

Hacking the stress system: the stress response as an adaptive trait that can work for or against you

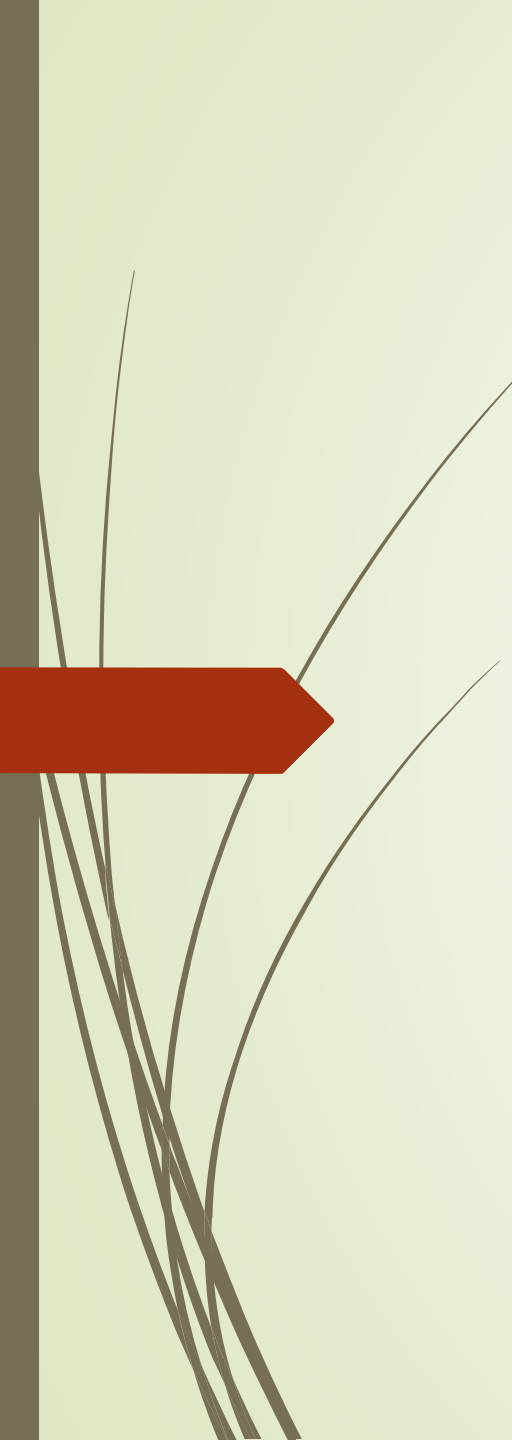
VI - Depression, PTSD and anxiety

Disclaimer

**I am not a psychiatrist! I am a scientist!
Especially if you are experiencing
mental health issues, do NOT try
anything we discuss today without first
consulting your specialist. Non of the
information discussed today reflects
individual differences and your
personal physician can judge much
better than I ever can, how this
information may affect you.**



Major Depression



A point in advance: major depression is every bit as much a biological disorder as diabetes or hypothyroidism. However, rather than glucose transportation or thyroid function being disrupted, the mechanism that allows you to pull yourself together after a stressful event does not work properly. Major depression is therefore often described as learned helplessness.



Major depression: symptoms

- Psychological symptoms
 - Sadness / grief
 - Learned helplessness
 - No motivation
 - Anxiety
 - Elevated pain susceptibility
 - Suicidal thoughts
 - Loss of interest in social interactions
 - Loss of interest in hobbies
- Slower speech
- Early morning wakening
- Constipation
- Altered appetite
- Diminished sex drive



Neurobiology of depression




- ▶ Disruption of
 - ▶ Noradrenalin
 - ▶ Psychomotor symptoms
 - ▶ Dopamine
 - ▶ Motivation / anhedonia
 - ▶ Serotonin
 - ▶ Grief
- ▶ Neuro inflammation
 - ▶ Omega 3 supplementation
 - ▶ Antibiotics



Careful: Thyroid hormone deficiency

- ▶ Thyroid deficiency may lead to downstream effects which look a lot like impression.
 - ▶ Make sure you consume sufficient iodine!
 - ▶ Check thyroid function (Hashimoto's)
 - ▶ Sleep deprivation



Stress may trigger and deepen a major depression

- Strong evidence for a genetic predisposition
- Stress triggers the disorder (3-4 episodes of severe prolonged stress)
- After the first episode it becomes much easier to trigger the next one




Quitting: Noradrenaline / dopamine: and internal reward

- ▶ Dopamine appears to balance noradrenaline function and buffer the tendency to quit.
 - ▶ Keep us on the right path
 - ▶ Internal reward
 - ▶ Slicing time (milestones), small rewards
 - ▶ Altruism (perhaps a downstream effect of oxytocin)




Learned helplessness as an adaptive trait in the face of unescapable suffering

- High stress
 - Low motivation to avoid suffering (dopamine deficiency)
 - Diminished ability to find internal peace if disaster can strike at any moment
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Post-Traumatic stress disorder




Where Major depression can be understood as learned helplessness, PTSD seems to be the desperate struggle to keep control

- ▶ Highly adaptive under certain high-stress conditions
- ▶ The confrontation with the realisation that one is never safe
- ▶ High comorbidity with depression (it's a state that cannot be maintained for long)

Mac Gillavry, D. W., & Ullrich, D. (2020). A novel theory on the predictive value of variation in the β -endorphin system on the risk and severity of PTSD. *Military Psychology*, 1–14. <https://doi.org/10.1080/08995605.2020.1730111>

Hoge, C. W. (2010). *Once a warrior always a warrior: Navigating the Transition from Combat to Home— Including Combat Stress, PTSD, and mTBI*. Guilford: Globe Pequot Press.



Post-traumatic stress disorder: Main Symptoms

- Reaction to extreme stressor
 - Acute or repeated exposure
- Hypervigilance
- Flashbacks and intrusive thoughts
- Nightmares

American Psychiatric Association. (2013). Diagnostic and statistical manual of mental disorders (5th. In *American Journal of Psychiatry*. <https://doi.org/10.1176/appi.books.9780890425596.744053>
World Health Organization. (1993). *The ICD-10 classification of mental and behavioural disorders: Diagnostic criteria for research* (pp. 155–157). pp. 155–157. [https://doi.org/10.1002/1520-6505\(2000\)9:5<201::AID-EVAN2>3.3.CO;2-P](https://doi.org/10.1002/1520-6505(2000)9:5<201::AID-EVAN2>3.3.CO;2-P)

Neuromodulators involved in PTSD

- Noradrenaline & the locus coeruleus
- Dopamine
- β -endorphin / Galanin
 - Morphine
 - Post-combat delayed onset
- Oxytocin

van Zuiden, M., Frijling, J. L., Nawijn, L., Koch, S. B. J., Goslings, J. C., Luitse, J. S., ... Olff, M. (2017). Intranasal Oxytocin to Prevent Posttraumatic Stress Disorder Symptoms: A Randomized Controlled Trial in Emergency Department Patients. *Biological Psychiatry*, 81(12), 1030–1040. <https://doi.org/10.1016/j.biopsych.2016.11.012>

Ullrich, D., & Mac Gillavry, D. W. (2021). Mini-review : A possible role for galanin in post-traumatic stress disorder. *Neuroscience Letters*, 756(May), 135980. <https://doi.org/10.1016/j.neulet.2021.135980>

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Pan, X., Kaminga, A. C., Wen, S. W., & Liu, A. (2018). Catecholamines in post-traumatic stress disorder: A systematic review and meta-analysis. *Frontiers in Molecular Neuroscience*, 11(December). <https://doi.org/10.3389/fnmol.2018.00450>



Anxiety: the stress system in
overdrive




Anticipation gone to extremes

- Anxiety is largely related to the excessive anticipation of potential stressors in the future.
- Elevated noradrenaline
- Increased limbic system (emotional processing) activity
 - Amygdala!
 - Thalamus
 - Hypothalamus
 - Hippocampus
 - Prefrontal cortex

Griez, E., Faravelli, C., Nutt, D., & Zohar, D. (Eds.). (2001). *An Introduction to Clinical Management and Research*. Chichester: John Wiley & Sons, Ltd.



Feedback loops to reduce anxiety

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- Benzodiazepams / Valium
 - Meditation
 - Yoga