



Foundational Principles of Natural Hormone Replacement Therapy

Dr. Kristy A. Prouse MD, FRCSC (OB/GYN)

©January 2019 Institute for Hormonal Health Professional Development Division Inc.



Financial Disclosure: Nothing to Declare



Overview:

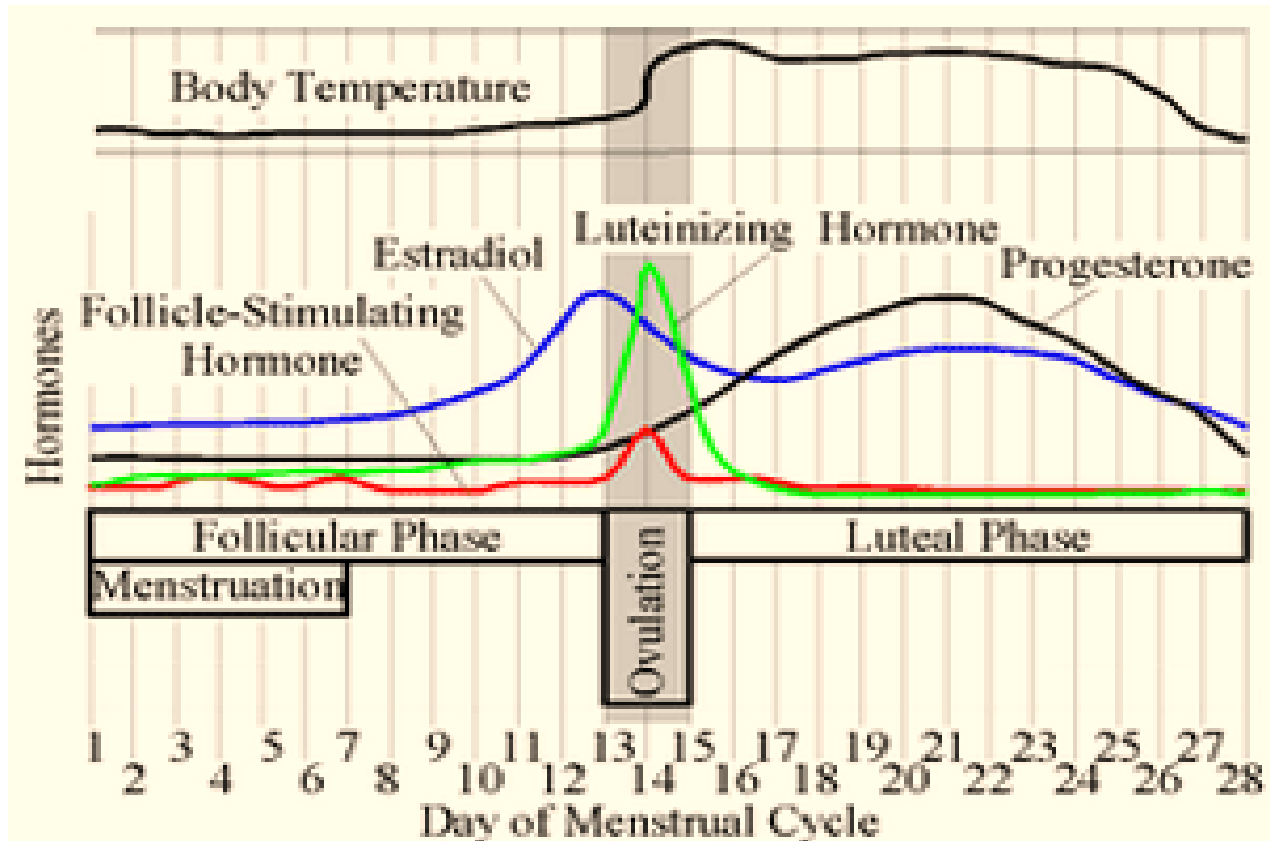
- Physiology of sex hormone imbalances
- How is symptomatology helpful in diagnosis and treatment?
- What you need to know about the conventional medical approach...
- ...and the best way to approach sex hormone deficiencies
- Conventional HRT vs BHRT: What's the deal?
- Principles of Prescribing
- The How-Tos of Practical Prescribing: Estrogen, Progesterone, Estriol
- When is it too much hormone? Symptoms of Over-replacement
- Troubleshooting dosing of BHRT
- When to refer the complicated patient



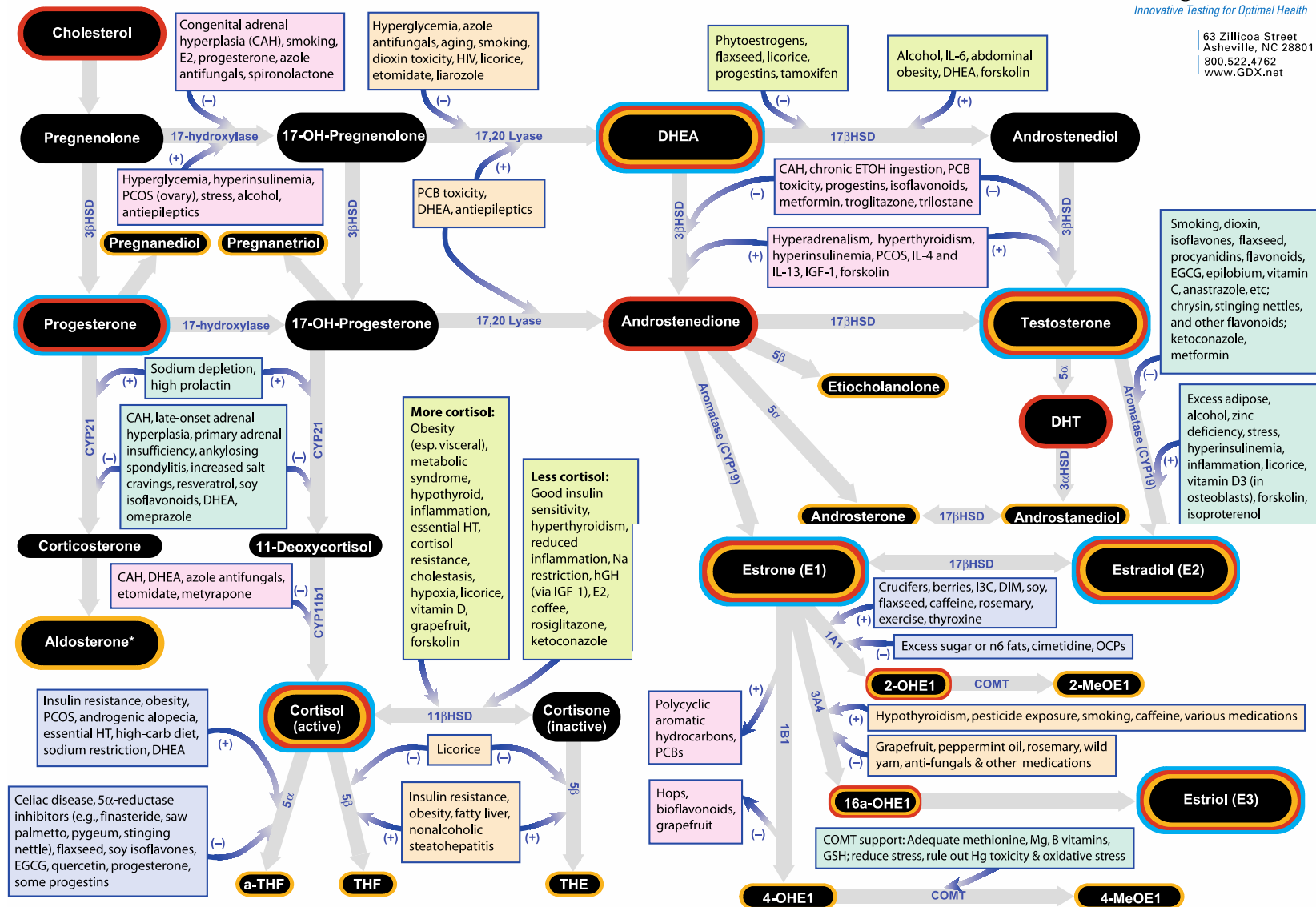
Physiology of Sex Hormones



Physiology of the Menstrual Cycle

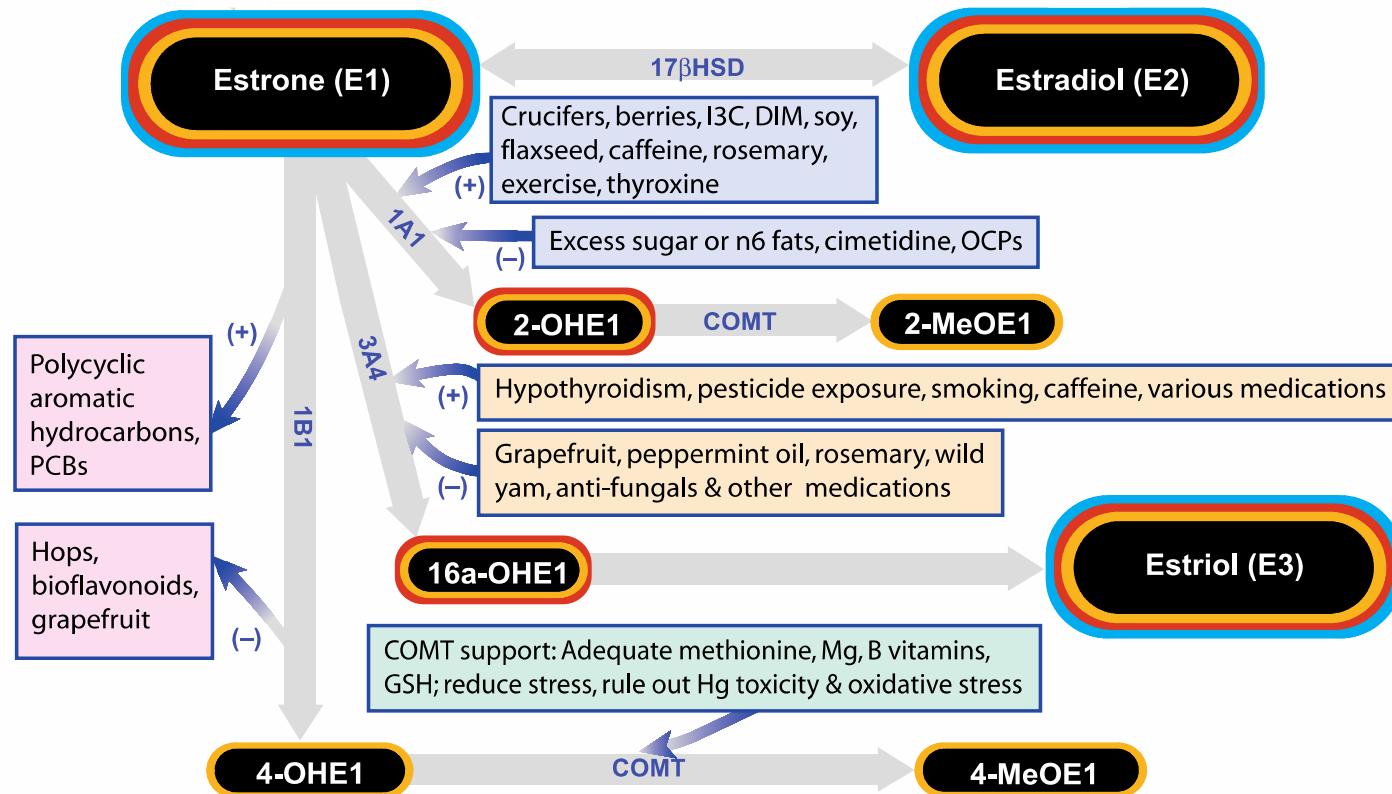


Steroidogenic Pathways



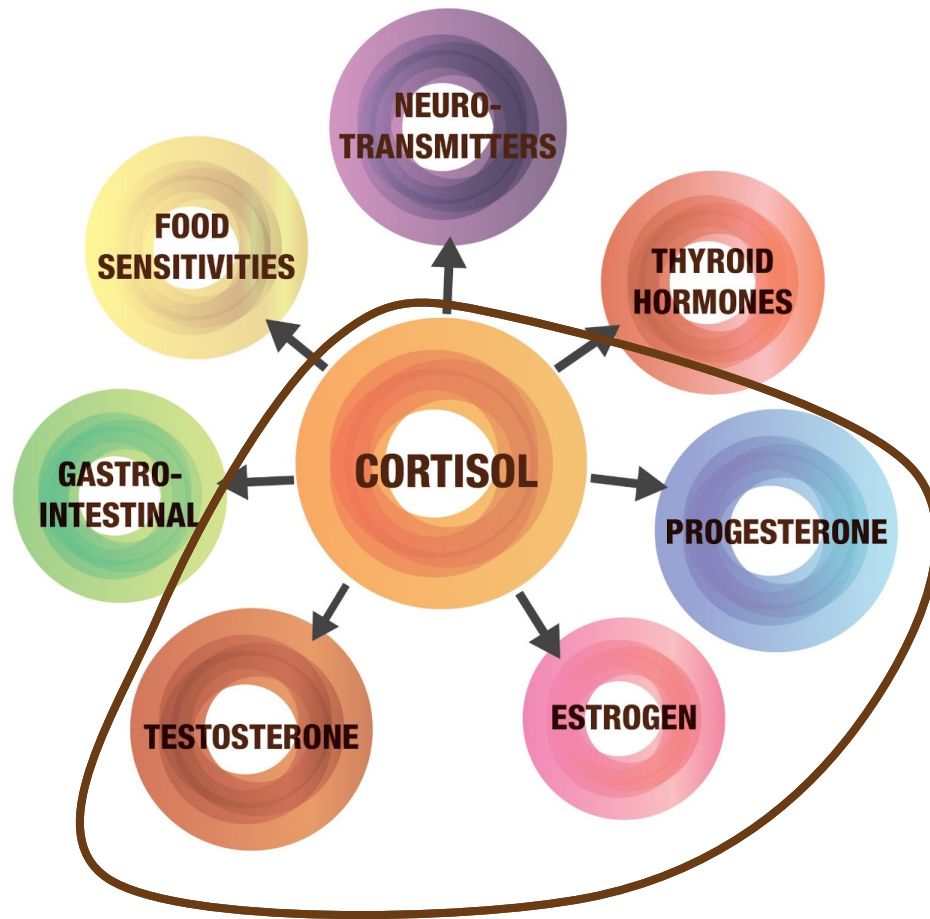


Estrogen Metabolism





Cortisol-Sex Hormone Connection





Classification and Symptomatology of Sex Hormone Imbalances



Sex Hormone: Classification Imbalance & Deficiencies

IMBALANCES

- Estrogen Dominance
- Relative Estrogen Dominance
- Elevated Androgens

DEFICIENCIES

- Estrogen
- Progesterone
- Testosterone





Sex Hormone Deficiencies: Symptomatology

LOW ESTROGEN/PROGESTERONE

- Hot flashes
- Memory decline
- Anxiety
- Insomnia
- Weight gain
- Vaginal dryness
- Painful intercourse
- Decreased libido
- Crawly skin
- Frequent bladder infections

LOW TESTOSTERONE (FEMALE)

- Decreased libido
- Decreased arousal/orgasm
- Decreased nipple sensation
- Loss of muscle mass
- Loss of vitality
- Decreased exercise tolerance





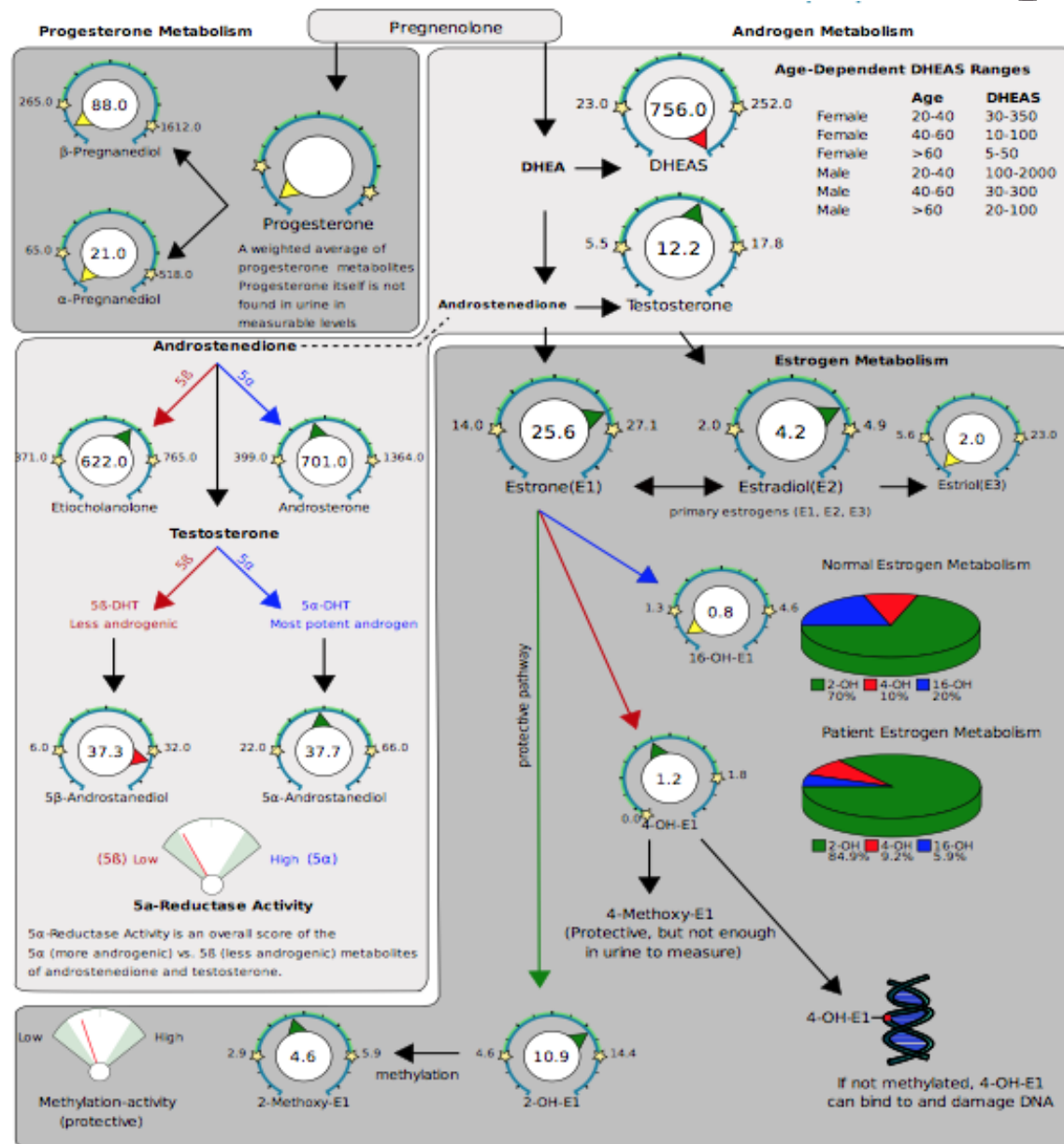
Diagnosis



Sex Hormones: Diagnosis

- Symptomatology: Imbalances/Deficiencies
- Physical Examination: breast and pelvic
- Saliva: historically gold standard for baseline
- Dried urine spot: baseline or for those on hormone replacement
- Bloodwork underestimates hormone levels

	Baseline Testing (no HRT)		With Hormone Replacement Therapy	
	Sex Hormones	Adrenal	Oral Progesterone	Transdermal Cream
Serum				
Saliva				
24 Hour Urine				
Dried Urine				
Notes on Dried Urine Testing	Dried urine testing is comprehensive and convenient.		Dried urine testing can offer insight into dosing.	Proceed with caution. No testing is highly reliable for creams. Urine may be best for high doses (>1mg E2, 200mg Pg, 500mg T) or with alcoholic gels and saliva for low doses of creams





Treatment Approach



Sex Hormone Imbalance: Treatment Approach

- Eliminating the Cause
 - Dietary/Lifestyle modifications
- Help the body to heal itself
 - Dietary/Lifestyle modifications
- Sex Hormone Support
 - Herbal Remedies
- Bioidentical Hormone Replacement (BHRT) vs. conventional HRT



Sex Hormone Imbalance: Treatment Approach

Herbal Remedies

ESTROGEN DOMINANCE

- Indole-3-Carbinol
- Calcium D-glucarate
- DIM

LOW ESTROGEN

- Black cohosh
- Borage oil
- Evening primrose oil

LOW PROGESTERONE

- Chasteberry
- Wild yam



ELEVATED TESTOSTERONE

- Progesterone or Chasteberry
- Saw palmetto
- Pygeum Bark
- Fenugreek
- Nettle Root

LOW TESTOSTERONE

- Maca
- Tribulus Terrestris
- Zinc
- L-arginine



Diagnosis and Treatment

CONVENTIONAL APPROACH

- No testing for hormone levels needed to prescribe
- *IF* tested (blood)-limitations
- No follow up testing with treatment
- No balancing of estrogen, progesterone, testosterone
- Progesterone not used if woman has hysterectomy
- Synthetic hormones commonplace

INTEGRATIVE MEDICINE APPROACH

- Always test hormone levels
- Dried urine hormone testing with 4 pt cortisol
- Follow up testing recommended (annually)
- Balancing of estrogen, progesterone, testosterone key
- Progesterone used if deficient even without a uterus
- Bio-identical hormones only



BHRT vs CHRT



Definitions

- **Bio-identical Hormones:** have a chemical structure identical to endogenous human hormones but are chemically synthesized such as progesterone, estriol, estradiol, and testosterone.
- **Non-bio-identical hormones:** are not structurally identical to human hormones and may either be chemically synthesized such as MPA or derived from a nonhuman source such as CEE.
- **Progesterone:** our own endogenous hormone.
- **Progestins:** synthetic chemicals that mimic the effects of progesterone by binding to progesterone receptors e.g., MPA
- **Progestogens:** an umbrella term for both progesterone and progestins.



Sex Hormone Imbalances: Treatment Approach

Bio-Identical Hormone Replacement Therapy (BHRT)

- BiEst (Estradiol/Estriol) TD
- Estriol pv
- Progesterone po/TD
- Testosterone TD
- 7-keto DHEA SL/TD
- DHEA SL/TD





Health Canada Approved Hormones

Type/Source	Brand Name	Bioidentical?
17 Beta-estradiol/plant	(Estrace) po/pv	Yes
17 Beta-estradiol/plant	(Esclim) TD patch	Yes
17 Beta-estradiol/plant	(Climara) TD patch	Yes
17 Beta-estradiol/plant	(Estraderm) TD patch	Yes
17 Beta-estradiol/plant	(Vivelle) TD patch	Yes
17 Beta-estradiol/plant	(Estrogel) TD gel	Yes
17 Beta-estradiol/plant	(Estrasorb) TD cream	Yes
17 Beta-estradiol/plant	(Estring) vaginal ring	Yes
Estradiol acetate	(Femring) vaginal ring	Yes
Estradiol hemihydrate	(Vagifem)vaginal tablet	Yes
Progesterone micronized	(Prometrium) po	Yes
Progesterone	(Procheive 4%) vaginal gel	Yes



Health Canada Approved Hormones

Type/Source	Brand Name	Bioidentical?
Conjugated equine estrogens	(Premarin)po/pv	No
Ethinyl estradiol	(Estinyl)po	No
Medroxyprogesterone acetate (MPA)	(Provera)po	No
Norgestrel	(Ovrette) po	No
Norethindrone	(Micronor) po	No
CEE and MPA	(Prempro) po	No
Ethinyl acetate and norethindrone acetate	(FemHRT) po	No
17 beta estradiol and norgestimate	(Combipatch) TD	No
17 beta estradiol and levonorgestrel	(Climara Pro) TD	No



Comparison of Bio-identical Hormones vs. Conventional Hormone Replacement

	Bio-identical Progesterone	Synthetic Progestins
Clinical Efficacy	best	good
Physiologic actions on breast tissue	protective	negative
Risk of breast cancer	protective	increased
Risk of CVD	protective	increased
Risk of DVT	none	none

	Bio-identical E2/E3-TD	Synthetic Estrogens
Breast cancer risk	protective	increased
DVT risk	none	increased



Prescribing Bio-identical Hormones

Principals of Prescribing BHRT

- 
-
- Start low and go slow
 - Balance
 - Physiologic ranges
 - Understand downstream hormones
 - Avoid down regulation of receptors
 - Conversion dosing TD->po (x4-5)
 - Perform baseline investigations



Baseline Investigations

- ✓ Trans-vaginal pelvic ultrasound (TVUS)
- ✓ PAP smear
- ✓ Mammogram or thermography (>50yo)
- ✓ Bone Mineral Density (BMD)(>50yo)

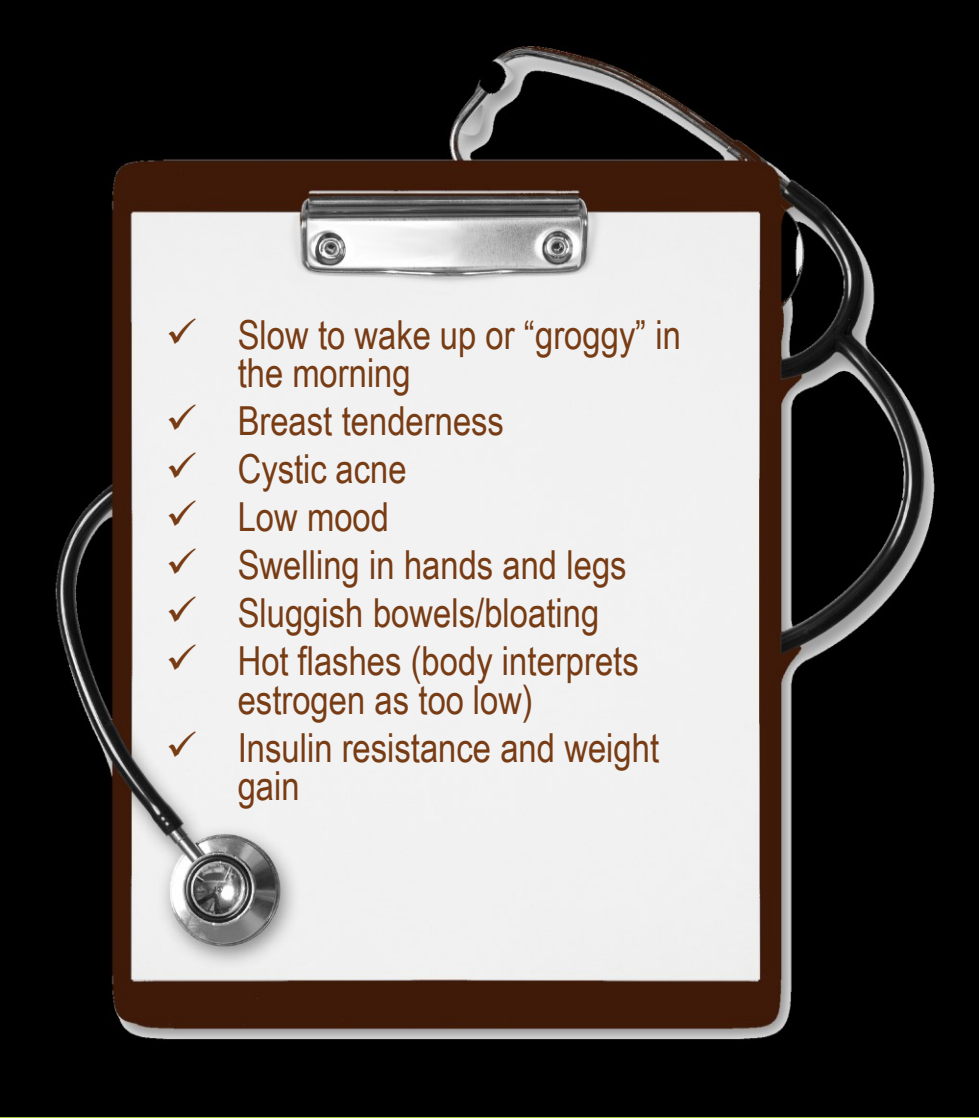


Progesterone



- Used to balance estrogen and to support cortisol production
- TD or po applications most common
- Prescribed orally for sleep/anxiety
- Given cyclically to menstruating woman
- Given daily (with a day of rest) to post-menopausal woman
- Initiate dose according to test results and titrate to symptoms

Over-replacement of Progesterone

- 
- ✓ Slow to wake up or “groggy” in the morning
 - ✓ Breast tenderness
 - ✓ Cystic acne
 - ✓ Low mood
 - ✓ Swelling in hands and legs
 - ✓ Sluggish bowels/bloating
 - ✓ Hot flashes (body interprets estrogen as too low)
 - ✓ Insulin resistance and weight gain

Typical Progesterone Dosing

Post-Menopausal



- Compounded progesterone 75-175mg SR po qhs Monday to Saturday

**increments of 25mg

- Compounded progesterone cream 10-30 mg TD qhs Monday to Saturday

**increments of 5mg

BiEst

- Used to replace estrogen when objectively deficient
- Combination of estradiol and estriol (20:80) or (50:50)
- Transdermal application only
- Given daily (with a day of rest) to post-menopausal woman
- Initiate dose according to test results and titrate to symptoms
- Must be balanced with progesterone



Over-replacement of Estrogen

- 
- Irritability/weepiness
 - Acne
 - Breast tenderness
 - Swelling
 - Post-menopausal vaginal bleeding

Typical BiEST Dosing

Post-menopausal

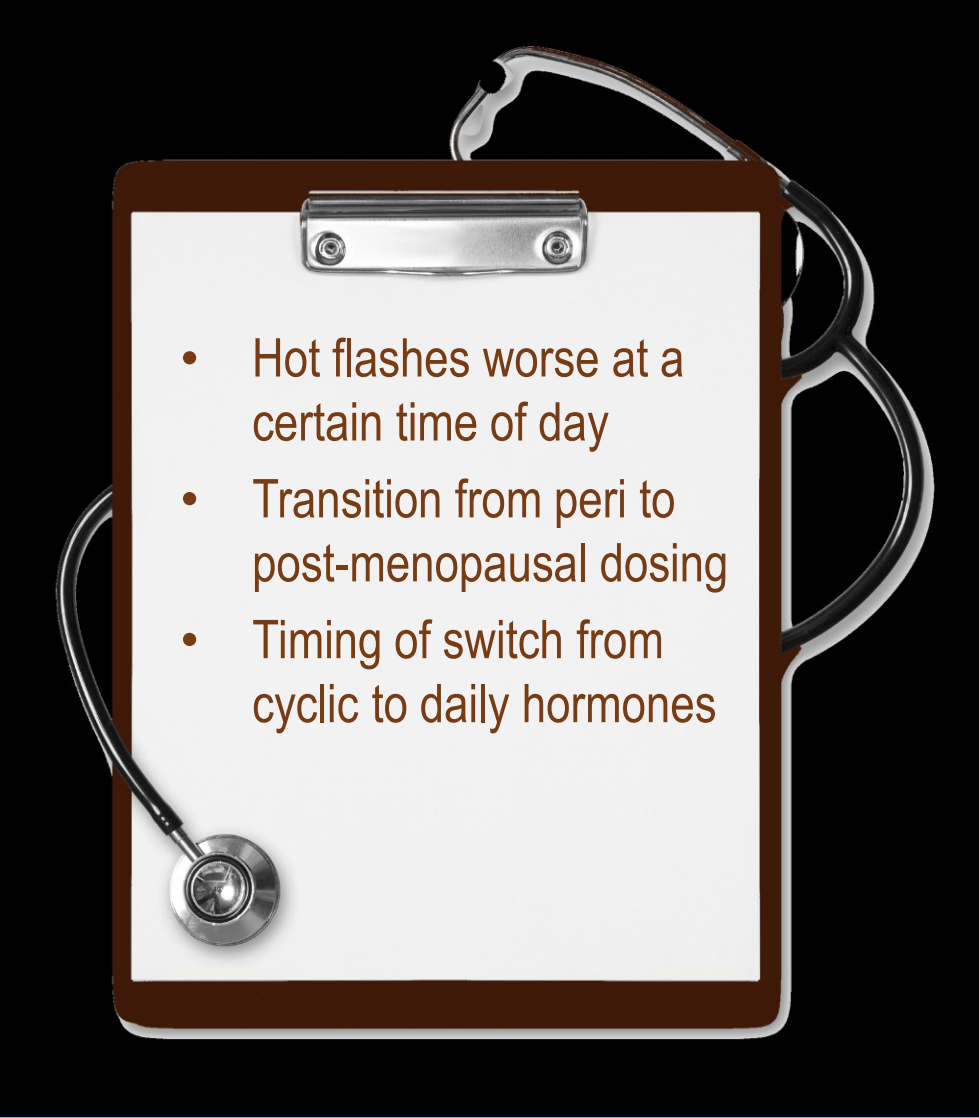


- Compounded BiEST cream 80:20 (E3/E2) 0.5-1.75mg TD BID Monday to Saturday

**Increments of 0.5mg

- Surgical menopause may require higher doses to manage symptoms

Troubleshooting Estrogen

- 
- Hot flashes worse at a certain time of day
 - Transition from peri to post-menopausal dosing
 - Timing of switch from cyclic to daily hormones

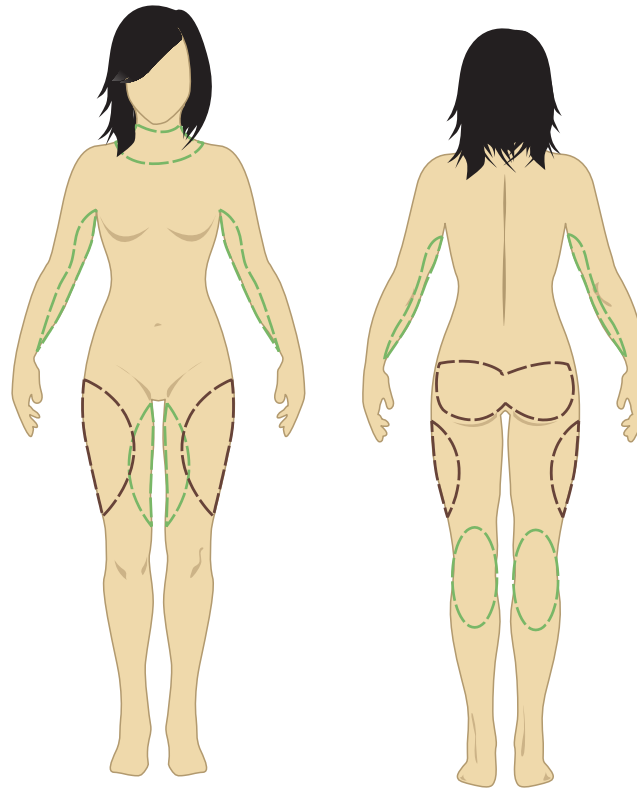
Estrogen/Progesterone Application Sites



Progesterone



Estrogen



Estriol for Vaginal Atrophy



- Used for vaginal atrophy resulting in bothersome vaginal dryness, painful intercourse or recurrent urinary tract infections
 - Transvaginal application
 - Administered in tapering doses at bedtime
 - May be transferred to sexual partner
 - May initiate without testing based on symptoms and physical examination
-
- Be aware of the modern strategies for urogenital atrophy treatment including laser therapy and photobiomodulation

Typical Vaginal Estriol Dosing

- 
- Compounded Estriol vaginal cream 0.25-0.5 mg qhs x 2 weeks, then twice weekly x 2 weeks, then once per week x 2 months and then as needed

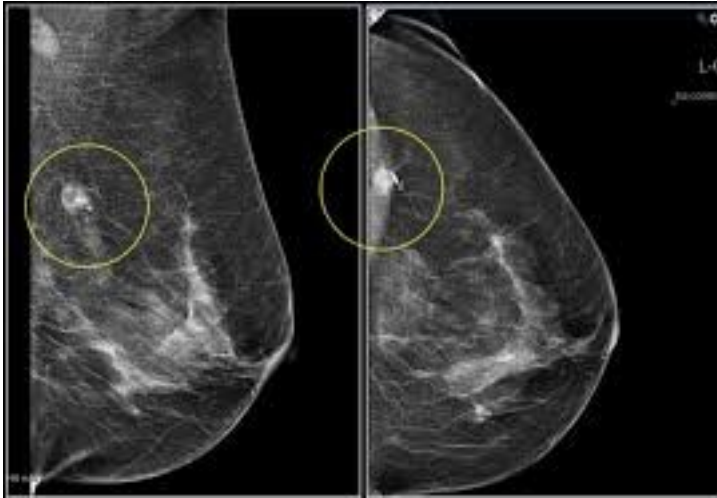
Delivery Methods

- Carrier bases
- Application site
- Suppositories (E_3)





The Complicated Patient: Know when to refer



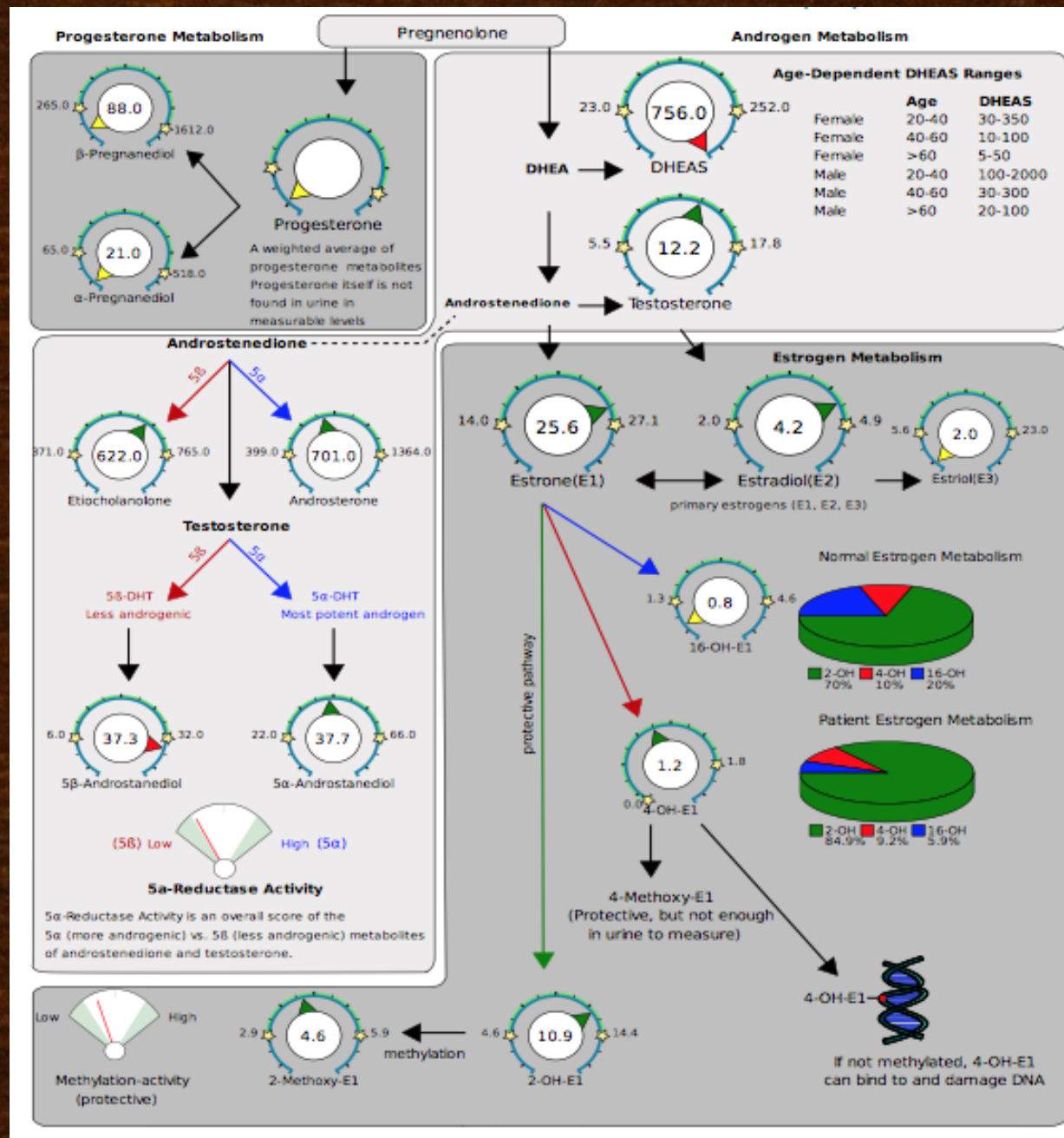
- ✓ Post menopausal bleeding
- ✓ Thickened endometrium on trans-vaginal pelvic ultrasound
- ✓ Breast cancer history
- ✓ Urogenital atrophy unresolved with systemic and local estrogen



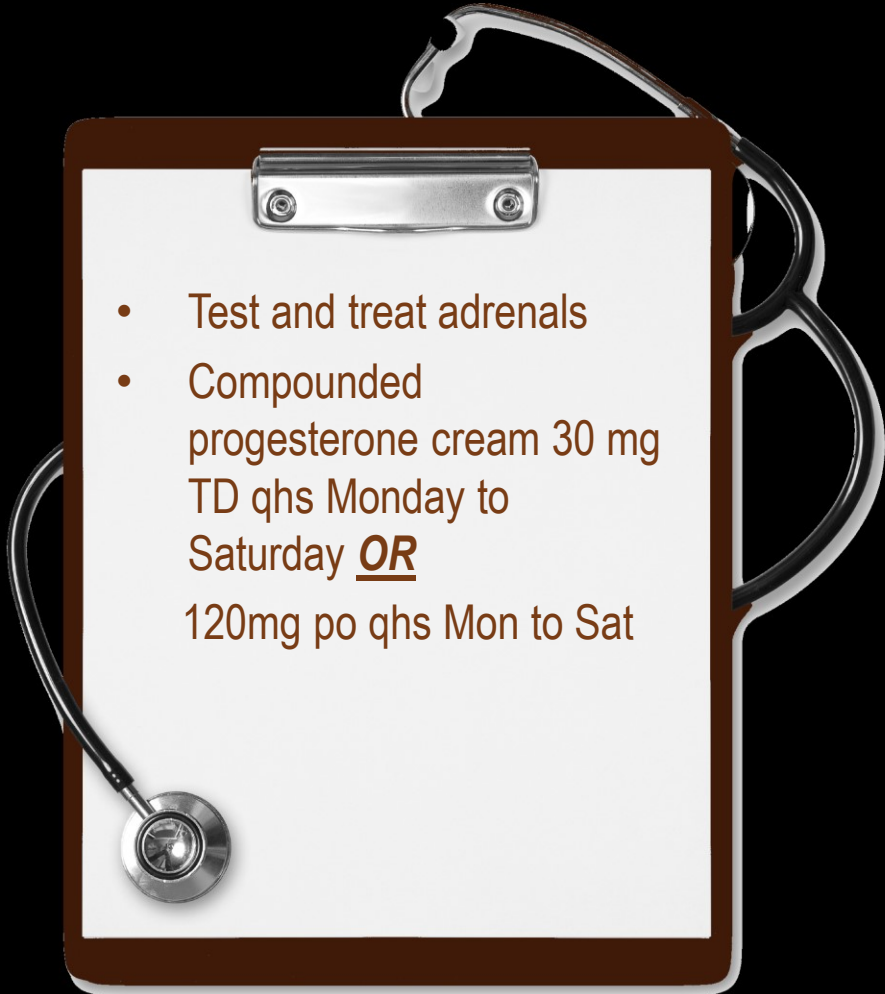
Sex Hormone Case Studies

CASE STUDY #1

47 yo peri-menopausal female not on any hormone replacement therapy complaining of heavy periods with breast tenderness in the week before her period. She is complaining of new onset anxiety and sleeplessness.

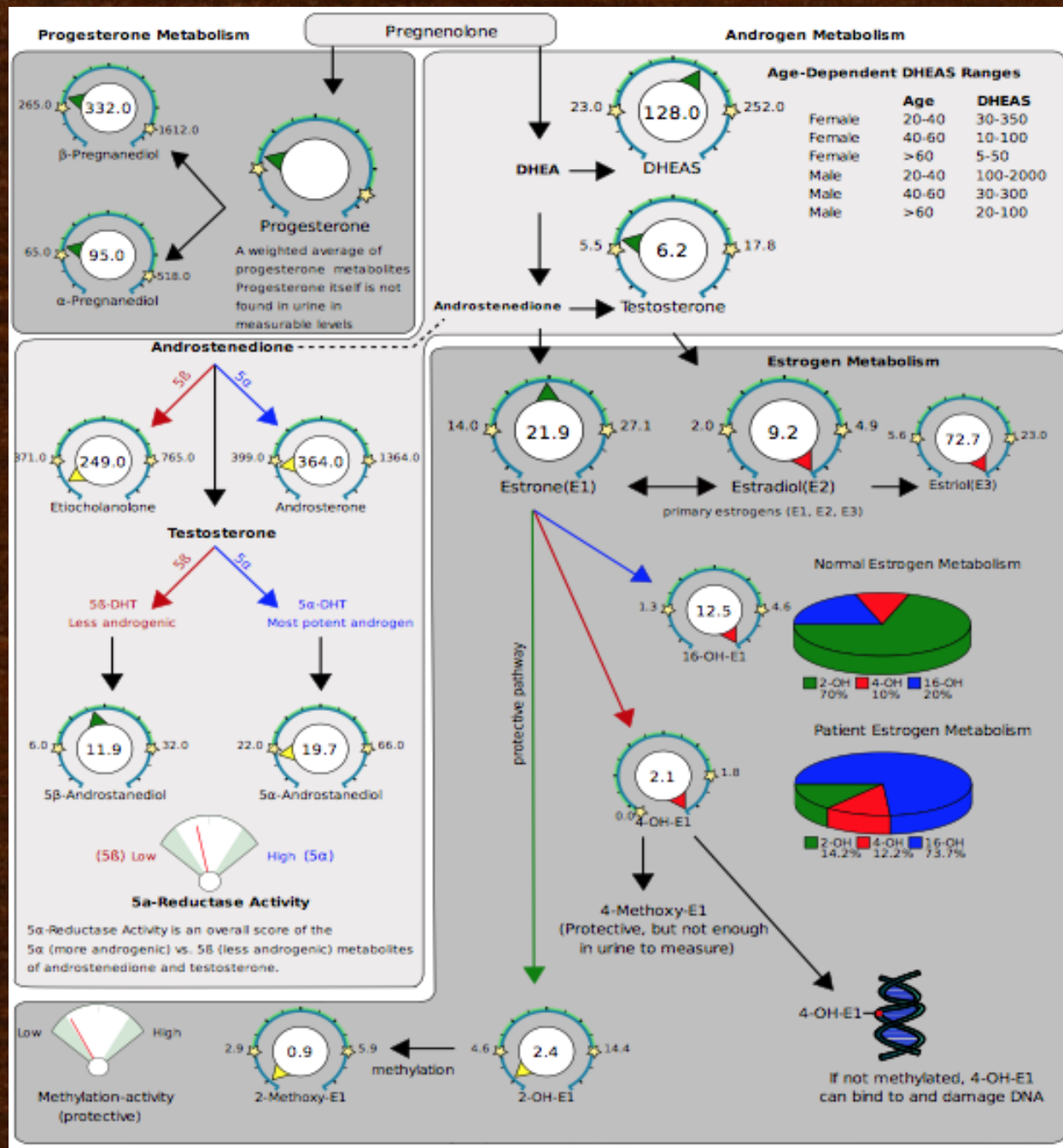


Relative Estrogen Dominance

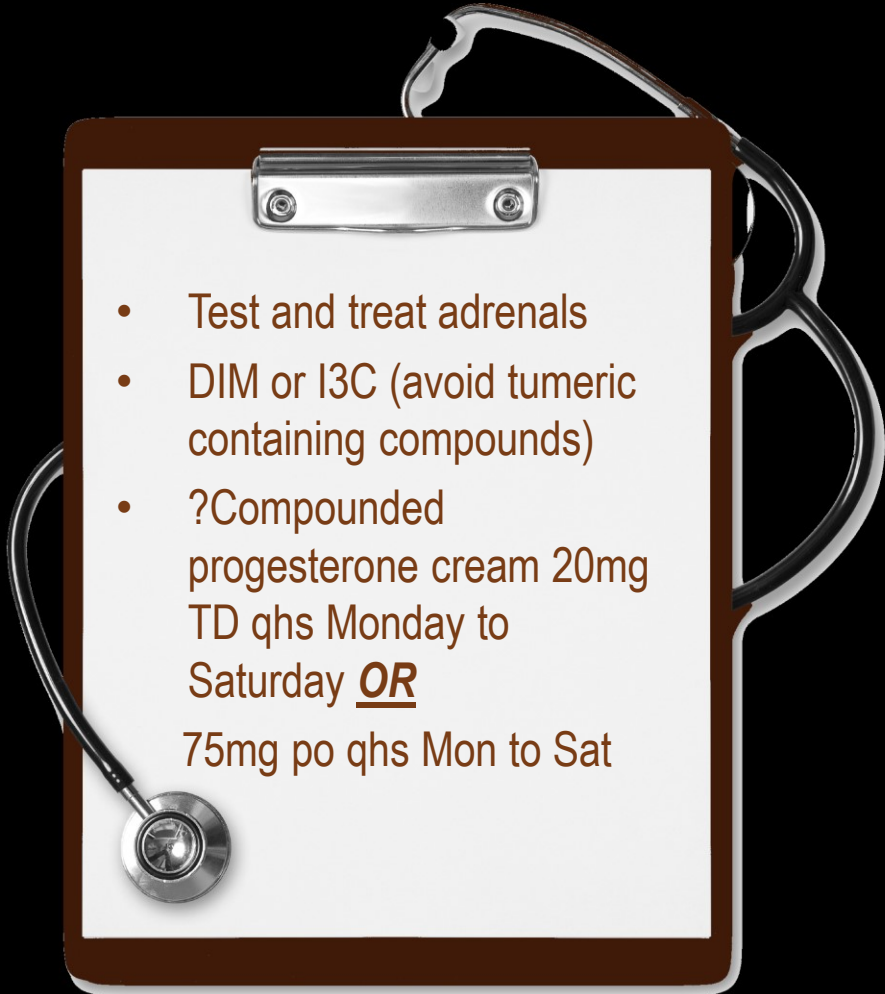
- 
- Test and treat adrenals
 - Compounded progesterone cream 30 mg TD qhs Monday to Saturday **OR** 120mg po qhs Mon to Sat

CASE STUDY #2

47 yo peri-menopausal female not on any hormone replacement therapy complaining of heavy and painful periods with PMS symptoms in the week before her period (28d cycle).

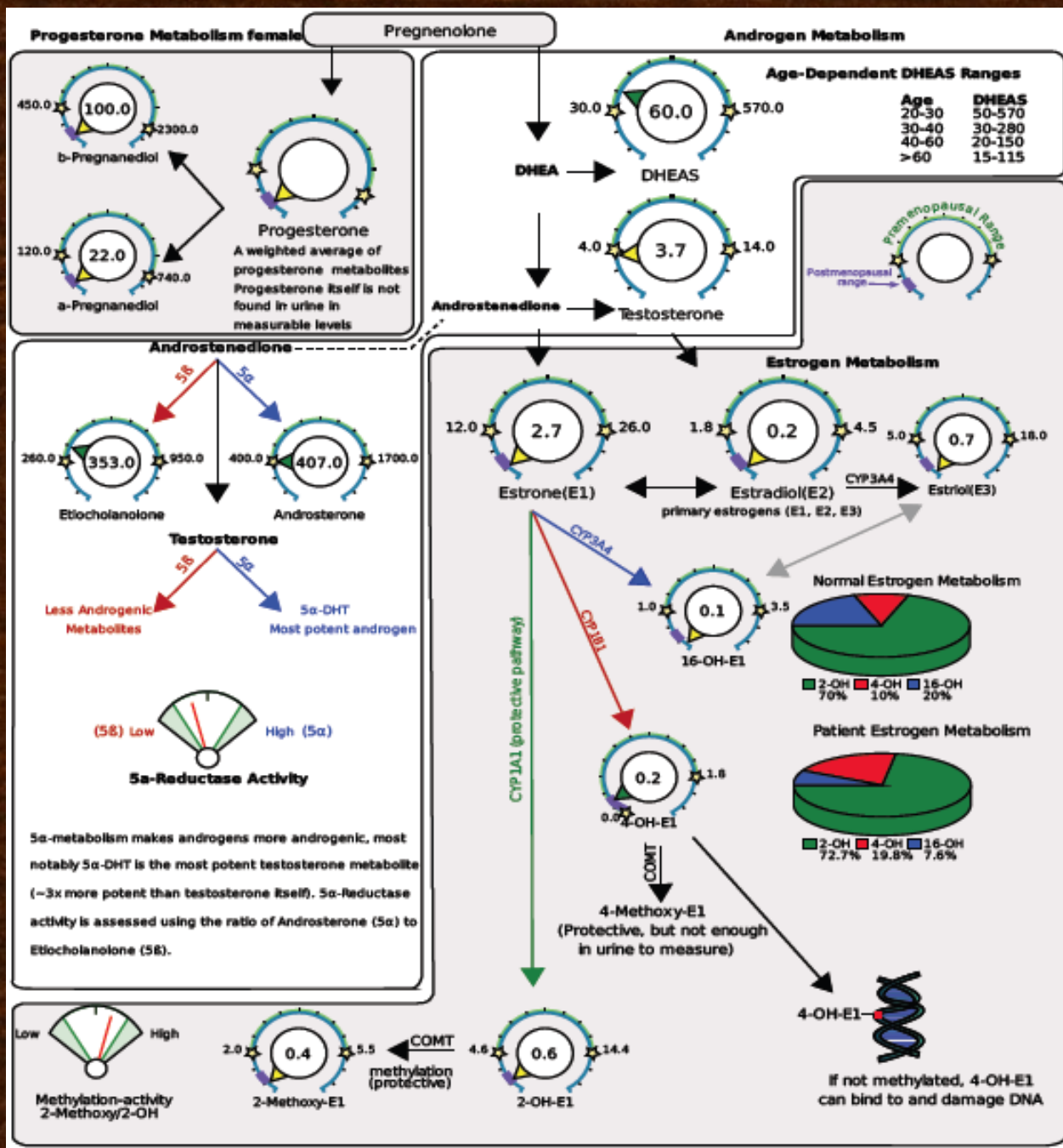


Estrogen Dominance

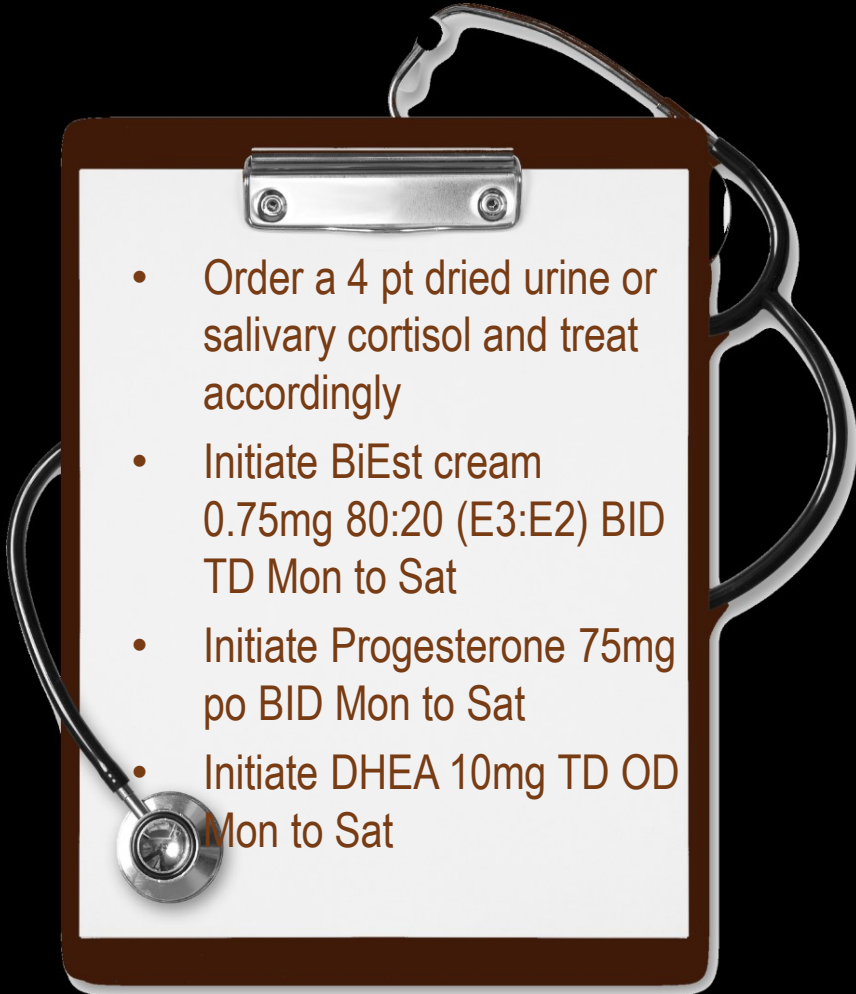
- 
- Test and treat adrenals
 - DIM or I3C (avoid tumeric containing compounds)
 - ?Compounded progesterone cream 20mg TD qhs Monday to Saturday **OR** 75mg po qhs Mon to Sat

CASE STUDY #3

55 yo post-menopausal female not on any hormone replacement therapy complaining of hot flashes, irritability, awakening between 2-4am, low libido, vaginal dryness and painful intercourse.

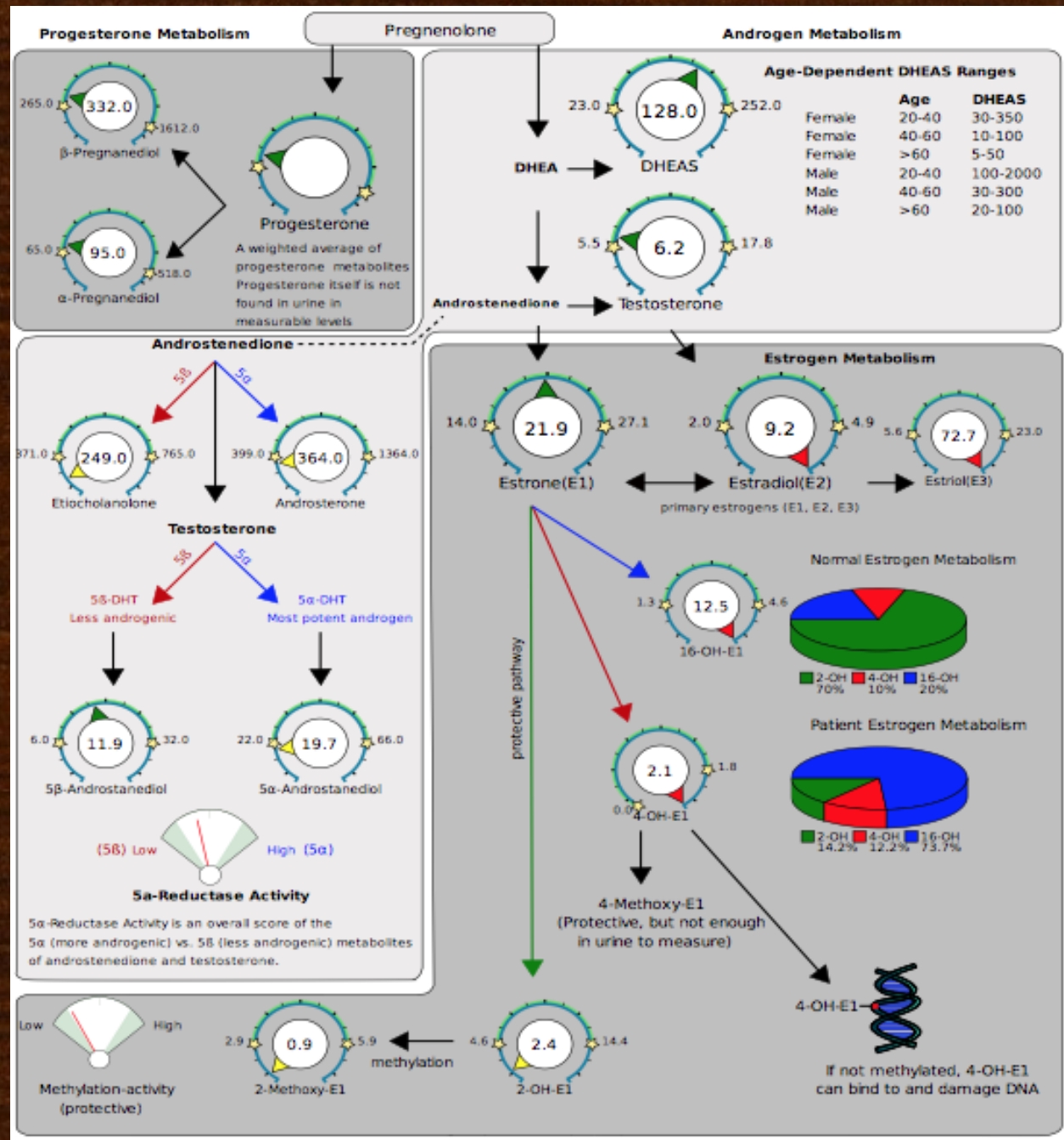


Estrogen/Progesterone/Testosterone Deficiency

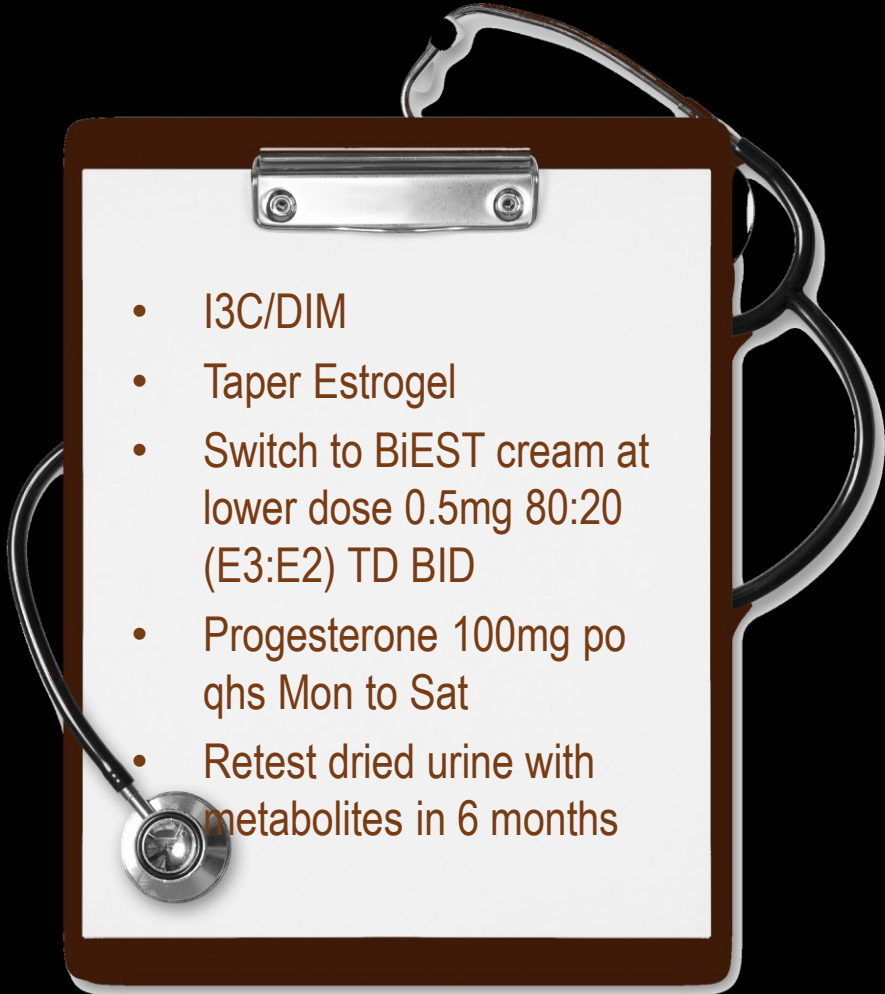
- 
- Order a 4 pt dried urine or salivary cortisol and treat accordingly
 - Initiate BiEst cream 0.75mg 80:20 (E3:E2) BID TD Mon to Sat
 - Initiate Progesterone 75mg po BID Mon to Sat
 - Initiate DHEA 10mg TD OD Mon to Sat

CASE STUDY #3

47 yo post-menopausal female started on Estrogel 1 pump and Prometrium 100mg po by her family doctor without testing. She is largely asymptomatic however with questioning she does note occasional breast tenderness. She has not had any PV bleeding and endometrial thickness is 4mm.



Over-replacement of Estrogen

- 
- I3C/DIM
 - Taper Estrogel
 - Switch to BiEST cream at lower dose 0.5mg 80:20 (E3:E2) TD BID
 - Progesterone 100mg po qhs Mon to Sat
 - Retest dried urine with metabolites in 6 months



Conclusion: Foundational Principles of Natural Hormone Replacement Therapy