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Xerostomia:

Recognition and Management

By Philip C. Fox, DDS, FDS, RCSEd

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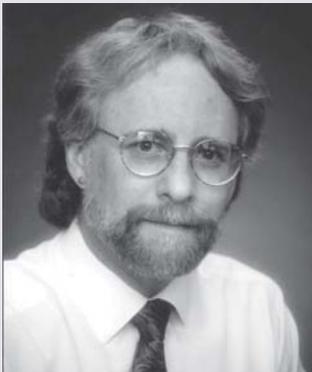
Introduction

Xerostomia (dry mouth) is the subjective feeling of oral dryness. It is important to recognize that xerostomia is not a diagnosis, but a symptom with multiple possible causes. Although dry mouth is most frequently associated with altered salivary gland function, there are other etiologies for this oral issue. It is important to perform a complete evaluation of the patient with dry mouth, determining, if possible, the cause of the xerostomia so that appropriate management can be instituted in a timely manner. The patient with xerostomia who has salivary gland hypofunction is at risk for many oral complications and it is critical to institute appropriate preventive measures. Xerostomia may also be a consequence of systemic disease, and early recognition may aid in treatment. This monograph will review methods of evaluation for xerostomia and practical measures for management of the patient with dry mouth.

Epidemiology and Etiology of Xerostomia

Xerostomia is a common oral concern for many patients. It is estimated that up to 10 percent of the general population experiences persistent oral dryness.^{1,2} Xerostomia is more frequent with increasing age, and over 25 percent of elders complain of daily dryness.³ That percentage is even higher in institutionalized elders. However, it should be recognized that xerostomia is not a result of aging per se, and should not be dismissed as such. Salivary function in healthy, non-medicated elders does not vary significantly from younger individuals. It is believed that the increased frequency of dryness complaints with aging is a result of systemic disease and medication use, both more common in elders and both associated with xerostomia.⁴

As noted earlier, there are non-salivary causes of xerostomia. These include conditions in which there are alterations of oral mucosal sensation or central cognitive changes, such as



Philip C. Fox, DDS, FDS, RCSEd, received his BA and DDS degrees from Columbia University and completed training in oral and maxillofacial surgery at Harlem Hospital Center, New York, N.Y. and a fellowship in oral medicine at the National Institutes of Health in Bethesda, Md. He was a member of the National Institute of Dental and Craniofacial Research, National Institutes of Health, for 22 years, serving as chief of the Clinical Investigations Section, Gene Therapy and Therapeutics Branch, and as clinical director of the Intramural Research Program. At present, he is a visiting scientist at the Department of Oral Medicine, Carolinas Medical Center, in Charlotte, N.C., and an independent biomedical consultant focusing primarily in the area of clinical trial design and analysis. He is a diplomate of the American Board of Oral Medicine.

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following a cerebral vascular accident (stroke). Dryness has been associated with depression, even in the absence of recognizable changes in salivary function. The most common causes of xerostomia, however, are conditions or circumstances that result in alterations in salivary gland function, either quantitative or qualitative.¹

The most frequent cause of dry mouth complaints is the use of prescription drugs. There are hundreds of pharmaceuticals that have xerostomia as a side effect.^{5,6} The salivary glands are stimulated strongly by cholinergic agents. Therefore, anti-cholinergic medications, such as antihistamines, are most likely to cause decreased salivary output and xerostomia.⁵ However, many other classes of medications, including sedatives, antipsychotics, antidepressants and diuretics are associated with xerostomia. Interestingly, many herbal preparations may induce complaints of oral dryness, and patients should always be questioned about non-prescription drug use.⁷ Examples of commonly used preparations include members of the Labiatae family (specifically all the salvias), capsicum, garlic, Ginkgo biloba, and St. John's Wort (*Hypericum perforatum*). Herbal preparations with strong diuretic properties, such as stinging nettle (*Urtica dioica*) and dandelion (*Taraxacum officinale*) also may lead to complaints of dry mouth. Caffeine is well recognized as a cause of xerostomia.⁸ As many as 25 million persons in the United States may experience xerostomia as a result of medication use.

The most frequent cause of dry mouth complaints is the use of prescription drugs.

There are also medical therapies that result in xerostomia. Radiotherapy to the head and neck region that includes the salivary glands in the treatment fields can lead to significant and persistent dry mouth. This is an expected complication at exposure levels above 5200 cGy.^{9,10} Systemically administered radionuclides, such as ¹³¹I for treatment of thyroid cancer, are a recognized cause of dry mouth, particularly when re-treatment is required.¹¹ Bone marrow transplantation also may have xerostomia as a side effect, due to the salivary gland involvement in graft-versus-host disease.¹²

Another major cause of xerostomia is systemic disease. There are a large number of conditions that can affect salivary gland function and lead to complaints of dryness, including diabetes, thyroid disorders, cystic fibrosis and connective tissue diseases.¹³ The most prominent of the connective tissue diseases is Sjögren's syndrome, an autoimmune condition that may affect up to 4 million people in the United States. Xerostomia is one of the hallmark symptoms in Sjögren's syndrome, experienced by over 95 percent of patients.¹⁴ Patients experience a number of dryness complaints in addition to xerostomia. These include eye, nose, throat, skin and vaginal dryness. Since complaints of oral dryness typically do not appear until salivary function has been reduced by approximately 50 percent,¹⁵ more than a single gland must be affected

Table I. Systemic Conditions that May Have Xerostomia as a Symptom

- Autoimmune connective tissue disorders
 - Sjögren's syndrome, primary and secondary
- Granulomatous diseases
 - sarcoidosis, tuberculosis
- Graft-versus-host disease
- Cystic fibrosis
- Bell's palsy
- Diabetes (poorly controlled)
- Amyloidosis
- Human immunodeficiency virus infection
- Thyroid disease
 - hypo- and hyper-thyroidism
- Late stage liver disease
- Affective disorder
- Eating disorders and malnutrition
 - anorexia, bulimia, dehydration

for xerostomia to manifest. This emphasizes that dryness resulting from salivary dysfunction is usually indicative of systemic involvement. Table I provides a fuller listing of systemic conditions with xerostomia as a possible symptom.

Diagnosis of Xerostomia

IT IS CRITICAL THAT ORAL health care practitioners recognize xerostomia and salivary gland dysfunction in their patients, as saliva plays an essential role in protection of oral hard and soft tissues and support of oral functions.¹⁶ One should follow a systematic assessment approach to the patient with xerostomia: listen for and elicit symptomatic complaints, examine for oral signs of salivary gland dysfunction and evaluate salivary gland function.

The most frequently voiced complaint is a feeling of dryness of all the oral mucosal surfaces, including the throat. Additionally, patients may report difficulty in chewing, swallowing, or speaking. A need to drink fluids to help swallow while eating or an inability to swallow dry foods is also common. Many patients will carry fluids at all times for comfort and to aid in speaking and swallowing. Pain is often reported, and the oral mucosa may be sensitive to spicy or coarse foods, which may limit the patient's diet and enjoyment of meals. Patients may report that their lips and the oral mucosa 'stick' to their teeth. The salivary glands may swell intermittently or chronically. Removable denture wearers commonly report problems with retention and mucosal damage due to lack of lubrication. When denture adhesives are used, they may not fully develop their adhesive and retentive properties through failure to hydrate adequately. Patients should also be questioned concerning dryness in other areas. Eye, throat, nasal, skin or vaginal dryness, in addition to xerostomia, may be indicative of a systemic condition, such as Sjögren's syndrome.



Figure 1. Dry mouth associated with Sjögren's syndrome. The tongue is dry and pale and has lost papillation of the surface.

It is recognized that the severity of xerostomia is poorly correlated with salivary gland function. Some individuals tolerate marked (or almost complete) lack of salivation with few complaints, while for others, modest decreases in salivary output may trigger severe complaints of dry mouth and oral discomfort. The reasons for this poor correlation are unclear and reflect a lack of knowledge about the elements necessary for oral comfort. Although subjective dryness does not correlate well with measurable salivary gland dysfunction, some symptoms have been found to have predictive value.¹⁷ When xerostomia is reported, patients should be questioned in greater detail about its nature, frequency and duration. If questioning reveals difficulties with oral activities dependent on adequate salivation, such as chewing and swallowing, it is likely that salivary function is diminished.

Clinical Assessment of Xerostomia

On examination, the patient with xerostomia due to reduced salivary gland function usually has obvious signs of mucosal dryness.¹ The lips may be cracked, peeling and atrophic; the buccal mucosa pale and corrugated in appearance; and the tongue smooth and reddened with loss of papillation (Figure 1). The oral mucosa may appear reddened, thinner and more fragile. There is often a marked increase in erosion and dental caries, particularly at the gingival margin, and even cusp tip involvement. The decay may be rapid and progressive even in the presence of excellent oral hygiene (Figure 2). One should consider whether the caries' history and current condition are consistent with the patient's oral hygiene. Candidiasis, most frequently of the erythematous form, is frequent and may contribute to mucosal sensitivity. The salivary glands should be examined for enlargement, changes in texture, and pain and also to determine if saliva can be expressed from the main excretory ducts. The saliva should be clear, watery and copious. A cloudy exudate may be a sign of bacterial infection, although some patients with very low salivary output will have opaque secretions that are sterile.



Figure 2. The lips, tongue and all mucosal surfaces are dry in this patient with Sjögren's syndrome. Note also the erosion and the presence of epithelial debris on the teeth, a sign of diminished salivary secretions.

While inter-individual variability makes it difficult to determine a "normal" value for salivary output, there is agreement on the minimal values necessary to consider salivary output normal. Unstimulated whole saliva flow rates of <0.1 mL/min are considered abnormally low and indicative of marked salivary hypofunction.²¹ It is important to recognize that output higher than that amount does not guarantee that function is normal. For some individuals, this may be a markedly reduced value. However, this value represents a generally accepted lower limit of normal and is a useful guide for the clinician. Using a well-defined, standardized, and clearly documented procedure for collection will allow meaningful comparisons to published material and with repeat measures in an individual over time.

The composition of the collected saliva may be analyzed (sialochemistry). Over 60 components have been identified in saliva.²² There is increasing interest in salivary diagnostics for systemic and oral diseases.²³ Saliva is used currently for determining blood alcohol levels, smoking and drugs of abuse as well as for HIV testing and detection of exposure to other microbes. Using newer genomic and proteomic techniques, researchers have begun to identify possible salivary biomarkers for a number of oral and systemic conditions. This is a rapidly expanding area of investigation.

Salivary function and gland integrity also can be examined with technetium (Tc) pertechnetate scintiscanning. Uptake of intravenously injected ^{99m}Tc tracer from the blood into the glands and subsequent secretion into the oral cavity can be monitored with a gamma camera. Salivary scintiscans correlate well with salivary function and can be used in clinical studies to monitor changes in function following interventions.²⁴

Tumors and structural alterations in the salivary glands can be visualized with sialography, ultrasonography, MRI or CT imaging.²⁵ These techniques are useful to detect salivary stones (sialoliths), duct blockage, constriction or damage, tumors and

In addition to these physical effects on the oral cavity, a patient's quality of life is severely impacted.^{18,19} Since adequate salivary function is important to support alimentation and communication, salivary gland dysfunction and xerostomia may adversely affect some of our most critical activities of daily living.

Salivary gland function should be assessed in the patient with xerostomia. This may be done easily in an office setting by determining the total unstimulated output of saliva, termed the whole saliva flow rate. Whole saliva is the mixed fluid contents of the mouth. Using the expectoration model, the patient allows saliva to accumulate in the mouth and then expectorates into a pre-weighed graduated cylinder every 60 seconds, for 5 to 15 minutes.²⁰ To insure an unstimulated sample, patients should refrain from eating, drinking, oral hygiene or smoking for 90 minutes prior

cysts. These are not functional measures, but are useful for diagnosis of salivary dysfunctions.

Biopsy of the salivary glands can be done as well. The most common procedure is biopsy of the minor salivary glands of the lower lip. Minor glands are abundant in this region and can be removed with minimal morbidity.²⁶ Characteristic changes are seen in these glands with Sjögren's syndrome. The labial minor salivary gland biopsy is considered the best sole diagnostic criterion for the salivary component of Sjögren's syndrome.²⁷

Laboratory studies may be helpful in the diagnosis of Sjögren's syndrome. Patients often have characteristic autoimmune alterations in the blood, including the presence of autoantibodies directed against SS-A (Ro) and SS-B (La), antinuclear antibodies and elevated immunoglobulins.

Using a systematic evaluation approach, the cause of dry mouth complaints can be identified in about 80 percent of patients. The remainder of patients are characterized as having idiopathic xerostomia, but still require palliative care for symptomatic relief.

Dental Hygienists' Role and Management of Xerostomia

THE DENTAL HYGIENIST HAS AN important role in recognition and management of the patient with xerostomia. With careful attention and regular management, most patients will achieve increased oral comfort and oral complications can be minimized.

Management may be viewed as a series of goals: to provide symptomatic relief of dryness complaints; to institute appropriate preventive measures; to treat any ongoing oral problems; if salivary function is decreased, to stimulate remaining function; and to address the underlying disease process. Management of xerostomia is a team process, involving the dentist, the dental hygienist and the patient's physicians. Effective communication is essential for optimal patient care.

Management starts with methods to relieve symptoms. These approaches are useful for any patient with dry mouth complaints, regardless of the cause. Frequent sips of water will help to relieve dryness, ease swallowing, hydrate tissues and cleanse the mouth. As noted earlier, many patients with dry mouth carry water with them at all times, and this should be encouraged. Oral rinses and mouthwashes, gels, sprays and artificial salivas are frequently used and may reduce discomfort and improve function transiently. The dental hygienist should advise the patient to use oral care products that have been specifically formulated for a dry mouth; that is, without alcohol, using only mild detergents and flavorings, having a neutral or alkaline pH, containing no sugar and with added lubricants. The use of chewing gums, lozenges, candies or mints may provide symptomatic relief by stimulating salivation, but they must be sugar-free and non-acidic. Products containing xylitol as a sweetening agent should be encouraged, as xylitol has been shown to have anti-caries properties.

For dry lips, a hydrating cream or ointment may help relieve symptoms. Use of products with aloe vera or vitamin E should be encouraged. Highly flavored toothpastes should be avoided because they may irritate the dry mucosa and cause pain or sensitivity. A diet that contains moisture-rich foods and not hot or spicy foods may be more comfortable for the patient. Alcohol and caffeine may produce further drying and should be avoided or used only in limited quantities. In addition to its other negative health consequences, smoking may be drying and irritating to the mucosa and should be avoided. An increase in environmental humidity is exceedingly important. Patients, especially in the wintertime, often experience a worsening of their symptoms. Use of room humidifiers, particularly at night, may improve discomfort markedly.

Preventive measures are necessary to minimize oral hard- and soft-tissue damage. Patients should be seen by the dental hygienist on a frequent recall schedule of at least three times per year, until symptoms are minimized and dental caries is controlled. They must be instructed in and practice vigilant home oral hygiene. Patients should brush with fluoride toothpaste after each meal. At a minimum, rinsing the mouth immediately after eating to remove food debris is beneficial. Advice on diet is critical. Hygienists must stress the importance of extremely low sugar intake, the use of sugar-free substitutes and the avoidance of sticky, carbohydrate-rich foods to minimize the caries risk. Patients should also be aware of the increased risk of tooth wear. Patients must be careful not to consume acidic foods and beverages in excess (e.g. carbonated sodas) in an effort to relieve symptoms. Remineralizing mouth rinses are now available and their use should be encouraged.

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Patients with dry mouth have an increased risk of fungal infections. Consumption of sugar-free yogurt containing active yeast cultures may help to control oral fungal populations. If a fungal infection is present, appropriate antifungal treatments should be started. Due to the salivary dysfunction, treatment may be prolonged and re-treatment is frequently required. A therapeutic agent should be selected that does not contain sugar as a flavoring agent, as is common in many antifungal lozenges. Nystatin rinses may be prepared by a pharmacist from powder without the added flavoring ingredient(s).

Dental caries management is based on the severity of salivary dysfunction and the observed caries rate. All patients with xerostomia should use some supplemental fluoride, in addition

to a fluoride-containing toothpaste. Selection of the type, dose and frequency of application should be made in consultation between the dentist, the dental hygienist and the patient. Often a combination of office-applied and home-based fluoride treatments is optimal.

Stimulating salivary gland flow is effective for relief of dryness symptoms and provides the myriad benefits of natural saliva. Chewing will stimulate salivary flow effectively, as will sour tastes. The combination of chewing and taste, as provided by gums or mints, can be very effective in relieving symptoms for patients who have remaining salivary function. As noted, however, patients must be told not to use products containing sugar as a sweetener. Electrical stimulation of the salivary glands has been attempted, but application is a challenge and the results have been modest at best. Presently, newer devices are being investigated and show promise for the future.²⁸

Systemic drugs, termed secretagogues, may be prescribed. The Food and Drug Administration (FDA) has approved two medications for the relief of dry mouth symptoms in Sjögren's syndrome: pilocarpine (Salagen[®]) and cevimeline (Evoxac[®]).²⁹⁻³¹ These medications are parasympathomimetic, muscarinic agonists, which induce a transient increase in salivary output and statistically significant improvement in complaints of oral dryness. Common side effects of both medications include sweating, flushing, urinary incontinence and gastrointestinal discomfort. While rarely severe or serious, side effects are frequent, which may limit the usefulness of these agents in some individuals. Parasympathomimetics are contraindicated in patients with uncontrolled asthma, narrow-angle glaucoma, and acute iritis and should be used with caution in patients with significant cardiovascular disease, Parkinson disease, asthma and chronic obstructive pulmonary disease. Pilocarpine is recommended at a dosage of 5 mg up to four times daily, while cevimeline is prescribed at 30 mg three times daily. These medications are widely used and provide significant relief of dryness complaints in many patients. Pilocarpine has also been approved for treatment of xerostomia related to head and neck radiotherapy.

A number of different approaches have been attempted to address the underlying disease process and therefore improve salivary gland dysfunction, particularly in Sjögren's syndrome. Current studies are targeting specific biologic pathways to modify the underlying autoimmunity.

Very low dose interferon-alpha, 150 or 450 IU once or three times a day orally, has been studied in primary Sjögren's syndrome. Use of 150 IU interferon-alpha three times daily resulted in increased stimulated whole salivary flow rates at 12 weeks compared to placebo.³² A subsequent phase 3 trial at this dose found increased unstimulated salivary function compared to placebo at 24 weeks. However, the co-primary endpoints of stimulated whole saliva flow and oral dryness were not significantly improved relative to placebo,³³ and further clinical studies will be necessary.

Another approach has been to modify the inflammatory cytokine pathway, specifically targeting tumor necrosis factor alpha (TNF α). Anti-TNF α agents, including infliximab,

etanercept and thalidomide, which have shown benefit in rheumatoid arthritis and lupus erythematosus, have shown no significant efficacy in randomized controlled trials in Sjögren's syndrome.³⁴⁻³⁶

At present, there is interest in modulation of B-cell activation. As there is well-documented B-cell hyperreactivity in Sjögren's syndrome, this approach may have an impact on the underlying disease. Rituximab is a humanized monoclonal antibody that binds specifically to the CD20 antigen, which is present on B lymphocytes and is involved in cell activation. In an open-label study, improvements in symptoms of dry mouth and salivary gland function were demonstrated.³⁷ Although these results are promising, a randomized controlled trial is necessary before recommendation of this treatment option for Sjögren's syndrome. A recent open-label trial looked at an anti-CD22 monoclonal antibody and reported very promising results in Sjögren's syndrome.³⁸ Further studies are anticipated.

Management of xerostomia requires a team approach. The dentist and dental hygienist should work together closely to minimize the patient's dryness complaints and to preserve and protect the oral tissues. Patient education is key.

Management of xerostomia requires a team approach. The dentist and dental hygienist should work together closely to minimize the patient's dryness complaints and to preserve and protect the oral tissues. Patient education is key. The dental team needs to explain the oral sequelae of dry mouth and the problems and issues patients may encounter in their daily lives. The oral condition needs to be assessed and monitored closely, and issues such as oral hygiene care and diet should be emphasized. Consultation with the patient's physician is indicated if, for example, the xerostomia appears to be related to medication use. In this case, discussion of alternative medications or dosing regimens to minimize xerostomia is indicated. If systemic disease is suspected as the underlying cause of salivary dysfunction, patients should be referred to their physician or an appropriate specialist without delay. Ongoing review and monitoring of the patient's oral and general health are important for optimal management.

Summary and Conclusions

Awareness and recognition of xerostomia are essential in order to help patients minimize dryness symptoms, to institute preventive measures and to limit oral complications. The dental hygienist has the opportunity to ask every patient if they are

experiencing dry mouth. In particular, complaints of dryness while eating, or difficulty swallowing dry foods, or the necessity of using liquids to ease swallowing are important clues that salivary function may be impaired. As part of a routine oral examination, one should examine the oral cavity carefully for signs of salivary gland dysfunction. Findings such as an increase in caries activity, mucosal alterations, infection or salivary gland enlargement may indicate salivary dysfunction. Evaluation should be conducted proactively at each patient visit. Early recognition will minimize damage and dysfunction and allow appropriate management to begin. Although the salivary dysfunction may be irreversible, preventive measures and conservative treatments can avoid or limit mucosal breakdown, infections and permanent damage to teeth. Adequate symptomatic relief is possible with local palliative and systemic measures in many patients. Appropriate management of symptoms and increasing saliva output may help patients feel more comfortable and improve their quality of life.

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