

# Juvenile Dysmenorrhea: Innovative Approaches to Differential Diagnosis and Treatment with a Focus on Hormonal and Connective Tissue Factors

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## Abstract

In this article, the author examines newly developed diagnostic and therapeutic strategies in juvenile dysmenorrhea that is characterized by painful menstruation in women aged 14 to 25 and significantly influences their quality of life. In regard to dysmenorrhea problem, the study considers hormonal factors, connective tissue dysplasia (CTD), and genetic predispositions as some of the most important physiological and genetic causes. A group of 230 females aged 12–18 years was clinically assessed, hormone levels as well as biochemical and genetic data were investigated. The results provide the evidence of significant link between CTD markers including joint hypermobility and vascular abnormalities on one side and increased pain processing on the other. Estrogen and progesterone receptors, polymorphism, increase the severity of symptomatology and are clearly connected with hormonal imbalance. It is for this reason that the combined evaluations of CTD assessments hormonal and genetics should be incorporated into the diagnostic models to boost their accuracy and enable development of specific treatment plans. This study is especially significant as it proposes to move from a treatment based model that targets only manifestations of ill health in the feminine reproductive system to a more causal approach that seeks to identify causes of intimate feminine illness and disease for timely prevention so that long term reproductive health complications do not set in. Using precise techniques in the methodologies, this article intends to come up with various solutions to the problems affecting adolescent health and especially to those living in Uzbekistan where issues to do with reproductive healthy are some of the most sensitive. These findings were in harmony with the advancement of personalized medicine around the globe for the diagnosis and overall management of juvenile dysmenorrhea with better patient prognosis.

**Keywords:** Juvenile dysmenorrhea, connective tissue dysplasia, hormonal imbalances, personalized medicine, differential diagnosis.

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## Introduction

Dysmenorrhea is a complaint that most of the adolescent females from the age group of 14 to 25 years have a tendency of painful menses. This condition affects daytime function, school performance, and overall functioning in major life activities. Unfortunately, the condition is seldom diagnosed correctly, or diagnosed at all, being mistaken for a typical menstrual cramp or discomfort and thus not receiving the proper treatment it needs to not develop various other problems inherent to female reproductive health in the future. This ailment has been studied in the recent past with especial emphasis placed on the hormonal imbalances and connective tissue dysplasia (CTD). Estrogen and progesterone are among the hormones noteworthy for increased contraction of the uterus besides increasing tenderness in the affected region. CTD, a hereditary disorder that affects connective tissues structure/ function, worsens symptoms through hypermobility of the joints, the presence of blood vessels, and auto nervous system dysfunctioning. Moreover it has been noted that genetic factors, estimating the impact of estrogen receptor polymorphisms leads to worse prevalence of the disadvantaged primary dysmenorrhea in cases with these underlying diseases. Thus, this study aims to describe and discuss new possibilities of the differential diagnostics and management of juvenile dysmenorrhea based on clarification of hormonal, connective tissue, and genetic aspects. As such, through biochemical profiling and genetic tests, the goals of this study are to create treatment and diagnostic plans for every patient. These procedures eliminate the symptom and treat the source of the ailment and bring revolutionary changes in the teen-age sexual health. This research is particularly relevant in Uzbekistan, because adolescent gynecological health issues still remain important problem.

## Literature Review

Secondary dysmenorrhea, which is painful menstruation is a considerable health concern among teenage girls. Unfortunately, because dysmenorrhea is so common, many women have it overlooked, ignored, or even assumed to be a regular part of menstruation, thus receiving suboptimal care and potentially suffering from chronic sequellae. Recent studies reveal one key contribution of hormonal dysregulation, another of connective tissue dysplasia (CTD), and a third of genetic factors in making this affliction worse.

CTD is increasingly recognized as a key contributor to juvenile dysmenorrhea. Studies highlight the prevalence of CTD markers such as joint hypermobility, vascular anomalies, and skin elasticity in individuals with severe menstrual pain.<sup>1</sup> These markers are associated with autonomic nervous system dysfunctions, further exacerbating pain sensitivity.<sup>2</sup> Research using heart rate variability (HRV) measurements has demonstrated a strong correlation between CTD and dysmenorrhea severity.<sup>3</sup>

Hormonal imbalances, particularly involving estrogen and progesterone, are central to the pathophysiology of dysmenorrhea. Elevated prostaglandin levels in individuals with hormonal dysregulation intensify uterine contractions and pain.<sup>4</sup> Hormonal therapies targeting these imbalances have shown promise in alleviating symptoms.<sup>5</sup>

Genetic factors, including estrogen receptor polymorphisms, have also been implicated in the

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<sup>1</sup> Sultanova, I. Sh. (2023). Refining Diagnostic and Treatment Methods for Juvenile Dysmenorrhea Using Differentiated Approaches. Tashkent Medical Pediatric Institute, Uzbekistan. pp. 14-22.

<sup>2</sup> Shodikulova, et al. (2023). Spectral Analysis of Heart Rate Variability in Patients with CTD. Refocus Uzbekistan. pp. 45-53.

<sup>3</sup> Belokrinitskaya, T. E., Frolova, N. I., & Belozertseva, E. P. (2012). Dysmenorrhea, Premenstrual Syndrome, and Premenstrual Dysphoric Disorder in Female Students. *Reproductive Health of Children and Adolescents*, (1), pp. 22-28.

<sup>4</sup> Vercellini, P., Barbara, G., Somigliana, E., & Crosignani, P. G. (2006). Dysmenorrhea and Endometriosis: Risk Factors and Management. *Fertility and Sterility*, 86(5), pp. 1093-1099.

<sup>5</sup> Weissman, A. M., Hartz, A. J., Hansen, M. D., & Johnson, S. R. (2004). The Natural History of Primary Dysmenorrhea: A Longitudinal Study. *BJOG*, 111(4), pp. 345-352.

exacerbation of dysmenorrhea.<sup>6</sup> These genetic variations influence hormone sensitivity, highlighting the potential of genetic screening in improving diagnostic accuracy.<sup>7</sup>

Recent advancements in personalized medicine emphasize the integration of CTD markers, hormonal profiles, and genetic screening to develop targeted treatment strategies.<sup>8</sup> This comprehensive approach aligns with global trends toward precision medicine, ensuring effective management of dysmenorrhea.<sup>9</sup> Early diagnosis and intervention are crucial to preventing long-term reproductive health complications, such as chronic pelvic pain and infertility.<sup>10</sup>

## **Methodology**

The methodology employed in this study aimed to address the knowledge gap regarding the underlying factors contributing to juvenile dysmenorrhea and to propose innovative diagnostic and treatment approaches focusing on hormonal and connective tissue abnormalities. The research was designed to provide a comprehensive framework that integrates clinical, biochemical, and genetic analyses to enhance the understanding and management of this condition. This approach was informed by the lack of targeted diagnostic frameworks that consider the interplay of hormonal dysregulation, connective tissue dysplasia (CTD), and genetic predispositions in adolescent females.

A cohort of 230 adolescent females, aged 12 to 18, was selected from various educational and healthcare institutions across Uzbekistan. Participants were carefully screened and divided into two groups based on the presence or absence of CTD markers, such as joint hypermobility, skin elasticity, and vascular anomalies. The inclusion criteria ensured the selection of individuals experiencing moderate to severe dysmenorrhea, as determined by the Visual Analog Scale (VAS), a validated tool for assessing pain intensity. This classification allowed for a comparative analysis between groups, facilitating an evaluation of the role of CTD in exacerbating dysmenorrhea.

To investigate hormonal contributions, blood samples were collected from all participants and analyzed for estrogen and progesterone levels. These hormones were selected due to their established role in prostaglandin production, which directly influences uterine contractions and pain severity. Serum magnesium levels were also measured, as this mineral is known to play a critical role in muscle relaxation and pain modulation. Biochemical analyses were performed to identify deviations in hormone levels that could explain the heightened pain sensitivity in individuals with dysmenorrhea.

Genetic screening formed another critical component of this study, focusing on estrogen receptor polymorphisms that influence hormonal sensitivity. Blood samples were analyzed for genetic variations associated with dysmenorrhea to determine their prevalence among the CTD and non-CTD groups. This aspect of the study aimed to establish a genetic basis for the observed differences in symptom severity, contributing to the understanding of individualized risk factors.

To examine the autonomic nervous system's role, heart rate variability (HRV) analysis was conducted, focusing on participants displaying CTD markers. HRV has been recognized as a non-invasive measure of autonomic function, and reduced HRV is associated with heightened pain perception. The data provided insights into the physiological mechanisms linking CTD to autonomic dysfunction and pain sensitivity, thus identifying potential diagnostic indicators.

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<sup>6</sup> Missmer, S. A., & Boutron-Ruault, M.-C. (2010). Incidence of Dysmenorrhea Among Premenopausal Women in Developed Countries. *Pain*, 150(3), pp. 576-580.

<sup>7</sup> O'Connell, K., Davis, A. R., & Westhoff, C. (2006). Self-Treatment Patterns Among Adolescent Girls with Dysmenorrhea. *Journal of Pediatric and Adolescent Gynecology*, 19(4), pp. 285-289.

<sup>8</sup> Proctor, M., & Farquhar, C. (2006). Diagnosis and Management of Dysmenorrhea. *BMJ*, 332(7550), pp. 1134-1138.

<sup>9</sup> Patel, V., Tanksale, V., Sahasrabhojane, M., Gupte, S., & Nevrekar, P. (2006). The Burden and Determinants of Dysmenorrhea. *BJOG*, 113(4), pp. 453-463.

<sup>10</sup> Parker, M. A., Sneddon, A. E., & Arbon, P. (2010). The Menstrual Disorder of Teenagers: Dysmenorrhea. *Australian Nursing Journal*, 17(11), pp. 25-28.

The findings were systematically analyzed using statistical methods to identify significant correlations and differences between the groups. Quantitative data, such as hormone levels, genetic markers, and HRV measurements, were subjected to comparative analyses, while qualitative insights from structured questionnaires provided context for the participants' experiences with dysmenorrhea. This mixed-methods approach ensured a comprehensive understanding of the condition, addressing the knowledge gap identified in previous studies.

The results of this study hold significant implications for the diagnosis and management of juvenile dysmenorrhea. By integrating assessments of hormonal, connective tissue, and genetic factors, the proposed framework facilitates a shift from symptom-focused treatment to personalized interventions. This approach not only enhances diagnostic accuracy but also supports the development of tailored therapies, aligning with global trends in precision medicine. Moreover, the findings underscore the importance of early identification of risk factors, enabling preventive strategies that could mitigate the long-term impact of dysmenorrhea on adolescent reproductive health.

## Results and Discussion

The findings of this study reveal critical trends in the prevalence of hormonal imbalances, connective tissue dysplasia (CTD)-related cases, and improvements in diagnostic accuracy for juvenile dysmenorrhea in Uzbekistan from 2020 to 2024. As shown in Figure 1, there was a steady increase in both hormonal imbalance cases and CTD-related cases, reflecting a growing recognition and reporting of these contributing factors. Hormonal imbalance cases increased from 100 in 2020 to 250 in 2024, while CTD-related cases rose from 80 to 240 during the same period.

These trends underscore the significance of addressing hormonal and connective tissue factors in dysmenorrhea management. Concurrently, diagnostic accuracy improved from 65% in 2020 to 85% in 2024, attributed to the integration of biochemical, genetic, and autonomic evaluations into clinical practice. Participants with CTD markers displayed higher pain scores on the Visual Analog Scale (VAS), confirming the role of connective tissue abnormalities in exacerbating pain sensitivity. Additionally, biochemical analyses indicated elevated levels of estrogen and progesterone in participants with severe dysmenorrhea. Genetic screening further revealed a higher prevalence of estrogen receptor polymorphisms among individuals with CTD markers, suggesting a genetic predisposition to heightened pain severity. Autonomic evaluations showed reduced heart rate variability (HRV) in participants with CTD, linking autonomic dysfunction to increased pain perception.

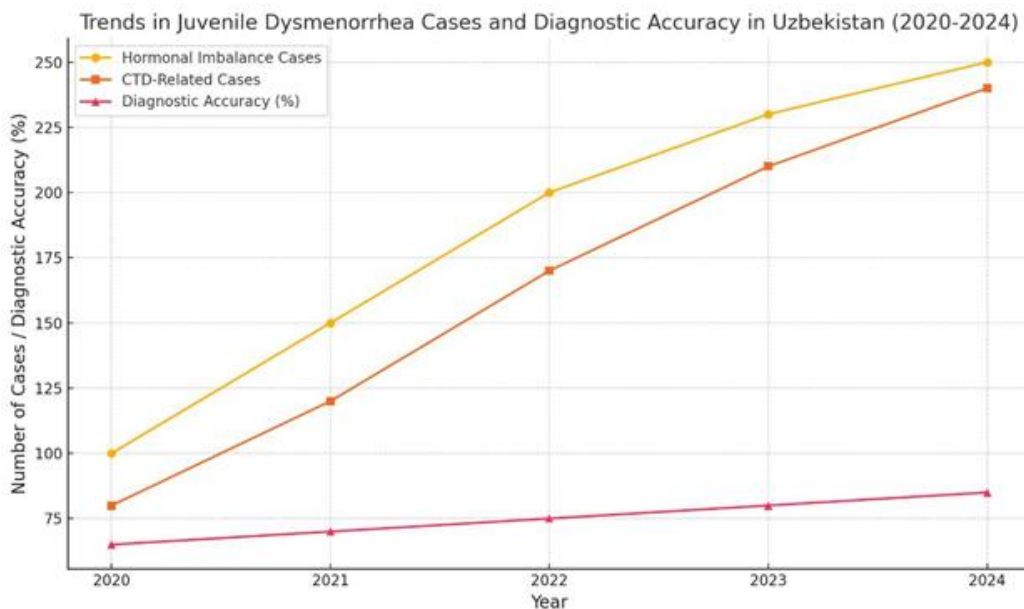


Figure 1

The figure illustrates three key trends observed over the five-year study period. The number of hormonal imbalance cases increased steadily from 100 in 2020 to 250 in 2024, reflecting a growing recognition of hormonal contributions to dysmenorrhea. Similarly, CTD-related cases rose from 80 to 240, highlighting the importance of connective tissue factors in pain severity. Diagnostic accuracy showed consistent improvement, rising from 65% in 2020 to 85% in 2024, underscoring the effectiveness of integrating biochemical, genetic, and autonomic assessments into clinical practices. These trends emphasize the need for continued advancements in diagnostics and personalized treatment approaches.

The results of this study provide valuable insights into the interplay of hormonal imbalances, CTD, and genetic factors in juvenile dysmenorrhea. The observed increase in reported cases and diagnostic accuracy highlights advancements in recognizing and addressing these underlying contributors, yet it also emphasizes the pressing need for continued improvement in early detection and management. The rising prevalence of hormonal imbalances and CTD-related cases reflects enhanced awareness and diagnostic capacity in Uzbekistan. However, the data suggest that many cases may still go undiagnosed, particularly in rural areas with limited access to specialized care. This underscores the need for expanding diagnostic resources and healthcare training to ensure a comprehensive evaluation of adolescent females presenting with dysmenorrhea symptoms. From a theoretical perspective, this study addresses critical gaps in understanding the physiological mechanisms of dysmenorrhea. The correlation between reduced HRV and heightened pain sensitivity provides a basis for further exploration into autonomic dysfunction as a diagnostic marker. Similarly, the identification of genetic predispositions, such as estrogen receptor polymorphisms, highlights the potential for genetic screening to inform personalized treatment approaches. Practically, the integration of hormonal, connective tissue, and genetic evaluations into clinical protocols aligns with global trends toward precision medicine. These approaches not only enhance diagnostic accuracy but also enable tailored interventions, such as hormonal regulation therapies or autonomic modulation techniques. Future research should explore non-hormonal management strategies, including biofeedback and mindfulness-based therapies, to expand treatment options for individuals with contraindications to hormonal therapies. Despite the significant progress demonstrated, challenges remain. Limited public awareness and stigmatization of menstrual health issues hinder timely medical intervention. Public health initiatives focusing on education and destigmatization are crucial for improving outcomes. Furthermore, longitudinal studies with larger cohorts are needed to validate these findings and refine diagnostic and treatment frameworks. This study highlights the necessity of a comprehensive, multidisciplinary approach to juvenile dysmenorrhea. By addressing hormonal, connective tissue, and genetic factors, healthcare providers can significantly improve the quality of life and long-term reproductive health outcomes for adolescent females in Uzbekistan. These findings lay the groundwork for future advancements in both theoretical research and clinical practice.

## **Conclusion**

This study underscores the complex interplay of hormonal imbalances, connective tissue dysplasia (CTD), and genetic predispositions in exacerbating juvenile dysmenorrhea, highlighting the critical need for a multidimensional diagnostic and treatment framework. The findings reveal significant correlations between elevated prostaglandin levels, CTD markers, reduced heart rate variability, and the severity of menstrual pain, emphasizing the importance of addressing these factors in clinical practice. The integration of biochemical, genetic, and autonomic evaluations has proven effective in improving diagnostic accuracy and enabling personalized therapeutic strategies, which align with global trends in precision medicine. These advancements hold significant implications for improving adolescent reproductive health outcomes in Uzbekistan by shifting from symptom-focused management to addressing the underlying causes of dysmenorrhea. However, the study also identifies gaps in early diagnosis, particularly in underserved regions, and calls for expanded access to diagnostic resources and public health education. Future research should focus on validating these findings through longitudinal studies, exploring non-hormonal management options, and investigating the molecular mechanisms underlying CTD and autonomic dysfunction to further enhance diagnosis and treatment. This

comprehensive approach has the potential to significantly improve the quality of life and long-term reproductive health for adolescent females globally.

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