

CHRONIC PAIN

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ANNOTATSIYA:

Chronic pain is a complex, multifaceted condition that affects millions of individuals worldwide, often disrupting daily functioning and reducing quality of life. Unlike acute pain, which serves as a warning signal of injury, chronic pain persists for months or even years, frequently without an identifiable cause. The pathophysiology of chronic pain involves complex interactions between the peripheral and central nervous systems, neuroplasticity, and psychosocial factors. This review examines the current understanding of chronic pain mechanisms, its impact on physical and mental health, and explores contemporary treatment approaches, ranging from pharmacological interventions to non-pharmacological therapies, including cognitive-behavioral therapy and physical rehabilitation. Despite advancements in pain management, many individuals continue to experience insufficient relief, highlighting the need for personalized, multimodal approaches. Understanding the biopsychosocial model of chronic pain and integrating it into clinical practice remains crucial for improving patient outcomes and enhancing quality of life.

Chronic pain: mechanisms and management approaches

Chronic pain is defined as pain that persists for more than three months or continues beyond normal tissue healing time. It is now considered a complex disease rather than a simple symptom, involving central sensitization, neuroinflammation, and maladaptive neural plasticity within the central nervous system (CNS).

Central sensitization refers to an abnormal increase in the excitability of neurons in the spinal cord and brain, leading to heightened pain responses even to non-painful stimuli. Neuroinflammation—an inflammatory process occurring within the CNS—is driven by activated microglia and astrocytes that release pro-inflammatory cytokines such as IL-6, IL-1 β , and TNF- α . These mediators amplify pain signaling and maintain chronic pain states.

Epidemiological data show that chronic pain affects approximately 20-25% of adults worldwide, representing one of the leading causes of disability. The persistence of pain is maintained not only by peripheral nociceptor activation but also by long-term changes in the CNS, including dysregulated descending inhibitory pathways and altered neurotransmitter systems such as glutamate and GABA.

Pharmacological Interventions in Pain Management

Pharmacological treatment remains the cornerstone of chronic pain management, though its efficacy is often limited by tolerance, side effects, and incomplete relief. Common drug classes include:

- Non-steroidal anti-inflammatory drugs (NSAID)-reduce peripheral inflammation and nociceptor sensitization.
- Opioid analgesics- act on mu-opioid receptors to inhibit pain transmission but pose a risk of dependence.
- Antidepressants-modulate serotonergic and noradrenergic pathways and may reduce neuroinflammatory mediators.
- Anticonvulsants-inhibit calcium channels to reduce neuronal hyperexcitability.

According to a systematic review, pharmacological interventions show measurable efficacy in reducing pain intensity, particularly when combined with other modalities. However, treatments are largely symptomatic and do not fully address the underlying neurobiological mechanisms such as central sensitization and glial activation.

Novel pharmacological targets under investigation include glial modulators, ion-channel blockers, and neuroimmune agents that may suppress neuroinflammatory pathways.

Non-Pharmacological Therapies

Non-pharmacological therapies are critical components of chronic pain management and often provide long-term benefits without the risks associated with pharmacological agents. These include:

-Physical rehabilitation- structured exercise programs, stretching, aerobic conditioning, and physiotherapy improve muscular strength, mobility, and pain tolerance.

-Psychological therapies- Cognitive Behavioral Therapy (CBT), mindfulness, and Acceptance and Commitment Therapy (ACT) reduce pain catastrophizing, anxiety, and depression associated with chronic pain.

-Complementary and alternative therapies- acupuncture, yoga, tai chi, and transcutaneous electrical nerve stimulation (TENS) are shown to modulate peripheral and central pain mechanisms.

Recent neuroimaging studies indicate that physical and psychological interventions can reduce central sensitization markers, including decreased activation in pain-processing brain regions and reduced levels of pro-inflammatory cytokines. For example, CBT has been shown to increase endogenous opioid release and decrease inflammatory mediators such as IL-6 and TNF- α .

Psychosocial Impact of Chronic Pain

Chronic pain extends beyond physiological dysfunction- it has significant psychosocial consequences. Patients often experience depression, anxiety, social isolation, loss of employment, and reduced quality of life. The biopsychosocial model recognizes pain as an interaction between biological, psychological, and social factors.

Psychosocial stress can exacerbate pain perception through neuroendocrine pathways involving cortisol and the hypothalamic-pituitary-adrenal (HPA) axis. Conversely, psychological interventions and social support systems improve coping mechanisms and reduce perceived pain intensity. Studies have demonstrated that targeting psychosocial factors significantly improves overall functional outcomes.

Multimodal and Interdisciplinary Approaches

Modern pain management emphasizes multimodal approaches- combining pharmacological,non-pharmacological,physical,and psychosocial interventions.According to a review in Research, Society and Development journal, integrative approaches yield superior outcomes compared with single-modality treatments.

Multimodal programs may include:

- Pharmacotherapy
- Exercise-based physical rehabilitation
- Psychological therapies
- Patient education and self-management strategies

Guidelines from the U.S CDC and WHO recommend multimodal,interdisciplinary pain management to minimize opioid use and enhance long-term recovery.

Combining physical rehabilitation with psychosocial education improves both pain reduction and functional ability,leading to better quality of life.

Neuroinflammation:A Central Mechanism

Neuroinflammation is increasingly recognized as a key driver of chronic pain persistence.Activated glial cells release pro-inflammatory mediators that sensitize nociceptive pathways and alter neuronal connectivity.Animal and human studies show that suppressing neuroinflammation- via pharmacological or behavioral interventions - reduces chronic pain symptoms and enhances CNS homeostasis.

Moreover,emerging therapies targeting microglial activation and cytokine modulation represent promising dorections for chronic pain research.The integration of these findings underscores the importance of addressing neuroimmune mechanisms alongside conventional treatment.

Conclusion

Chronic pain is a multifactorial condition involving complex interactions between peripheral and central neural processes,neuroinflammatory signaling,and psychosocial influences.

- pharmacological therapies remain essential but insufficient alone.
- Non-pharmacological therapies and physical rehabilitation provide complementary benefits,targeting both body and mind.

-Multimodal,interdisciplinary approaches yield the most effective and sustainable outcomes.

-Addressinf neuroinflammation and CNS plasticity may lead to novel,mechanism-based treatments for chronic pain.

A patient-centered model that integrates pharmacological,physical,and psychological therapies is now considered the gold standard for chronic pain management.

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