



Case Report

Effects of mix herbs topical and oral therapies on diabetic foot ulcers: a case report

Haryanto ¹, Tisa Gusmiah, Lestari Makmuriana, Wuriani

¹Nursing Study Program, Department of Medical Surgical Nursing, STIK Muhamadiyah Pontianak, Central Kalimantan, Indonesia

ARTICLE INFORMATION

Received: Desember 02, 2019
Revised: January 07, 2020
Available online: January 17, 2020

KEYWORDS

Diabetic foot ulcer; Topical dressing; Mix herb; Wound healing

CORRESPONDENCE

Phone: +6282149803923
E-mail: haryanto@stikmuhtk.ac.id

ABSTRACT

Background: Indonesia is a tropical country with lots of potential-therapy resource plants, including those for wound care like honey, sea cucumber, black seed and aloe vera. There are other complementaries used by oral such as *Radix Rehmanniae Preparata*, *Panax Ginseng* and *Gypsum Florosum*.

Case presentation: This study case reported the effectiveness of complementary herbs by topical and oral for two patients with diabetic foot given on the wound area.

Conclusion: The result revealed that mix herbs (*radix rhamanniae*, *panax ginseng* and *gypsum florosum*) by oral and topical (natural honey, black seed and sea cucumber) could heal diabetic foot ulcers.

INTRODUCTION

At present, the use of herbs and botanical in medicine have been increasingly widespread. Botanicals and herbs could be used for wound healing, both topically and orally. They always are used as an alternative and complementary therapy. Many studies revealed that botanicals and herbs have therapeutic effects. *Aloe vera* is a medicinal plant well known for its therapeutic benefits and has been used for medical purposes for a long time. The therapeutic effect of *Aloe vera* is caused by its contains, namely vitamins, minerals, enzymes, amino acids, natural sugars. In addition, *Aloe vera* also has bioactive compounds such as antimicrobial, anti-inflammatory, antioxidant, antifungal, and antiseptic¹. Numerous studies revealed that this plant was effective to be topical for burns.

A similar effect is also found in honey. Numerous studies have been proven that it has a therapeutic effect. The liquid has been used in wound care as a topical treatment. It could stimulate tissue regeneration, debriding necrotic, reducing edema, and promoting wound healing^{2,3}. In addition, it has many advantages for wound healing, such as

angiogenesis, decreasing wound area, and re-epithelization on acute and chronic⁴⁻⁶.

Another herb is *Black Seed*. The Black Seed in Arabic has long been used as an alternative therapy. It has antioxidant, antimicrobial, antidiabetic, and antibacterial effects^{7,8}. *Sea cucumber*, or was known as gamat. In vivo studies using animals, it was reported that sea cucumber has antibacterial, angiogenesis, antioxidant, and anti-inflammatory effects⁹.

Other herbs commonly used for traditional Chinese medicinal herb is *Radix Rehmanniae Preparata* (*Shu Dihuang*). *Radix Remanniae Preparata* is produced from the root of *Rehmannia glutinosa Libosch* (*Dihuang*). Its already has known and widely used by some countries such as China, Japan, Korea, and other Asian countries¹⁰. Due to *radix* contains various compounds such as polysaccharides, iridoid glycoside, phenol glycoside ionone, flavonoid, amino acid, inorganic ions, and microelement, so it has an anti-fatigue effect¹¹. Other herbs is *Panax ginseng C. A. Meyer* (*ginseng*). *Panax ginseng* contains 15% water-soluble polysaccharides (WGP). As a result, ginseng has pharmaco-

<https://doi.org/10.30595/medisains.v17i3.6039>

©(2019) by the Medisains Journal. Readers may use this article as long as the work is properly cited, the use is educational and not for profit, and the work is not altered. More information is available at [Attribution-NonCommercial 4.0 International](https://creativecommons.org/licenses/by-nc-sa/4.0/).

logical effects, such as anti-tumor, antioxidant, and hypoglycaemic activities¹²⁻¹⁴. Also, in Chinese traditional medicine, ginseng is used for the development of physical strength, particularly severe fatigue patients¹⁵. The last herb in this report is *Gypsum florosum*. It is always administered with radix for a cough with a high temperature¹⁶. It has anti-hypertensive effect¹⁷ and asthma¹⁸.

The herb could be applied by both topical and oral. However, no clinical trial has been reported combining between topical and oral on diabetic foot ulcer (DFU's). In this study, we will report combining herbs application by topical and oral on two cases, diabetic foot ulcer in the clinical setting.

CASE PRESENTATION

Case 1

Mrs. L, 52 years old, female, government employee, living in Parit. H. Husin 2, Pontianak, is a diabetic patient (blood glucose value: 435 mg/dl), treated at the Kitamura clinic, and reported suffering from necrotic wounds, edema, and infection of the dorsal at the left foot (Figure 1). Based on the medical record, it is found that the physical assessment recorded a good general condition and consciousness. The first wound assessment was on April 2, 2019, DFUS=16, and BWAT=36, and the wound area was 11.09 cm x 13.09 cm. The treatment was with saline solution at 0.9%, followed by the application of the modern dressing (duoderm gel, alginate, cutisorb, and allevin) and complimentary dressing (natural honey, black seed, and sea cucumber). The wound was debrided based on the wound bed. Changing the dressing was done on a regular daily basis. Besides, the care was given for the affected area by the systematic use of medication to control glucose levels, antibiotic, and herbal via oral. Interestingly after 29 weeks, the wound was healed, indicated by DFUS=0 and BWAT=11 (Figure 2).

Case 2

Mrs. W, 58 years old, female, living in Darma putra, Pontianak, is a diabetic type II patient (blood glucose: 424 mg/dl), treated at the Kitamura clinic, and reported to suffer from sleepy, fatigue. He had a wound on the dorsal left foot 3 months ago with slough and necrotic (Figure 3). Based on the medical record, it is reported that the mechanism of injury was traumatic. No data was found about pain, previous amputation, monofilament test, vibration sensation, and temperature perception. The first wound assessment was on June 5, 2019, DFUS=21 and BWAT=34, and the wound area was 24.39 cm x 18.91 cm. The treatment was done using the saline solution at 0.9%, followed by the application of the modern dressing (ribbon gauze, intrasite gel, stomahesive, duoderm gel) and complimentary dressing (natural honey and sea cucumber). The changing of dressing was every 3-day and debrided based on wound

bed condition. Besides, the care was also provided for the affected area, such as the systematic use of medication to control glucose levels, antibiotic, and herbal via oral. After 24 weeks, the wound was healed, showing DFUS=0 and BWAT=12 (Figure 4).

DISCUSSION

The wound infection is caused by the existence of bacteria, particularly on chronic wound¹⁹. There are two factors influencing wound healing namely local factors (oxygenation, infection, foreign body, and venous insufficiency) and systemic factors (age and gender, sex, hormones, stress, ischemia, diseases, uremia, obesity medications, chemotherapy, alcoholism, smoking, and Immunocompromised conditions), and nutrition.

DFUs were a common complication of diabetes. Of 84% of all diabetes done lower-leg amputations¹⁹. There are complex pathophysiology mechanisms in diabetes patients, so influencing the wound healing process of both DFUs and acute wounds. Hypoxia is one of a cause of nonhealing injuries such as DFUs, venous, and pressure ulcer wounds²⁰. This condition may be caused by both insufficient perfusion and insufficient angiogenesis. In addition, it will be increasing the levels of oxygen radical related to an increased inflammatory response by hypoxia²¹.

All patients were controlled by antibiotics. One of the nutrients that the most influencing wound healing was protein. The capillary formation, fibroblast proliferation, proteoglycan synthesis, collagen synthesis, and wound remodeling will be inhibited related to protein deficiency. Besides, the inadequacy of protein also affected the immune system, with resultant decreased leukocyte phagocytosis and increased susceptibility to infection²². The major protein component of connective tissue was collagen. Compounds of collagen are composed essentially of glycine, proline, and hydroxyproline. Lysine, proline, and co-factors such as ferrous iron and vitamin C are required to collagen synthesis. Deficiencies of these co-factors will resulted delayed wound healing²³.

In this present study, we evaluated the wound healing process using BWAT and DFUS. Patient 1 (total score of DFUS 16 to 0 and BWAT 36 to 11) and Patient 2 (total score of DFUS 21 to 0 and BWAT 34 to 12). The result has shown that wounds were significantly healed. This healing is reached because all patients used complementary therapies, both topical (natural honey, black seed, and sea cucumber) and oral. Many studies show that natural honey does accelerate wound healing. This could be due to the varying pH of different types of honey, unusual composition, and floral source of the honey used²⁴.



Figure 1. Initial wound



Figure 2. After 29 week



Figure 3. Initial wound



Figure 4. After 24 week

Using the mixture of black seed oil, honey on wound healing could enhance the healing in the experiment in rabbit²⁵, but clinically no data was recorded. Meanwhile, using sea cucumber on wound healing for two weeks has been proven to reduce wound area, but no data was recorded about the healing time²⁶.

In this case, all patients got complementary therapy (a combination of *radix rehmanniae*, *panax ginseng*, and *gypsum florosum*) via oral. *Panax ginseng* promotes vitality and stamina and maintains general well being. It contains constituents (the ginsenosides) that are structurally related to the corticosteroids. Corticosteroids have main functions in regulating the immune and endocrine systems that are mediated by the HPA axis²⁷. The actions of *radix rhamanniae* root include regulating blood sugar, blood pressure and immune functions²⁸ and the last *gypsum florosum* has actions of clearing the excess heat and good for stomach.²⁹

CONCLUSIONS AND RECOMMENDATION

In this study case, the wound areas have been healed well, and they are much smaller than the initial condition. The wound area treated is different from the previous studies. However, this study has revealed that mix herbs both by

oral (*radix rehmanniae*, *panax ginseng* and *gypsum florosum*) and topical (natural honey, black seed, and sea cucumber) could successfully heal diabetic foot ulcers. Surely, this study has certain limitations and other studies involving larger samples and a more extended period to make sure the effectiveness of the treatment for diabetic ulcers.

REFERENCES

1. Sahu PK, Giri DD, Singh R, Pandey P, Gupta S, Shrivastava AK. Therapeutic and medicinal uses of aloe vera: a review. *Pharmacology & Pharmacy*. 2013;4(08):599-610. doi.org/10.4236/pp.2013.48086.
2. Molan P. Not all honeys as the same for wound healing. *Eur Tissue Rep Soc Bull*. 2002;9(1):5-6.
3. Van Der Weyden EA. The use of honey for treatment of two patients with pressure ulcers. *Br J Community Nurs*. 2013;8(12):S14-S20. doi.org/10.12968/bjcn.2003.8.Sup6.12553.
4. Haryanto, Sugama J, Nakatani T and Urai T. Effectiveness of Indonesia Honey toward acceleration of wound healing: An experimental study in mice (Pilot Study). *Proceeding EPUAP Annual Conference*. Birmingham United Kingdom: 1-3 September 2010.
5. Haryanto, Sugama J, Nakatani T. Effectiveness of Indonesian Honey on the Acceleration of Cutaneous

- Wound Healing: An Experimental Study in Mice. *WOUNDS*. 2012;24(4):110-119.
6. Nakajima Y, Mukai K, Nasruddin, Komatsu E, Iuchi T, Kitayama Y, Sugama J and Nakatani T. Evaluation of the effect of honey on acute phase deep burn wounds. *Evidence-Based Complementary and Alternative Medicine*. 2013;1-20. doi.org/10.1155/2013/784959
 7. Al-Mutheffer EH. The effect of local application of blackseed (nigella sativa) oil on wound healing in rabbits. *Al-anbar. J, Vet. Sci*. 2010;3(1): 90-97.
 8. Khalaf RA. Effect of Locally Applied Black-seed Oil and Honey Mixture on Wound Healing. *Inter J Scien & Tech Res*. 2013;2(12):31-34.
 9. Zohdi RM, Zakaria ZA, Yusof N, Mustafa NM, Abdullah MN. Sea cucumber (*Stichopus Hermanii*) based hydrogel to treat burn wounds in rats. *J of Biomed Mat Res*. 2011; 98(1):30-37. doi:10.1002/jbm.b.31828.
 10. Tan W, Yu K, Liu Y, Ouyang M, Yan M, Luo R and Zhao X. Anti-fatigue activity of polysaccharides extract from *Radix Rehmanniae Preparata*. *Inter J Bio Macro*. 2012 ;50(1):59-62. doi: 10.1016/j.ijbiomac.2011.09.019.
 11. Tomoda M, Kato S, Onuma M. Water-soluble constituents of *rehmanniae radix*. I. Carbohydrates and acids of *Rehmannia glutinosa* f. *hueichingensis*. *Chem Pharm Bull*. 1971;19(11):1455-1460.
 12. Konno C, Murakami M, Oshima Y, Hikino H. Isolation and hypoglycemic activity of panaxans Q, R, S, T and U, glycans of *Panax ginseng* roots. *Journal of Ethnopharmacology*. 1985;14(1):69-74. doi.org/10.1016/0378-8741(85)90030-3.
 13. Shin HJ, Kim YS, Kwak YS, Song YB, Kim YS, Park JD. 2004. Enhancement of antitumor effects of paclitaxel (taxol) in combination with red ginseng acidic polysaccharide (RGAP). *Planta Medica*. 2004;70(11):1033-1038.
 14. Luo DH, Fang BS. Structural identification of ginseng polysaccharides and testing of their antioxidant activities. *Carbohydrate Polymers*. 2008;72(3):376-381. doi: 10.1016/j.carbpol.2007.09.006.
 15. Saito H, Yoshida Y, Takagi K. Effect of *Panax ginseng* root on exhaustive exercise in mice. *The Japanese Journal of Pharmacology*. 1974;24(1):119-127. doi: 10.1254/jjp.24.119
 16. Wang J, Wang P, Xiong X. Current Situation and Re-understanding of Syndrome and Formula Syndrome in Chinese Medicine. *Intern Med*. 2012;2(3):1-5. doi:10.4172/2165-8048.1000113
 17. Wang J, Xiong XJ. Control strategy on hypertension in Chinese medicine. *Evidence-based Complementary and Alternative Medicine*. 2012, Article ID 284847, 6 pages. Volume 2012. doi.org/10.1155/2012/284847.
 18. Wang X, Zhang H, Chen L, Shan L. Liquorice, a unique "guide drug" of traditional Chinese medicine: A review of its role in drug interactions. *J Ethnopharmacol*. 2013;150(3):781-790. doi: 10.1016/j.jep.2013.09.055
 19. Vincent AM, Russell JW, Low P, Feldman EL. Oxidative stress in the pathogenesis of diabetic neuropathy. *Endocr Rev*. 2004;25(4):612-628.
 20. Brem H, Tomic-Canic M. Cellular and molecular basis of wound healing in diabetes *J Clin Invest*. 2007;117(5):1219-1222. Doi: 10.1172/JCI32169
 21. Tandara AA, Mustoe TA. Oxygen in wound healing more than a nutrient. *World J Surg*. 2004;28(3):294-300. doi: 10.1007/s00268-003-7400-2
 22. Mathieu D, Linke JC, Wattel F. Non-healing wounds. In: Handbook on hyperbaric medicine, Mathieu DE, editor. Netherlands: Springer. 2006;pp. 401-427.
 23. Gogia PP. Physiology of wound healing. In: Clinical wound management. Gogia PP, editor. Thorofare, NJ: Slack Incorporated. 1995;pp 8-12.
 24. Campos A.C, Groth A.K, Branco A.B. Assessment and nutritional aspects of wound healing. *Curr Opin Clin Nutr Metab Care*. 2008;11(3):281-288. doi: 10.1097/MCO.0b013e3282fbd35a.
 25. Hashim M, et al. The efficacy and safety of natural honey on the healing of foot ulcers: a case series. *WOUNDS*. 2015;27(4):103-114.
 26. Budiharto I, Pranggono, Emmy H, Kurniawan T, Haryanto. Effect of sea cucumber extract on diabetic foot ulcers. *World Council of Enterostomal Therapists Journal*. 2016;36(1):34-39.
 27. Smith, Sean M, and Wylie W Vale. The role of the hypothalamic-pituitary-adrenal axis in neuroendocrine responses to stress. *Dialogues in clinical neuroscience*. 2006; 8(4):383-395.
 28. Panossian A, Wagner H. Stimulating effect of adaptogens: an overview with particular reference to their efficacy following single dose administration. *Phytother Res*. 2005;19(10):819-838. doi: 10.1002/ptr.1751.
 29. Jin Y, Huang H, Li-Jiang Z. Introduction to Chinese Medica. *World Century Compedium to TCM*.2013; 3. World Scientific Publishing.