

Symptom Management Algorithm

Oral Fungal Infections

In Adults with Cancer

About the Oral Fungal Infections Algorithm

- The purpose of this algorithm is to outline the best clinical practice for pharmacological interventions of oral fungal infections. The patient must be properly diagnosed through conventional testing, including but not limited to: exam findings, visual exam, culture and sensitivity from oral swabs, blood cultures, X-rays, and biopsies as indicated
- This algorithm does not negate the need for consultation with the prescriber, dental provider, infectious disease specialist and/or pharmacist

Considerations for All Patients

- Good oral care is important to prevent and decrease oral complications, to maintain normal function of the oral tissues, to maintain comfort, and to reduce the risk of local and systemic infection. See the basic oral care tables (pages 4 and 5)
- Significant risk factors for the development of oral complications include the type of cancer, type of cancer treatments, cumulative doses of chemotherapy or radiation treatment (current or prior), method of delivery, and duration of treatment
- Predisposing medical, dental, and lifestyle factors such as uncontrolled diabetes, pre-existing autoimmune conditions, polypharmacy, tobacco use, alcohol use, oral hygiene, and dental health may increase severity of oral complications
- Oral complications can significantly affect the patient's morbidity, ability to tolerate treatment, and overall quality of life
- Rigorous assessment, diagnosis and early intervention are important in preventing and decreasing oral complications; this includes the assessment of nutritional status and adequacy of oral intake
- A large variety of medications may cause oral complications. Consultation with the prescriber, dental provider, and/or pharmacist is recommended
- Some pharmaceuticals may be unaffordable, and alternatives should be offered where possible
- If odontogenic or periodontal infection is suspected, consultation with a oncology team is strongly recommended

Risk Factors, Signs and Symptoms, and Prevention

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Risk Factors		
Neutropenia/immunosuppression	Tobacco use	
Dry mouth	 Poor oral hygiene 	
• Dehydration	 Malnutrition 	
Antibiotic course	• Denture use	
Corticosteroids (topical, inhaled, systemic)		

Prevention

Non-Pharmacological

 Follow meticulous oral care, see the basic oral care tables (pages 4 and 5)

Pharmacological—see dosing table (page 3)

 Fluconazole to reduce oral fungal colonization in patients receiving cancer therapy

Signs and Symptoms

- Pseudomembranous candidiasis: white curd like, removable with pressure, erythematous (red) oral mucosa, burning pain, taste changes, foul taste when not eating
- Hyperplastic candidiasis: hyperkeratotic white patch, cannot be removed with wiping
- Erythematous candidiasis: denuded (bald) and intensely erythematous oral mucosa, burning sensation, pain, and taste changes
- Angular cheilitis: erythema (rash), fissuring and crusting of the commissure (corner of mouth)
- Median rhomboid glossitis, mid-dorsum depapillated tongue with possible 'kissing' contact lesion on the hard palate



Figure 1: Pseudomembranous oral candidiasis 1



Figure 4: Hyperplastic candidiasis



Figure 2: Pseudomembranous oral candidiasis 2



Figure 6: Angular cheilitis



Figure 3: Erythematous oral candidiasis



Figure 5: Coated tongue, suspected oral candidiasis overgrowth

Management

Pharmacological—See dosing table below for details

- Chlorohexidine can be complimentary to systemic or topical therapy, with the exception of patients on active head and neck radiation therapy. See the basic oral care tables (pages 4 and 5)
- Topical agents are preferable to systemic agents for managing mild intraoral fungal infections, due to lower risk of side effects and drug interactions.
- If topical agents are not well tolerated, or the response rate is poor, use systemic agents
- Consider antifungal prophylaxis with antimicrobial therapy if patient is at risk of oral candidiasis
- Mild oropharyngeal candidiasis: clotrimazole or nystatin suspension may be used as first line treatment
- Moderate to severe oropharyngeal candidiasis: systemic fluconazole may be used as first line treatment
- Patients with acute or chronic renal impairment may need reductions of fluconazole or topical therapy
- For fluconazole refractory disease, itraconazole or posaconazole are recommended
- For ongoing refractory cases, voriconazole and amphotericin-B may be used.

Follow-Up and Ongoing Monitoring

 Seek dental care in addition to prescribing medication. For ongoing infections not responding to prescribed treatment, request the assistance of an infectious disease specialist or consider a swab for culture and sensitivity, if feasible

Antimicrobial Stewardship and Opioid Use

Antimicrobial Stewardship

- Antimicrobial stewardship is the appropriate use of antimicrobials to optimize clinical outcomes, combat resistant infections, avoid adverse drug events, and minimize costs. It means that patients get the right antibiotic, only when needed and for only as long as needed
- Visit Choosing Wisely Canada and The Canadian Association for Hospital Dentists for best practices of antimicrobial stewardship

Pharmacological Dosing Table for Management of Oral Fungal Infections

Rx	Dispense	Dose and Route
Fluconazole 100mg	15 tablets	 Take 2 tablets (200mg) initially with a meal, then 1 tablet a day with a meal until finished LU Code: 203
Fluconazole 400mg		 400mg by mouth, daily for hematopoietic stem cell transplantation (HSCT) fungal prophylaxis For autologous HSCT patients: fluconazole stops with engraftment For allogenic HSCT patients: continue fluconazole for several months
Nystatin suspension 100,000 units/mL	250mL	 Swish 5 to 10mL in mouth for 30 seconds and swallow four times a day for 14 days For patients with dentures: soak dentures in denture bath with solution and rinse
Ketaconazole 2% cream	30g	Apply to corner of mouth 7 to 14 times a day

^{*}Extended use of antimicrobials may cause secondary oral candidiasis, an adjuvant antifungal therapy should be considered

Basic Oral Care Tables

Flossing

Basic	 Patients who have not flossed routinely before cancer treatment should not begin flossing at this time Patients with mouth cancers, trismus, dysphagia, and/or dysgeusia may not be able to floss; use of interproximal brushes can replace flossing Floss at least once daily Waxed floss may be easier to use and minimize trauma to the gums
Intensified	Continue with basic plan until discomfort becomes too great
End of Life	Discontinue flossing if patient chooses

Discontinue flossing if:

• Gums bleed for longer than two minutes

Restart flossing if:

• Platelet count is >20x10⁹ cells/L, or as instructed by cancer care team

Brushing

Basic	 Use a small, ultra-soft-headed, rounded-end, bristle toothbrush (an ultrasonic toothbrush, may be acceptable) Rinse toothbrush in hot water to soften the brush before using Use a prescription strength fluoride toothpaste. Spit out the foam but do not rinse mouth Use a fluoridated toothpaste and re-mineralizing toothpaste containing calcium and phosphate Brush tongue gently from back to front, using a sweeping motion Rinse brush after use in hot water and allow to air dry Change toothbrush when bristles are not standing up straight Brush within 30 minutes after eating and before bed. Ensure the gingival portion of the tooth and periodontal sulcus (where the tooth and gums meet) are included Consider topical anesthetics (e.g. viscous lidocaine 2% or viscous xylocaine 2%, 2-5 mL) before brushing and eating to minimize pain With continuous pain, a regularly prescribed oral analgesic allows for more thorough tooth brushing
Intensified	 Encourage patient to continue brushing through treatment phase even when it causes discomfort If bleeding occurs, encourage gentler brushing Use a non-flavoured, non-alcoholic chlorhexidine gluconate (CHX) 0.12% rinse to aid in plaque control, 2 times a day after meals If unable to continue brushing with a toothbrush, use a moist gauze or foam swab Discontinue use of toothpaste if it is too astringent and dip toothbrush in bland rinse If there has been an oral infection, use a new toothbrush after infection has resolved If unable to tolerate brushing, seek assistance from nursing or dental staff
End of Life	 Continue with basic and intensified mouth care plan, if possible Instead of moist gauze may use a foam brush soaked in CHX

Discontinue brushing if:

• Gums bleed for longer than two minutes

Restart brushing if:

• Platelet count is >20x10⁹ cells/L, or as instructed by cancer care team

Bland rinse:

• 1 teaspoon salt, 1 teaspoon baking soda, 4 cups of water

Lidocaine alternative:

• Dyclonine 0.5 or 1% rinse (5 mL every 6 to 8 hours, swish and swallow) as needed for pain

Patients with head and neck cancers:

- Brushing may not be appropriate in the area of tumour involvement
- Consult with a dentist
- Patients should be assessed for the use of daily fluoride tray

Patients with dentures:

- Remove dentures, plates and prostheses before brushing
- Brush and rinse dentures after meals and at bedtime
- Remove from mouth nightly (at least 8 hours per 24 hours) and soak in bland rinse
- Leave dentures out as much as possible during radiation therapy
- Patients who have had head and neck surgery should not wear dentures post-surgery unless assessed by a dental specialist or head and neck surgeon, to prevent

Rinsing

Basic	 Rinse the oral cavity with a bland rinse vigorously, at least twice a day to maintain mouth moisture, remove the remaining debris and toothpaste, and reduce the accumulation of plaque and infection Use a bland rinse to increase oral clearance for oral hygiene maintenance and improved patient comfort. Following emesis, rinse with bland rinse immediately to neutralize the mouth If allergic to lidocaine, dyclonine 0.5 or 1% rinse (5 mL every 6 to 8 hours, swish and swallow) may be used as needed for pain
Intensified	 Rinse in place of brushing if patient is unable to brush Seek dental care where possible for removing plaque In addition to rinsing twice a day, encourage rinsing every 1 to 2 hours while awake and every 4 hours through the night if awake, to minimize complications of decreased saliva If unable to clean using moist gauze, or foam swab, consider rinsing via syringe if platelet count >20x10⁹ cells/L
End of Life	 Continue with basic and intensified mouth care plan Consider sialagogues in instances of dry mouth for pharmacotherapy relief (pilocarpine, and anethole trithione)

Moisturizing the Oral Cavity

Basic	 Moisturize the mouth with water, artificial saliva products, or other water soluble lubricants Apply lubricant after each cleaning, at bedtime, and as needed. Water-based lubricant needs to be applied more frequently Frequent rinsing as needed with basic mouth rinse Patients may suck on xylitol lozenges (up to 6 grams a day), xylitol containing popsicles, or xylitol containing gum
Intensified	 Continue with basic mouth care plan with increased frequency and intensity Increase frequency of bland mouth rinse to every hour
End of Life	 Continue with basic mouth care plan with increased frequency and intensity, as needed Use a steam vaporizer at night May use a cool mist humidifier at night, but use should be weighed against the risk for fungal infection

Lip Care

Basic	 To keep lips moist and avoid chapping and cracking, use water soluble lubricants, lanolin (wax-based), or oil based lubricants (mineral oil, cocoa butter) Water soluble lubricants should be used inside and outside the mouth, and may also be used with oxygen (e.g. products compounded with Glaxal base or Derma base) Apply lubricant after each cleaning, at bedtime, and as needed. Water-based lubricants need to be applied more frequently
Intensified	Continue with basic mouth care plan with increased frequency and intensity
End of Life	 Continue with basic mouth care plan with increased frequency and intensity, as needed May use a cool mist humidifier at night, but use should be weighed against the risk for fungal infection

Miscellaneous

Basic	 Dental evaluation and treatment as indicated prior to cancer therapy is desirable to reduce risk for local and systemic infections from odontogenic sources for hematologic, solid or head and neck cancers
Intensified	Continue with basic mouth care plan with increased frequency and intensity
End of Life	Continue with basic mouth care plan with increased frequency and intensity, as needed

Patients with dentures:

- After removing dentures, rinse mouth thoroughly with rinse solution
- Brush and rinse dentures after meals and at bedtime
- Rinse with rinsing solution before placing in mouth
- Remove from mouth nightly (at least 8 hours per 24 hours) and soak in rinsing solution

Bland rinse:

• 1 teaspoon salt, 1 teaspoon baking soda, 4 cups of water

Avoid:

- Club soda due to the presence of carbonic acids
- Commercial mouthwashes with hydroalcoholic base or astringent properties

Avoid:

- Glycerin or lemon-glycerin swabs as they dry the mouth
- Acidic or minty mouth products, if they burn

Avoid:

- Touching any lip lesions
- Oil based lubricants on the inside of the mouth
- Petroleum based products

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