



# Molecular Signals and Biological Indicators in Pain Evaluation and Clinical Practice

Jonas Feldman\*

*Department of Anesthesiology and Pain Therapy, Central European University of Health Sciences, Vienna, Austria*

## DESCRIPTION

Pain is a complex physiological and psychological experience that varies widely among individuals, making objective assessment difficult in many clinical situations. Traditional approaches depend heavily on self-reporting tools such as numerical rating scales and visual analogs methods, which, while useful, are inherently subjective. This has led to increasing interest in measurable biological indicators that can provide more objective insight into the presence, intensity, and mechanisms of pain. Pain biomarkers refer to measurable substances or physiological changes that correlate with pain states, offering potential for improved diagnosis, monitoring, and treatment decisions.

Biomarkers associated with pain arise from multiple biological systems, including the nervous, immune, and endocrine systems. These markers can be detected in blood, cerebrospinal fluid, saliva, or even through imaging techniques. They often reflect underlying processes such as inflammation, nerve injury, or alterations in neurotransmitter activity. For example, inflammatory cytokines like interleukin-6 and tumor necrosis factor-alpha are frequently elevated in individuals experiencing chronic inflammatory pain. Their presence provides insight into ongoing immune responses that may be contributing to pain perception and persistence.

Neuropathic pain, which results from injury or dysfunction in the nervous system, has its own distinct biomarker profile. Substances such as substance P, calcitonin gene-related peptide, and nerve growth factor have been linked to changes in nerve signaling and heightened sensitivity. Elevated levels of these molecules are often associated with increased excitability of pain pathways, contributing to symptoms such as burning sensations or electric shock-like pain. Identifying these markers can assist clinicians in distinguishing neuropathic pain from other types, which is important because treatment strategies often differ significantly.

Advancements in neuroimaging have also contributed to the identification of pain-related biomarkers. Functional magnetic

resonance imaging and positron emission tomography can detect changes in brain activity associated with pain processing. Regions such as the anterior cingulate cortex, insula, and prefrontal cortex show altered activation patterns in individuals experiencing pain. These imaging findings provide a non-invasive method to observe how the brain interprets pain signals and how this interpretation may change over time or in response to treatment.

Genetic and epigenetic factors also play a role in pain perception and variability. Certain gene polymorphisms influence how individuals respond to pain stimuli or metabolize analgesic medications. For instance, variations in genes related to opioid receptors or enzymes involved in drug metabolism can affect both pain sensitivity and treatment outcomes. Epigenetic modifications, such as Deoxyribonucleic acid (DNA) methylation, may alter gene expression in response to environmental factors, contributing to chronic pain development. These findings highlight the importance of considering individual biological differences when evaluating pain and selecting therapies.

Hormonal influences further complicate the landscape of pain biomarkers. Cortisol, often referred to as the stress hormone, can influence pain sensitivity and inflammatory responses. Abnormal cortisol patterns have been observed in individuals with chronic pain conditions, suggesting dysregulation of the hypothalamic-pituitary-adrenal axis. Similarly, sex hormones such as estrogen and testosterone may modulate pain perception, which could partly explain observed differences in pain prevalence and intensity between genders.

The integration of multiple biomarkers into clinical practice offers opportunities for more accurate pain assessment. Instead of relying on a single marker, combining data from inflammatory, neural, and hormonal indicators may provide a more comprehensive understanding of a patient's pain profile. This approach can support more precise diagnoses and help guide treatment decisions, particularly in complex cases where pain has multiple contributing factors.

**Correspondence to:** Jonas Feldman, Department of Anesthesiology and Pain Therapy, Central European University of Health Sciences, Vienna, Austria, E-mail: jonas.feldman@ceuhs.at

**Received:** 27-Feb-2026, Manuscript No. JPMME-26-31519; **Editor assigned:** 02-Mar-2026, Pre QC No. JPMME-26-31519; **Reviewed:** 16-Mar-2026, QC No. JPMME-26-31519; **Revised:** 23-Mar-2026, Manuscript No. JPMME-26-31519; **Published:** 30-Mar-2026, DOI: 10.35248/2684-1320.26.12.379

**Citation:** Feldman J (2026). Molecular Signals and Biological Indicators in Pain Evaluation and Clinical Practice. *J Pain Manage Med.* 12:379.

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## CONCLUSION

Pain remains a multifaceted experience influenced by biological, psychological, and social factors. While biomarkers cannot fully capture this complexity, they offer valuable tools for improving

understanding and management. With ongoing advancements in biomedical research and technology, the role of biomarkers in pain medicine is likely to expand, contributing to more precise and individualized care strategies.