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Review Article

Herbal Antispasmodics with Magnesium: A Novel Approach to Dysmenorrhea Management

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ABSTRACT

Dysmenorrhea, also known as menstruation discomfort, affects more than 70% of reproductive-age women worldwide, considerably impairing their quality of life and everyday function. Current treatment options rely mostly on nonsteroidal antiinflammatory medicines (NSAIDs), which provide symptom relief but are frequently linked with gastrointestinal discomfort, renal side effects, and hormone imbalance with prolonged usage. There is an increasing need for natural, safer, and more patientfriendly options. Herbal antispasmodics like Zingiber officinale (Ginger), Foeniculum vulgare (Fennel), and Valeriana officinalis (Valerian) have demonstrated effective outcomes in alleviating uterine spasms and inflammation by inhibiting COX enzymes and promoting smooth muscle relaxation, as evidenced by clinical studies and metaanalyses. At the same time, magnesium, a vital mineral, has shown effectiveness in easing menstrual pain by influencing prostaglandin production and calcium ion movement, thus relaxing uterine muscles and reducing cramps. Effervescent tablets present an innovative drug delivery system that guarantees quick onset of effects, better bioavailability, pleasant taste, and increased patient adherence, particularly during episodes of acute pain. The use of magnesium combined with standardized herbal antispasmodic agents in an effervescent tablet form has not been investigated in current market offerings, rendering this method both novel and commercially viable. This analysis examines the scientific foundation, pharmacological principles, formulation approaches, an current market scenario for this new herbal-magnesium effervescent dosage form, setting the stage for future investigations, product innovation, and potential nutraceutical or AYUSH-oriented commercialization.

INTRODUCTION

Dysmenorrhea:-

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Dysmenorrhea, commonly known as menstrual discomfort or cramps, is one of the most widespread gynecological issues that affects women of reproductive age. It is marked by cramping pain in the lower abdomen, often accompanied by symptoms such as nausea, fatigue, headaches, diarrhea, and mood swings, which can significantly affect daily activities and overall quality of life. It is estimated that worldwide, more than 70% of adolescent and young adult females experience dysmenorrhea, with varying levels of intensity and recurrence.(1)

Dysmenorrhea is broadly classified into two types:-

Primary dysmenorrhea is characterized by menstrual pain without any detectable pelvic condition. It generally starts in adolescence, a few years after the onset of menstruation, and is thought to be linked to an excess production of uterine prostaglandins, especially prostaglandin $F2\alpha$ (PGF2 α). These prostaglandins lead to strong uterine contractions, diminish uterine blood flow (causing ischemia), and heighten the perception of pain. Elevated levels of prostaglandins can also lead to systemic symptoms such as diarrhea and nausea.(2)

Secondary dysmenorrhea arises from existing pelvic or reproductive system disorders, including endometriosis, adenomyosis, pelvic inflammatory disease (PID), fibroids, or the use of intrauterine devices (IUDs). It usually develops later in life and tends to escalate if the underlying issue remains unaddressed.(2)

The pathophysiology of dysmenorrhea is characterized by a complex interaction of hormonal, inflammatory, and neurological components. A decrease in progesterone levels before menstruation triggers the shedding of the endometrium, resulting in the release of

arachidonic acid, which starts the production of prostaglandins and leukotrienes. These biochemical agents increase uterine tone, cause hypercontractility, and amplify pain sensitivity by activating peripheral pain receptors and pathways within the central nervous system.(2)

Factors that also contribute to dysmenorrhea include:-

- A deficiency in magnesium, which increases the excitability of uterine muscles.
- Psychological stress and emotional turmoil.
- Lack of physical activity or a sedentary lifestyle.
- Being overweight or underweight.
- Early onset of menstruation, extended menstrual periods, or heavy menstrual bleeding.
- Although many women use NSAIDs and oral contraceptives, side effects are common, leading some to seek natural alternatives. This has spurred interest in complementary and alternative medicine (CAM) approaches, especially herbal treatments and nutritional supplements.

This review examines an innovative method for managing dysmenorrhea by combining herbal antispasmodics with magnesium in an effervescent tablet formulation, providing a quicker-acting, natural, and user-friendly solution for menstrual pain relief.

Enhanced Section: Global Burden & Lifestyle Impact of Dysmenorrhea:-

Dysmenorrhea is not only common in medical settings but also has significant social ramifications. It impacts 45–95% of menstruating women worldwide, with the highest occurrence noted among adolescents and young adults. A survey in India indicated that over 72% of college-

aged women suffer from moderate to severe menstrual pain, often resulting in missed school, college, or workdays, as well as a notable decline in productivity and concentration.(3)

Research from Japan found that one-third of women experiencing dysmenorrhea miss at least one day of work each month, and the situation is even more critical in developing nations due to inadequate access to pain relief and menstrual hygiene products.(4)

Additionally, lifestyle elements such as unhealthy diets, stress, insufficient physical activity, and irregular sleep patterns have been shown to exacerbate the severity and duration of menstrual cramps. Women who lead sedentary lifestyles, experience high levels of anxiety, or have low magnesium intake are particularly vulnerable to more intense symptoms.(5)

The effect of lifestyle on dysmenorrhea:-

Specifically, a lack of physical activity can lead to uterine ischemia and reduced blood flow in the pelvic area, while elevated stress levels can increase cortisol, heightening pain sensitivity through inflammation. A study conducted by Unsal et al. (2010) found that students who engaged in unhealthy habits, such as skipping meals and poor sleep practices, reported significantly greater menstrual pain compared to those who led more balanced lifestyles. These results suggest that comprehensive management of dysmenorrhea should encompass just medication for relief, but also lifestyle modifications as a preventive approach.(6)

Scientific Basis for Herbal and Magnesium Use:-

The use of herbal antispasmodics and magnesium for treating dysmenorrhea is supported by growing pharmacological and clinical research. For instance, Zingiber officinale (ginger) inhibits cyclooxygenase (COX) enzymes, mimicking the action of NSAIDs but without adverse effects on the stomach. Foeniculum vulgare (fennel) works through phytoestrogens and facilitates smooth muscle relaxation, whereas Valeriana officinalis acts centrally as a GABA agonist to alleviate cramps linked to anxiety.

Magnesium operates at a neuromuscular level by decreasing intracellular calcium and reducing uterine hypercontractility. A randomized placebocontrolled trial conducted by Ziaei et al. (2001) found that magnesium supplementation notably decreased pain intensity scores over two consecutive menstrual cycles. Consequently, these substances provide complementary mechanisms—anti-inflammatory, antispasmodic, and muscle-relaxant—making them potential candidates for combination therapy.(7)

Necessity for Innovative Formulation **Methods:**-

Even though there is growing awareness around menstrual health, the majority of existing pharmaceutical and herbal solutions for dysmenorrhea continue to depend on traditional dosage forms like capsules, syrups, and tablets, which frequently face issues related to delayed onset, poor taste, or gastrointestinal discomfort. Furthermore, a significant number of patients favor non-hormonal, plant-based alternatives; however, the current market lacks rapid-acting, user-friendly solutions tailored for menstrual pain relief.

Effervescent tablets present an innovative solution. They facilitate quick disintegration and absorption, providing faster relief and better adherence from patients, particularly during intense pain episodes. A 2020 review by Kalita et

al. highlighted that effervescent delivery systems enhance both pharmacological efficacy and patient satisfaction, especially in situations requiring immediate action, such as spasmodic or musculoskeletal pain. The combination of standardized herbal antispasmodics with magnesium in this format has yet to be commercialized, making this formulation a unique and viable concept in the realm of menstrual wellness.(8)

Pathophysiology of Dysmenorrhea:-

Primary dysmenorrhea is characterized by painful menstruation that occurs without any underlying pelvic conditions. It usually affects young women, particularly adolescents, and is strongly associated with hormonal fluctuations that trigger a series of uterine and neurochemical responses leading to discomfort.

1. Hormonal Triggers and Prostaglandin Overproduction

Towards the conclusion of the luteal phase, levels of progesterone decrease, prompting endometrial cells to release arachidonic acid, which is then transformed into prostaglandins (PGF₂α and PGE₂) through the cyclooxygenase (COX) pathway. The rise in prostaglandins heightens uterine contractions, diminishes blood circulation, and increases the sensitivity of pain receptors. (9) Women experiencing dysmenorrhea have prostaglandin levels in menstrual fluid that are 3–4 times higher than those who do not have pain.(10)

2. Uterine Hypercontractility and Ischemia

Heightened levels of $PGF_2\alpha$ lead to forceful, frequent, and disorganized uterine contractions, resulting in increased intrauterine pressure. This elevation causes vasoconstriction of spiral arteries,

leading to diminished blood supply to the uterus, which induces ischemia and discomfort. The ischemic effects are similar to angina-like chest pain in terms of mechanism, as both scenarios initiate anaerobic metabolism and the accumulation of lactic acid.(11)

3. The Function of Vasopressin and Leukotrienes

Some women experience elevated levels of vasopressin, a hormone that encourages vasoconstriction in the uterus and increases the activity of PGF₂\alpha. Leukotrienes, particularly LTC₄ and LTD₄, are generated via the lipoxygenase pathway and play a role in smooth muscle contraction and inflammation, especially women who do not respond well to NSAIDs.(12,13)

4. Peripheral and Central Sensitization

Frequent intense contractions and episodes of ischemia can lead to hyperalgesic priming, where peripheral nociceptors become overly sensitive. Over time, central sensitization develops, resulting in structural alterations in brain regions that regulate pain, causing even slight uterine contractions to be perceived as painful.(11)

5. Secondary Dysmenorrhea

This type results from pathological conditions such as endometriosis, adenomyosis, fibroids, or pelvic inflammatory disease. These conditions lead to abnormal growth of endometrial or uterine tissue, increased inflammation, and pain that may begin earlier and last longer than typical menstrual cramps.(14)

Herbal Antispasmodics for Managing Dysmenorrhea:-

Table.1 Commonly studied herbal antispasmodics, their active constituents, and mechanisms of action.



Sr.	Common	Botanical	Active	Mechanism of action	Reference
No.	name	name	constituent		
1.	Ginger	Zingiber	Gingerols,	Inhibits COX and LOX	Ozgoli et al.,
		officinale	Shogaols	→ reduces prostaglandin	2009; Rahnama
				synthesis $\rightarrow \downarrow$ uterine	et al., 2012;
				contractions	Daily et al., 2015
2.	Fennel	Foeniculum	Anethole,	Phytoestrogenic effect; inhibits	Modarres et al.,
		vulgare	Fenchone,	uterine contractions by blocking	2011; Delaram
			Estragole	oxytocin receptors	et al., 2011
3.	Valerian	Valeriana	Valerenic acid,	Modulates GABA receptors →	Mirabi et al.,
		officinalis	Iridoids,	muscle relaxant and CNS calming	2011; Sadeghi
			Flavonoids	effect	Aval et al., 2014
4.	Chamomile	Matricaria	Apigenin,	Anti-inflammatory, COX-2	Sharifi et al.,
		chamomilla	Bisabolol,	inhibition, mild sedative; relieves	2014; Srivastava
			Flavonoids	cramps and mood swings	et al., 2010
5.	Peppermint	Mentha	Menthol,	Calcium channel blocker;	Nahidi et al.,
		piperita	Menthone,	antispasmodic effect on uterine	2009; Uritu et
			Flavonoids	and intestinal smooth muscle	al., 2021

Dysmenorrhea, especially its primary type, is often linked to heightened uterine contractions driven by increased prostaglandin levels, which lead to ischemia and pain. A range of natural agents known as herbal antispasmodics show promise in alleviating this condition by influencing smooth muscle tone, decreasing inflammation, and regulating hormonal responses. These herbs have a strong reputation in various traditional medical systems, including Ayurveda, Persian, and Chinese medicine, and are gaining more backing from clinical studies and scientific research.

Mechanisms of Action:-

Herbal antispasmodics exert their effects through various pharmacological pathways:

- **Prostaglandin Suppression:** Numerous herbs inhibit COX and LOX pathways, resulting in decreased levels of PGF₂α and PGE₂.
- Calcium Channel Inhibition: Some herbs diminish the availability of intracellular calcium in uterine smooth muscle cells, facilitating relaxation.

- Modulation of Neurotransmitters: Certain herbs influence pathways involving GABA, serotonin, or adrenergic systems to promote smooth muscle relaxation and provide sedative effects.
- **Phytoestrogenic Effects:** Some herbs interact with estrogen receptors, modifying the tone of the uterine muscle.

Commonly Studied Herbal Antispasmodics:-

1. Ginger (Zingiber officinale)

Ginger is among the most extensively researched herbal remedies for dysmenorrhea. Its primary active compounds, gingerols and shogaols, work by inhibiting the production of prostaglandins through COX and LOX pathways, which helps to alleviate uterine spasms.

Clinical Support:-

In a randomized controlled trial, ginger (250 mg taken four times daily) demonstrated effectiveness comparable to that of mefenamic acid and ibuprofen in diminishing the intensity of menstrual pain (Ozgoli et al., 2009). A meta-analysis



conducted by Daily et al. (2015) further validated the effectiveness of ginger in various RCTs.l



Fig.2 Ginger (Zingiber officinale)

2. Fennel (Foeniculum vulgare)

Fennel contains anethole, a compound with estrogenic properties that also inhibits uterine muscle contractions by acting on oxytocin receptors and calcium channels.

Clinical Support:-

Modarres et al. (2011) found fennel extract (30 drops every 4 hours) significantly reduced menstrual pain intensity compared to placebo. It also improved associated symptoms like bloating and nausea.



Fig.3 Fennel (Foeniculum vulgare)

3. Valerian (Valeriana officinalis)

Valerian exerts a GABAergic effect, promoting central and peripheral muscle relaxation. Its components such as valerenic acid reduce uterine spasms through modulation of GABA-A receptors.

Clinical Support:

A double-blind RCT showed that valerian significantly reduced pain severity in young women over two menstrual cycles (Mirabi et al., 2011). Sadeghi Aval et al. (2014) also noted reduced reliance on rescue analgesics in valerian users.



Fig.4 Valerian (Valeriana officinalis)

4. Chamomile (Matricaria chamomilla)

Chamomile is rich in apigenin and bisabolol, which are flavonoids that have anti-inflammatory properties and mild sedative effects. These compounds inhibit the release of prostaglandins and decrease the contractions of the uterus.

Clinical Evidence:

According to Sharifi et al. (2014), women who drank chamomile tea during their menstrual periods experienced significantly reduced pain and improved emotional health. The effectiveness of chamomile was found to be similar to that of NSAIDs, but with fewer side effects.



Fig.5 Chamomile (Matricaria chamomilla)



5. Peppermint (Mentha piperita)

Peppermint oil is rich in menthol, which acts as a calcium channel blocker, thereby relaxing uterine and gastrointestinal smooth muscles. Though more commonly used for IBS, it shows potential in dysmenorrhea.

Clinical Support:

A small RCT by Nahidi et al. (2009) indicated that peppermint significantly reduced dysmenorrhea scores in students after one cycle of use.



Fig.6 Peppermint (Mentha piperita)

Role of Magnesium in Dysmenorrhea Management:-

Magnesium is essential for neuromuscular functioning, vascular regulation, and balancing prostaglandins, all of which are significant in the development of dysmenorrhea. Its role as a natural antagonist to calcium helps to alleviate uterine muscle contractions, while its properties that reduce inflammation and promote blood vessel dilation aid in relieving pain.

Mechanism of Action

 Magnesium inhibits calcium channels, leading to lowered intracellular calcium concentrations in uterine smooth muscle cells, which results in reduced muscle contractions and cramps (Walker et al., 1998).

- It also suppresses the synthesis of prostaglandin $F_2\alpha$, a key factor in increased uterine contractility (Zahradnik et al., 2010).
- Deficient magnesium levels are linked to heightened pain sensitivity, alterations in mood, and fatigue during menstruation (Gartlehner et al., 2019).

Clinical Evidence

- A randomized, double-blind, placebocontrolled trial conducted by Ziaei et al. (2001) found that women who received 250 mg of magnesium daily from day 15 of their menstrual cycle until menstruation reported significantly lower menstrual pain intensity when compared to those given a placebo.
- In a similar study, Ylikorkala et al. (1977) showed that taking magnesium supplements resulted in a quantifiable reduction in uterine contractions and a decrease in prostaglandin levels in menstrual fluid.
- A systematic review by Gartlehner et al. (2019) concluded that magnesium is effective for treating mild-to-moderate primary dysmenorrhea, demonstrating few side effects and being well-tolerated.

Critical Assessment of Clinical Trials:

Although the clinical studies analyzed show favorable results for both herbal remedies and magnesium in treating dysmenorrhea, several limitations are significant. Numerous trials have small participant numbers, brief durations (1–2 menstrual cycles), or lack a placebo comparison, which could influence the statistical reliability. For example, the magnesium study conducted by Ziaei et al. (2001) was encouraging but only involved 50 subjects. Likewise, studies on valerian and chamomile do not maintain consistency in extract dosages and types of formulations. These deficiencies stress the necessity for widespread,



multicentric studies to validate effectiveness, refine dosages, and assess against conventional NSAIDs.

A recent meta-analysis reaffirmed that ginger is comparably effective as NSAIDs while resulting in fewer gastrointestinal issues (Daily et al., 2020). Another review emphasized the potential of herbs from the Lamiaceae family, such as peppermint, in managing pain, supporting their use in treating dysmenorrhea (Uritu et al., 2021). Furthermore, a double-blind randomized trial found magnesium supplementation significantly alleviated pain intensity and enhanced mood scores among teenagers experiencing primary dysmenorrhea (Shahnazi et al., 2022). (15)

Commercial Herbal Products for Dysmenorrhea:-

Several herbal formulations are presently sold in both Indian and international markets for alleviating primary dysmenorrhea and menstrual discomfort. These products typically incorporate herbs known for their antispasmodic, anti-inflammatory, or hormonal-balancing properties. They come in various forms such as capsules, syrups, teas, tonics, and granules; however, the absence of effervescent tablets indicates an opportunity for your formulation concept to fulfill this gap.

Tabel.2 Marketed herbal formulations for menstrual pain relief.

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Sr.No	Brand Name	Manufacturer	Formulation	Key Herbal	Claim/Use		
1.	EveCare	Himalaya	Syrup/	Ashoka (Saraca indica),	Menstrual		
		Wellness	Capsule	Lodhra (Symplocos	regulation, relief from		
			_	racemosa), Dashamoola	cramps		
2.	MyTanix	Aimil	Syrup	Shatavari, Ashoka,	Relieves pain, regulates		
	Syrup	Pharmaceuticals		Lodhra, Guduchi	cycles		
3.	Femiza	Solumiks	Tablet	Shatavari, Ashoka,	Supports uterine health,		
	Tablets	Herbaceuticals		Lodhra, Nagarmotha	manages pain		
4.	Gynocare	Ayusya	Capsule	Lodhra, Ashoka,	Alleviates menstrual		
	Capsules	Naturals		Shatavari, Kumari	discomfort		
5.	Herbocare	Morpheme	Capsule	Ginger, Fennel,	Natural pain relief from		
	Dysmeno	Remedies	_	Chamomile, Valerian	cramps and bloating		
6.	MenstCare	Tea Treasure	Tea	Hibiscus, Chamomile,	Relaxing tea to reduce		
	Herbal Tea	Herbal		Fennel, Peppermint	spasms and irritability		
7.	Moha Pain	Charak Pharma	Topical Oil	Nirgundi, Eucalyptus,	Applied topically to		
	Oil		_	Pudina, Ajwain	ease menstrual cramps		

Global Strategies for Herbal Treatment of Dysmenorrhea:

Across various cultures, herbal treatments for dysmenorrhea differ according to their traditional medical practices. In Traditional Chinese Medicine (TCM), Angelica sinensis and Chuanxiong are frequently utilized to enhance circulation and alleviate stagnation. Iranian traditional medicine highlights Zataria multiflora,

fennel, and cinnamon, with multiple clinical studies affirming their antispasmodic properties. European herbal medicine, particularly in Germany, often incorporates Valerian, Chamomile, and Melissa officinalis, which are regulated by the German Commission E Monographs. These international approaches emphasize the cross-cultural acknowledgment of herbal methods and support the idea of innovative delivery systems such as effervescent tablets.

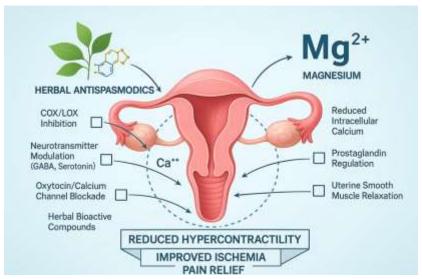


Fig.7 Synergistic role of herbal antispasmodics and magnesium in Reducing uterine hypercontractility and menstrual pain.

Benefits of Merging Herbal Antispasmodics with Magnesium for Managing Dysmenorrhea:

The simultaneous use of herbal antispasmodics and magnesium creates a synergistic and multifaceted approach to address primary dysmenorrhea. Each element offers unique pharmacodynamic effects that, when combined, may deliver better therapeutic results with fewer adverse effects compared to traditional NSAIDs.

1. Complementary Antispasmodic Effects

Herbal remedies such as ginger, fennel, and valerian alleviate uterine muscle contractions by suppressing prostaglandin production or modulating GABAergic transmission. Magnesium acts as a calcium channel blocker, promoting relaxation of the uterine muscles.(16,17)

2. Combined Anti-Inflammatory Actions

Both herbal ingredients (e.g., chamomile, ginger) and magnesium work to inhibit inflammatory mediators:

Herbs suppress COX-2, nitric oxide, and cytokines

• Magnesium lowers prostaglandin F2α levels and influences neuroinflammation.(18)

3. Nutraceutical Synergy

While herbal medicines offer active relief, magnesium addresses micronutrient deficiencies by supporting:

- Muscle relaxation
- Serotonin regulation
- Energy metabolism during menstruation. (19,20)

4. Enhanced Safety and Tolerability

In comparison to NSAIDs, this combination is:

- Non-ulcerogenic
- Hormone-free
- Less likely to result in renal or gastrointestinal side effects.

5. Increased Compliance through Effervescent Format

Effervescent tablets can:

Enhance taste and speed of onset



• Ensure quicker absorption

• Improve acceptability among younger individuals.(21)

Table3.Compared to conventional NSAIDs, herbal antispasmodics combined with magnesium offer a safer, Hormone-free, and patient-friendly option for dysmenorrhea management

Parameter	NSAIDs(Conventional)	Herbal Antispasmodics+ Magnesium (Prosposed)
Mechanism	Inhibit COX enzymes → ↓	Herbs: COX inhibition,
	prostaglandins	oxytocin/GABA modulation, anti- inflammatory;
		Magnesium: calcium channel blockade, ↓ PGF₂α
On set of	Fast (30–60 min)	Moderate (herbs slower, magnesium faster when
action		combined)
Side effect	GI irritation, ulcers, renal risk,	Minimal GI effects, hormone-free, nutrient support
	hormonal imbalance (long-term)	
Safety	Long-term risks	Generally safe, well tolerated
Patient	Moderate (bitter, requires food)	Better (natural, holistic, fewer side effects)
compliance	· · · · · · · · · · · · · · · · · · ·	
Innovation	Conventional	Integrative & novel

Patient-Centered Perspective:-

Adherence to treatment is a crucial factor influencing the success of therapy. Research indicates that women, particularly adolescents and young adults, often favor natural, hormone-free solutions for menstrual discomfort because of their worries about the long-term safety of NSAIDs and hormonal contraceptives. Interventions that are pleasant in taste, easy to administer, and holistic tend to enhance both compliance and overall satisfaction.(12,13)

Challenges and Limitations of Herbal Formulations for Dysmenorrhea:-

Despite the encouraging effectiveness of herbal remedies in alleviating menstrual discomfort, various obstacles impede their widespread acceptance, large-scale clinical validation, and consistent therapeutic outcomes. It is essential to recognize these challenges when contemplating the incorporation of herbal antispasmodics into contemporary dosage forms such as effervescent tablets.

1. Lack of Standardization

The active ingredient content in herbal extracts can differ considerably due to variations in plant origin, timing of harvest, extraction methods, and storage conditions. This inconsistency affects the potency, safety, and clinical effectiveness.(22)

2. Limited Clinical Evidence and Lack of Large-Scale Trials

While many herbs have traditional and empirical backing, only a few have been subjected to expansive, multicentric randomized controlled trials to confirm their efficacy and safety for treating primary dysmenorrhea. Small participant numbers and the absence of placebo controls diminish the reliability of many current studies.(23)

3. Herb-Drug Interactions

Numerous herbs can interact with prescribed medications through enzyme inhibition or activation (e.g., CYP450). This is particularly concerning for people simultaneously taking NSAIDs, hormonal contraceptives, or anticoagulants during their menstrual cycle.(24)



4. Delayed Onset of Action

In comparison to synthetic pain relievers like ibuprofen or mefenamic acid, herbal formulations may require longer to produce discernible effects due to slower pharmacokinetics and lower bioavailability in plasma.(25)

5. Regulatory and Quality Control Issues

Herbal formulations generally fall under less rigorous regulatory categories (such as nutraceuticals or supplements), leading to variability in product quality, claims, and safety information, unlike synthetic drugs.(26)

6. Poor Solubility or Taste Profile in Certain Dosage Forms

Many herbal extracts have a bitter flavor, are poorly soluble, or may be unstable in watery or effervescent bases, presenting challenges in formulation. Stabilizers or taste-masking agents might be necessary, particularly in innovative delivery forms like effervescent tablets.(27)

Literature Gaps and Opportunities for Future Reviews:-

- Although there is a significant amount of research on herbal antispasmodics and magnesium for alleviating menstrual pain, several gaps in the literature persist:
- Few investigations have looked into the combination of herbal and mineral elements within a single delivery system.
- There are currently no clinical trials or reviews that have assessed these combinations in an effervescent tablet format.
- The absence of standardized dosing protocols across herbal studies impacts their reproducibility.

• There is a necessity for systematic reviews or meta-analyses that compare herbal treatments with NSAIDs regarding clinical outcomes.

Future Perspectives and Broader Considerations:-

The combination of herbal antispasmodics and magnesium in an effervescent dosage form represents an innovative and patient-focused method for managing dysmenorrhea. Although this idea has not yet been commercialized, it paves the way for several future advancements:

- 1. **New Application:** Effervescent tablets that include herbal and mineral components may provide a quicker onset of action, improved taste, and greater adherence compared to conventional forms.(21)
- 2. **Interdisciplinary Opportunities:** The development of formulations could gain from knowledge in pharmacognosy, synthetic biology, and research on gut microbiota to enhance bioavailability and tailor treatments to individual needs.(19)
- 3. Challenges and Ethical Considerations:
 Concerns such as standardization, solubility,
 long-term safety, and interactions between
 herbs and drugs need to be addressed for
 clinical dependability.(19)
- 4. Research in Clinical and Formulation Areas: Additional studies are required to corroborate the effectiveness of herbal-magnesium combinations in effervescent forms through randomized controlled trials and comparative studies.
- 5. **Regulatory Considerations:** Understanding classifications under AYUSH (India), DSHEA (USA), or EMA (Europe) is vital for international commercialization.(23)
- 6. **Patient Perspective:** Young women and adolescents who favor natural and user-friendly treatments might find effervescent



formats more appealing due to their improved taste and convenience.(21,27)

- 7. **Economic Potential:** Innovations combining herbal and magnesium elements could decrease reliance on NSAIDs, provide safer alternatives, and present cost-effective options for relief from menstrual pain.(26,27)
- 8. Develop standardized herbal extracts with measurable markers (e.g., gingerols in ginger, anethole in fennel, valerenic acid in valerian).
- 9. Carry out multicenter randomized controlled trials that directly compare herbal and magnesium therapy to NSAIDs.
- 10. Examine pharmacokinetic and pharmacodynamic interactions between herbs and magnesium as well as conventional medications.
- 11. Investigate the modulation of gut microbiota as a factor influencing the effectiveness of herbs and magnesium.
- 12. Consider individualized treatment strategies based on nutritional status and genetic factors.(11,29)

CONCLUSION:

Dysmenorrhea continues to be a significant clinical and quality-of-life issue for women worldwide, typically managed with NSAIDs and hormonal therapies that may produce unwanted side effects. This review underscores the potential therapeutic benefits of herbal antispasmodics like ginger, fennel, valerian, and chamomile, alongside the vital mineral magnesium, which offer diverse mechanisms such as prostaglandin inhibition, muscle relaxation, and anti-inflammatory properties.

The Idea of merging these components into a single effervescent tablet offers a novel and patient-friendly strategy with the capability to improve the speed of action, taste, and adherence, particularly among younger women. Although the

formulation has yet to be created, this concept introduces an encouraging path for forthcoming research and innovation.

Nevertheless, existing literature highlights significant gaps, such as the lack of standardized herbal preparations, a shortage of large-scale trials, and no combination studies involving both herbal and mineral agents in advanced delivery systems. Addressing these shortcomings through well-structured clinical investigations and formulation research will be crucial in translating this novel idea into a safe, effective, and commercially feasible solution for menstrual health.

REFERENCES

- 1. Ju H, Jones M, Mishra G. The prevalence and risk factors of dysmenorrhea. BJOG. 2014;121(6):678–686.
 - https://doi.org/10.1111/1471-0528.12328
- 2. Iacovides S, Avidon I, Baker FC. What we know about primary dysmenorrhea today: A critical review. Hum Reprod Update. 2015;21(6):762–778.
 - https://doi.org/10.1093/humupd/dmv039
- 3. Sharma A, Taneja DK, Sharma P, Saha R. Problems related to menstruation and their effect on daily routine of students of a medical college in Delhi, India. Asia Pac J Public Health. 2019;22(2):234–241.
- 4. Ortiz MI. Primary dysmenorrhea among Mexican university students: Prevalence, impact, and treatment. Eur J Obstet Gynecol Reprod Biol. 2010;152(1):73–77.
- 5. Chia CF, et al. Dysmenorrhea among adolescent girls and its impact on school attendance. Aust Fam Physician. 2013;42(11):765–769.
- 6. Unsal A, Ayranci U, Tozun M, Arslan G, Calik E. Prevalence of dysmenorrhea and its effect on quality of life among university students. Ups J Med Sci. 2010;115(2):138–



- 145. https://doi.org/10.3109/03009730903457218
- 7. Ziaei S, Zakeri M, Kazemnejad A. A randomized controlled trial of vitamin E in the treatment of primary dysmenorrhea. BJOG. 2001;108(9):1181–1183. https://doi.org/10.1111/j.1471-0528.2001.00287.x
- 8. Kalita S, Choudhury A, Dey B. Effervescent drug delivery systems: A review. J Drug Deliv Ther. 2020;10(2):174–178. https://doi.org/10.22270/jddt.v10i2.3971
- 9. Osayande AS, Mehulic S. Diagnosis and initial management of dysmenorrhea. Am Fam Physician. 2014;89(5):341–346.
- Harel Z. Dysmenorrhea in adolescents and young adults: Etiology and management. J Pediatr Adolesc Gynecol. 2006;19(6):363– 371.
 - https://doi.org/10.1016/j.jpag.2006.09.001
- 11. Adeney KL, Chang AM. The impact of dysmenorrhea on quality of life among young women in Australia. Br J Nurs. 2007;16(16):976–980. https://doi.org/10.12968/bjon.2007.16.16.270 45
- 12. Bajalan Z, Moafi F, MoradiBaglooei M, Alimoradi Z. Mental health and primary dysmenorrhea: A systematic review. J Psychosom Obstet Gynaecol. 2019;40(3):185–194. https://doi.org/10.1080/0167482X.2018.1470 624
- 13. Zondervan KT, Becker CM, Missmer SA. Endometriosis. N Engl J Med. 2018;378(13):1244–1256. https://doi.org/10.1056/NEJMra1707158
- 14. Uritu CM, Mihai CT, Stanciu GD, et al. Medicinal plants of the family Lamiaceae in pain therapy: A review. Pain Res Manag. 2021;2021:2731697. https://doi.org/10.1155/2021/2731697

- 15. Ziaei S, Zakeri M, Kazemnejad A. Oral magnesium for primary dysmenorrhea: A randomized controlled trial. BJOG. 2001;108(9):867–870. https://doi.org/10.1111/j.1471-0528.2001.00220.x
- 16. Rahnama P, Montazeri A, Huseini HF, Kianbakht S, Naseri M. Effect of ginger on pain relief in primary dysmenorrhea. BMC Complement Altern Med. 2012;12:92. https://doi.org/10.1186/1472-6882-12-92
- 17. Daily JW, Zhang X, Park S. Efficacy of ginger for alleviating primary dysmenorrhea: A systematic review. Pain Med. 2015;16(12):2243–2255. https://doi.org/10.1111/pme.12853
- 18. Walker AF, De Souza MC, Vickers MF. Magnesium supplementation alleviates premenstrual symptoms. J Womens Health. 1998;7(9):1157–1165. https://doi.org/10.1089/jwh.1998.7.1157
- Ylikorkala O, Dawood MY. New concepts in dysmenorrhea. Am J Obstet Gynecol. 1977;128(7):723–730. https://doi.org/10.1016/0002-9378(77)90738-6
- 20. Rao S, Kulkarni PK, Karunakar AK. Effervescent dosage forms: A review. Int J Pharm Sci Res. 2021;12(4):1921–1927.
- 21. Kunle OF, Egharevba HO, Ahmadu PO. Standardization of herbal medicines: A review. Int J Biodivers Conserv. 2012;4(3):101–112.
 - https://doi.org/10.5897/IJBC11.163
- 22. Bent S. Herbal medicine in the United States: Review of efficacy, safety, and regulation. J Gen Intern Med. 2008;23(6):854–859. https://doi.org/10.1007/s11606-008-0632-y
- 23. Fugh-Berman A. Herb-drug interactions. Lancet. 2000;355(9198):134–138. https://doi.org/10.1016/S0140-6736(99)06415-0



- 24. Williamson EM. Synergy and other interactions in phytomedicines. Phytomedicine. 2001;8(5):401–409. https://doi.org/10.1078/0944-7113-00060
- 25. Ekor M. The growing use of herbal medicines: Safety and monitoring challenges. Front Pharmacol. 2014;4:177. https://doi.org/10.3389/fphar.2013.00177
- 26. Kulkarni PK, Rao S, Karunakar AK. Effervescent dosage forms: Formulation challenges and innovations. Int J Drug Dev Res. 2021;13(2):10–16.
- 27. De Sanctis V, Silliman A, Elsedfy H, Soliman N, El Kholy M. Dysmenorrhea in adolescents and young adults: A global review. Acta Biomed. 2016;87(3):233–246.
- 28. Dawood MY. Primary dysmenorrhea: Advances in pathogenesis and management. Obstet Gynecol. 2006;108(2):428–441. https://doi.org/10.1097/01.AOG.0000230214.26638.0c
- 29. Ozgoli G, Goli M, Moattar F. Comparison of ginger, mefenamic acid, and ibuprofen for pain in dysmenorrhea. J Altern Complement Med. 2009;15(2):129–132. https://doi.org/10.1089/acm.2008.0311
- 30. Modarres M, et al. The effect of fennel on pain intensity in dysmenorrhea. J Pediatr Adolesc Gynecol. 2011;24(3):189–192. https://doi.org/10.1016/j.jpag.2010.10.007
- 31. Delaram M, Forouzandeh N, Mirzakhani K. The effect of fennel on primary

- dysmenorrhea. Iran J Nurs Midwifery Res. 2011;16(Suppl 1):S70–S73.
- 32. Mirabi P, Dolatian M, Mojab F, Majd HA. Effects of valerian on severity of dysmenorrhea. J Complement Integr Med. 2011;8(1). https://doi.org/10.2202/1553-3840.1396
- 33. Sadeghi Aval S, Akhondzadeh S, Ahmadi Abhari S. Valerian for primary dysmenorrhea: A randomized trial. Complement Ther Clin Pract. 2014;20(2):81–85. https://doi.org/10.1016/j.ctcp.2013.10.004
- 34. Sharifi F, Simbar M, Mojab F, Majd HA. Chamomile vs mefenamic acid in dysmenorrhea. Iran J Obstet Gynecol Infertil. 2014;17(97):1–9.
- 35. Srivastava JK, Shankar E, Gupta S. Chamomile: A herbal medicine of the past with a bright future. Mol Med Rep. 2010;3(6):895–901. https://doi.org/10.3892/mmr.2010.377
- 36. Nahidi F, Kariman N, Simbar M, Mojab F. Effect of Mentha piperita on primary dysmenorrhea. J Reprod Infertil. 2009;10(3):163–169.

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