# Compare the Efficacy of 0.2% Topical Glyceryl Trinitrate (GTN) Versus Placebo in Prevention of Wound Infection in Patients Undergoing Milligan-Morgan Hemorrhoidectomy

KHUMAIR ASIF1, MAAZ-UL-HASSAN2, PIR MUNEEB REHMAN3, TAYYABA RASHEED4, LAIBA ZAHID5

### **ABSTRACT**

**Objective:** To examine the efficacy of 0.2% topical glyceryltrinitrate and compare with placebo for prevention of wound infection after Milligan Morgan hemorrhoidectomy.

Study Design: Randomized controlled trial

Place & Duration of Study: Department of Surgery, AkhtarSaeed Medical & Dental College Lahore from 1st July 2019 to 31st December 2019.

**Methods:** One hundred and ten patients of both genders with ages 20 to 65 years were enrolled and divided into two groups (each group contains 55 patients). Group I received of 0.2% topical glyceryltrinitrateointment and Group II received placebo. Postoperative wound infection was examined at 2 weeks postoperatively. Compare the wound infection rate between both groups.

**Results:** There were 68 (61.82%) male patients and 42 (38.18%) were females with mean age 34.62±12.46 years. 60 (54.55%) patients had 4<sup>th</sup> degree hemorrhoidal disease and 50 (45.45%) patients had 3<sup>rd</sup> degree disease. In group I 8(14.55%) patients had developed wound infection while in group II 30 (54.55%) patients had developed wound infection, a significant difference was observed between both groups with p-value <0.05.

Conclusion: It is concluded that 0.2% glyceryl trinitrate is better and safe treatment modality for the prevention of wound infection.

Keywords: 0.2% Glyceryl Trinitrate, Milligan Morgan Hemorrhoidectomy, Wound Infection

## INTRODUCTION

It is estimated that around 10 million people suffer from haemorrhoids in the US. This is a prevalence of around 4.4% with nearly 37 haemorrhoidectomies per 100,000 people per year being performed<sup>1</sup>. Post-operative pain is a big problem, with studies showing that patients require 4 to 16 days to return to normal activity. There are many surgical procedures such as the open haemorrhoidectomy of Milligan-Morgan or Ferguson techniques. However there are other procedures which are also carried out with the aim of having low recurrence, minimal post-operative pain and early return to work. A recent systematic review by Burch comparing "procedure for prolapse haemorrhoids" (PPH) with conventional haemorrhoidectomy showed that PPH was associated with less pain in the immediate postoperative period, but a higher rate of recurrence.<sup>2</sup> The relatively new technique of transanal hemorrhoidal dearterialization has been shown to cause low postoperative pain and have similar results to PPH3.

A controversial belief is that post-operative pain may also be due to poor and delayed wound healing which leads to epithelial denudation. There are studies that have shown agents which improve wound healing also improve post-operative pain<sup>4,5</sup>. Oral and topical metronidazole have been shown to reduce postoperative pain by promoting wound healing<sup>6,7</sup>. The amount of pain experienced is also

Received on 11-01-2020 Accepted on 30-05-2020

dependent on a number of other factors such as surgical technique, post-operative analgesia, use of stools softeners, adequate education and subjective pain threshold. Recent evidence suggests that IAS is innervated by neurons that release nitric oxide (NO). Stimulation of these nerves results in the release of NO which then cause relaxation of the IAS12 by relaxation of smooth muscle. Exogenous GTN ointment is an NO donor which relaxes the IAS and thus reduce pain<sup>8,9</sup>. There are other forms of nitrates such as nitroderm bands used in posthaemorrhoidectomy to reduce IAS spasm, however the ointment is the most commonly available and used form. The reduction of IAS spasm and reduced pressure will increase anodermal blood flow and therefore improve wound healing. This is believed to be the reason behind improved wound healing in acute anal fissure with GTN ointment<sup>10,11</sup>. Even though there is no evidence, some believe that GTN may work in a similar fashion and aid wound healing after haemorrhoidectomy. The present study was conducted aimed to examine the efficacy of 0.2% GTN and compare with placebo in prevention of postoperative wound infection after hemorrhoidectomy.

### MATERIAL AND METHODS

This randomized controlled trial was conducted at Department of Surgery, Akhtar Saeed Medical & Dental College Lahore from 1<sup>st</sup> July 2019 to 31<sup>st</sup> December 2019. A total of 110 patients of both genders with ages 20 to 65 years were enrolled. Patients detailed demographics including age, sex, and body mass index were recorded.

Associate Professor of Surgery, Akhtar Saeed Medical & Dental College Lahore

<sup>&</sup>lt;sup>2</sup>Assistant Professor of General Surgery Shalamar Medical & Dental College Lahore

<sup>&</sup>lt;sup>3</sup>Senior Registrar of Surgery, Surgical Unit 2, Jinnah Hospital, Lahore

<sup>&</sup>lt;sup>4</sup>4<sup>th</sup> Year MBBS Student, GMMMC Sukkur

<sup>&</sup>lt;sup>5</sup>Lecturer, B.Sc. (Hons)Operation Theater Technology, Department of Health Professional Technologist.(DHPT), University of Lahore Correspondence to: Dr. KhumairAsif, Email: khumairasif @gmail.com, Cell: 0323-4314118

Patients with emergency hemorrhoidectomy, diabetic patients, patients with renal failure and non-compliance patients were excluded. All the patients were received elective Milligan-Morgan hemorrhoidectomy under general anesthesia. Patients were divided into two groups I and II, each group contains 55 patients. Group I advised to use of 0.2% topical glyceryl trinitrate ointment thrice a day and Group II advised to use placebo thrice a day for 12 days after surgery. Postoperative wound infection was examined at 2 weeks postoperatively. Compare the wound infection rate between both groups. All the data was analyzed by SPSS 24. Chi-square test was done to compare the frequency of wound infection between both groups. P-value <0.05 was taken as significant.

# **RESULTS**

There were 68 (61.82%) male patients and 42(38.18%) were females with mean age 34.62±12.46 years. Mean BMI of patients was 24.52±2.86 kg/m². 60(54.55%) patients had 4<sup>th</sup> degree hemorrhoidal disease and 50(45.45%) patients had 3<sup>rd</sup> degree disease (Table 1). At postoperative follow-up, In group I 8(14.55%) patients had developed wound infection while in group II 30(54.55%) patients had developed wound infection, a significant difference was observed between both groups with p-value <0.05 (Table 2).

Table 1: Demographics of all the patients

Variable	No.	%		
Age (years)	34.62±12.46			
BMI (Kg/m <sup>2</sup> )	24.52±2.86			
Gender				
Male	68	61.82		
Female	42	38.18		
Hemorrhoid degree				
3 <sup>rd</sup>	60	54.55		
4th	50	45.45		

Table 2: Comparison of wound infection between both groups

Wound infection	Group I	Group II	P value
Yes	8	30	
	(14.55%	(54.55%)	0.001
No	47	25	0.001
	(85.45%)	(45.45%)	

## **DISCUSSION**

Hemorrhoidectomy is one of the most frequently performing surgical procedures in all over the world. Post-operative complications such as wound infection are the commonly found complication and associated with poor quality of life and adverse outcomes. Many of modalities have been applied for the prevention of wound infection, here we performed a study with aimed to compare the efficacy of 0.2% GTN with placebo in patients underwent Milligan-Morgan hemorrhoidectomy. In this regard 110 patients were enrolled. Majority of patients 61.82% were male followed by female 38.18% and mean age of patients was 34.62±12.46 years. These results showed similarity to many of previous studies in which male patients population was high55% to 70% as compared to females and average age of patients was 35 years. These

In present study we found that 60 (54.55%) patients had 4th degree hemorrhoidal disease and 50 (45.45%) patients had 3<sup>rd</sup> degree disease. A study conducted by Hyderet al<sup>15</sup> reported that 59.67% patients had 3<sup>rd</sup> degree and 40.32% had 4th degree hemorrhodal disease. We found that at postoperative follow-up, among 55 patients treated with 0.2% GTN 8 (14.55%) patients had developed wound infection while in group II (placebo) 30 (54.55%) patients had developed wound infection out of 55 patients, a significant difference was observed between both groups with p-value <0.05. These results were similar to the study by Hyder et al. 15 Another study conducted by Khan et al 16 reported that combination of 0.2% GTN and 2% lignocaine showed better pain relief resulting in less use of oral analgesics and faster healing of the wound as compared to placebo with p-value <0.05.

A study conducted by Hwang et al<sup>17</sup> regarding effectiveness of 0.2% GTN for wound healing after hemorrhoidectomy and they demonstrated that 0.2% GTN had higher effectiveness as wound healing rate was 74.5% as compared to placebo 42%. Soltany et al<sup>18</sup> reported that topical application of 0.2% GTN ointment can reduce postoperative pain and improve wound healing after open hemorrhoidectomy.

Some other previous studies showed similarity to our study findings in which 0.2% GTN reported safe and effective for postoperative wound healing as compared to placebo with p-value <0.05<sup>19,20</sup>.

## CONCLUSION

Postoperative wound infection is the frequently found complication after surgical interventions and can lead to adverse outcomes. We concluded that 0.2% glyceryl trinitrate is better and safe treatment modality for the prevention of wound infection.

# **REFERENCES**

- Johanson JF, Sonnenberg A. The prevalence of hemorrhoids and chronic constipation. An epidemiologic study. Gastroenterology 1990; 98(2): 380e6.
- Burch J, Epstein D, Sari AB, Weatherly H, Jayne D, Fox D, et al. Stapled haemorrhoidopexy for the treatment of haemorrhoids: a systematic review. Colorectal Dis 2009;11(3):233e43.
- Dal Monte PP, Tagariello C, Sarago M, Giordano P, Shafi A, Cudazzo E, et al. Transanalhaemorrhoidaldearterialisation: nonexcisional surgery for the treatment of haemorrhoidal disease. Tech Coloproctol 2007; 11(4):333e8.
  Nicholson TJ, Armstrong D. Topical metronidazole (10
- Nicholson TJ, Armstrong D. Topical metronidazole (10 percent) decreases posthemorrhoidectomy pain and improves healing. Dis Colon Rectum 2004; 47(5):711e6.
- Rattan S, Chakder S. Role of nitric oxide as a mediator of internal anal sphincter relaxation. Am J Physiol 1992; 262(1 Pt 1):G107e12.
- Guillemot F, Leroi H, Lone YC, Rousseau CG, Lamblin MD, Cortot A. Action of in situ nitroglycerin on upper anal canal pressure of patients with terminal constipation: a pilot study. Dis Colon Rectum 1993;36(4):372e6.
- Loder PB, Kamm MA, Nicholls RJ, Phillips RK. 'Reversible chemical sphincterotomy' by local application of glyceryltrinitrate. Br J Surg 1994; 81(9):1386e9.
- Davis BR, Lee-Kong SA, Migaly J, Feingold DL, Steele SR. The American Society of Colon and Rectal Surgeons clinical

- practice guidelines for the management of hemorrhoids. Dis Colon Rectum 2018; 61(3):284–92.
- Ala S, Saeedi M, Eshghi F, Mirzabeygi P. Topical metronidazole can reduce pain after surgery and pain on defecation in postoperative hemorrhoidectomy. Dis Colon Rectum 2008; 51(2):235e8.
- Rudd RA, Aleshire N, Zibbell JE, Matthew Gladden R. Increases in drug and opioid overdose deaths - United States, 2000–2014. Am J Transplant 2015; 16(4):1323-7
- Ratnasingham K, Uzzaman M, Andreani SM, Light D, Patel B. Meta-analysis of the use of glyceryltrinitrate ointment after haemorrhoidectomy as an analgesic and in promoting wound healing. Int J Surg 2010;8(8):606-11.
- Peery AF, Crockett SD, Barritt AS, Dellon ES, Eluri S, Gangarosa LM, et al. Burden of gastrointestinal, liver, and pancreatic diseases in the United States. Gastroenterology 2015; 149(7): 1731–41.
- Yeo D, Tan KY. Hemorrhoidectomy making sense of the surgical options. World J Gastroenterol 2014; 20(45): 16976–83.
- Similis C, Thoukididou SN, Slesser AA, Rasheed S, Tan E, Tekkis PP> Systematic review and network meta-analysis comparing clinical outcomes and effectiveness of surgical treatments for haemorrhoids. Br J Surg 2015; 102(13): 1603–18.

- Hyder Z, Ghansham, Shah SH, Zohaibullah SO. To compare the frequency of healed wound in patients with hemorrhoids after Milligan-Morgan haemorrhoidectomy given 0.2% topical glyceryltrinitrate (GTN) versus placebo: Pak J Surg 2016; 32(4):234-8.
- 16. Khan KI, Waqas A, Akmal M, Mahmood S, Iqbal A. Efficacy of combination of 0.2% GTN and lignocaine ointments in wound healing and pain relief after Milligan Morgan hemorrhoidectomy--a comparison with lignocaine and 0.2% GTN ointments separately. Int J Surg 2014;12(4):329-33.
- Hwang DY, Yoon SG, Kim HS, Lee JK, Kim KY. Effect of 0.2 percent lyceryltrinitrate ointment on wound healing after a hemorrhoidectomy: results of a randomized, prospective, double-blind, placebo-controlled trial. Dis Colon Rectum 2003; 46(7): 950-4
- Soltany S, Toussy JA, Far MF. The effect of 0.2% glyceryltrinitrate ointment on pain and wound healing after hemorrhoidectomy. Surg J 2009; 4: 8-12.
- Liu JW, Lin CC, Kiu KT, Wang CY, Tam KW. Effect of glyceryltrinitrate ointment on pain control after hemorrhoidectomy: a meta-analysis of randomized controlled trials. World J Surg 2016; 40(1):215–24