

## A Complication of Radiotherapy-Oral Mucositis: An Updated Review for Healthcare providers

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### Abstract

*Oral mucositis (OM) is a debilitating condition characterized by erythema, edema, and ulcerative lesions of the oral mucosa, commonly arising as a complication of radiotherapy (RT), chemotherapy, or hematopoietic stem cell transplantation (HSCT). It significantly impacts patients' quality of life, causing severe pain, nutritional deficiencies, and increased risk of infections, often necessitating treatment modifications. This review aims to provide healthcare providers with an updated understanding of the etiology, pathophysiology, epidemiology, and management strategies for OM, emphasizing evidence-based interventions to mitigate its impact on patients undergoing cancer therapy. This study aims to evaluate the main role of healthcare providers, dentists, and radiologists. The review synthesizes current literature on OM, focusing on its pathogenesis, clinical presentation, diagnostic evaluation, and therapeutic approaches. It highlights the five-phase model of OM development and discusses grading scales such as CTC/AE, WHO, and OMAS for assessment. Management strategies, including basic oral care, cryotherapy, low-level laser therapy (LLLT), and chemoprotective agents like palifermin, are evaluated. OM is a multifactorial condition influenced by treatment intensity, patient-specific factors, and mucosal vulnerability. Preventive measures, such as basic oral care and cryotherapy, significantly reduce OM severity. LLLT and palifermin show promise in high-risk populations, though safety concerns remain. Pain management, including topical morphine rinses, improves patient comfort and adherence to therapy. Effective management of OM requires a multidisciplinary approach, integrating preventive strategies, early detection, and tailored interventions. Healthcare providers must prioritize patient education, optimize treatment protocols, and employ evidence-based therapies to minimize OM-related morbidity and enhance treatment outcomes.*

**Keywords:** Oral Mucositis, Radiotherapy, Chemotherapy, Low-Level Laser Therapy, Palifermin, Cryotherapy, Pain Management, Multidisciplinary Care.

### Introduction

Oral mucositis represents a profoundly debilitating pathological condition marked by the presence of erythema, edema, and ulcerative lesions affecting the oral mucosal lining [1]. This condition frequently arises as a consequential complication associated with therapeutic interventions such as radiation therapy (RT) targeting the head and neck region, systemic chemotherapy, combined chemoradiotherapy protocols, and hematopoietic stem cell transplantation (HSCT) [2]. The clinical manifestations of oral mucositis often include severe pain, which can significantly compromise oral intake, necessitating, in certain instances, the administration of parenteral nutrition to maintain adequate caloric and nutritional support [3].

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Furthermore, the ulcerative lesions characteristic of this condition disrupt the integrity of the mucosal barrier, thereby predisposing patients to an elevated risk of localized or systemic infections, which can further exacerbate the clinical course [4]. In cases of severe oral mucositis induced by chemotherapy, the associated morbidity may necessitate modifications to subsequent treatment cycles, including dose reduction or delays in administration. Such alterations in therapeutic regimens can have profound implications for patients, adversely affecting their overall quality of life and potentially compromising treatment efficacy, thereby worsening long-term prognostic outcomes [5]. The interplay between oral mucositis and its broader systemic consequences underscores the critical need for effective management strategies to mitigate its impact on both patients' well-being and therapeutic success.

### *Etiology*

Oral mucositis is a prevalent and clinically significant complication observed in patients undergoing radiation therapy (RT) directed at the head and neck region, chemotherapy for the treatment of solid tumors or lymphomas, and high-dose myeloablative chemotherapy administered as part of conditioning regimens prior to hematopoietic cell transplantation. The incidence and severity of oral mucositis exhibit considerable variability depending on the specific chemotherapeutic agents utilized [3]. Agents that interfere with DNA synthesis, particularly those targeting the S-phase of the cell cycle, such as 5-fluorouracil, methotrexate, and cytarabine, are associated with a notably high incidence of oral mucositis [5]. Additionally, other classes of chemotherapeutic drugs, including anthracyclines, mTOR inhibitors, alkylating agents, and antimetabolites, have been implicated in an elevated risk of inducing this condition [6][7]. The pathogenesis of oral mucositis is closely linked to the unique biological characteristics of the oral mucosa. The basal epithelial layer of the oral mucosa is characterized by a high rate of cellular turnover, rendering it particularly vulnerable to the cytotoxic effects of radiation and chemotherapy [1]. Radiation injury to this rapidly proliferating tissue results in cellular damage and apoptosis, while the impaired regenerative capacity of the oral mucosa further exacerbates the condition, leading to the development of mucosal ulcerations and inflammation [1]. The disruption of the mucosal barrier not only causes significant pain and functional impairment but also predisposes patients to secondary infections, which can complicate the clinical course and delay recovery. Collectively, these mechanisms underscore the multifactorial etiology of oral mucositis, highlighting the interplay between treatment-related toxicity and the intrinsic vulnerability of the oral mucosa.

### *Epidemiology*

Oral mucositis is a common and clinically significant adverse effect observed in patients undergoing cancer treatment, with its prevalence and severity influenced by various factors. Studies indicate that between 20% to 40% of patients with solid tumors who receive chemotherapy develop mucositis, typically manifesting within five to fourteen days following the initiation of treatment [8]. The incidence and severity of this condition are highly variable and depend on several factors, including the specific chemotherapeutic agents used, the cumulative number of chemotherapy cycles, the dosage of chemotherapy administered, and individual patient characteristics [5]. Notably, patients undergoing myeloablative conditioning regimens as part of hematopoietic stem cell transplantation (HSCT) are at a significantly higher risk of developing oral mucositis, with the condition being almost universally observed in this population [9]. Research has demonstrated that patients receiving high-dose chemotherapy or undergoing bone marrow transplantation face a 76% risk of developing mucositis, underscoring the profound impact of treatment intensity on mucosal toxicity [10]. Furthermore, radiation-induced oral mucositis (RIOM) is an inevitable complication in head and neck cancer patients receiving altered fractionation radiotherapy, with a 100% incidence rate reported in this population [10]. Additional risk factors contributing to the development of oral mucositis include poor nutritional status and inadequate oral hygiene, both of which exacerbate mucosal vulnerability. Interestingly, younger patients may also exhibit a higher incidence of oral mucositis, potentially due to the higher proliferative activity of their mucosal tissues [9]. These epidemiological insights highlight the multifactorial nature of oral mucositis and emphasize the need for tailored preventive and therapeutic strategies to mitigate its impact on patients undergoing cancer treatment.

### *Pathophysiology*

The pathophysiology of oral mucositis (OM) resulting from radiation therapy (RT), chemotherapy, or combined chemoradiotherapy is a multifaceted process that unfolds through a sequential series of biological events. This process is best described by the five-phase model proposed by Sonis, which delineates the progression of OM into distinct stages: initiation, signaling, amplification, ulceration, and healing [11]. Each phase contributes to the development and resolution of mucosal injury, highlighting the dynamic interplay between cellular damage, inflammatory responses, and tissue repair mechanisms. The initial phase, initiation, is characterized by direct tissue injury caused by the cytotoxic effects of RT or chemotherapy. These treatments induce DNA damage and generate reactive oxygen species (ROS), leading to the apoptosis of basal epithelial cells, which are critical for maintaining the integrity of the oral mucosa [3]. In the subsequent signaling phase, the ROS not only perpetuate cellular death but also activate key inflammatory pathways, including nuclear factor-kappa B (NF- $\kappa$ B), which further exacerbates tissue damage [3]. The amplification phase involves the upregulation of pro-inflammatory cytokines, such as tumor necrosis factor-alpha (TNF- $\alpha$ ), interleukin-1 beta (IL-1 $\beta$ ), and interleukin-6 (IL-6), amplifying the inflammatory cascade and contributing to widespread mucosal injury [11]. During the ulceration phase, the mucosal barrier becomes compromised, resulting in painful ulcerations that are often colonized by microbial flora, increasing the risk of secondary infections. This phase is marked by significant inflammation and clinical symptoms, including pain and difficulty in oral intake. Finally, the healing phase involves the proliferation and differentiation of epithelial cells, facilitated by growth factors and signaling molecules, leading to the restoration of mucosal integrity [11]. Understanding this intricate pathophysiology is essential for developing targeted interventions to prevent or mitigate the severity of oral mucositis in patients undergoing cancer therapy.

### *History and Physical*

Radiotherapy-induced oral mucositis is a consequence of direct tissue injury caused by radiation beams, typically manifesting during the third week of treatment [8] and persisting for a duration ranging from 7 to 98 days [10]. This condition initially presents as acute inflammation affecting the oral mucosa, tongue, and pharyngeal tissues [10]. The most pronounced damage is observed in the soft palate, followed by the hypopharynx, floor of the mouth, buccal mucosa, tongue, and lips [1]. The associated pain is often intense, necessitating the temporary discontinuation of radiotherapy in many cases [8]. Additionally, patients frequently report significant difficulty in speaking and eating, alongside an increased tendency for gingival bleeding during oral hygiene practices. Chemotherapy-induced oral mucositis generally emerges within five to fourteen days following the initiation of treatment [8]. It initially presents as mucosal erythema, which progressively deteriorates into erosion and ulceration. The ulcerative lesions may be covered by a white fibrinous pseudomembrane. The distribution of these ulcers is predominantly confined to non-keratinized oral surfaces, including the buccal mucosa, lateral and ventral aspects of the tongue, and the soft palate [12]. In immunocompromised patients, including those undergoing hematopoietic stem cell transplantation, oral mucositis begins to resolve as the absolute neutrophil count improves. The restoration of normal immune function facilitates mucosal healing, leading to gradual resolution of inflammation and ulceration.

### *Evaluation*

The evaluation of oral mucositis primarily relies on a thorough clinical history and a detailed physical examination, as laboratory tests and radiographic imaging are generally not useful in diagnosing this condition. However, if ulcerative lesions are observed on the hard palate, attached gingiva, or dorsum of the tongue, it is essential to obtain cultures to rule out viral or fungal infections, which can mimic or exacerbate mucositis. The severity of oral mucositis is assessed using well-defined grading scales, several of which have been developed to standardize its evaluation and facilitate consistent reporting in clinical and research settings [4]. One of the most widely used tools is the Common Terminology Criteria for Adverse Events (CTCAE), developed by the National Cancer Institute (NCI). This scale evaluates mucositis based on two components: a clinical examination and a functional/symptoms-based assessment. The functional/symptoms-based exam grades mucositis from Grade 1 (asymptomatic or mild symptoms with no dietary modifications required) to Grade 5 (death). Similarly, the clinical exam ranges from Grade 1

(mucosal erythema) to Grade 5 (death), with intermediate grades reflecting increasing severity, such as patchy ulcerations, confluent ulcers, and tissue necrosis [4].

Another commonly used scale is the World Health Organization (WHO) Oral Mucositis Scale, which integrates both subjective symptoms and objective clinical findings. This scale ranges from Grade 0 (no mucositis) to Grade 4 (ulcers preventing alimentation). It is particularly useful for its simplicity and ability to capture both functional impairment and clinical severity. The Oral Mucositis Assessment Scale (OMAS) is an objective tool that quantifies erythema and ulceration at nine specific sites within the oral cavity. This scale has demonstrated high interobserver reproducibility and a strong correlation between clinical findings and patient-reported symptoms, making it a reliable tool for both clinical practice and research [13]. Notably, studies by Sonis et al. have shown that concomitant symptomatic measurements may be unnecessary when using OMAS, as the scale itself effectively captures the severity of mucositis [13]. Additionally, the Eastern Cooperative Oncology Group (ECOG) Common Toxicity Criteria are frequently employed in oncology trials to assess mucositis and other treatment-related toxicities. These criteria provide a standardized framework for evaluating adverse events, ensuring consistency across clinical studies. In summary, the evaluation of oral mucositis involves a combination of clinical assessment and standardized grading scales, such as CTCAE, WHO, OMAS, and ECOG criteria. These tools enable healthcare providers to accurately diagnose, grade, and monitor mucositis, facilitating timely interventions and improving patient outcomes. The choice of scale often depends on the clinical context, with some scales emphasizing objective findings (e.g., OMAS) and others incorporating functional impairment (e.g., WHO, CTCAE).

### *Treatment and Management*

Clinicians employ various strategies to manage oral mucositis in patients undergoing cancer therapy [14]. The following treatment modalities aim to alleviate symptoms, prevent complications, and enhance patient quality of life.

#### *Basic Oral Care*

Basic oral care (BOC) is a cornerstone in managing oral mucositis, as recommended by the 2020 Multinational Association of Supportive Care in Cancer and the International Society of Oral Oncology (MASCC/ISOO) clinical guidelines [2]. BOC encompasses interventions by both patients and healthcare providers to maintain oral hygiene, prevent infections, and reduce discomfort. These measures include mechanical cleaning of teeth, the use of mouth rinses, and lubrication of the oral mucosa [2]. Frequent assessment of oral tissues before and during cancer therapy is crucial for minimizing infection risks and identifying early signs of mucositis [15]. Adhering to oral hygiene protocols significantly reduces the severity and duration of mucositis by limiting microbial colonization and preventing secondary infections [2][16]. Prophylactic dental interventions, such as extracting compromised teeth and addressing carious lesions before cancer therapy initiation, have been shown to reduce mucositis risk by over 25%, particularly in high-risk individuals [8].

#### *Mechanical Cleaning*

Enhancing mechanical cleaning is essential for oral mucositis prevention. Patients should be encouraged to brush their teeth more frequently using a soft-bristled toothbrush, replacing their toothbrush regularly, and using interdental cleaning techniques [1].

#### *Oral Rinses*

Non-medicated oral rinses, including saline water and sodium bicarbonate solutions, should be used every four hours to maintain oral hygiene and comfort [1]. The MASCC/ISOO guidelines recognize these rinses as safe and effective, despite limited clinical data, due to their ability to increase oral clearance and alleviate mucosal irritation [2]. However, chlorhexidine rinses are not recommended for preventing radiotherapy-induced oral mucositis due to insufficient supporting evidence [2][1].

### *Anti-inflammatory Agents*

The MASCC/ISOO guidelines endorse benzydamine mouthwash for preventing oral mucositis in patients with head and neck cancer receiving moderate-dose radiotherapy (less than 50 Gy) and those undergoing radiotherapy-chemotherapy (RT-CT) regimens [2].

### *Hydration and Lubrication of the Oral Mucosa*

Maintaining adequate hydration is critical for mucosal health. Patients should avoid irritants such as tobacco, alcohol-containing beverages, and alcohol-based mouth rinses [1][8]. To alleviate oral dryness, lubricating agents such as mousses and topical barrier gels may be applied to the mucosal surfaces [1].

### *Dietary Considerations*

Dietary modifications play a crucial role in reducing mucosal irritation. Patients should avoid foods that may exacerbate mucositis or pose a risk of trauma during mastication, including spicy, sharp, or hard foods [8]. A soft, nutrient-rich diet is recommended to promote healing and minimize discomfort.

### *Pain Management*

Effective pain management is essential for improving patient comfort and adherence to cancer therapy. The MASCC/ISOO guidelines recommend the use of topical morphine 0.2% mouth rinse for pain relief in patients undergoing radiotherapy and chemotherapy for head and neck cancer [2]. Additionally, various oral rinses have been developed to manage mucositis-related pain. One such formulation is "magic mouthwash," which typically contains an anesthetic, an antacid, and diphenhydramine, with possible inclusion of steroids and antifungal agents [15]. However, a study involving 26 patients receiving concurrent RT and CT revealed that those using topical morphine 0.2% mouth rinse experienced a shorter duration of severe pain and required fewer systemic analgesics compared to patients using magic mouthwash containing lidocaine, diphenhydramine, and magnesium aluminum hydroxide [15].

### *Low-Level Laser Therapy (LLLT)*

Photobiomodulation (PBM), also known as low-level laser therapy (LLLT), is an emerging treatment that utilizes localized light energy to induce biological responses in tissues [2]. Preclinical studies have demonstrated that PBM enhances tissue regeneration and mitigates inflammation [15]. Clinical trials further support the efficacy of LLLT in reducing mucositis severity among patients undergoing chemoradiotherapy before hematopoietic stem cell transplantation [15]. The MASCC/ISOO guidelines recommend intraoral LLLT for oral mucositis prevention in the following patient populations:

- Adults undergoing hematopoietic stem cell transplantation conditioned with high-dose chemotherapy, with or without total body irradiation [2].
- Adults receiving head and neck radiation therapy without concurrent chemotherapy, considering the safety implications for patients with oral cancer [2].
- Adults undergoing radiotherapy-chemotherapy for head and neck malignancies, with careful consideration of potential risks for cancerous tissues [2].

While clinical data supports LLLT in reducing mucositis severity, *in vitro* studies indicate that this therapy may activate pro-tumorigenic pathways in malignant cells [15]. Therefore, LLLT should not be applied directly to cancerous tissue, and patients undergoing this treatment require close monitoring [15]. Managing oral mucositis in cancer patients requires a multidisciplinary approach integrating preventive and therapeutic strategies. Basic oral care, including meticulous oral hygiene, prophylactic dental interventions, and mechanical cleaning, plays a crucial role in reducing the incidence and severity of mucositis. Non-medicated oral rinses, hydration, and lubrication further contribute to symptom relief. Pain management strategies,

particularly the use of topical morphine rinses, have demonstrated effectiveness in alleviating discomfort. Low-level laser therapy presents a promising adjunct therapy; however, clinicians must carefully evaluate its safety in patients with active malignancies. By implementing these evidence-based interventions, healthcare providers can improve patient outcomes and minimize the impact of oral mucositis on cancer therapy adherence.

### *Cryotherapy*

Cryotherapy is a preventive strategy used to mitigate the development of oral mucositis in patients undergoing chemotherapy. It involves the application of a cold source, such as ice chips or cold water, within the oral cavity during the administration of cytotoxic agents. The mechanism of action is primarily based on the induction of vasoconstriction in the oral mucosa, which reduces blood flow to the tissue and limits the delivery of chemotherapeutic drugs to the area. This, in turn, minimizes the exposure of the basal epithelial cells to the cytotoxic effects of the drugs, thereby reducing the risk of mucosal injury [17]. Additionally, the cooling effect may lower the metabolic activity of the basal epithelial layer, rendering these cells less susceptible to damage caused by chemotherapy agents [18]. However, the protective effects of cryotherapy are transient, as the cooling of the oral mucosa is only temporary. Consequently, this intervention is most effective when used in conjunction with short-duration chemotherapy protocols or with cytotoxic agents that have a short half-life [2]. The Multinational Association of Supportive Care in Cancer and International Society of Oral Oncology (MASCC/ISOO) guidelines recommend the use of oral cryotherapy for 30 minutes during the administration of bolus 5-fluorouracil, a chemotherapeutic agent known to cause mucositis [2]. Furthermore, cryotherapy is also recommended for patients receiving high-dose melphalan as part of conditioning regimens for autologous hematopoietic stem cell transplantation (HSCT), as it has been shown to reduce the incidence and severity of oral mucositis in this population [2].

### *Chemoprotective Agent*

Palifermin, a recombinant keratinocyte growth factor, is another effective intervention for preventing and managing severe oral mucositis. It functions as a chemoprotective agent by stimulating the proliferation, differentiation, and migration of epithelial cells, thereby enhancing the repair and regeneration of the oral mucosa. Palifermin is specifically recommended for patients undergoing autologous HSCT who are at high risk of developing severe oral mucositis (Grade 3 or higher) [2]. Clinical studies have demonstrated that palifermin significantly reduces both the incidence and duration of severe mucositis in this patient population, improving their overall quality of life and treatment outcomes.

### *Zinc Supplementation*

Zinc is an essential trace element that plays a critical role in tissue repair and possesses antioxidant properties, making it a potential candidate for the prevention of oral mucositis. The 2014 MASCC/ISOO guidelines initially recommended systemic zinc supplementation for oral cancer patients undergoing radiotherapy or chemoradiation, based on its potential to enhance mucosal healing and reduce oxidative stress [19]. However, in the 2020 update of the guidelines, the panel revised this recommendation, stating that no definitive guidelines could be established due to insufficient evidence supporting its efficacy [2]. This reversal highlights the need for further research to clarify the role of zinc in the prevention and management of oral mucositis. In summary, cryotherapy and chemoprotective agents like palifermin are well-supported interventions for preventing oral mucositis in high-risk patients, while the role of zinc supplementation remains uncertain and requires further investigation. These strategies underscore the importance of tailored approaches to managing oral mucositis based on the specific treatment regimen and patient population.

### *Differential Diagnosis*

The differential diagnosis of oral mucositis includes various infectious and dermatologic conditions that may present similar clinical manifestations. Although infections do not directly cause oral mucositis, they can complicate its presentation and necessitate additional treatment. Viral infections, such as herpes simplex

virus (HSV), can lead to ulcerative lesions that resemble mucositis. Similarly, fungal infections, particularly oral candidiasis, may develop concurrently, exacerbating symptoms and requiring antifungal therapy. Systemic inflammatory diseases with dermatologic manifestations, including systemic lupus erythematosus (SLE) and rheumatoid arthritis, can also contribute to oral ulcerations that mimic mucositis. These conditions often present with other systemic symptoms, which can aid in differentiation. Additionally, oral squamous cell carcinoma (OSCC) must be considered in patients with non-healing ulcers or persistent mucosal lesions, especially in individuals with risk factors such as tobacco and alcohol use. A biopsy may be required to confirm the diagnosis of OSCC and distinguish it from radiation- or chemotherapy-induced mucositis. Nutritional deficiencies, particularly zinc deficiency, have been associated with oral lesions resembling mucositis. Zinc plays a crucial role in epithelial repair and immune function, and its deficiency can impair wound healing and mucosal integrity. Patients with suspected nutritional deficiencies may benefit from dietary assessment and zinc supplementation as part of their management plan.[20] In clinical practice, a thorough history, physical examination, and diagnostic testing are essential to differentiate oral mucositis from these conditions. Accurate diagnosis ensures appropriate treatment and prevents unnecessary interventions or delays in managing the underlying cause of the patient's symptoms.

### *Prognosis*

The prognosis of oral mucositis largely depends on its severity and the underlying cause. In uncomplicated cases, mucositis is often self-limited and can be effectively managed with symptomatic treatment and palliative care, including pain relief, oral rinses, and nutritional support [8]. However, in more severe cases, particularly among head and neck cancer patients undergoing chemoradiotherapy, the condition can have significant implications for both treatment outcomes and quality of life. Nearly 20% of these patients develop high-grade mucositis, which may necessitate hospitalization and lead to delays or modifications in their cancer therapy [15]. Such interruptions not only disrupt the continuity of treatment but also worsen the overall prognosis by potentially reducing the efficacy of the therapeutic regimen. Additionally, severe mucositis contributes to increased healthcare resource utilization, including prolonged hospital stays, the need for parenteral nutrition, and management of secondary complications. These factors collectively underscore the importance of early intervention and preventive strategies to mitigate the impact of mucositis on patient outcomes and healthcare systems.

### *Complications*

Oral mucositis is associated with a range of complications that can significantly affect patient well-being and treatment efficacy. The severe pain caused by mucosal ulcerations often leads to a reduction in oral intake, resulting in nutritional deficiencies and a decline in overall quality of life. This can further complicate the clinical course, as inadequate nutrition impairs the body's ability to heal and withstand cancer treatment. Moreover, the disruption of the protective mucosal barrier increases the risk of local and systemic infections, which can escalate to life-threatening conditions such as septicemia [21]. To address these risks, many chemotherapy protocols incorporate prophylactic measures, including oral care regimens and antibiotics, to prevent the onset of mucositis and its associated complications. These strategies highlight the critical need for integrated supportive care in managing mucositis and minimizing its adverse effects on treatment outcomes.

### *Patient Education*

Patient education plays a pivotal role in the prevention and early management of oral mucositis. Before initiating chemotherapy or radiotherapy, patients should be informed about the signs and symptoms of mucositis, such as oral pain, erythema, and ulcerations, to facilitate early diagnosis and intervention [8]. Educating patients about alarming symptoms, including difficulty swallowing, severe pain, or signs of infection, is essential to ensure timely medical attention and prevent complications. Additionally, patients should be counseled on the importance of maintaining basic oral hygiene, avoiding irritants such as alcohol-based mouthwashes or spicy foods, and adhering to a soft, non-acidic diet to minimize mucosal irritation. Providing detailed guidance on these preventive measures empowers patients to take an active role in their

care, potentially reducing the incidence and severity of mucositis and improving their overall treatment experience.

### *Enhancing Healthcare Team Outcomes*

Effective management of oral mucositis requires collaboration among healthcare professionals, including oncologists, dentists, nurses, and pharmacists. Since oral mucositis affects up to 91% of patients undergoing radiotherapy for head and neck cancer and 20% to 40% of those receiving chemotherapy for solid tumors, proactive measures are essential to minimize complications and improve patient outcomes. Oncologists and primary care providers play a central role in educating patients about the signs and symptoms of oral mucositis before initiating chemotherapy or radiotherapy. Early identification enables timely interventions, reducing the severity and duration of mucositis-related discomfort. Clinicians should assess patient risk factors, including pre-existing oral conditions, immunosuppression, and comorbidities, which may worsen mucositis. Dentists are key members of the healthcare team, as oral health optimization before cancer therapy can significantly reduce mucositis risk. Prophylactic dental evaluations, including extracting compromised teeth and restoring carious lesions, decrease the likelihood of infection and mucosal breakdown. Studies indicate that such interventions lower mucositis incidence by over 25% in high-risk patients. Nurses play an essential role in patient education and monitoring. They guide patients on oral hygiene practices, advise against irritant exposure (e.g., tobacco and alcohol), and reinforce dietary modifications. Nurses also assess mucositis progression, ensuring early referral for additional supportive care when needed. Pharmacists contribute by optimizing medication regimens. They recommend appropriate pain management strategies, including topical analgesics like morphine mouthwash or systemic analgesics for severe cases. They also assist in selecting safe and effective oral rinses, avoiding agents like chlorhexidine, which lacks evidence for mucositis prevention. A multidisciplinary approach enhances patient outcomes by integrating preventive measures, early detection, and targeted interventions. Regular interprofessional communication ensures coordinated care, minimizing the impact of oral mucositis on cancer patients [22].

### *Role of Radiologists in Oral Mucositis*

Radiologists play a critical role in the diagnosis, prevention, and management of oral mucositis, particularly in patients undergoing radiotherapy for head and neck cancers. Their primary responsibility is to optimize radiation therapy protocols to minimize mucosal damage while ensuring effective tumor control. Through imaging techniques such as computed tomography (CT) and magnetic resonance imaging (MRI), radiologists assist in treatment planning by identifying high-risk areas, assessing tumor burden, and mapping radiation fields to avoid unnecessary exposure to healthy tissues. Advanced radiation delivery techniques, such as intensity-modulated radiotherapy (IMRT) and proton therapy, have been developed with the guidance of radiologists to enhance dose precision and reduce toxicity to the oral mucosa. In addition to treatment planning, radiologists play a key role in early detection and monitoring of radiation-induced oral mucositis. Imaging modalities such as positron emission tomography (PET) scans can be used to evaluate inflammatory responses in the oral mucosa and differentiate mucositis from tumor recurrence or infection. Radiologists also collaborate with oncologists and other specialists to assess treatment-related complications, including secondary infections and fibrosis, which may arise from persistent mucosal injury. Their ability to interpret imaging findings helps guide clinical decisions regarding supportive care interventions, such as the initiation of cryotherapy, low-level laser therapy, or chemoprotective agents. Furthermore, radiologists contribute to research and clinical trials aimed at reducing radiation toxicity and improving patient outcomes. By analyzing imaging biomarkers and exploring novel radioprotective strategies, they help refine treatment protocols that balance efficacy with minimizing adverse effects. Interdisciplinary collaboration between radiologists, oncologists, dentists, and supportive care specialists is essential in mitigating the impact of oral mucositis and enhancing the overall quality of life for cancer patients. As imaging technologies continue to evolve, the role of radiologists in the prevention and management of oral mucositis will remain integral to advancing cancer care [22].



## Conclusion

Oral mucositis (OM) remains a significant challenge in cancer care, particularly for patients undergoing radiotherapy, chemotherapy, or hematopoietic stem cell transplantation. Its multifactorial etiology, involving direct mucosal injury, inflammatory cascades, and compromised tissue repair, underscores the complexity of its management. The clinical impact of OM is profound, with severe pain, nutritional deficiencies, and increased infection risk often leading to treatment interruptions, reduced quality of life, and poorer prognostic outcomes. This review highlights the importance of a multidisciplinary approach to OM management, emphasizing preventive strategies and evidence-based interventions. Basic oral care, including meticulous hygiene and prophylactic dental interventions, forms the foundation of OM prevention. Cryotherapy, particularly during bolus 5-fluorouracil administration, has demonstrated efficacy in reducing OM severity by limiting mucosal exposure to cytotoxic agents. Low-level laser therapy (LLLT) and chemoprotective agents like palifermin offer promising results in high-risk populations, though their use requires careful consideration of safety and patient-specific factors. Pain management is a critical component of OM care, with topical morphine rinses emerging as an effective option for alleviating severe discomfort. Additionally, patient education plays a pivotal role in early detection and adherence to preventive measures, empowering patients to actively participate in their care. The integration of advanced radiation techniques, such as intensity-modulated radiotherapy (IMRT) and proton therapy, guided by radiologists, further minimizes mucosal toxicity while maintaining therapeutic efficacy. Collaborative efforts among oncologists, dentists, nurses, and pharmacists are essential for optimizing OM management, ensuring timely interventions, and improving patient outcomes. Despite advancements in OM management, gaps in knowledge persist, particularly regarding the role of zinc supplementation and the long-term safety of emerging therapies like LLLT. Future research should focus on refining preventive strategies, exploring novel therapeutic agents, and enhancing interdisciplinary collaboration to address the evolving needs of cancer patients. In conclusion, OM is a complex and multifactorial condition that demands a proactive, patient-centered approach. By leveraging evidence-based interventions, fostering multidisciplinary collaboration, and prioritizing patient education, healthcare providers can significantly mitigate the burden of OM, enhancing both treatment efficacy and patient quality of life.

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## التهاب الغشاء المخاطي الفموي كمضاعفة للعلاج الإشعاعي: مراجعة محدثة لمقدمي الرعاية الصحية

### الملخص

**الخلفية:** التهاب الغشاء المخاطي الفموي (OM) هو حالة منهكة تتميز بالحمى، والوذمة، والآفات القرحية في الغشاء المخاطي الفموي، وغالبًا ما تحدث كمضاعفة للعلاج الإشعاعي (RT)، والعلاج الكيميائي، أو زراعة الخلايا الجذعية المكونة للدم (HSCT). يؤثر OM بشكل كبير على جودة حياة المرضى، مما يسبب ألمًا شديدًا، ونقصًا غذائيًا، وزيادة خطر العدوى، مما يستدعي في كثير من الأحيان تعديل العلاج.

**الهدف:** تهدف هذه المراجعة إلى تزويد مقدمي الرعاية الصحية بفهم محدث لأسباب OM، وآليات المرضية، وانتشاره، واستراتيجيات التعامل معه، مع التركيز على التدخلات المبينة على الأدلة لتخفيف تأثيره على المرضى الخاضعين لعلاج السرطان. تهدف هذه المراجعة لتقييم دور العاملين بالمجال الصحي والأطباء وأطباء الأسنان وأخصائي الأشعة.

**الأساليب:** تجمع هذه المراجعة بين أحدث الأدبيات حول OM، مع التركيز على الفيزيولوجيا المرضية، والعرض السريري، والتقييم التشخيصي، والنهج العلاجية. تسلط الضوء على النموذج الخماسي لتطور OM، وتناقش مقاييس التقييم مثل CTCAE، و WHO، و OMAS. كما يتم تقييم استراتيجيات العلاج، بما في ذلك العناية الفموية الأساسية، والعلاج بالتبريد، والعلاج بالليزر منخفض الشدة (LLLT)، والعوامل الوقائية الكيميائية مثل باليفيرمين.

**النتائج:** يعد OM حالة متعددة العوامل تتأثر بكثافة العلاج، والعوامل الخاصة بالمرضى، وقابلية الغشاء المخاطي للإصابة. تقلل التدابير الوقائية، مثل العناية الفموية الأساسية والعلاج بالتبريد، بشكل كبير من شدة OM. كما يُظهر كل من LLLT وباليفيرمين نتائج واعدة في الفئات عالية الخطورة، رغم استمرار المخاوف بشأن الأمان. يساهم التحكم في الألم، بما في ذلك استخدام غسولات المورفين الموضعية، في تحسين راحة المرضى والالتزام بالعلاج.

**الخلاصة:** يتطلب التعامل الفعال مع OM نهجًا متعدد التخصصات يدمج بين الاستراتيجيات الوقائية، والكشف المبكر، والتدخلات المصممة وفقًا لحالة المريض. يجب على مقدمي الرعاية الصحية إعطاء الأولوية لتتقيف المرضى، وتحسين بروتوكولات العلاج، وتطبيق العلاجات المبينة على الأدلة لتقليل مضاعفات OM وتحسين نتائج العلاج.

**الكلمات المفتاحية:** التهاب الغشاء المخاطي الفموي، العلاج الإشعاعي، العلاج الكيميائي، العلاج بالليزر منخفض الشدة، باليفيرمين، العلاج بالتبريد، إدارة الألم، الرعاية متعددة التخصصات.