



The role of female hormones and mental health

Dr Louise Newson BSc(Hons) MBChB(Hons) MRCGP FRCGP





Declarations

- No financial conflicts
- Director of Newson Health
- Director of Newson Health Research and Education (not-for-profit)
- Founder of Newson Health Menopause Society
- Director of Balance Ltd free balance menopause app
- Founder of The Menopause Charity
- Member of the UK Government Menopause Taskforce
- Visiting Fellow of Murray Edwards College, Cambridge University
- I take HRT...

Stratford-upon-Avon











Previous experience of menopause care



stated that they had been refused HRT by their GP



already taking HRT



had been offered antidepressants



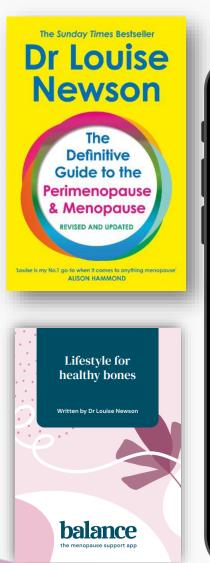


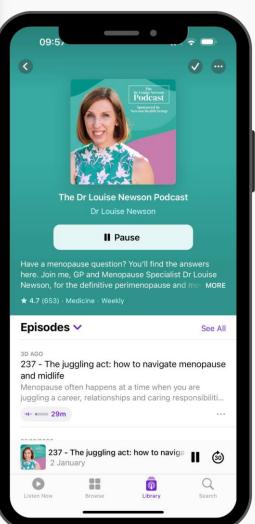
Newson Health is more than just a clinic

- Clinic
- Balance app
- Education
- Research
- Outreach work
- Media
- Social media
- Podcast







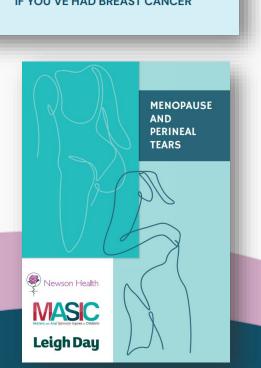


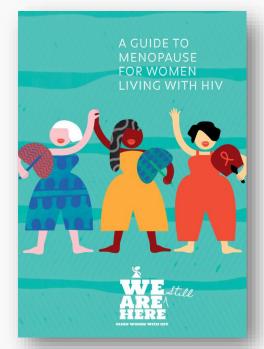




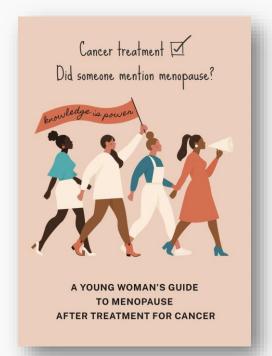


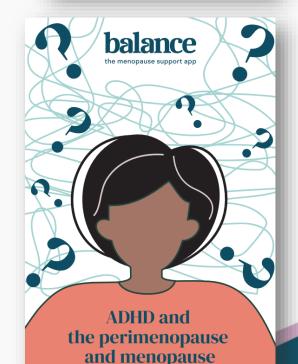












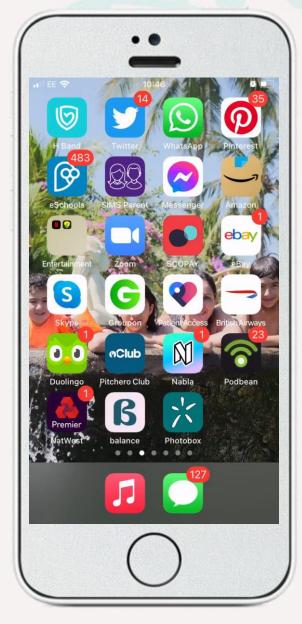


Written by Dr Louise Newson & Ovarian Cancer Action



Mental health and emotional wellbeing in the perimenopause and menopause





Free balance app

- The basics on perimenopause and menopause
- Journal insights
- Expert articles
- Log symptoms regularly





- Now aged 42
- Past 3 years:
 - Poor sleep
 - Anxiety
 - Low mood
 - More argumentative
 - Self harming
 - Memory problems
 - Not wanting to socialise





- Diagnosed with depression
- Given sertraline
 - Not improve her mood
- Mirtazepine added
 - Sleep slightly better
- Anxiety and low mood gradually worsened
- Prescribed quetiapine





- Worsening migraines and headaches
- Memory awful
- Constantly tired
- No self-motivation
- Muscle and joint pains
- Recurrent urinary tract infections
- Some incontinence at times





- Given painkillers / strong medication for her migraines
- Numerous antibiotics
- Referred to hospital for numerous tests
 - Brain scan
 - Heart investigations
 - X-rays
 - Bladder scans





All tests normal

- Starting drinking more alcohol
- "I feel that I am now addicted to painkillers because of my symptoms - I use them to help with pain but also to reduce my stress, anxiety and insomnia"



Laura – What is her diagnosis?



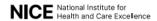
Perimenopause

+/- Clinical depression



What should or could have been prescribed for Laura?

NICE / IMS / ESHRE Guidelines





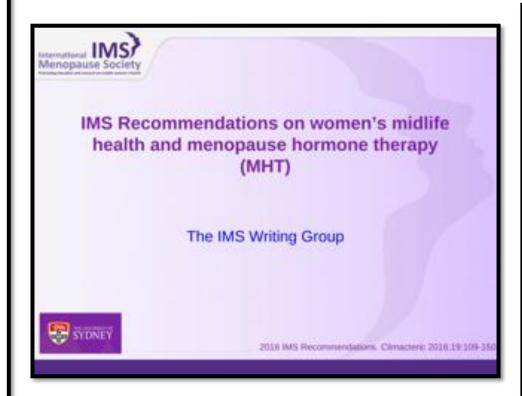
Menopause: diagnosis and management

NICE guideline

Published: 12 November 2015 Last updated: 5 December 2019

www.nice.org.uk/quidance/ng23

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Management of women with premature ovarian insufficiency

Guideline of the European Society of Human Reproduction and Embryology

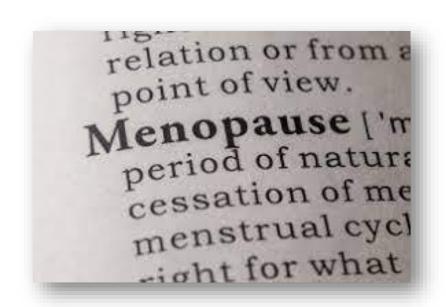
POI Guideline Development Group

December 2015





- One year since last menstrual period
- End of a woman's reproductive age
- Consider PMS / PMDD / Perimenopause
- When levels of oestrogen (oestradiol), progesterone and testosterone decline



"Women's Hormone Insufficiency (WHI)"

The problem





The influence of oestrogen

Brain

Anti-inflammatory

Improves blood flow in brain

Improves mood and reduces anxiety

Helps with learning

Body temperature control

Improves memory and concentration

Increases levels of other neurotransmitters including dopamine, serotonin, acetylcholine, noradrenaline, melatonin

Improves sleep

Increases connections between brain cells

Improves energy

Heart

Controls heart rate

Keeps endothelium (cells lining interior surface of blood vessels) healthy

Lowers blood pressure

Liver

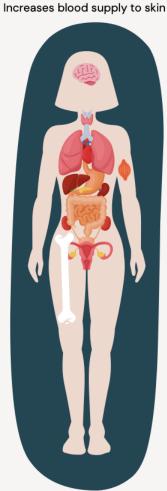
Improves cholesterol regulation

Improves glucose metabolism

Increases breakdown of fat Improves liver function

Skin

Increases collagen production
Reduces moisture loss
Improves elasticity



Bones

Increases bone mineral density

Reduces inflammation in ioints

Increases muscle strength Improves flexibility Lubricates joints

Joints and muscles

Anti-inflammatory

Muscle strength and flexibility

Joint lubrication

Bowel

Maintains function

Maintains balance of friendly bacteria Reduces heartburn

Nerves

Improves nerve transmission

Bladder

Reduces risk of infection Improves bladder function

Vagina/vulva

Increases lubrication

Maintains balance of friendly bacteria in vagina Keeps tissues healthy



The Influence of progesterone

Brain

Helps brain cells to communicate better, which helps improve mood, memory and brain health

Helps nerve functioning

Breasts

Tempers the effect of oestrogen and reduces breast cysts

Immune system

Reduces inflammation

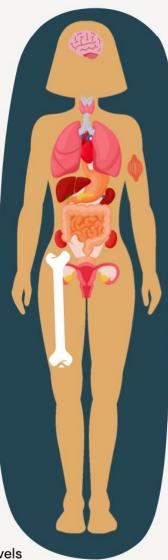
Lowers risk of autoimmune disease

Bones

Builds bone

Metabolism

Regulates blood sugar levels



Promotes sleer

Promotes sleep

Relieves anxiety

Helps use fat for energy

Muscles

Stimulates growth of new muscle

Reduces muscle spasm

Reproductive/ sexual function

Regulates menstruation

Supports pregnancy

Reduces bleeding





The influence of testosterone

Eye health

Improves meibomian gland function and lubrication

Reduces dry eyes

Cardiovascular health

Lowers triglyceride and cholesterol

Improves cardiac capacity and output makes your heart stronger and more efficient

Improves endothelial function - helps the lining of your blood vessels work better, increasing blood flow

Circulation

Red blood cell production

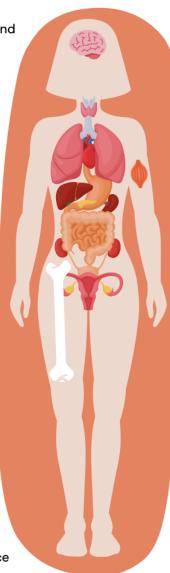
Reproductive and sexual function

Libido, arousal and orgasm

Urogenital health

Improves urinary symptoms including urgency and incontinence

Reduces symptoms related to vaginal dryness and soreness



Brain function

Improves concentration

Improves memory, verbal learning and spatial abilities

Sleep quality improves

Mood

Psychological wellbeing Improves energy

Muscle

Improves muscle mass and strength

Metabolism

Maintains normal metabolic function (blood pressure, lipids, glucose metabolism)

Bone health

Increased bone mineral density

Bladder

Reduces risk of infection

Improves bladder function

Vagina/vulva

Increases lubrication

Keeps tissues healthy





How do we diagnose perimenopause and menopause?





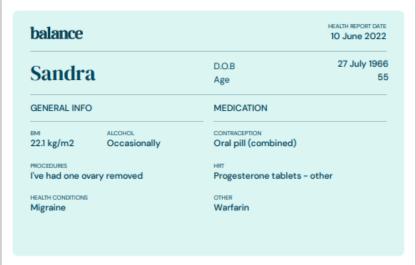
Hormone blood tests are not helpful

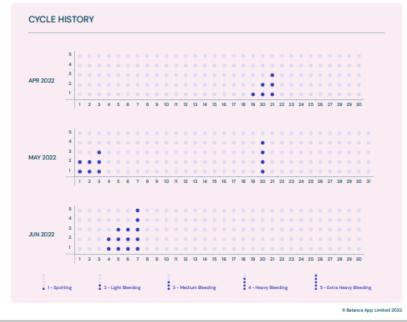




Women often make the diagnosis when they have the right information

Download the balance Health Report®









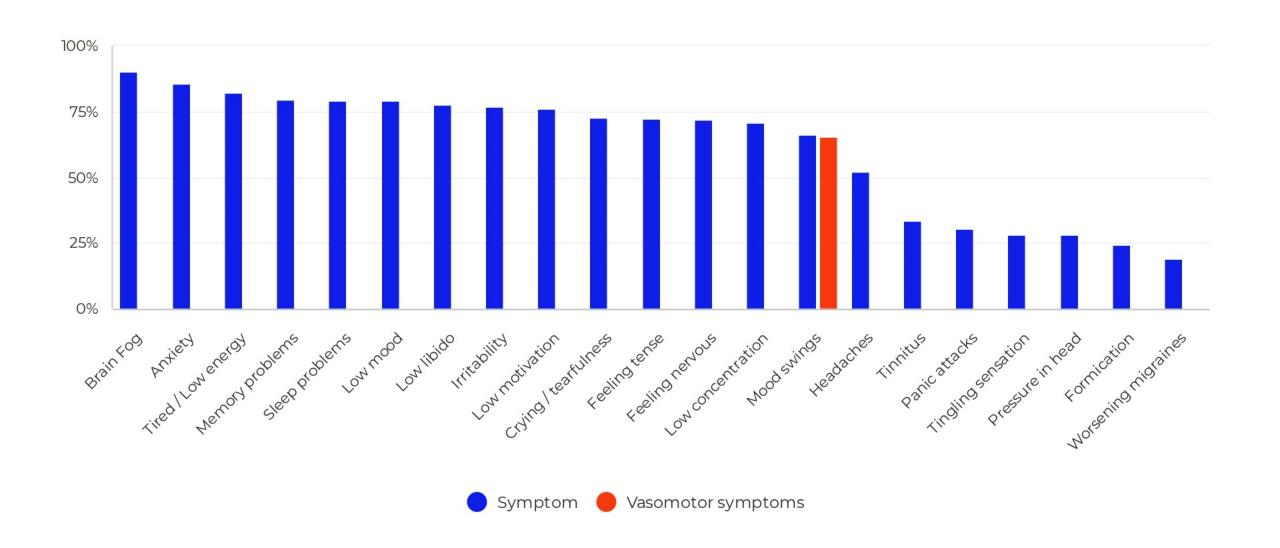


What are the symptoms of perimenopause and menopause?





Symptoms affecting brain / nervous system



Quotes from women



I don't know who I am anymore This is just not like me

I can't live like this anymore I feel like a black fog is hanging over me all the time

I feel like I have lost myself

I don't like who I have become

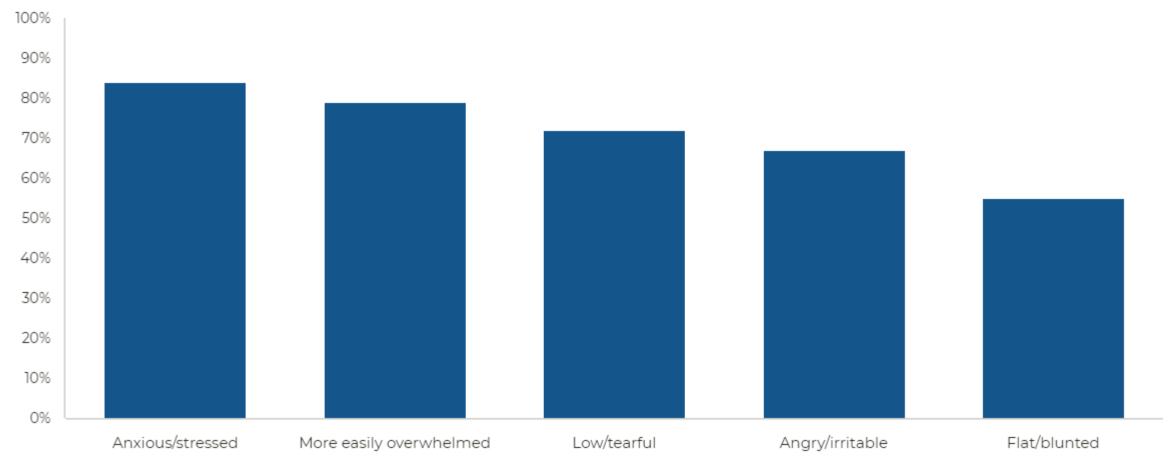
I feel like a dead person who is just existing



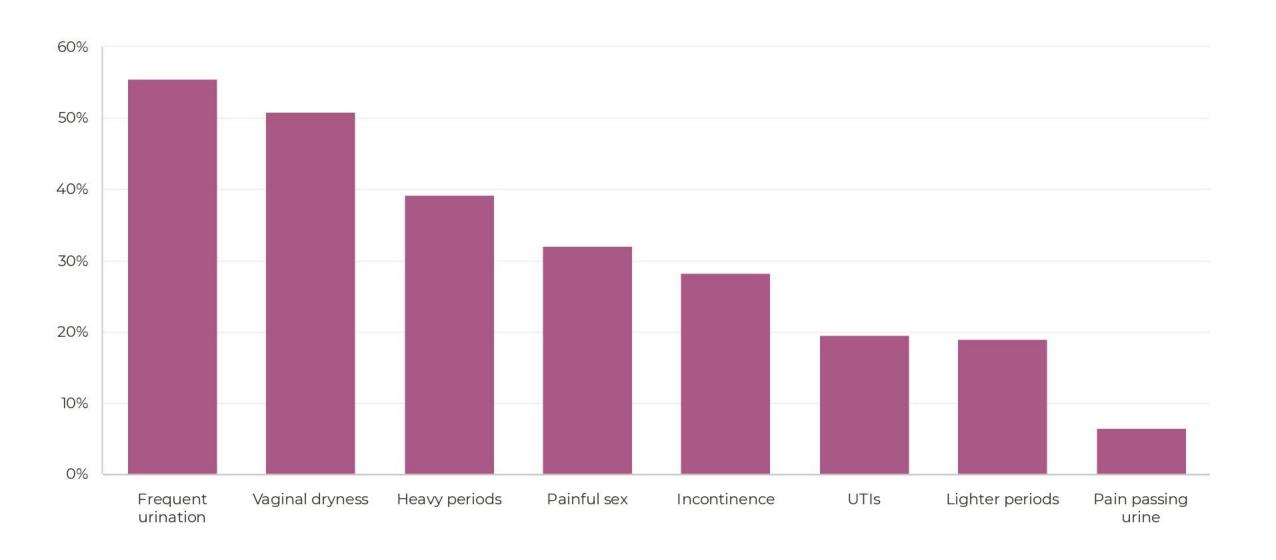
Mental Health

Change in mood and emotions

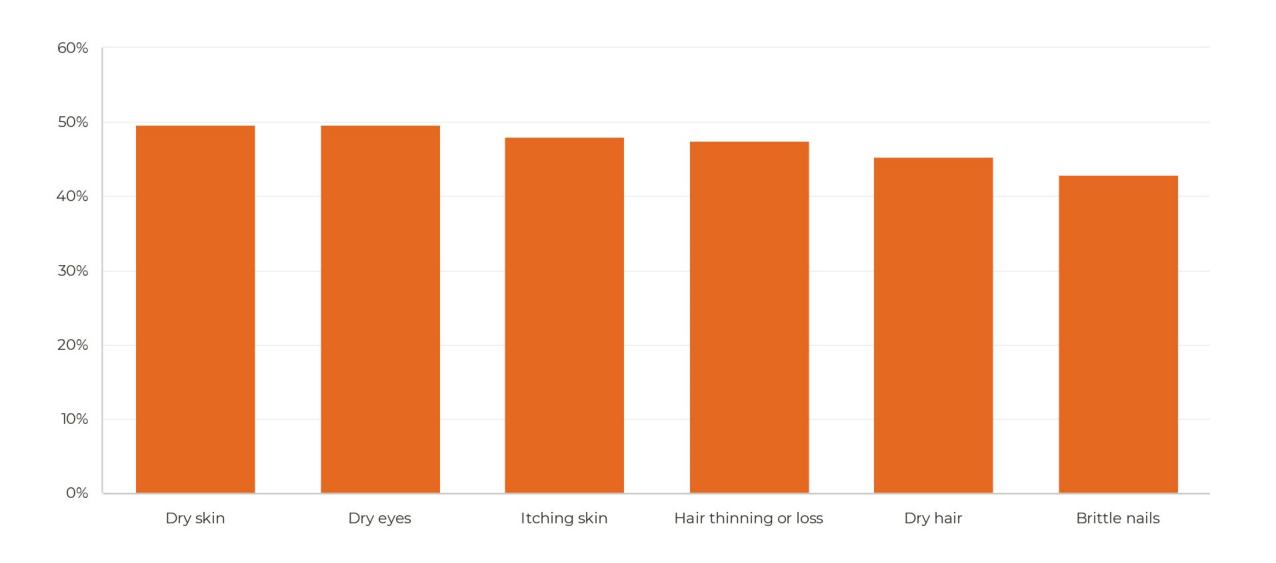




Symptoms affecting genitourinary system



Symptoms affecting skin, hair and nails



Oestradiol and female brain

"Nature's psychoprotectant"



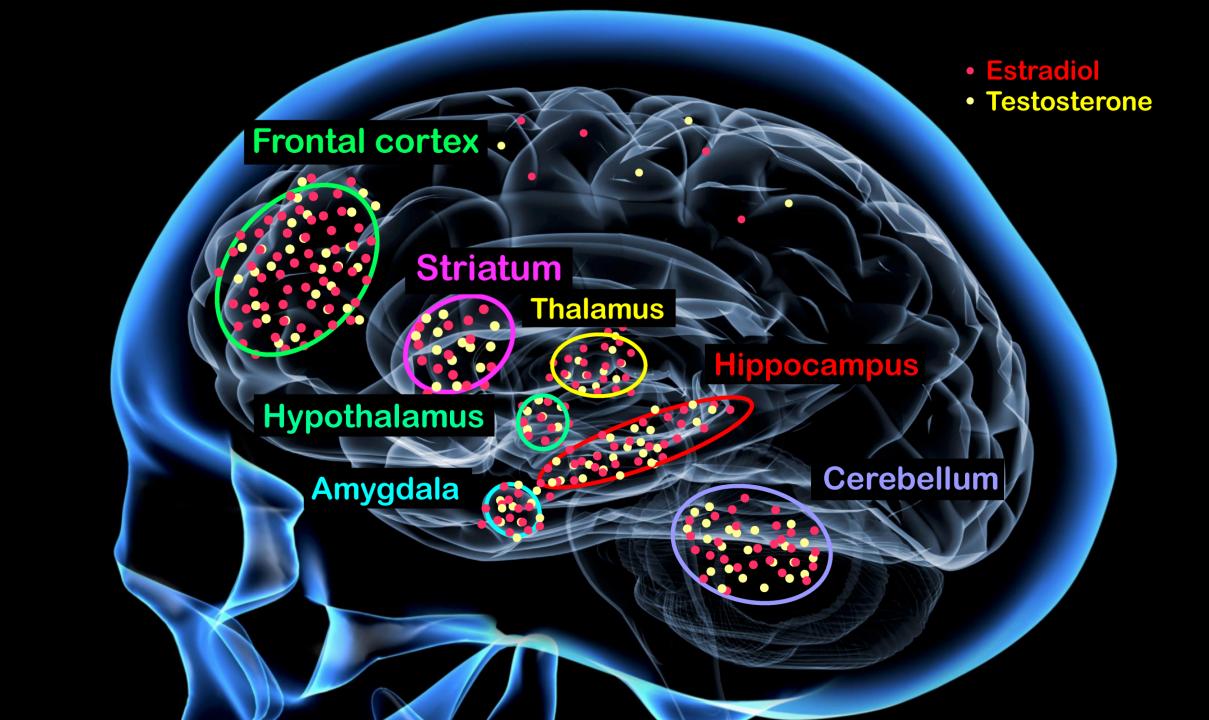
- Promote neural sprouting and myelination
- Enhance synaptic density and plasticity
- Facilitate neuronal connectivity
- Act as an anti-inflammatory and antioxidants
- Inhibit neural cell death
- Improve cerebral flow
- Improve glucose metabolism
- Improve mitochondrial function



Oestradiol and neurotransmitters

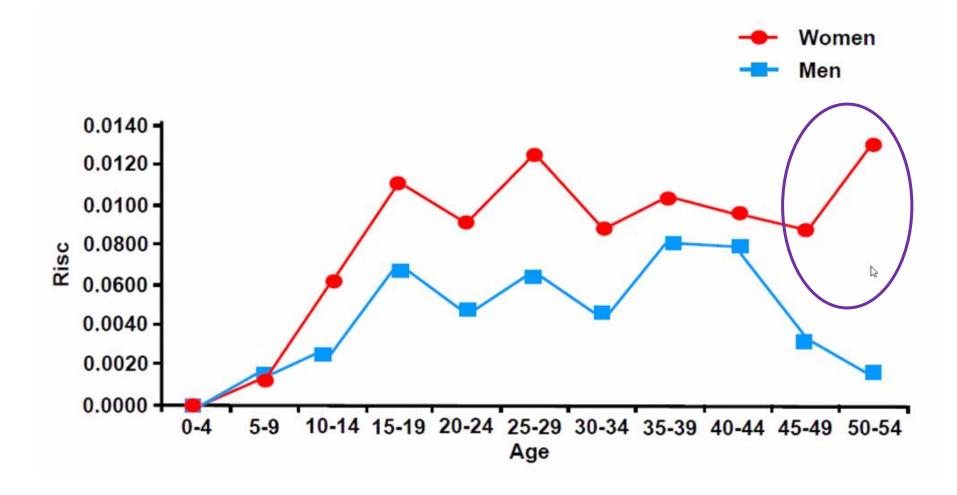


- Oestradiol can increase levels of these important neurotransmitters:
 - Acetylcholine
 - Dopamine
 - Glutamate
 - GABA (Gamma-aminobutryic acid)
 - Serotonin
 - Norepinephrine (noradrenaline)
 - Epinephrine (adrenaline)
 - Endorphins



Depression during life





Perimenopausal depression

- 16-fold increase in depression in women aged 45-52
- 7-fold increase in suicide in women aged 40-50



Diseases associated with menopause



- Cardiovascular disease
- Cognitive decline and dementia
- Type 2 diabetes
- Osteoporosis
- Clinical depression
- Kidney disease
- Cancer
- Obesity
- Earlier death

PERIMENOPAUSE / MENOPAUSE AND ADDICTION SURVEY

- On line survey
- Social media
- 1,178 respondants



'I had many perimenopausal symptoms, saw so many different doctors. Not one of them made the connection and I was clueless. I had no idea I was perimenopausal.

I was prescribed painkillers for my migraines, sleeping tablets for my sleepless nights, antiacids for my indigestion, antidepressants for my depression and anxiety. I was told I had bad hygiene for my recurring cystitis.

I was broken, lost, confused. I hated what I'd become. I lost way my humour, my lust for life. I considered ending it all.

I had dark thoughts. Worse time of my life. I thought I was going mad. Prescription drugs and therapy didn't help me.'



Reasons for drinking more alcohol during perimenopause or menopause

70% ½ ½ ½ ½ ½ ½ ½ ½ ½ ½

Due to increased anxiety, stress or depression

29% % % % % % % % % % %

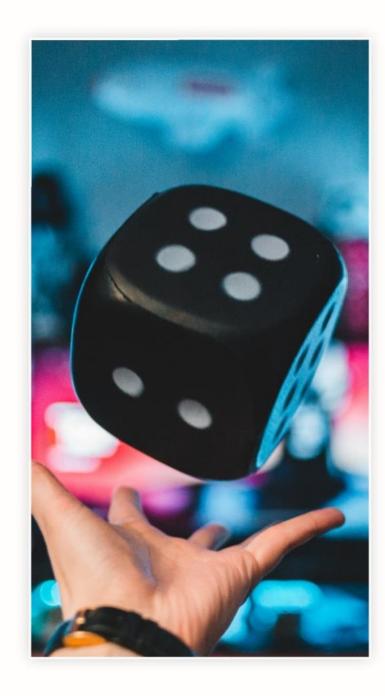
Drank to manage their menopausal symptoms

13% % % % % % % % % % %

To increase their confidence

5% % % % % % % % % % %

Due to lack of access to perimenopause / menopause treatments



Gamble more

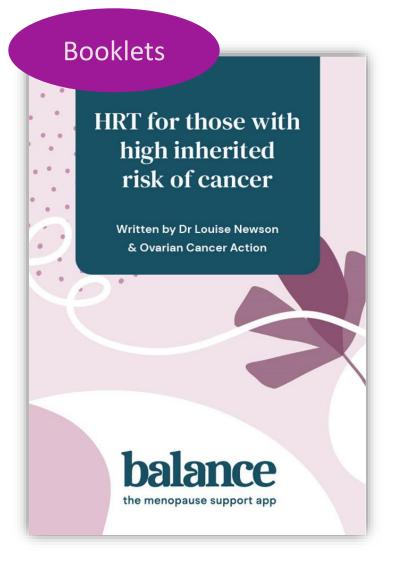


"IT ACTS AS A DISTRACTION FROM MY SYMPTOMS"

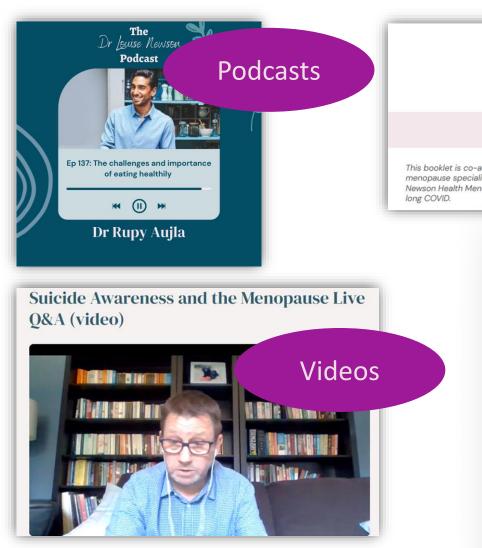
Petra



Treatment for perimenopause



Patient Information

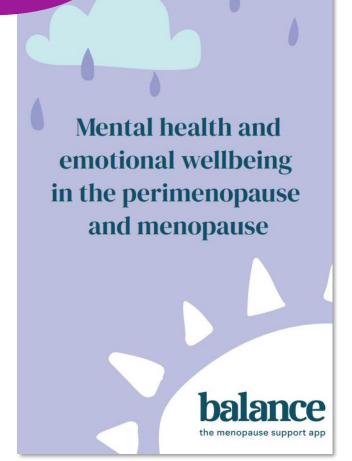






Patient Information

Booklet





HRT - Hormone Replacement Therapy

- Not a "one size fits all"
- Numerous different preparations
- Oestrogen / Progesterone / Testosterone
- Pills / Patches / Gels / Spray

















Does HRT treat depression?

- Double blinded randomised placebo controlled trial giving perimenopausal women 100mcg oestradiol patches or placebo for 12 weeks
- Remission of depression was achieved in 68% of women treated with oestradiol compared to 20% in placebo group



Efficacy of Estradiol for the Treatment of Depressive Disorders in Perimenopausal Women

A Double-blind, Randomized, Placebo-Controlled Trial

Claudio de Novaes Soares, MD, PhD: Osvaldo P, Almeida, MD, PhD: Hadine Ioffe, MD: Lee S, Cohen, MD

Background: Results of previous studies suggest that estrogen improves somatic and mild depressive symptoms experienced by perimenopausal women. This study investigated the efficacy of 17B-estradiol for the treatment of clinically significant depressive disorders in endocrinologically confirmed perimenopausal women.

Methods: Perimenopausal women (aged 40-55 years, with irregular menstrual periods and serum concentrations of follicle-stimulating hormone > 25 IU/L), meeting criteria for major depressive disorder, dysthymic disorder, or minor depressive disorder, according to DSM-IV, were randomized to receive transdermal patches of 17B-estradio (100 µg) or placebo in a 12-week, double-blind, placeboontrolled study. A 4-week washout period followed the 12-week treatment phase. Outcome measures were the Montgomery-Åsberg Depression Rating Scale and Blatt-Kupperman Menopausal Index scores.

of Psychiatry, University of São Paulo Medical School,

ñao Paulo, Brazil (Dr Soares

(Dr Almeida); and Perinatal

and Reproductive Psychiatry

Clinical Research Program,

Hospital, Harvard Medical School, Boston (Drs Soares,

Massachusetts Genera

and Behavioral Science.

Australia, Perth

Results: Fifty women were enrolled in the study; 26 me DSM-IV criteria for major depressive disorder, 11 for dysthymic disorder, and 13 for minor depressive disorder treated with 17β-estradiol compared with 5 (20%) in the placebo group (P = .001). Subjects responded similarly o estradiol treatment, regardless of DSM-IV diagnosis Patients treated with estradiol sustained antidepressan benefit of treatment after the 4-week washout period, al though somatic complaints increased in frequency and intensity. Treatment was well tolerated and adverse events were rare in both groups.

Conclusion: Transdermal estradiol replacement is an ef fective treatment of depression for perimenopausal

Arch Gen Psychiatry. 2001;58:529-534

common during the transition to menopause.1-3 Cross-sectional surveys describe high rates of de pressive symptoms among women treated in menopause clinics.45 We recently reported depressive disorders (major depressive disorder [MDD], dysthymic disorder, and minor depressive disorder) in 30 of 101 endocrinologically confirmed perimenopausal women attending a gynecological clinic.^{6,7} Unlike clinic-based surveys, community-based studies found that perimenopause may be a period of risk for mood disturbance for some women but does not necessarily represent a time of risk for major depression.8-10

The use of estrogen replacement for the treatment of menopausal symptoms has been shown to enhance "psychological wellbeing, "11 However, clinical studies using diverse forms of estrogen replacement for the treatment of depression produced mixed results. Three estrogen treatment studies in perimenopausal and newly postmenopausal women failed to demonstrate superiority over placebo for the treatment of

transdermal patches of 17β-estradiol13,16 and case series in which patients were treated with sublingual estradiol17 suggested that estrogen improves mood in women with postpartum depression and severe premen

See also page 537

leagues18 described that perimenopausal women with major (n=8) or minor deflushes experience greater relief of depres sive symptoms with estrogen than with placebo. However, data confirming these eliminary findings are lacking, particularly in a larger group of perimenopausa women with major depression.

termine the efficacy of 17B-estradiol for the treatment of depressive disorders in endocrinologically confirmed perimenopausal women. Based on the results of pre vious reports, 15,16 we hypothesized that the use of transdermal 17B-estradiol would have a greater antidepressant benefit than

(REPRINTED) ARCH GEN PSYCHIATRY/VOL 58, JUNE 2001 WWW.ARCHGENPSYCHIATRY.COM



Oestradiol and antidepressants

- Oestrogen can enhance the efficacy of antidepressant medication in menopausal women
- 17β-oestradiol can augment the modulatory effects of escitalopram on hippocampal levels of brain-derived neurotrophic factor and serotonin reuptake transporter



What are the benefits of women taking HRT??





- Improves symptoms!
- For the majority of women, the benefits of taking HRT outweigh any risks
- Reduces risk of:
 - Cardiovascular disease
 - Cognitive decline and dementia
 - Type 2 diabetes
 - Osteoporosis
 - Clinical depression
 - Kidney disease
 - Cancer
 - Obesity
 - Earlier death
- Only around 14% menopausal women in UK take HRT

Cochrane Database Syst Rev. 2015;(3):CD002229 Alzheimers Dement. 2021;7(1):e12174 Menopause. 2016;23(4):461–70 JAMA Psychiatry. 2018;75(2):149–157 Climacteric. 2022;25(4):362–368 Endocr Rev. 2017;38(3):173–188

The bigger problem



of women globally receive hormones



Laura

- Started on 75mcg Evorel patch changing twice a week with 200mg Utrogestan at night for 2 out of 4 weeks
- Also Estring inserted (changed every 3 months)
- Happier
- Sleeping better
- Calmer
- Physically more active
- Periods lighter and more regular
- No more UTIs



"Most of my symptoms have completely improved. I am enjoying my life again and feel happy. I no longer have joint pains and I can sleep all night. I have not felt this good and well for many years.

I have now stopped taking so many other medications."





"Take Home" Messages



- Diagnosis of perimenopause and menopause is clinical
- Symptoms can be very debilitating
- HRT is safe
- Antidepressants work better in oestrogenised women
- Consider HRT with testosterone





RCPsych's online learning resource for mental health professionals

Mental health during the perimenopause and menopause



Find out more

CPD

Credits

This module looks at the perimenopause and menopause in the context of mental health, the hormonal and neurotransmitter changes that occur, perimenopausal mood disorders and prescribing hormone replacement therapy (HRT). We are pleased to offer this module free to those without a paid subscription.

By Dr Abbie Laing, Dr Gina Waters, Dr Louise Newson and Dr Olivia Jones



Designed to enhance your knowledge of the perimenopause and menopause Downloaded by more than 30,000 people worldwide since 2021, we're back with a fresh new look and exciting new content on a new platform Our course is designed for everyone – regardless of gender or profession. Whether you're a woman looking for information to help you make the right decisions, or you're a partner, friend, or colleague who simply wants to know more.