Perimenopause: Taming the Heat in this Proinflammatory Life Phase

By Lorna Driver-Davies Nutritional Therapist, Naturopath, Functional Medicine Practitioner

Integrative & Personalised Medicine Congress June 2023, London.

Introduction (1)

Perimenopause is more than just an endocrinological transition.

I welcome you to assess this transition as a shift in neuro-immuno-endocrinology.

If we can regulate this hormonal transition better, by understanding its impact on the immune system, we can then better protect neurological health during perimenopause.

And through into menopause where the risks become even more serious and potentially long lasting.

We also need to...

Change the **narrative** of Perimenopause – it is a significant **prelude** to Menopause. A distinct, very different phase to Menopause.

The very 'experience' of menopause' may be dictated or mapped out by Perimenopause.

Introduction (2)

"Although primarily viewed as a reproductive transition, the symptoms of perimenopause are largely neurological in nature.... a hypometabolic state associated with neurological dysfunction can develop". (*1)

Neurological symptoms that emerge during perimenopause are indicative of disruption in multiple oestrogen-regulated systems:

Such as...thermoregulation, sleep, cognitive and sensory processing, mood, general psychology, neurotransmitter production and regulation and more. Significant nervous system dysfunction and 'stress'.

For some women, this hypometabolic state might increase the risk of developing neurodegenerative diseases later in life." (*2)

"Emerging evidence is showing that peri-menopause is pro-inflammatory and disrupts oestrogen-regulated neurological systems...". (*3)

A Functional approach...

My expertise over 13 years of clinical work has focused particularly on gynaecological nutrition, with particular interest and specialism in 'immunologically challenged' disease.

Such as Endometriosis, Polycystic Ovarian Syndrome and immuno-endocrinological overlaps such as autoimmune thyroid disease. And of course, the fascinating perimenopause to menopause transition.

Immune modulation = addresses hormone function/ stability

Hormone modulation = addresses immune function/stability

Hormone & immune modulation = addresses neurological, cognitive and psychological, & cardio-metabolic health.

Key Take Aways...

- 1. It's the trio of support...
- Oestrogen (and oestrogen metabolism) and other steroid hormone support
- Combined with neuro protective nutrition
- Combined with immunological support (anti-inflammatory support)
- Make sure you assess the cardio-metabolic picture
- 2. Application of hormone support and replacements (medical, natural, botanical).

3. Check inflammatory markers or associated markers NOT just hormones. E.g Ferritin, HBA1C, lipid peroxidase, thyroid antibodies, thyroid reverse T3, C-RP, organic acids, toxicity (environmental functional medicine) melatonin, cholesterol lipoproteins and more...

3. Methylation assessment: genetics, neurotransmitter production (nervous system health), regulation of inflammation, antioxidant production. Nutrition for methylation (folate, B12, TMG and so forth).

4. Other medicines or herbs to support - omega 3 (NF-KB (Nuclear factor kappa B) expression of cytokines regulated by, steroid hormone production) and Vitamin D are critical. We still have mass population low levels.

Botanicals

Shatavari (*asparagus racemosus*) Dong quai (*angelica sinensis*) Black cohosh (*cimicifuga racemose*) Saffron (*Crocus sativus L*) Agnus castus (vitex agnus castus)

Ashwagandha (withania somnifera) Siberian Ginseng *(eleutherococcus senticosus)* Botanical Mushrooms

Nervous system herbs

Nootropics Berberine (*hydrastis various*) Echinacea *Antioxidant plant sources*





Thank you for listening

As practitioners we all have opportunity to 'change' below what this study showed!

A 2021 study commissioned by the UK government shows that of the 70% of those that experienced perimenopausal symptoms in their 30s and 40s, nine in 10 (90%) failed to recognise the immediate link to their fluctuating and declining hormones, instead attributing symptoms to ageing, stress, anxiety or depression.

How to find Lorna:

www.lornadriverdavies.com

www.wildnutrition.com (Head of Nutrition)

References (1):

1. Perimenopause as a neurological transition state. Nat Rev Enocrinol 2015 Jul;11(7):393-405. doi: 10.1038/nrendo.2015.82. Epub 2015 May 26. Roberta D Brinton et al.

2. Perimenopause as a neurological transition state. Nat Rev Enocrinol 2015 Jul;11(7):393-405. doi: 10.1038/nrendo.2015.82. Epub 2015 May 26. Roberta D Brinton et al.

3. The peri-menopause in a woman's life: a systemic inflammatory phase that enables later neurodegenerative disease. Journal of Neuroinflammation. volume 17, Article number: 317 (2020). Micheline McCarthy et al.

4. Dynamic Neuroimmune Profile during Mid-life Aging in the Female Brain and Implications for Alzheimer Risk. I science 2020 Dec 18; 23(12): 101829. Aarti Mishra et al.

5. Proinflammatory and anti-inflammatory cytokine changes related to menopause. 2014 Jun;13(3):162-8. doi: 10.5114/pm.2014.43818. Epub 2014 Jun 30. Andrei Mihai Malutan et al.
6. Long COVID risk - a signal to address sex hormones and women's health. The Lancet, Volume 11 100242, 2021. Stuart Stewart et al (Louise Newson).

7. Autoimmune Disease in Women: Endocrine Transition and Risk Across the Lifespan. Front Endocrinol (Lausanne). 2019; 10: 265. Maunil K Desai et al.

8. GABA Regulates Release of Inflammatory Cytokines From Peripheral Blood Mononuclear Cells and CD4+ T Cells and Is Immunosuppressive in Type 1 Diabetes. E BioMedicine 2018 Apr; 30: 283–294. Amol K Bhandage et al. et al.

References (2):

9. Sex and Gender Differences in Alzheimer's Disease Dementia. <u>Psychiatr Times. 2018 Nov; 35(11):</u> <u>14-17. Michelle M. Mielke</u>.

10. The peri-menopause in a woman's life: a systemic inflammatory phase that enables later neurodegenerative disease. Journal of Neuroinflammation. volume 17, Article number: 317 (2020). Micheline McCarthy et al.

11. Brain functional changes in perimenopausal women: an amplitude of low-frequency fluctuation study. Menopause 2021 Jan 11;28(4):384-390. doi: 10.1097/GME.0000000000001720. <u>Ningning Liu</u> et al.

12. The experience of perimenopausal distress: examining the role of anxiety and anxiety sensitivity. J Psychosom Obstet Gynaecol 2016;37(1):26-33. doi: 10.3109/0167482X.2015.1127348. Ljiljana Muslić et al.

13. Brain functional changes in perimenopausal women: an amplitude of low-frequency fluctuation study. Menopause 2021 Jan 11;28(4):384-390. doi: 10.1097/GME.0000000000001720. <u>Ningning Liu</u> et al.

14. The SWAN Song: Study of Women's Health Across the Nation's Recurring Themes. <u>Obstet Gynecol</u> <u>Clin North Am. 2011 Sep; 38(3): 417-423.</u> doi: <u>10.1016/j.ogc.2011.05.001</u>. <u>Nanette Santoro</u>, MD et al.

References (3):

15. Neurotrophic and neuroprotective actions of estrogen: basic mechanisms and clinical implications. Steroids 2007 May;72(5):381-405. doi: 10.1016/j.steroids.2007.02.003. <u>Darrell W Brann</u> et al.

16. Invited Review: Estrogens effects on the brain: multiple sites and molecular mechanisms. J Appl Physiol (1985). 2001 Dec;91(6):2785-801. doi: 10.1152/jappl.2001.91.6.2785. <u>B S McEwen</u>.

17. Females with ADHD: An expert consensus statement taking a lifespan approach providing guidance for the identification and treatment of attention-deficit/ hyperactivity disorder in girls and women. <u>BMC</u> <u>Psychiatry.</u> 2020; 20: 404. Susan Young et al.

 Dopamine-Dependent Cognitive Processes after Menopause: The Relationship between COMT Genotype, Estradiol, and Working Memory. <u>Neurobiol Aging. 2018 Dec; 72: 53-61.</u> Julie A Dumas et al.
Estradiol and tryptophan depletion interact to modulate cognition in menopausal women. Neuropsychopharmacology 2006 Nov;31(11):2489-97. doi: 10.1038/sj.npp.1301114. Epub 2006 Jun 7. <u>Zenab Amin</u> et al.

20. Inflammatory Responses in Brain Ischemia. <u>Curr Med Chem. 2015; 22(10): 1258–1277.</u> doi: 10.2174/0929867322666150209154036. Masahito Kawabori et al.

21. Cardiovascular Risk in Perimenopausal Women. Curr Vasc Pharmacol 2019;17(6):591-594. doi: John C Stevenson et al.

22. Perimenopause and Cognition. <u>Obstet Gynecol Clin North Am. 2011 Sep; 38(3): 519-535.</u> doi: <u>10.1016/j.ogc.2011.05.007</u>. <u>Gail A. Greendale</u> et al.