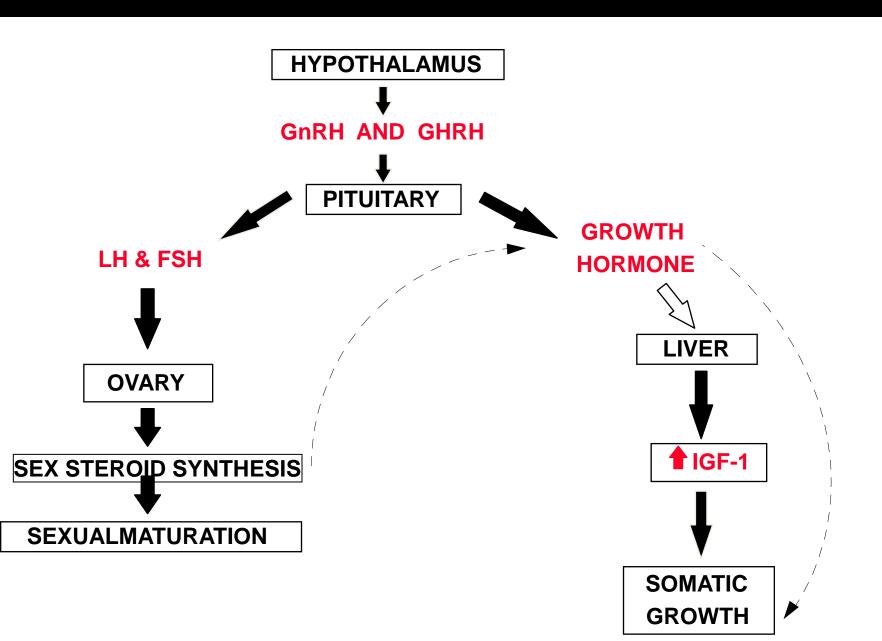
- estrogen induces secondary sex characteristics:
  - growth of pelvis
  - deposit of subcutaneous fat
  - growth of internal reprod. organs, external genitalia
- androgen release by adrenal glands increases >
  growth of pubic hair, lowering of voice, growth of
  bone, increased secretion from sebaceous glands

- LH and FSH release increases ~10 yrs. of age
- spermatogenesis; androgen secretion
- adrenals also secrete androgens
- androgens initiate growth of sex accessory structures (e.g. prostate), male secondary sex characteristics (facial hair, growth of larynx)

- androgens causes retention of minerals in body to support bone and muscle growth
- Sertoli cells also secrete some estrogen

# Sleep dependent nocturnal rise in LH





# Staging of pubertal development (Tanner)

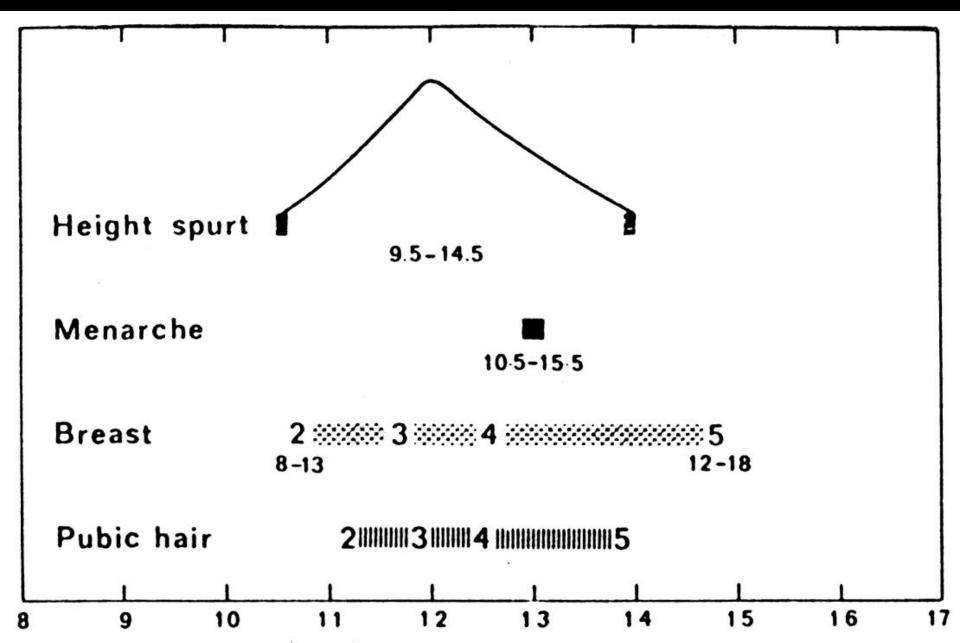
# Pubertal development is classified according to the Tanner standard – 5 different stages

- Girls:  $b\underline{reast}$  ( $B_{1-5}$ ), pubic hair ( $Pu_{1-5}$ ), axillary hair ( $A_{1-5}$ ), menarche
- Boys:  $testicular\ volume > 4\ ml\ (Te)$ , penis enlargement  $(G_{1-5})$ , pubic hair  $(Pu_{1-5})$ , axillary hair  $(A_{1-5})$ , spermarche

#### Monitoring of the pubertal growth acceleration

- growth velocity is 2-3 times greater than prepubertal
- sexual dimorfism in pubertal growth

#### Sequence of normal puberty in girls



### Normal pubertal development

	Boys	Girls
Age of start	12,5	11,5
(yrs)	(10 - 14)	(9 - 13)
First sign of puberty	G2 (testicular volume up to 4 ml)	B2
Growth velocity (cm/yr)	10,3 (Tanner III-IV)	9,0 (Tanner II-III)
Duration of puberty (yrs)	$3,2 \pm 1,8$ (adult size of testis )	2,4 ± 1,1 (menarché)

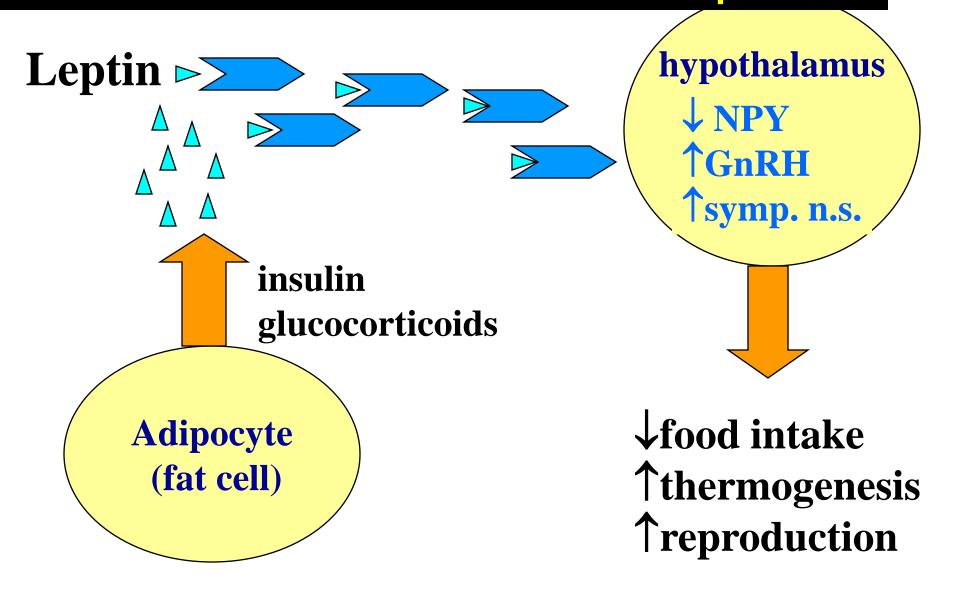
### Timing of Puberty

- trend toward earlier puberty exists within Western Europe and USA
- examination of lifestyle changes may give clues regarding mechanisms inducing onset
- one of the contributing factors: nutrition

#### **Nutrition**

- Critical body weight must be attained before activation of the reproductive system".
- even though age of menarche is decreasing, the average body weight of menarche remains the same
- earlier puberty due to improvement of nutrition, living conditions, healthcare?
- evidence supporting hypothesis:
  - obese girls go through early menarche
  - malnutrition is associated with delayed menarche
  - primary amenorrhea common in lean female athletes
  - "bodyfat" setpoint very noticeable in girls with fluctuating body weight due to anorexia nervosa

# Potential involvement of Leptin:



#### **Pubertal disorders**

A. Precoccious puberty

B. Delayed puberty

#### PRECOCIOUS PUBERTY

Precocious onset of puberty is defined as occurring younger than 2 SD before the average age

Girls <8 years old Boys <9 years old

- 1. Gonadotrophin-dependent (true / central )
- 2. Gonadotrophin-independent

# Gonadotrophin-dependent precocious puberty

- (true / central)
- Intra-cranial lesions (tumours, hydrocephalus, CNS malformations
- Gonadotrophin secreting tumours v. rare

# Gonadotrophin-independent precocious puberty

- Precocious pseudopuberty
- No spermatogenesis or ovarian development
- FSH & LH suppressed
- Congenital adrenal hyperplasia (CAH)
- Sex steroid secereting tumours
  - adrenal or ovarian

#### **Delayed puberty - definition**

#### Initial physical changes of puberty are not present

- by age 13 years in girls
   (or primary amenorrhea at 15.5-16y)
- by age 14 years in boys

#### Pubertal development is inappropriate

the interval between first signs of puberty and menarche in girls/completion genital growth in boys is > 5 years

# Causes of delayed puberty

- Gonadal failure (Hypergonadotrophic hypogonadism)
  - Turner's Syndrome
  - Post-malignancy chemo / radiotherapy / surgery
  - Polyglandular autoimmune syndromes
- Gonadal deficiency
  - Congenital hypogonadotrophic hypogonadism (+anosmia)
  - Hypothalamic/pituitary lesions (tumours, post-radiotherapy)
  - Rare gene mutations inactivating FSH/LH or their receptors

#### Turner syndrome

Karyotype 45,X (45,X/46,XX, structural abnormalities of X chromosome)

Short stature (final height 144-146 cm)

Gonadal dysgenesis

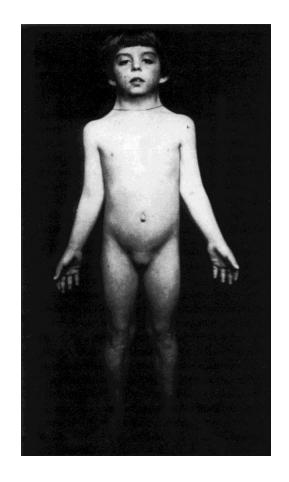
Skletal abnormalities

Cardiac and kidney malformation

Dysmorfic face

No mental defect Impairment of cognitive function)

Therapy: growth hormone, sex hormone substitution



H. Tuner, 1938