APPLICATION OF CADEXOMER IODINE POWDER TO MINIMIZE BIOFILMS IN DIABETIC FOOT ULCER PATIENTS (DFU) AT GRIYA AFIAT WOUND CARE CLINIC: CASE REPORT

Moh Gifari S¹, Uyunul Jannah¹, Takdir Tahir^{1*)} & Saldy Yusuf¹

¹School of Nursing, Fakulty Of Nursing, Hasanuddin Universit. Jl. Perintis Kemerdekaan Kampus Tamalanrea Makassar - Indonesia

Abstract

Every wound in diabetics has the potential to experience a setback in healing into a chronic wound so that the wound also has the potential to form into biofilm. One of the modern dressings commonly used to suppress the growth of biofilm is Cadexomer Iodine Powder. However, there is still limited research that addresses directly related to the effectivity of Cadexomer Iodine Powder towards wound healing. Therefore, this report aims to evaluate effectiveness Cadexomer Iodine Powder on wound healing with biofilm problems at Griya Afiat Makassar's wound care clinic. The study aimed to identify the effectiveness of Cadexomer Iodine Powder in minimizing biofilms in diabetic foot ulcer (DFU) patients. This study was conducted on one patient who was followed prospectively for 32 days from 10 July to 10 August 2019 with 10 treatments. The dressing used for wound care is Cadexomer Iodine Powder. To evaluate the reduction in the number of biofilms performed by the expert wound when treatment continues. Meanwhile, to evaluate the wound healing process, the modified Bates-Jensen Wound Assessment was used. This report shows a decrease in biofilm from the first treatment to the last treatment with the number of biofilms in the first treatment (+++) to a few (+) in the 10th treatment. The treatment also took a short period of time which is 32 days with a wound condition at the beginning of treatment which is 30% of slough and 70% of granulation experiencing a very swift healing in the 10th treatment to 40% of epithelium and 60% of granulation. Cadexomer Iodine Powder is very effective in the process of wound treatment which can be seen through the reduction of biofilms. In addition, wound healing is much better with shorter treatment time as a proof.

Keywords: Diabetic Foot Ulcer (DFU); Cadexomer Iodine; Modern Dressing.

*) Corresponding author: Email: takdirtahir@unhas.ac.id

1. Introduction

Diabetes mellitus is a lifetime condition that is one of the leading causes of death in the world(Kartika, 2017). In Indonesia itself, the high prevalence of diabetics makes Indonesia one of the top 10countries with highest Diabetes Mellitus (DM) prevalence(Yusuf et al., 2016). The tendency of the prevalence of chronic wounds which is DM in Indonesia has increased from 2007 which was 1.1% to 2.1% in 2013(Kemenkes, 2013). One of complication that often occurs in patients with Diabetes Mellitus is diabetic foot injury (International Diabetes Federation, 2017).

Diabetic Foot Wound (DFW) often encountered and feared because the treatment is often disappointing and end in amputation, even death (Langi, 2013). The results of research conducted at endocrine clinics in several regional hospitals in eastern Indonesia showed that from 249 registered patient, the prevalence of risk factors for diabetic foot injury was found to be 55.4%. Meanwhile the prevalence of diabetic foot injury itself is 12% (Yusuf et al., 2016).

Diabetic ulcerwill cause an infection if not treated properly. Majority of patients come with advanced stage diabetic foot ulcers, because 93% of the lesions have a III-V degree according to Wagner classification, in addition lesions usually happens with a state of advanced infection, because 60% of the lesions are grade 3-4 on aPEDIS scale (Cervantes-García & Salazar-Schettino, 2017). To see the severity of the wound also to predict healing of the wound, we can use a classification system and a wound evaluation instrument (Rasyid, Yusuf, & Tahir, 2018). The results showed that most of the patients who visited the wound treatment clinic were patients with the type of chronic wounds(Risma, Tahir, & Yusuf, 2018).

Every non-healing chronic wound must contain biofilm, the results of a study of 185 chronic wounds state that the prevalence of biofilms in chronic wounds is 78.2% (Malone, Bjarnsholt, et al., 2017). Beside that, Malone et al (2017) in his research entitled "Effect of cadexomer iodine on the microbial load and diversity of chronic non-healing diabetic foot ulcers complicated by biofilm in vivo" stated that based on scanned electron microscopy and fluorescence in situ hybridization confirmed the presence of biofilms in all research samples consisting of 17 people with diabetic foot ulcers.

Wound treatment can be done with traditional and modern dressings. Traditional medicine has antioxidant. anti-inflammatory, antibacterial activity in wound treatment (Nurfiah, Tahir, & Yusuf, 2017). One of traditional medicine that is often used is dragon fruit (Tahir et al., 2017). While for modern dressings that are commonly used include Cadexomer iodine (CI) has supreme benefit compared to various wound dressing of the Pseudomonas aeruginosa biofilm in the ex vivo model (Wanna et al., 2017). Previous studies referred only to the effectiveness of Cadexomer Iodine (CI) which was done with laboratory experiments, but still limited research was based on the direct effectiveness of cadexomer lodine powder on the healing of a patient with Diabetic ulcer.

The description above is the background of researchers to observe the development of diabetes foot injuries in patients with biofilm wound problems and the treatment process carried out in patients with modern wound care using Cadexomer Iodine Powder in Griya Afiat Wound Care Clinic. This research can be the basis for further research in observing wound healing. In this study, a patient with a biofilm wound problem was followed for 32 days.

2. Method

This research is a study case which carried out prospectively at Griya Afiat Wound Care Clinic. The method of collecting samples in this study is Nonprobability Sampling by using an Purposive Sampling technique where researchers select a sample directly by considering the inclusion and exclusion criteria appropriate to the research's purpose. The process of treatment and evaluation of wound healing is carried out from 10 July 2019 to 10 August 2019. The type of dressing used is 3 gram of Cadexomer Iodine Powder0.9%. The dressing was sprinkled evenly onto the wound surface. Treatments were carried out 2 times a week with an interval of 3 days. Initial measurements were made at the 1st treatment in the first week while the final measurements were made at the 10th treatment in the 5th week

To evaluate biofilm in wounds, biofilm characteristics evaluation instruments were used. Evaluations are divided into 5 categories consisting of very many categories (++++), many categories (+++), medium categories (++), few categories (+), and none (-). Determination of the category is done bywound expert when treatment was undergoing. Meanwhile, to evaluate wound treatment, the used instrument was modified *Bates Jensen Wound Assessment Tool* (Bates, 2001).

3. Result and Discussion

Table 1: Client's Demography						
Demography						
Age	: 51 y.o.					
Sex	: Female					
Occupation	: Housewives					
Education	: High School					
Ethnicity	: Buginese					
Religion	: Islam					
Marital Status	: Married					

Table 1 show that the patient Mrs. H51 years old. Latest education is high school with occupation as housewives. A Buginese with Islam as her.

Table 2: Injury histories				
History of injuries				
Onset	Wounds occurred around 3 months ago			
Cause	Client stated that the wound originally			
	caused by the skin of left foot's toe got			
	blistered. It never recovered completely and			
	resulting in the swollen and redden instep			
Prior	Client went to clinics several times but only			
Treatments	given blushes and the wound just cleaned			
	with NaCl at home			
Treatment	Currently client intensively treated in Griya			
received	Afiat Wound Care Clinic. Client was treated			
	with Antimicrobial Silver Ag, Hidrofobic,			
	and Cadexomer Iodine Powder			

Table 2 show that patients had diabetes about 3 years ago. Mrs. H wound is*diabetic foot ulcer* (DFU) which was originally caused by a blistered big toe. The wound did not recover and eventually the instep became swollen and red which later became a new wound on the patient's instep. The patient had taken medication at clinic but it was not continued because the patient said that he only received blushes and had to clean the wound himself at home. Currently, patients choose to take care the wound at the Griya clinic and now undergoing modern wound treatment at Griya Afiat Wound Care Clinic.

Table 3 showed a decrease in biofilm from the first treatment to the end of treatment with the number of biofilms in the first treatment (+++) to a few (+) in the 10th treatment. While for the dominant dressing used is Cadexomer Iodine Powder to overcome the problem of biofilms in wounds.

Table 4 showed the result of thetreatment for 5 weeks, there was a decrease in the amount of slough from 30% in the first week to 0% in the 5th week. While epithelial tissue increased from 0% in the first week, to 40% in the 5th week. The size of the wound also reduced from 5 x 3 x 0.75 cm in the first week to 2.5 x 1 x 0.2 in the 5th week. Available on: http://nursingjurnal.respati.ac.id/index.php/JKRY/index Jurnal Keperawatan Respati Yogyakarta, 7(3), September 2020, 150 - 154

	Technique of Wound							
Treatment	Primery Dressing	Secondary Tertiary Dressing Dressing		Edges Of Wound Management	Characteristics of Biofilm			
1	Antimicrobial Silver + Hidrogel	Low Adherent	Gauze, Hypafix	Zink Zalp	Slough, Maserasi, biofilm (+++)			
2	Antimicrobial Silver + Hidrogel	Low Adherent	Gauze, Hypafix	Zink Zalp	Slough, Maserasi, biofilm (+++)			
3	Antimicrobia Cadexomer Iodine Powder + Hydrofobik	Low Adherent	Gauze, Hypafix	Zink Zalp	Biofilm (+++)			
4	Antimicrobia Cadexomer Iodine Powder + Hydrofobik	Low Adherent	Gauze, Hypafix	Zink Zalp	Biofilm (+++)			
5	Antimicrobia Cadexomer Iodine Powder + Hydrofobik	Low Adherent	Gauze, Hypafix	Zink Zalp	Biofilm (++)			
6	Antimicrobia Cadexomer Iodine Powder + Hydrofobik	Low Adherent	Gauze, Hypafix	Zink Zalp	Biofilm (++)			
7	Antimicrobia Cadexomer Iodine Powder + Hydrofobik	Low Adherent	Gauze, Hypafix	Zink Zalp	Biofilm (++)			
8	Antimicrobia Cadexomer Iodine Powder + Hydrofobik	Low Adherent	Gauze, Hypafix	Zink Zalp	Biofilm (+)			
9	Antimicrobia Cadexomer Iodine Powder + Hydrofobik	Low Adherent	Gauze, Hypafix	Zink Zalp	Biofilm (+)			
10	Antimicrobia Cadexomer Iodine Powder + Hydrofobik	Low Adherent	Gauze, Hypafix	Zink Zalp	Biofilm (+)			

Table 3: Characteristics of Biofilm and The Technique of Wound

Table 4: Evaluation of Wound Treatment

	Care to									
Wound Healing Evaluation	Week 1		Week 2		Week 3		Week 4		Week 5	
	1	2	3	4	5	6	7	8	9	10
Clinical										
Appearance :										
Necrotic	0%	0%	0%	0%	0%	0%	0%	0%	0%	0%
Slough	30%	20%	10%	5%	5%	0%	0%	0%	0%	0%
Granulation	70%	80%	90%	95%	95%	100%	95%	85%	80%	60%
Epithelization	0%	0%	0%	0%	0%	0%	5%	15%	20%	40%
Size	5 x 3 x	4 x 3 x	4 x 3 x	4 x 3 x	4 x 3 x	4x 2.7 x	4x 2.5 x	4 x 2 x	3 x 1.5	2.5 x 1
	0.75 cm	0.75 cm	0.5 cm	0.5 cm	0.5 cm	0.5 cm	0.2 cm	0.2 cm	x 0.2	x 0.2
Eksudat :										
Volume	High	Hight	High	Hight	Medium	Medium	Medium	Medium	Medium	Low
Viscositas	Medium	Medium	Medium	Low						
Odour	Low	Low	Low	Low	Low	-	-	-	-	-
Edges of Wound	Healthy	Healthy	Healthy	Healthy	Healthy	Healthy	Healthy	Healthy	Healthy	Healthy
Paint	0	0	0	0	0	0	0	0	0	0
Infection Status	Local	Local	Local	Local	Local	Local	Local	Local	Local	Local
	Infection	Infection	Infectio							
			n	n	n	n	n	n	n	n
Edema	-	-	-	-	-	-	-	-	-	-

Biofilm plays a crucial role in postponing the healing of wounds (Mori et al., 2019). This report shows that Cadexomer Iodine Powder proven effective to overcome the problem of biofilms in wounds that can be seen through a decrease in biofilms from the first treatment to the last treatment with the number of biofilms in the first treatment (+++) decrased to (+) in the 10th treatment. This is in accordance with the research of Roche et al (2019)which states that the effectiveness of

Cadexomer Iodine has been proven effective towards biofilms in various in vitro models, towards biofilms of Staphylococcus aureus that are resistant to methicillin in rat wounds, and clinically in diabetic foot ulcers which got complicated by biofilms. In addition, the effectiveness of Cadexomer Iodine was also explained in the research of Malone et al (2017)regarding the effect of Iodine Cadexomer on microbial burden in chronic diabetic foot ulcers which got complicated by biofilms which stated off the seventeen samples participating in the research, eleven participants showed a log 10 reduction in microbial load after treatment (range 1-2 log 10) compared to six patients who experienced a reduction of <1 log 10. Reduction in microbial load correlated with reduction of wound proteases before and after treatment (P = 0.03). Matthew Malone et al (2019) also explains in his study that 14 of the 18 (78%) samples with diabetic foot ulcer related complications of chronic biopilm infections achieve a reduced load of microbes with a cadexomer iodine.

This report also shows the swift treatment of diabetic foot wounds which is 32 days with the wound condition at the beginning of 30% of slough treatment and 70% of granulation experiencing very swift treatment in the 10th treatment to 40% of epithelium and 60% of granulation. This is confirmed by the results of research conducted by Gifari (2018) which states that the average length of treatment conducted in Griya Afiat Wound Care Clinicwhich is 62 days with the highest frequency of wound types namely chronic wounds of 86.2%. Yunus (2015) in his study also mentions that the average length of time care for injuries in patients at the etn centre's makassar wound clinic is 1-24 weeks at percentage of 96.5%. The delay in the healing process itself can result from many factors between the age, gender, pattern of life, treatment of wounds, and the presence of disease. Various disease processes can affect a chain of events that are involved in healing wounds and lead to incurable chronic wounds (Han & Ceilley, 2017). Diabetic is one of the factors inhibitory to the healing process. In diabetics, normal development through phases of wound repair is impaired, leading to a state of continuous inflammation and dysfunctional epitel of injury (Dekker et al., 2019).

In addition, in the first and second treatment, combination of Antimicrobial Silver and а Hydrophobics were used as a primary dressing to overcome the problem of sloughing on the patient's wound. Ariyanta (2014) in his research entitled "Preparasi Nanopartikel Silver Dengan Metode Reduksi dan Aplikasinya Sebagai Antibakteri Penyebab Luka Infeksi" observed the application of synthesis of Silver Nanoparticles on wound dressing with the best soaking time for 36 hours. The performance of the soaking results in inhibiting bacterial growth was evaluated through the activity tests on bacteria that cause infection, which are Eschericia coli, Bacillus subtilis, and Staphylococcus *aureus*. Quantitative test results show that by soaking for 36 hours the percentage of bacterial reduction reaches 100%. Indrayati, Koto, & Mulyadi (2018) in his research also mentioned that there are significant differences between hydrophobic (CutimedSorbact®) and Silver (Acticoat TM). Usage of Silver (Acticoat TM) have a better response, evidenced by the high cure rate and the time needed to heal faster than expected.

As for the supporting dressing, low adherent and gauze is used to create a moist ambiance in the wound area and to avoid the gauze to stick to the wound surface. The balanced moisture in sores facilitated the growth of cells and the proliferation of collages in a healthy noncellular matrix (Kartika, 2015). Understanding the concept of healing a moist wound and choosing a proper dressing is a key concept to support the healing process (Handayani, 2016). The dressing is made from knitted synthetic rayon, the top layer is Non-Woven to prevent bacteria from entering and to make sure exudate sealed-in, the surface is not sticky to the wound, and can provide an ideal ambiance for wound treatment (Maryunani, 2015).

4. Conclusion

Cadexomer Iodine Powder is very dependable in treating patients with biofilm problem on wound. Cadexomer Iodine Powder can suppress the growth of biofilms and accelerate the process of wound healing in patients with underlying biofilm wounds

5. References

- Ariyanta, H. A. (2014). I Silver Nanoparticles Preparation by Reduction Method and its Application as Antibacterial for Cause of Wound Infection. Jurnal MKMI, 1, 36–42.
- Bates, B. (2001). Bates-Jensen Wound Assessment Tool. 5–8.
- Cervantes-García, E., & Salazar-Schettino, P. M. (2017). Clinical and surgical characteristics of infected diabetic foot ulcers in a tertiary hospital of Mexico. *Diabetic Foot and Ankle*, 8(1). https://doi.org/10.1080/2000625X.2017.13672
- 10 den Dekker, A., Davis, F. M., Kunkel, S. L., & Gallagher, K. A. (2019). Targeting epigenetic mechanisms in diabetic wound healing. *Translational Research*, 204, 39–50. https://doi.org/10.1016/j.trsl.2018.10.001
- Gifari S, M., Tahir, T., Jafar, N., & Yusuf, S. (2018). Gambaran Karakteristik Luka Dan Perawatannya Di Klinik Perawatan Luka Griya Afiat Makassar. *Skripsi*.
- Han, G., & Ceilley, R. (2017). Chronic Wound Healing: A Review of Current Management and Treatments. *Advances in Therapy*, 34(3), 599–610. https://doi.org/10.1007/s12325-017-0478-y
- Handayani, L. T. (2016). Studi Meta Analisis Perawatan Luka Kaki Diabetes Dengan Modern Dressing. 6(2), 149–159.
- Indrayati, N., Koto, Y., & Mulyadi, B. (2018). Penyembuhan Ulkus Diabetik dengan Aplikasi Antimikrobial Wound Dressing Silver (ActicoatTM). Jurnal Ilmiah Ilmu Keperawatan Indonesia, 8(04), 508–515.

https://doi.org/10.33221/jiiki.v8i04.174

- International Diabetes Federation. (2017). Eighth edition 2017. In *IDF Diabetes Atlas, 8th edition*. https://doi.org/http://dx.doi. org/10.1016/S0140-6736(16)31679-8.
- Kartika, R. W. (2015). Perawatan Luka Kronis dengan Modern Dressing. *CDK-230*, 42(7), 546–550.
- Kartika, R. W. (2017). Pengelolaan gangren kaki Diabetik. Continuing Medical Education, 44(1), 18–22.
- Kemenkes. (2013). *RISET KESEHATAN DASAR*. Jakarta: Badan Penelitian dan Pengembangan Kesehatan.
- Langi, Y. A. (2013). Penatalaksanaan Ulkus Kaki Diabetes Secara Terpadu. *Jurnal Biomedik* (*Jbm*), 3(2), 95–101. https://doi.org/10.35790/jbm.3.2.2011.864
- Malone, M., Bjarnsholt, T., McBain, A. J., James, G. A., Stoodley, P., Leaper, D., ... Wolcott, R. D. (2017). The prevalence of biofilms in chronic wounds: A systematic review and meta-analysis of published data. *Journal of Wound Care*, 26(1), 20–25. https://doi.org/10.12968/jowc.2017.26.1.20
- Malone, M., Johani, K., Jensen, S. O., Gosbell, I. B., Dickson, H. G., McLennan, S., ... Vickery, K. (2017). Effect of cadexomer iodine on the microbial load and diversity of chronic nonhealing diabetic foot ulcers complicated by biofilm in vivo. *Journal of Antimicrobial Chemotherapy*, 72(7), 2093–2101. https://doi.org/10.1093/jac/dkx099
- Malone, M., Schwarzer, S., Radzieta, M., Jeffries, T., Walsh, A., Dickson, H. G., Micali, G., & Jensen, S. O. (2019). Effect on total microbial load and community composition with two vs six-week topical Cadexomer Iodine for treating chronic biofilm infections in diabetic foot ulcers. *International Wound Journal*, 16(6), 1477–1486.

https://doi.org/10.1111/iwj.13219

- Maryunani, A. (2015). *Perawatan Luka Modern* (pp. 160–161). pp. 160–161. Jakarta: In Media.
- Mori, Y., Nakagami, G., Kitamura, A., Minematsu, T., Kinoshita, M., Suga, H., Kurita, M., Hayashi, C., Kawasaki, A., & Sanada, H. (2019). Effectiveness of biofilm-based wound care system on wound healing in chronic

wounds. *Wound Repair and Regeneration*, 27(5), 540–547. https://doi.org/10.1111/wrr.12738

- Nurfiah, Tahir, T., & Yusuf, S. (2017). Aktifitas Zat Aktif Berbasis Tanaman Tradisional Indonesia Dalam Penyembuhan Luka. Jurnal Keperawatan Muhammadiyah, 2(2), 104–107.
- Rasyid, N., Yusuf, S., & Tahir, T. (2018). Study Literatur: Pengkajian Luka Kaki Diabetes. Jurnal Luka Indonesia, 4(2), 123–137.
- Risma, Tahir, T., & Yusuf, S. (2018). Gambaran Karakteristik Luka dan Perawatannya di Ruangan Poliklinik Luka di RS Dr. Wahidin Sudirohusodo Makassar. Jurnal Luka Indonesia, Vol. 4(3)(November), 153–163.
- Roche, E. D., Woodmansey, E. J., Yang, Q., Gibson, D. J., Zhang, H., & Schultz, G. S. (2019).
 Cadexomer iodine effectively reduces bacterial biofilm in porcine wounds ex vivo and in vivo. *International Wound Journal*, *16*(3), 674–683. https://doi.org/10.1111/iwj.13080
- Tahir, T., Bakrib, S., Patellongic, I., Amand, M., Miskade, U. A., Maryunisf, M., ... Hasriyani. (2017). Evaluation of Topical Red Dragon Fruit Extract Effect (Hylocereus Polyrhizus) on Tissue Granulation and Epithelialization in Diabetes Mellitus (DM) and Non-DM Wistar Rats: Pre Eliminary Study. International Journal of Sciences - Basic and Applied Research, 32(1), 309–320.
- Wanna, A. G. B., Noble, J. H., Carlson, M. L., Gifford, H., Dietrich, M. S., Haynes, D. S., & Dawant, B. M. (2017). Cadexomer Iodine provides superior efficacy against bacterial wound biofilms in vitro and in vivo. In *The Laryngoscope*.

https://doi.org/10.1002/acr.22212

Yunus, B. (2015). Faktor-Faktor Yang Mempengaruhi Lama Penyembuhan Luka Pada Pasien Ulkus Etn Centre Makassar Bahri Yunus. *Uin-Alauddin.Ac.Id.* Retrieved from http://repositori.uin-

alauddin.ac.id/1333/1/Bahri Yunus.pdf

Yusuf, S., Okuwa, M., Irwan, M., Rassa, S., Laitung, B., Thalib, A., ... Sugama, J. (2016). Prevalence and Risk Factor of Diabetic Foot Ulcers in a Regional Hospital, Eastern Indonesia. (January), 1–10.