COMPARATIVE STUDY OF GLYCERYL TRINITRATE OINTMENT VERSUS LATERAL INTERNAL SPHINCTEROTOMY IN MANAGEMENT OF CHRONIC ANAL FISSURE.

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ABSTRACT... Objectives: To compare efficacy of topical 0.2% glyceryl trinitrate ointment with lateral internal sphincterotomy in treatment of patients with chronic anal fissure. Study Design: Randomized Control Trial. Setting: Department of General Surgery, DHQ Teaching Hospital, Sahiwal. Period: From 20 September 2016 to 31 December 2017. Material & Methods: A total number of 60 patients included in this study and divided into two groups GTN and LIS group through computer generated randomization method. Three main outcome variables were assessed in this study; pain relief, healing of fissure and recurrence. SPSS version was used to analyze data. P value ≤ 0.05 was considered as significant. Results: A total number of 60 patients included in this study. Relief of pain GTN and LIS was (95%) and (86.7%) respectively. Fissure healing GTN and LIS was noted as (88.3%) and (83.3%) respectively. Recurrence was not found in any patient of the present study. Relief of pain was statistically significant. Conclusion: Topical application of GTN ointment for relief of pain is safe and effective method for relief of chronic anal fissure but pain relief is slow. On other hand lateral internal sphincterotomy is first line treatment and treatment of choice for chronic anal fissure when performed by an experienced surgeon.

Key words: Chronic Anal Fissure, Glyceryl Trinitrate, Lateral Internal Sphincterotomy, Pain Relief.

INTRODUCTION

Term anal fissure rendered to the painful condition due to mucosal tear of distal anal canal extending below the dentate line and ends at anal verge.¹,² Most common cause of anal fissure is trauma to anus of anal canal which affects normal bowel movements. Other causes include hard stool or recurrent episodes of diarrhea.³ Insertion of anima tip, rectal thermameter and ultrasonic probe may cause anal trauma. Anal fissures diagnosed in all age groups but young healthy adults diagnosed most likely without gender difference.⁴

Common symptoms of anal fissures include pain, per rectal bleeding, constipation and mucosal discharge.⁵ Predominantly fissures found in midline among them 90 percent found posterior and 10 percent interiorly.⁶ A fissure more than 6 weeks old with indurated margins on digital rectal examination, exposure of horizontal fibers, and fibrosis at basis and sentinel tag labeled as chronic anal fissure.⁷

Fissure also associated with anal spasm and stenosis of internal anal sphincter which reduced the blood flow and delayed the healing process.⁸ In 1951 Eisenhammer introduced internal sphincterotomy and later on Parks described lateral open spenecterotomy in 1967 for the management of anal fissures.⁹ After sometime closed spinecterotomy was also described. In is also a useful agent for relaxation of internal sphincter muscles which allows fast healing by increase of blood supply of this region.¹⁰
area.

METHODOLOGY
This randomized control trial was conducted in the department of general surgery, DHQ Teaching Hospital, Sahiwal from 20 September 2016 to 31 December 2017. Study was started after ethical approval from hospital ethical board and informed consent from patients. Non probability consecutive sampling technique was used. Total number 60 patients were included in the study after calculations. Sample size was calculated from WHO sample size calculator. Adult patients of age 20 years to 40 years diagnosed as anal fissures of more than 6 weeks were included in the study. Patients taking any kind of medicine for illness like ischemic heart disease, any immunity deficiency (human immune deficiency virus), pregnant women, anal fissures of bowel disease, Crohn’s disease ulcerative colitis and tuberculosis were excluded from the study. Patients were divided into two equal groups with computer generated randomization method, 30 patients in each group. Group GTN and group LIS, GTN group was treated with GTN 2% ointment for six weeks and LIS group treated surgically with lateral internal sphincterotomy method.

Pain was assessed with visual analogue scale score consist of 10 numbers 0 to 10. Zero score means no pain and score 10 represents severe pain. After start of treatment patients were asked to intake fibers diet and followed up at every 2 weeks initially and then after 3rd week till six weeks. At every visit patients were examined for fissure healing, pain relief and any type of complications. If fissure was healed patient asked to stop GTN application and continue fibrous diet. To see recurrence patients were followed up till 3 months after complete healing. In GTN group any patient who was not treated completely offered to avail LIS mode of treatment.

Collected data was entered in a computer software SPSS version 24 and analyzed for mean SD of numerical data like age and VAS score frequency percentages for categorical data like complete healing of anal fissures and recurrence rate. Chi square test was applied to see association of outcome variable. P value ≤ 0.05 was considered as significant.

RESULTS
A total number of n=60patients were included in this study, both gender. Gender distribution revealed as n=32 (53.3%) male and n=28 (46.7%) females, (Figure-1). The mean age was 26.42±3.80 years. Relief of pain GTN and LIS was n=57 (95%) and n=52 (86.7%) respectively. Fissure healing GTN and LIS was noted as n=53 (88.3%) and n=50 (83.3%) respectively. No recurrence was found in the present study. Headache was reported in n=20 (33.3%) patients of GTN group and only n=3 (5%) reported in LIS group. Relief of pain (p=0.002) and headache (p=0.000) were statistically significant. (Table-I).

| Author                      | Year | No. | Duration | GTN   | LIS   | P-Value |
|-----------------------------|------|-----|----------|-------|-------|---------|
| Relief of pain              | 2018 | 60  | 6 weeks  | (95%) | (86.7%) | p=0.002 |
| Fissure healing             | 2018 | 60  | 6 weeks  | (88.3%) | (83.3%) | p=0.321 |
| Recurrence                  | 2018 | 60  | 24 months | 0     | 0     | p=0     |
| Complication, headache      | 2018 | 60  | 24 months | n=20 (33.3%) | n=3 (5%) | p=0.000 |

Table-I. Comparison of outcome variables in both groups

![Figure-1](image-url)
DISCUSSION
Management of 6 weeks older anal fissures may include conservative and surgical interventions. GTN and LIS are two effective methods for the treatment of anal fissures in our study we compare both groups with respect pain relief, fissure healing and recurrence. Among these outcome variables LIS found to be more effective with respect to complete healing, for relief of symptoms and temporary relief GTN is effective. In a study conducted by Siddiqui et al\textsuperscript{11} reported that 84.85% of patients after internal sphincterotomy have complete healing and 35.84% of patients have complete healing after GTN ointment application. This shows supremacy of LIS group. P values was <0.001 a significant value. Complete healing was assessed at six week of follow up duration.

In another study conducted by Mishra et al\textsuperscript{12} reported that after treatment of six week GTN application 18 patients out of 20 patients were healed and after LIS 17 out of 20 patients were healed. Which shows better outcome in GTN group as compare to LIS group? This study is also comparable with our study. Only side effect reported in this was headache 25% patients’ complaint about mild headache in GTN group and 1 patient in LIS group was found with hematoma. A similar study was conducted by Bansal et al\textsuperscript{13} reported GTN is safe and effective mod of treatment and good alternative of all kinds of anal fissures treatment. But in case of GTN failure LIS can be used as treatment option. In this study 72% of patients treated with GTN healed fissures completely whereas in group LIS all patients 100% healed completely at six week of treatment.

In another study conducted by Leo et al\textsuperscript{14} complete healing 53.6% in GTN group and 86.7% in LIS group with P value 0.004 a significant value. While in our study complete healing was 88.3% and 83.3% respectively and P value was 0.321 also a non significant value. Study of Leo et al\textsuperscript{14} also comparable with our study. He reported relief of pain in 86.6% of patients and 96.6% of patients respectively. In our study relief of pain was 95% and 86.7% respectively.

Another study conducted by Muhammad et al\textsuperscript{15} reported relief of pain 56.66% and 96% in GTN and LIS group respectively whereas healing of complete fissures was reported 50% and 96% in GTN and LIS group respectively. This study is also comparable with our study with respect of all three outcome variables. Third variable that was observed in our study was recurrence.

Recurrence of anal fissures was not found in any patient of our study in both groups. A similar study was conducted by Oettle GJ et al\textsuperscript{16} and reported similar finding that recurrence was not observed in any patient in both GTN and LIS groups. This study is comparable with our study. Another study was conducted by Ellaban et al\textsuperscript{17} in 2010 and reported 2.5% recurrence in GTN group and no recurrence were reported in LIS group. This study is also comparable with our study. Recurrence was not found in study conducted by Leo et al\textsuperscript{14} in 2011. In study of Bansal et al\textsuperscript{13} which was conducted in 2015 reported 11% recurrence in GTN group and 8% in LIS group these findings also shows superiority of LIS.

Many authors who conducted studies on this topic reported similar findings about relief of pain, healing rate and recurrence rate and also reported that LIS is a treatment of choice when performed by some experienced and skilled surgeon.\textsuperscript{18,19,20}

CONCLUSION
Topical application of GTN ointment for relief of pain is safe and effective method for relief of chronic anal fissure but pain relief is slow. On other hand lateral internal sphincterotomy is first line treatment and treatment of choice of chronic anal fissure when performed by an experienced surgeon.

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| Sr. # | Author(s) Full Name       | Contribution to the paper                                      | Author(s) Signature |
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| 1     | Waqas Hussain Qureshi     | Conceived the idea and planned the study, acquisition of data. |                    |
| 2     | Zahid Sattar              | Drafting and writing of the manuscript, acquisition of data.   |                    |
| 3     | Akhtar Mahboob            | Drafting and writing of the manuscript, acquisition of data.   |                    |
| 4     | Aakif Yousaf              | Statistical analysis and interpretation of data.               |                    |
| 5     | Sajid Mukhtar             | Revision of the manuscript & editing of manuscript.            |                    |
| 6     | Waleeja Shamikha          | Revision of the manuscript & editing of manuscript.            |                    |