# Hormone Replacement Therapy

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# The influence of estrogen



### Brain

Body temperature control Memory Libido

### Heart

Heart rate Healthy lining of blood vessels Lower cholesterol

### Liver

Cholesterol regulation

#### **Bones**

Bone strength

### Skin

Collagen production Reduce moisture loss

### Joints and muscles

Anti-inflammatory Muscle strength Flexibility Joint lubrication

### **Bowel**

Maintains function Helps with gut microbiome

### Nerves

Nerve transmission

### Bladder

Reduces risk of infection Controls bladder function

### Vagina

Lubrication Reduces bacterial overgrowth



# Menopause – the basics

Average age 51, life expectancy 82 therefore 30% of life postmenopausal

80% of women suffer with sx: 25% of whom severe

Half of women do not see their GP

77% of women did not realise their sx were menopausal



# Symptoms

MSK sx

Debilitating!

**GSM** 

Low mood / anxiety

Sexual difficulties

Worsening migraines

**Palpitations** 

Brain fog / memory probs



## Diagnosis

CLINICAL!

 Do not order blood tests if >45 with sx of menopause

Only if <45, or on POP / Mirena</li>

Confirm with 2 x FSH >6wks apart

# Misdiagnosis

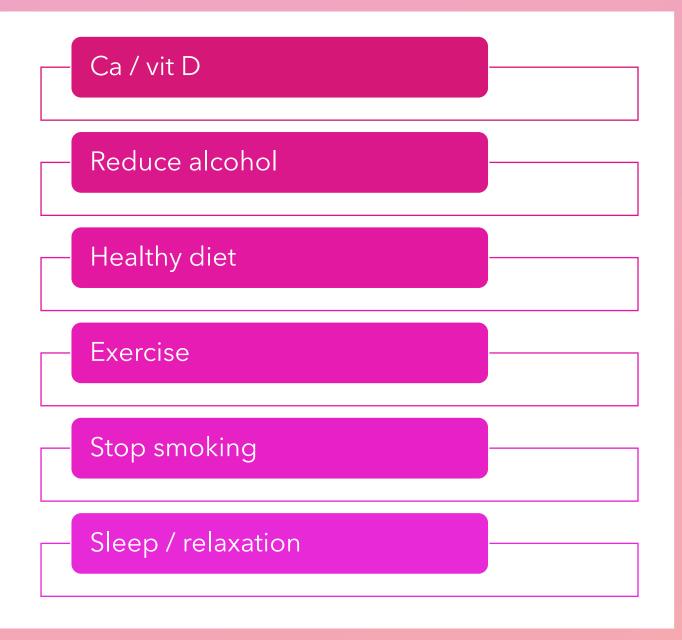
Fibromyalgia / palpitations / recurrent UTI / severe migraine / depression anxiety / memory problems

Wrong diagnosis  $\rightarrow$  wrong treatment  $\rightarrow$  significant side effects with lack of relief of sx + unnecc referrals & further tests

Economic consequences with time off work / early retirement



Mgt: lifestyle



# Or consider HRT...what are the benefits?

- Improves sx
- 1st line Rx and prevention of OP in women <60</li>
- Lower incidence of CVD in women starting within 10 yrs of menopause / younger than 60
- Improves skin turgor / collagen / thickness
- Reduces risk of: DMT2, dementia, obesity, depression, bowel ca
- And death! Reduces all cause mortality

# Sounds good! Why are we so worried?

The has sex timice a day? HRT rai cancer a stroke ris CHANEL Ductors halt trial of combine ned in Big massase' in risk of Cancer, coronal y choke

## HRT concerns

- Womens Health Initiative: was women on high dose HRT, many co-morbidities, preliminary data linked to press
- After 18 yrs f/up, using combined or oest only HRT was no asso with increased risk of all cause-mortality / CVD / cancer
- But what about breast cancer?
  - Risks mostly from synthetic progesterone, with no risk if on only oestrogen HRT or younger age & lower risk with micronised progesterone
  - Risks stop when HRT stops
  - No increased death from breast ca



# Understanding the risks of breast cancer

#### Difference in breast cancer incidence per 1,000 women aged 50-59.

Approximate number of women developing breast cancer over the next five years.

NICE Guideline, Menopause: Diagnosis and management November 2015

### 23 cases of breast cancer diagnosed in the UK general population



An additional four cases in women on combined hormone replacement therapy (HRT)

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Four fewer cases in women on oestrogen only Hormone Replacement Therapy (HRT)

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An additional four cases in women on combined hormonal contraceptives (the pill)

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An additional five cases in women who drink 2 or more units of alcohol per day

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Three additional cases in women who are current smokers

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An additional 24 cases in women who are overweight or obese (BMI equal or greater than 30)

Seven fewer cases in women who take at least 21/2 hours moderate exercise per week

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### Not great evidence

Fallacy of natural being safer ('appeal to nature fallacy')

Alternatives to HRT

SSRIs may help hot flushes if HRT CI, no evidence help low mood asso with meno.

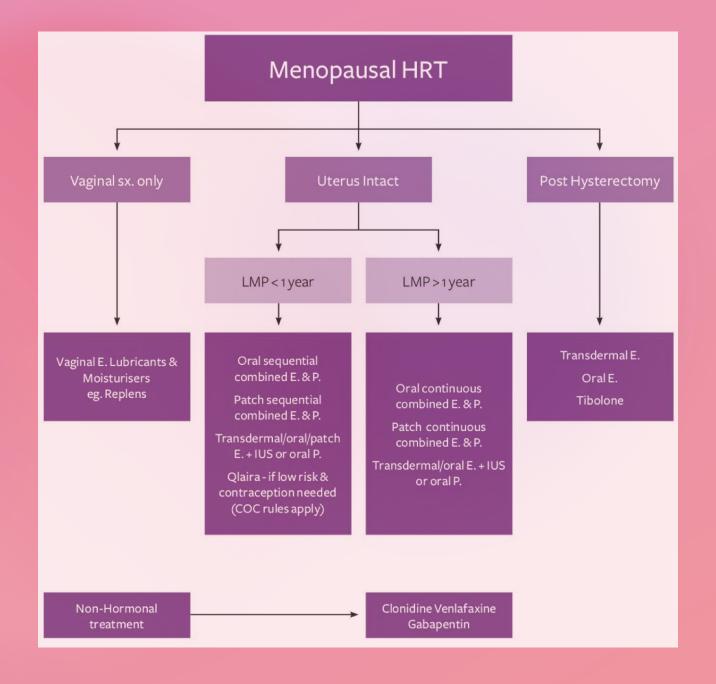
70% had been offered anti-dep's!

Bio-identical HRT is not regulared nor recommended

# Primary ovarian failure

- Needs HRT, but can still get preg, can continue if so
- Needs to cont until 51
- Higher dose oestrogen
- Testosterone often beneficial

# HRT prescribing



# Vaginal oestrogen

Maintain normal flora, lubrication, blood flow - can take months to provide relief

Safe to use as long as needed

Can give with HRT

May need lubricants too

Safe if non-active breast ca

Review after 3m



### No VTE risk

# Transdermal HRT

17 beta oestradiol - body identical, derived from yams

Patch, e.g., elleste, evorel, estradot - apply below waist

Gel, e.g., Oestrogel, Sandrena – upper outer arm or inner thigh

Lenzetto spray - inner forearm or inner thigh

Preferred over oral due to VTE risk and oral can lower libido

### Can improve CV / lipids / BP

No VTE risk

Micronised progesterone (utrogestan)

No breast ca risk for up to 5 yrs Rx

Can be px either 100mg ON each night (continuous) or 200mg for 14/28 (sequential)

Sedative, take on empty stomach

Mirena alternative if contraception needed

# Equivalent doses of oestrogen

Patch	Half a 25 microgram patch	25 micrograms	50 micrograms	75 - 100 micrograms
Gel - pump	½ pump	1 pump	2 pumps	3-4 pumps
Gel - sachet	½ of a 0.5mg sachet	0.5mg sachet	1 mg	1-2 mg
Spray		1 spray	2 sprays	3 sprays



# HRT side effects

Bloating (less likely MP)

Low mood

Breast tenderness

Unscheduled vaginal bleeding - usually settles 3-6m.

## When to stop?

- No maximum
- Can just abruptly stop
- Consider lower dose for older women

Women produce x3 amount pre-menopause > female androgen insufficiency

In NICE guidelines for use in menopause off-licence if low libido and HRT not enough

### Testosterone

Lots of health benefits to sex!

Using testosterone often also leads to improvement in mood, energy, concentration and stamina

Side effects from testosterone are very uncommon

## Bleeding on HRT

- Endometrial cancer is uncommon in women on HRT
- Consider first managing abnormal bleeding in sequential HRT with Tx acid or changing HRT, but Ix if persistently heavy / prolonged or persistent BTB >6m
- In CCHRT Ix if >6m be sensible to see if manage risk factors as risk of ca low.

# Use the Greene Climacteric Score to assess symptoms prior to consultation

Annual review, can be remote, but BP within a year req. (although HTN not CI for HTN; MI can lower BP)

Check smear and breast screening

Px 1 yr of HRT



# Take home messages

Stop ordering blood tests!

Consider menopause as a cause for any women 45-55 with new onset depression / anxiety

Women should be signposted to evidence-based and unbiased information about their menopause and its management <u>e.g.,</u> <u>www.menopausedoctor.co.uk</u> and <u>British Menopause Society</u> website

Transdermal oestrogen and micronised progesterone is the safest method of HRT

Women can stay on HRT as long as they need

Consider testosterone at review

Menopause. Hmmm. So, what you are saying is that I'm to look forward to ten years of hell... followed by death. Am I missing anything?

