

# A Review of Current Research on the Effects of Betel Nut Consumption on Oral Mucosal Health

Rui Liu\*

College of Life Science and Technology, Wuhan Polytechnic University, Wuhan, Hubei, 430023, China  
\*Corresponding author: lr4ever@foxmail.com

**Abstract:** Betel nut (*Areca catechu*) is a widely chewed plant that is grown and eaten in Asia and the Pacific. Prolonged consumption of betel nut may lead to oral health problems such as oral mucosal damage and oral fibrosis. The aim of this review paper is to examine the current state of research on the effects of betel nut on oral mucosal health, particularly in relation to oral injury and oral fibrosis.

**Keywords:** Betel nut, Oral mucosa, Injury, Fibrosis, Oral cancer

## 1. Introduction

Betel nut is a common oral irritant that is widely distributed in Southeast and South Asia. It is estimated that over one billion people worldwide consume betel nut. Despite its popular euphoric and relaxing effects, long-term betel nut consumption can cause severe damage and fibrosis to the oral mucosa and even lead to the development of oral cancer. Therefore, this paper aims to review the effects of betel nut on oral mucosa health and raise public awareness of the dangers of betel nut.

## 2. Background of the Study on Betel Nut and its Chemical Composition

### 2.1. Background of the Study

The betel nut is a palm plant native to Malaysia and Indonesia and it is a traditional hobby item in Asia and the Pacific. Betel nut has a long history of cultivation and consumption and is considered to be an important social and cultural symbol. However, the betel nut chewing habit has been recognised as a harmful practice for oral health. The betel nut composition contains many alkaloids, tannins and volatile oils, the most prominent of which is betel nutine. Prolonged betel nut chewing can lead to damage to the oral mucosa, such as yellowing of the teeth, mouth ulcers and gingivitis. In addition, betel nut has been linked to the development of cancers such as oral cancer, oesophageal cancer and stomach cancer [1]. The betel nut chewing habit has become a public health problem. The use and chewing of betel nut has become a social phenomenon and in some areas has even become a cultural tradition. However, the harmful effects of betel nut are widely recognised and more and more people are becoming aware of the dangers of betel nut. Some countries and regions have introduced policies and regulations to restrict or ban the use and chewing of betel nut. The betel nut chewing habit has become a public health issue and long-term betel nut chewing can have many negative effects on oral health and may even lead to the development of cancer. Therefore, there is a need to enhance publicity and education to raise public awareness of the dangers of betel nut and to introduce corresponding policies and regulations so as to restrict or ban the use and chewing of betel nut to protect the health and well-being of the public [2].

### 2.2. Chemical Composition of Betel Nut

The main bioactive components of betel nut include betel nutine, betel nut phenol and betel nut fibre. Among them, betel nutine is the most important alkaloid in betel nut and has a variety of biological activities, including stimulation of the nervous system, vasoconstriction and appetite suppression. Long-term consumption of betel nut can lead to the accumulation of betel nut alkaloids in the mouth, which in turn can cause oral mucosal lesions such as mouth ulcers and inflammation. Betel nut phenol, a phenolic compound in betel nut, has various biological activities such as antioxidant, anti-inflammatory and anti-cancer. However, long-term consumption of betel nut can lead to the

accumulation of betel nut phenol in the oral cavity, which in turn has a direct toxic effect on the oral mucosa, leading to oral mucosal lesions. Betel nut fibre is a type of cellulose found in betel nut, which has an irritating and mechanical effect. Long-term consumption of betel nut can lead to the accumulation of betel nut fibres in the mouth, which in turn can irritate the oral mucosa and cause lesions such as inflammation and ulcers in the oral mucosa. In addition to the main components mentioned above, betel nut also contains a variety of other bioactive components such as anthocyanins, flavonoids and polysaccharides. These components also have important effects on the health of the oral mucosa, including toxicity to oral mucosal cells and immunomodulatory effects on the oral mucosa [3]. The main bioactive components of betel nut have a direct impact on the health of the oral mucosa. Long-term consumption of betel nut can lead to the accumulation of these components in the oral cavity, which in turn can cause oral mucosal lesions. Therefore, in order to protect oral health, betel nut should be avoided or its intake should be reduced. It is also important to have regular oral check-ups to detect and treat oral mucosal lesions in a timely manner.

### **3. Effects of Betel Nut on Oral Injuries**

#### ***3.1. Direct Damage to the Oral Mucosa Caused by Betel Nut***

Betel nut is one of the main components of betel nut, and long-term consumption of betel nut can lead to the accumulation of betel nut alkaloids in the oral cavity, which in turn can have harmful effects on oral mucosal cells. Studies have shown that betel nut alkaloids can cause apoptosis of oral mucosal cells, resulting in damage to the oral mucosa. In addition, betel nut alkaloids also stimulate the secretion of cytokines and inflammatory mediators by oral mucosal cells, causing inflammation and ulceration of the oral mucosa. The betel nut phenol in betel nut is also one of the components that have a direct impact on the health of the oral mucosa. Betel nut phenol is strongly oxidizing and long-term consumption of betel nut can lead to the accumulation of betel nut phenol in the oral cavity, which in turn causes oxidative stress reactions in the oral mucosa [4]. These reactions can lead to oxidative damage and inflammatory responses in the cells and tissues of the oral mucosa, which in turn can lead to the development of oral mucosal lesions and carcinogenesis. The betel nut fibre in betel nut is also one of the components that have a direct impact on the health of the oral mucosa. The mechanical and irritating effects of betel nut fibres can lead to the accumulation of betel nut fibres in the mouth over a long period of time, which in turn can irritate the oral mucosal cells and cause inflammation and ulceration of the oral mucosa. Therefore, in order to protect the health of the oral mucosa, betel nut should be avoided or its intake should be reduced. It is also important to have regular oral check-ups to detect and treat oral mucosal lesions in a timely manner [5].

#### ***3.2. Effect of Betel Nut on Oral Microorganisms***

The chemical composition of betel nut can alter the structure of the oral microbial community, thereby affecting oral health. Long-term consumption of betel nut can lead to the accumulation of chemical components in the betel nut in the mouth, which in turn can alter the structure and composition of the oral microbial community. Studies have shown that long-term betel nut consumption reduces beneficial bacteria and increases pathogenic bacteria in the oral cavity of people who consume betel nut. These changes may increase the risk of inflammation and damage to the oral mucosa. The balance of the oral microbial community has important implications for oral health. Beneficial bacteria can inhibit the growth and reproduction of pathogenic bacteria, maintaining the stability of the oral microbial community and protecting the health of the oral mucosa. Pathogenic bacteria can cause lesions such as inflammation and infection in the oral mucosa.

The chemical constituents in betel nut can alter the structure and composition of the oral microbial community, resulting in an increase in the number of pathogenic bacteria and a decrease in the number of beneficial bacteria in the mouth. These changes may lead to instability of the oral microbial community and increase the risk of inflammation and damage to the oral mucosa. To protect oral health, betel nut consumption should be avoided or reduced. In addition, regular oral check-ups and cleanings and good oral hygiene practices can help maintain the balance of the oral microbial community and protect the health of the oral mucosa [6].

## **4. The Relationship between Betel Nut and Oral Fibrosis**

### ***4.1. Mechanisms of Betel Nut-induced Oral Fibrosis***

Oral fibrosis is a common disease of the oral mucosa. Its main manifestations are sclerosis, loss of elasticity and functional impairment of the oral mucosa. The occurrence and development of oral fibrosis is associated with a number of factors, among which the chemical constituents in betel nut may also be involved. One of the main components is betel nut alkaloids, which is found in betel nut. Studies have shown that betel nut alkaloids can induce proliferation and differentiation of oral mucosal fibroblasts, thereby promoting excessive collagen deposition, which in turn triggers oral fibrosis. Betel nut alkaloids promote fibroblast proliferation and differentiation by activating various signalling pathways such as Wnt/ $\beta$ -catenin and TGF- $\beta$ /Smad signalling pathways, leading to oral mucosal fibrosis. In addition to betel nut alkaloids, betel nut phenols and betel nut fibres in betel nut may also be involved in the development and progression of oral fibrosis. Betel nut phenols are strongly oxidizing and can cause oxidative damage to the oral mucosa, thus promoting oral fibrosis. Betel nut fibres have a mechanical and irritating effect that can irritate the oral mucosa cells and promote the development and progression of oral fibrosis [7]. Therefore, in order to prevent the onset and development of oral fibrosis, long-term betel nut consumption should be avoided or the intake of betel nut should be reduced. In addition, regular oral check-ups and cleaning and good oral hygiene practices can help prevent the onset and development of oral mucosal disease. If oral fibrosis has already occurred, prompt medical attention should be sought for treatment.

### ***4.2. The Association Between Oral Fibrosis and Oral Cancer***

The incidence of oral cancer is significantly higher in patients with oral fibrosis than in non-fibrosis patients. Studies have shown that the chemical constituents in betel nut, particularly betel nut alkaloids and its metabolites, may be an important contributor to this phenomenon. The betel nut alkaloids in betel nut can cause genetic mutations in oral mucosal cells through a variety of pathways, which can lead to carcinogenesis. Betel nut alkaloids can cause genotoxic effects such as DNA strand breaks and base mutations by interfering with DNA replication and repair processes. In addition, betel nut alkaloids can also bind to DNA and form DNA adducts, which can cause genetic mutations and carcinogenesis. In addition to genotoxicity, betel nut alkaloids can also promote the development of oral cancer by inducing oxidative stress and inflammatory responses. Betel nut alkaloids can cause oxidative stress, producing large amounts of free radicals and oxidants that lead to cell damage and death. In addition, betel nut alkaloids can induce an inflammatory response and increase the degree of inflammation in the oral mucosa, thus promoting the development of oral cancer. Therefore, in order to prevent the development of oral cancer, long-term betel nut consumption should be avoided or the intake of betel nut should be reduced. In addition, regular oral check-ups and cleaning and maintaining good oral hygiene habits can help prevent the occurrence and development of oral cancer. If oral cancer has already occurred, patients should seek medical treatment promptly to improve the cure and survival rates [8].

## **5. Prevention and Intervention Strategies**

### ***5.1. Stopping Betel Nut Chewing***

Abstaining from betel nut chewing is the most direct and effective way to reduce the impact of betel nut on oral mucosal health. Long-term betel nut consumption increases the risk of developing and developing oral cancer, therefore, quitting betel nut chewing is one of the most important measures to prevent oral cancer and individuals should consciously quit chewing betel nut. Quitting betel nut chewing requires sustained effort and commitment. There are a number of ways to help people quit chewing betel nut, including gradually reducing the amount of betel nut consumed, using chewing gum or candy instead of betel nut, and gradually reducing the number of chews, etc.

### ***5.2. Oral Health Care***

Regular oral check-ups and good oral hygiene habits can help prevent betel nut-induced oral mucosal damage and fibrosis. Firstly, it is very important to maintain good oral hygiene habits. Oral hygiene habits include brushing, rinsing, flossing and so on. These habits can help remove bacteria and food debris from the mouth and reduce the incidence of oral inflammation and infection. For betel nut

chewers, it is especially important to rinse your mouth with water promptly after chewing betel nut to wash off the residue and pigment to reduce irritation and damage to the oral mucosa. Secondly, it is also important to have regular oral check-ups. Betel nut chewers should visit their dentists regularly for oral check-ups to detect and treat oral diseases in a timely manner. A dentist can examine the changes in the oral mucosa and early signs of oral cancer and provide appropriate treatment advice and guidance. In addition, betel nut chewers should also pay attention to self-observation of changes in the oral mucosa. Therefore, if you notice any abnormal changes in the oral mucosa, such as changes in colour, shape and size, or uncomfortable symptoms such as pain and itching, you should consult your oral surgeon promptly and receive the appropriate treatment. Maintaining good oral hygiene habits and regular oral check-ups are one of the important measures to prevent betel nut-induced oral mucosal damage and fibrosis. Betel nut chewers should be aware of these measures and seek prompt medical attention and treatment to protect their oral health [9].

### **5.3. Drug Intervention**

It has been found that certain drugs such as antioxidants, anti-inflammatory drugs and immunomodulators may have a therapeutic effect on betel nut-induced oral mucosal damage and fibrosis. These drugs may prevent the development of oral cancer by reducing the irritation and damage to the oral mucosa caused by betel nut in different ways. Firstly, antioxidants can help reduce the oxidative damage to the oral mucosa caused by betel nut. Betel nut contains high levels of free radicals and reactive oxides, which cause oxidative damage to the oral mucosa and accelerate the fibrosis process of the oral mucosa. Antioxidants can protect the health of the oral mucosa by scavenging free radicals and reactive oxides and reducing oxidative damage to the oral mucosa. Secondly, anti-inflammatory drugs can help reduce the inflammatory response of betel nut on the oral mucosa. Betel nut contains high levels of alkaloids and organic acids, which cause inflammatory and immune responses in the oral mucosa and exacerbate the fibrotic process of the oral mucosa. Anti-inflammatory drugs can reduce the inflammatory response of the oral mucosa by inhibiting the inflammatory and immune response, thus preventing the development of oral cancer. Finally, immunomodulators can help regulate the function of the immune system, thereby reducing the immune response of betel nut on the oral mucosa. Betel nut contains a large amount of antigenic substances and immunostimulatory substances, which can lead to an immune and inflammatory response in the oral mucosa and exacerbate the fibrosis process of the oral mucosa. Immunomodulators can prevent the development of oral cancer by modulating the function of the immune system and reducing the immune response in the oral mucosa. However, the efficacy of these drugs needs further research and validation. At present, there is some controversy and uncertainty about the therapeutic effects of these drugs on betel nut-induced oral mucosal damage and fibrosis. Therefore, betel nut chewers should protect their oral health by maintaining good oral hygiene practices and regular oral check-ups, and use these drugs with caution under the guidance of a doctor.

### **5.4. Dietary Modifications**

Proper dietary modification can help prevent the development of oral cancer. In our diet, we should consume more vegetables and fruits rich in vitamins and minerals, and reduce the intake of foods high in fat, calories and sugar. Firstly, vegetables and fruits are rich in many vitamins and minerals, which are important for the prevention of oral cancer. For example, vitamin A helps maintain the health of the oral mucosa, vitamin C strengthens the immune system, vitamin E reduces the production of free radicals, and minerals such as selenium and zinc also help maintain the health of the oral mucosa. Therefore, we should consume more vegetables and fruits that are rich in these nutrients, such as carrots, broccoli, spinach and citrus fruits. Secondly, food intake high in fat, calories and sugar can increase the body's inflammatory response and oxidative damage, exacerbating the fibrotic process of the oral mucosa and increasing the risk of oral cancer. Therefore, the intake of these foods, such as candy, chocolate, crisps, fried chicken and other high-calorie and high-fat foods, should be reduced. In addition, nutrients such as protein, fat and carbohydrates should be consumed in moderation in the diet to ensure normal metabolism and immune function of the body. A varied diet with a reasonable mix of different types of food is recommended to control food intake and avoid overeating. We should adjust our diet structure appropriately to help prevent the occurrence of oral cancer. We should consume more vegetables and fruits rich in vitamins and minerals, reduce the intake of high-fat, high-calorie and high-sugar foods, and mix all kinds of foods reasonably, so as to ensure the normal metabolism and immune function of the body.

### **5.5. Government Awareness and Education**

Government publicity and education are important tools to prevent and control the problem of betel nut addiction. I will discuss several specific aspects below: Media publicity: The government can use various media publicity means, such as television, radio, newspapers, magazines and the Internet, to inform the public of the dangers of betel nut and ways to quit betel nut. These publicity tools can take various forms, such as news reports, public service announcements and posters, to make the public more aware of the health hazards of betel nut and to steer them away from it. There are also health education activities: the government can organise various health education activities, such as public lectures, seminars and health consultation activities, where the government can impart relevant health knowledge to the public, thereby raising public awareness of the hazards of betel nut and guiding the public away from it; Secondly, publicity by medical institutions: the government can promote the dangers of betel nut and ways to quit betel nut to patients through medical institutions. Doctors can provide relevant health education to patients during the consultation process, so that they can understand the health hazards of betel nut, and provide guidance and support to quit betel nut. Final policy measures: The government can adopt relevant policy measures to restrict the sale and use of betel nut, such as strengthening the regulation and control of betel nut sales and increasing the tax on betel nut. These policy measures can effectively restrict the use and sale of betel nut, thereby reducing public exposure to betel nut and reducing the risk of addiction to betel nut. Government publicity and education are important tools to prevent and control the problem of betel nut addiction. The government can inform the public of the dangers of betel nut and ways to quit betel nut through various means, such as media campaigns, health education activities, publicity by medical institutions, etc. In addition, the government can also adopt relevant policy measures to restrict the sale and use of betel nut, so as to reduce the public's exposure to betel nut and reduce the risk of betel nut addiction.

### **5.6. Seek Help from a Psychiatrist**

People who have been consuming betel nut for a long time may suffer from psychological effects such as anxiety and depression, and these emotional problems may affect the process of quitting betel nut. Therefore, psychological support is very important to help betel nut addicts deal with their emotional problems and increase their willpower to quit. The following are some possible methods of psychological support: Psychologists can provide psychotherapy tailored to the situation of the betel nut addict to help them deal with their emotional problems and increase their willpower to quit betel nut. Psychotherapy can use a variety of techniques such as cognitive behavioural therapy and supportive psychotherapy, which can be personalised to the betel nut addict's situation. Join a social support group such as an addiction support group: Joining a social support group such as an addiction support group can provide emotional support and practical advice to betel nut addicts. In these groups, people with betel nut addiction can share their experiences and views with other members, and provide support and encouragement to each other to better cope with the process of quitting betel nut. Finally do self-psychological support: betel nut addicts can also help themselves with some self-psychological support techniques such as relaxation exercises, breathing exercises, positive thinking, etc. These techniques can help betel nut addicts to relieve tension, adjust their mindset and increase their willpower to quit betel nut. In short, psychological support is very important to help betel nut addicts deal with their emotional problems and strengthen their willpower to quit betel nut. Betel nut addicts can seek help from a psychiatrist, join a social support group such as an addiction support group, or help themselves through self-psychological support techniques.

### **5.7. Medical Help**

Medical help is very important for people who are addicted to betel nut. Medical providers can offer services for the diagnosis and treatment of betel nut addiction, including guidance and support for quitting betel nut, treatment for oral mucosal damage and more. Here are some possible ways to get medical help: Diagnosis and assessment: A doctor can diagnose and assess a betel nut addict to understand the extent of their addiction, their physical condition and their psychological state, and to develop a personalised plan for quitting betel nut. The assessment process can include a variety of methods such as interviews, physical examinations and laboratory tests to ensure that the betel nut addict receives the most appropriate treatment.

## 6. Conclusion

The effects of betel nut on oral mucosal health have attracted widespread attention. This thesis summarises the effects of betel nut on oral injury and oral fibrosis and their potential mechanisms, providing a theoretical basis for further research on the effects of betel nut on oral health and the development of corresponding preventive measures. Future research should continue to focus on the effects of betel nut on oral health and explore more effective prevention and intervention strategies.

## References

- [1] Qiu Ruizhi, Wang Peiyu, Wang Yanru, et al. Effect of betel nut on autophagy in oral mucosa[J]. *Journal of Clinical Dentistry*, 2019, 35(10): 636-639.
- [2] Lin JH, Chen KWR, Tsai YL, et al. Effect of betel nutin on proliferation and apoptosis of oral mucosal epithelial cells[J]. *Journal of Modern Stomatology*, 2021, 35(2): 88-92.
- [3] Supharoek Samang, Weerasuk Bordin, Grudpan Kate, Ponghong Kraingkrai. Green Flow Injection Spectrophotometry with Betel Nut (*Areca catechu* L.) Natural Reagent for Indirect Determination of Ascorbic Acid [J]. *Journal of Analytical Chemistry*, 2022(10):77.
- [4] Franke Adrian A, Li Xingnan, Herzog Thaddeus A, Paulino Yvette C, Badowski Grazyna, Wilkens Lynne R, Lai Jennifer F. Salivary Areca and tobacco alkaloids for bioverification in the Betel Nut Intervention Trial (BENIT).[J]. *Drug testing and analysis*, 2022(1):15.
- [5] Dalisay Francis, Kawabata Yoshito, Buente Wayne, Pokhrel Pallav, Benitez Chantay, Herzog Thaddeus. Social media, peer norms, and betel nut susceptibility and use: Evidence from early adolescents in Guam [J]. *Frontiers in Communication*, 2022:33-36.
- [6] Deng Xia, Wang Qiang, Wang Xiaohui et al. Effects of betel nutrin on proliferation, apoptosis and cell cycle of oral cancer cells[J]. *Chinese Journal of Oral and Maxillofacial Surgery*, 2021, 19(1): 1-5.
- [7] Chen JF, Zhang L, Zhang WM, et al. Effect of betel nutin on the proliferation and cell cycle of human oral mucosa cells[J]. *Chinese Journal of Stomatology*, 2021, 56(5): 409-413.
- [8] Rabinovich O F, Rabinovich I M, Babichenko I I, Syomkin V A, Umarova K V, Shindich O I. [Optimization of the diagnosis of precancerous diseases of the oral mucosa and lips (Manganotti's abrasive precancerous cheilitis and erythroplakia)].[J]. *Stomatologiya*, 2021(2):100.
- [9] Nikoloudaki Georgia, Creber Kendal, Hamilton Douglas W. Wound healing and fibrosis: a contrasting role for periostin in skin and the oral mucosa. [J]. *American journal of physiology. Cell physiology*, 2020(6):318-320.