Stress mediated stirring up of the immune system



The immune system is generally activated by two kinds of signal: due to the recognition of pathogen and the release of 'damage signals' from the host cell. There is a growing body of evidence showing a third class of activating signal, in the form of stress that alters the status of the immune system.

A recently published research outlines the mechanisms of psychological stress influencing bowel inflammation (1). The group has shown the response of enteric nervous system (ENS) in modulating the brain derived psychological stress and triggering intestinal inflammation. Chronic stress causes the release of glucocorticoids from the adrenal gland that acts on glucocorticoid receptor expressed on enteric glial cells. The powered up glial cells then secrete molecules, including Csflthat results in recruitment and priming of TNF-producing monocytes to exacerbate gut inflammation. Thus, a cascade of events connects psychological stress to flaring up of intestinal inflammation, the elements of which may extend to other inflammatory diseases. This work emphasizes the importance of stress management in enhancing the efficacy of treating inflammatory diseases.

Earlier studies have shown that chronic stress can enhance the risk of developing autoimmune diseases such as rheumatoid arthritis and multiple schlerosis (2, 3). Importantly, childhood bullying have been associated with low-grade systemic inflammation in adulthood; and children exposed to

high psychological stress have been reported to have increased spontaneous secretion of IFN- γ and low IL10 (4, 5). In middleaged men, stressful events have been found to be associated with shorter leukocyte telomere length, a biomarker of cellular aging and this relation is explained by depression and low grade chronic inflammation (6).

Activities like aerobic exercise and meditation have shown to reduce the level of inflammatory markers in a few months time (7, 8). These findings highlight the importance of adopting behavioral interventions to modulate stress and inflammation.

Summary by Dr. Dimpu Gogoi

References:

- Schneider KM, Blank N, Alvarez Y, Thum K, Lundgren P, Litichevskiy L, et al. <u>The enteric nervous system relays</u> <u>psychological stress to intestinal inflammation</u>. *Cell* (2023).
- 2. Dhabhar FS <u>Effects of stress on immune function: the good, the bad, and the beautiful</u>. *Immunol Res* (2014) 58:193-210.
- 3. Karagkouni A, Alevizos M, Theoharides <u>TC Effect of</u> stress on brain inflammation and multiple sclerosis.

 Autoimmun Rev (2013) 12:947-953.
- 4. Copeland WE, Wolke D, Lereya ST, Shanahan L, Worthman C, Costello EJ <u>Childhood bullying involvement predicts low-grade systemic inflammation into adulthood.</u> *Proc Natl Acad Sci U S A* (2014) 111:7570-7575.
- 5. Carlsson E, Frostell A, Ludvigsson J, Faresjo M

 <u>Psychological stress in children may alter the immune</u>

 <u>response.</u> *J Immunol* (2014) 192:2071-2081.
- 6. Osler M, Bendix L, Rask L, Rod NH <u>Stressful life events</u> and leucocyte telomere length: Do lifestyle factors, somatic and mental health, or low grade inflammation mediate this relationship? Results from a cohort of <u>Danish men born in 1953.</u> Brain Behav Immun (2016)

- 58:248-253.
- 7. Nicklas BJ, Hsu FC, Brinkley TJ, Church T, Goodpaster BH, Kritchevsky SB, et al. Exercise training and plasma C-reactive protein and interleukin-6 in elderly people.

 J Am Geriatr Soc (2008) 56:2045-2052.
- 8. Pace TW, Negi LT, Adame DD, Cole SP, Sivilli TI, Brown TD, et al. <u>Effect of compassion meditation on neuroendocrine, innate immune and behavioral responses to psychosocial stress.</u> *Psychoneuroendocrinology* (2009) 34:87-98.