Route of administration

Progesterone Oral Incomplete effects

Side effects due to metabolites

Non-oral

Effective "physiological"

Progestin

Oral

Effective (genomic effects)

Variable side effects (non

genomic effects)

Non oral

Induces same side effects

- Molecule dependant
- Route independent

Progestins

- Progesterone: absorbed orally when micronized but poor bioavailability because of liver metabolism
- Synthetic progestins: man made molecules
 - Resist enzymatic degradation and remain active orally
 - Mimic effects on progesterone receptors (prevent hyperplasia)
 - Non genomic action unknown (side effects on central nervous system)
- Natural progesterone effective vaginally

Progestagens

- E2 alone induces endometrial hyperplasia and/or cancer
- Progesterone opposes the growth promoting effects of E2
 - Lowers E2 receptors
 - increases 17ß hydroxylase
- Hyperplasia and/or cancer never seen in ovulatory women
- Progestins completely revert risk

Estrogen treatment : Bone sparing doses

Estrogen preparations	Route of administration	Bone sparing or commonly used dose
Conjugated Equine Estrogen (Premarin®)	Oral	0.625 mg/24h
Micronized E2 (Estrace® and others)	Oral	1 - 2 mg/24h
E2 valerate (Progynova®)	Oral	0.01 mg/24h
Ethinyl-E2 (Estinyle® and others) (also constituant of most oral contraceptives)	Oral	0.05 mg/24h 3.5 or 7 day « patches »
Transdermal E2 « patches » (Estraderm® and other newer products)	Transdermal	1.5g of gel containing approx. 0.05 of E2

Estrogen therapy

• Daily production of E2 fluctuates through the cycle

• Early foll. phase 0.05 mg/24h

• Late foll. phase 0.5 mg/24h

• Luteal phase 0.3 mg/24h

• Should match early follicular phase levels and/or protect against osteoporosis

Introduction

- Menopause is a natural process
- Life expectancy after menopause has increased ("artificial" developments)
- Increase in life expectancy has lead to:
 - New expectations
 - New needs
- <u>Objective</u> ⇒ Treat symptoms
 - ⇒ Prevent consequences

High Risk Individuals

- First, "do not harm"
- When in doubt, opt for:
 - Physiological option