

King Khalid University Hospital
Department of Obstetrics & Gynecology
Course 482

AMENORRHOEA
Primary & Secondary



Objectives

- Define primary and secondary amenorrhea
- Explain the pathophysiology of amenorrhoea and identify the following types of primary amenorrhoea.
 - Amenorrhea with no breast development and sexual infantilism
 - Amenorrhea with breast development and müllerian anomalies
 - Amenorrhea with breast development and normal müllerian structures




- Explain the pathophysiology and identify the etiologies of secondary amenorrhea including:
 - • Pregnancy
 - • Hypothalamic causes
 - • Pituitary causes
 - • Ovarian causes
 - • Uterine causes
 - • Hyperandrogenism
- Describe the symptoms and signs of amenorrhea
- Outline a plan for investigation and management of amenorrhoea

PRIMARY AMENORRHOEA

1. No menstruation by the age of 14 years accompanied by failure to grow properly or develop sec. sexual characteristics.
2. No menstruation by age of 16 when growth and sexual development are normal.

SECONDARY AMENORRHOEA

- ★ Secondary absence of menses for six months (or greater than 3 times the previous cycle interval) in a women who has menstruated before.
 - ★ Pregnancy, lactation or hysterectomy must be excluded
 - ★ Prepubertal and post-menopausal conditions are also to be excluded as physiological causes
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CLINICAL APPROACH

There is a difference of opinion about the age at which **Primary Amenorrhoea** should be investigated → 18 yrs. often suggested.

Provided the patient has developed normal sec. sex. Characteristics and **cryptomenorrhoea** has been excluded.

While those patient with Primary amenorrhoea and sexual infantilism should be investigated at \cong age of 15 years or 16 years (may be earlier).

- Accurate, adequate **history** is essential to reach a firm diagnosis
- Specific questioning is necessary to establish diagnosis of Primary or Secondary amenorrhoea
- Is the amenorrhoea truly secondary (e.g. prev. menses were actually steroid – induced)
- Careful physical **examination** aids in reaching a fairly firm provisional diagnosis
- In minority, there is a need to go beyond simple out-patient **investigation.**



CAUSES OF AMENORRHOEA

- A. Disorder of outflow tract and or uterus
- B. Disorders of ovary
- C. Disorders of Ant. Pituitary
- D. Disorders of Hypothalamus



A. DISORDERS OF OUTFLOW TRACT & OR UTERUS

1. CRYPTOMENORRHOEA

Vaginal **atresia** or **imperforate hymen** → prevent menstrual loss from escaping.

FEATURES:

- ★ Prim. Amenorrhoea in a teenage girl with normal sexual development present

Complaining of:

- Intermittent lower abd. pain
- Possible difficulty of mict.
- Palpable lower abd. swelling (Haematometra)
- Bulging, bluish membrane at lower end of vagina (Haematocolpus).

MANAGEMENT:

INCISE MEMBRANE

2. ABSENCE OR HYPOPLASIA OF VAGINA:

FEATURES:

- ☛ Growth, develop, and ovarian function are usually normal.
- ☛ Uterus may be normal or rudimentary
- ☛ Renal anomalies (in 30%) or skeletal defects (in 10%) may be present.

MANAGEMENT:

Create a functional vagina by surgery or dilators

3. TESTICULAR FEMINIZATION: (Androgen Insensitivity)

- ✦ Phenotype is woman. Genotype is man (xy) → **testes** are present.
- ✦ Inherited by an X-linked recessive gene... (familial)
- ✦ Resulting in absence of cytosol androgen receptor

FEATURES:

- i. Growth and develop are normal (may be taller than average).
- ii. Breasts are large but with sparse glandular tissue and pale areola
- iii. Inguinal hernia in 50% of cases
- iv. Scanty, or no axillary and pubic hair
- v. Labia minora underdeveloped
- vi. Blind vagina, absent uterus, rudimentary fallopian tubes
- vii. Testes → in abd. or inguinal canal
- viii. **Normal levels of testosterone** are produced.. But no response to androgens (endog. or exogen)
- ix. No spermatogenesis
- x. There is ↑ incidence of testicular neoplasia (50%)

CONSIDER THE DIAGNOSIS IN A FEMALE CHILD:

1. With inguinal hernia
2. With 1^o amenorrhoea and absent uterus
3. When body hair is absent

MANAGEMENT:

- These patients are female.
- The gonads must be removed after puberty
→ then HRT started
- Rare cases of **incomplete test. feminization** do occur → have variable degrees of masculinization

4. ASHERMAN'S SYNDROME:

Sec. amenorrhoea following destruction of the endomet. by overzealous curettage → multip. Synechiae show up on “**Hystero**graphy”.

MANAGEMENT:

Under G.A. → breakdown intraut. Adhesions through hysteroscope → insert an IUCD to deter reformation → hormone therapy ($E_2 + P$)

5. INFECTION

e.g. Tuberculosis. Ut. Schistosomiasis



B. DISORDERS OF THE OVARIES

1. CHROMOSOMAL ABNORMALITIES

Turner's syndrome (45 x 0)

→ gonadal dysgenesis

FEATURES:

- i. Amenorrhoea (1^o, rarely 2^o)
- ii. Short stature
- iii. Failure of sec. sex. Develop
- iv. Webbing of the neck→
- v. ↑ carrying angle
- vi. Shield chest
- vii. Coartution of aorta
- viii. Renal collecting syst. defects

☛ **Streak ovaries present**

☛ **Gonadotrophins ↑↑**

☛ **↓ Estrgoens**

→ **Mosaic Chrom. Pattern**

(e.g. XO/XX) → lead to various degrees of gonadal dysgenesis and sec. amen. + premature menopause

→ If Y-Chrom is present in the genotype → risk of gonadal malig. makes gonadectomy advisable

2. GONADAL AGENESIS:

(Failure of gonadal develop):→ no other cong. abn.

3. RESISTANT OVARY SYNDROME

- ☛ A rare condition
- ☛ Normal ovarian develop and potential
- ☛ FSH ↑↑
- ☛ It may resolve spontaneously
- ☛ If hot flushes → Rx. With estrogen

4. PREMATURE MENOPAUSE:

Ovarian failure....due to

- i. Auto-immune dis. (associated with Addison's dis. ??)
- ii. Viral infection (e.g. mumps)
- iii. Cytotoxic drugs

5. PCOs:

- ☞ Mostly present with classical Stein-Leventhal syndrome (of oligomenorrhoea, obesity, hirsutism, and infertility)
- ☞ However a substantial group will have sec. amenorrhoea with no obesity or hirsutism
- ☞ Diagnosis is made by finding \uparrow LH/FSH ratio
- ☞ Confirmation is made by laparoscopy.
- ☞ USS \pm

C. DISORDERS OF PITUITARY

1. Pituitary Tumor causing “Hyperprolactinemia”

≅ 40% of women with hyperprolactinemia will have a pituitary adenoma

Pit. Fossa XR is necessary in all cases of amenorrhoea – particular 2^o.

FEATURES: In coned view:

- ★ Erosion of clinoid process
- ★ Enlarge of pituitary fossa
- ★ Double flooring of fossa

- If any of above features seen
- CT scan or MRI + Assessment of visual fields

MANAGEMENT:

- ★ **Bromocriptine (Dopamine agonist)**
 - **Suppres prolactin sec.**
 - **Correct estrogen deficiency**
 - **Permits ovulation**
 - **↓ Size of most prolactinomas**

- ★ **Surgical removal of tumor→**
 - ☞ **if extracellular manifestation (e.g. press. on optic chiasma) or if patient cannot tolerate or respond to medical Rx.**

2. OTHER CAUSE OF ↑ PROLACT.

- ♣ Drugs: e.g. phenothiazines, methyl-dopa, metoclopramide, anti-histamines, oestrogens and morphine.

3. CRANIOPHARYNGIOMA

- ♣ Other intracranial tumor

4. SHEEHAN'S SYNDROME

- ♣ Necrosis of ant. pituitary due to severe PPH
 - Pan – or partial hypopituitarism
- ♣ It is rare problem today due to better obstetric care and adequate blood transfusion

D. DISORDERS OF HYPOTHALAMUS

- ♣ Commonest reason for hypogonadotropic sec. amenorrhoea
- ♣ Often associated with stress e.g. in migrants, young women when leave home, university students
- ♣ Diagnosis by exclusion of pituitary lesions.
- ♣ Hormone therapy or ovulation induction is not indicated unless patient wishes to become pregnant

1. WEIGHT – LOSS ASSOCIATED AMENORRHOEA

A loss of > 10 kg is frequently associated with amenorrhoea

- i. In young women and teen ages girls become obsessed with their body image and starve themselves.
- ii. Jogger's amenorrhoea:
 - This is seen frequently in women training for marathon racing, in ballet dancers and other form of athletes.

CAUSES:

- ✧ + redistribution between proportion of body fat mass and body muscle mass.
 - ✧ May be also mediated by exercise related changes in β -endorphins
- iii. ANOREXIA NERVOSA
Associated with sec. amenorrhoea
(misnomer \rightarrow no loss of appetite)

2. AMENORRHOEA AND ANOSMIA:

rare cause of amenorrhoea of hypogonadotropic – hypo-gonadism.

(Counterpart in males is Kallman's syndrome)

POST-PILL AMENORRHOEA:

- ♣ There is no evidence that Est. prog. Contraceptive pills predispose to amenorrhoea.. once pill taking is ceased.
- ♣ An irregular men. cycle frequently precedes pill taking
- ♣ If this assumption of amenorrhoea being merely an after-effect of pill taking → many cases of hyperprolactinemia will be missed (1:5)
- ♣ And Premat. ovarian failure will be missed in 1:10 cases
- ♣ Once other causes are excluded, this type of ameno. Responds well to ovulation induction with Clomiphene citrate if preg. is desired.

INVESTIGATION OF AMENORRHOEA

1. S. Prolactin level and TFT
2. Karyotyping...if chrom. anomaly is suspected on clinical grounds
3. Progesterone withdrawal test....to check endog. estrogen.
e.g. Provera (medroxy-prog) → if bleeding PV=reactive endom. and patent outflow tract.

- If **PRL** is norm. + no galactorrhoea ---no need for further investigation for pituitary tumor
- If **GALACTOR** is present → further evaluation of pit. gland is necessary .. regardless of level of PRL and menstrual pattern
- If PRL is signific. elevated (excluding stress) → Radiology exam of pituitary to exclude tum.
- Visual fields assessment – if X-Ray abnormal
- **FSH & LH** level... especially if no withdrawal bleeding following prog. Challenge.
- ↓ LH (<5 IU/ml) → hypogonadotropic- hypogonadism
- ↑ FSH (>40 IU/ml) on successive readings → ovarian failure

If women < 35 years = premet. ovar. failure (menopause) → check **karyotype**. (if Y-Chrom + → high risk of gonadal malignancy)

4. USS:

Of uterus and ovaries → can be useful to investigating and monitor Rx. Of these women



FLOW CHAR FOR INVESTIGATING OF SEC. AMENORRHOEA

