Original Research Paper



General Surgery

ROLE OF DIOSMIN WITH HESPERIDIN IN TREATMENT OF VARICOSE VEINS AND HAEMORRHOIDS

| Dr. Sheldon George Mathias | Associate Professor, Department Of General Surgery, Father Muller Medical College, Mangalore |
|-------------------------------|--|
| Dr. Vinod Kumar | Associate Professor, Department Of General Surgery, Father Muller Medical College, Mangalore |
| Dr. Rohan D'souza | Associate Professor, Department Of General Surgery, Father Muller Medical College, Mangalore |

ABSTRACT Aims And Objectives: To evaluate the efficacy of diosmin plus hesperidin for the conservative management of hemorrhoids and varicose veins Introduction: The great and small saphenous veins, as well as their tributaries that run between the fascia and the skin, are involved in varicose veins in the lower extremities. Age and family history are key risk factors for varicose veins, which are defined as subcutaneous dilated, convoluted veins bigger than or equal to three millimeters. One of the most prevalent anorectal conditions is hemorrhoids split up into four grade levels. First and second grades are typically treated conservatively. Methodology: Retrospective study involving patients coming to tertiary care hospital for 2 years. Following factors were considered in the statistical analysis: demographic data, including sex, age, outcomes of disease and complications. Haemorrhoidal symptoms (pain, heaviness, bleeding, pruritus and anal discharge) and in the proctoscopic appearance of the haemorrhoids comparing baseline visit. Varicose veins assessed by Venous clinical severity score (VCSS) Results: 52 cases of varicose veins and 66 cases of hemorrhoids were monitored for two years at a tertiary care hospital. The . Out of the 66 patients, 41 had improved symptoms (62%), and 24 (36.4%) needed surgery in the end. In one instance, rectal prolapse occurred as shown in chart 3,4,5.0f the 52 patients with varicose veins, 42 (80.7%) had improved symptoms, while 9 (17.3%) needed surgery in the end. One (1.9%) developed necrotizing fasciitis. Conclusion: Hemorrhoidal symptoms, such as pain, heaviness, bleeding, pruritus, and anal discharge, as well as changes in the hemorrhoids' proctoscopic appearance when comparing the results of the initial visit to the last appointment eight weeks following diosmin hesperidin treatment. After receiving treatment for two months, venous symptoms decreased. Diosmin plus hesperidin for the conservative management of hemorrhoids and varicose veins has great efficacy. Aims -To asses effectiveness of diosmin in combination hesperidin in conservative treatment of varicose vein and haemorrhoids

KEYWORDS:

INTRODUCTION

varicose veins in the lower extremities involve the great and small saphenous veins and their tributaries between the fascia and the skin. Varicose veins are characterized by subcutaneous dilated, tortuous veins greater than or equal to three millimeters, involving the saphenous veins, saphenous tributaries, or non-saphenous superficial leg veins with age and family history considered important risk factors. Hemorrhoids is one of the most common anorectal disorders. Divided in to four grades. Grade one and two treated conservatively. Diosmin in combination with hesperidin has been found more effective than Diosmin alone on venous symptoms [1,2]. It has been found to be quite useful in the treatment of chronic venous insufficiency and hemorrhoids when combined with other flavonoids, particularly hesperidin. Its combination with hesperidin has been proven to be extremely helpful in the treatment of CVI and hemorrhoids, making it a good example of drug synergism. The most common symptoms of CVI include leg ache, sensation of heaviness or tension, nocturnal cramps, sensation of swelling, restless legs, and itching.. It also enhances the skin microcirculation's blood velocity. Daflon is not considered a cure for CVI because it does not address the underlying cause of the condition, it is used to alleviate the symptoms of the disease. Diosmin and Micronized Purified Flavonoid Fraction (MPFF) have been found to affect venous tone, lymphatic drainage, and microcirculation in CVI patients. They improve venous tone by blocking COMT (catechol-Omethyltransferase) from breaking down norepinephrine (noradrenaline) and so prolonging noradrenergic action. They enhance the number of functional lymphatics, lymphatic flow, capillary hematocrit, and red cell velocity while decreasing lymphatic channel width and intra-lymphatic pressure. They also prevent leukocyte adherence, intra-tissue movement, and the release of leukocyte (Lselectin) and endothelial (ICAM-1, VCAM-1) adhesion molecules, which protect microvascular permeability (Inflammatory mediators) [3,4]. Daflon treatment for four weeks has also shown improvement in the symptoms associated with hemorrhoids (pain, heaviness, bleeding, pruritus and anal discharge) [5,6,7]. Purified Diosmin has also been shown to reduce pain and bleeding. Diosmin hesperidin has also been reported to exhibit anti-oxidative property. Phlebotonics are known as vasoactive drugs, their mechanism of action is not scientifically well established despite the lack of studies examining their pharmacological and clinical properties. These drugs have been found to impact macrocirculation, such as enhancing venous tone [8], as well

as microcirculatory parameters such as capillary hyper-permeability [9]. Conservative treatment options are required to improve symptoms and prevent progression to higher degrees and complications. An operation is indicated when non-operative approaches have failed or complications have occurred [10, 11]. Topical agents like creams, lotions, and suppositories, which contain various ingredients (local anesthetics, corticosteroids, antibiotics, and anti-inflammatory drugs) have been employed for hemorrhoids management [11]. Although these agents help in improving symptoms, strong evidences supporting their true efficacy are lacking [10, 12].

MATERIALS AND METHODS

A single institute retrospective observational study was conducted where the varicose vein and haemorrohids patients treated with Diosmin in combination with hesperidin were reviewed from January 2022 to December2023. Institutional Ethical Committee (IEC) approval was obtained.

Inclusion criteria:varicose vein and haemorrhoids patients which were conservatively treated.

Exclusion criteria: complicated cases which are managed with surgery.

The parameters assessed Haemorrhoidal symptoms (pain, heaviness, bleeding, pruritus and anal discharge) and in the proctoscopic appearance of the haemorrhoids. Varicose vein assessed by Venous clinical severity score (VCSS). Following factors were considered in the statistical analysis: demographic data, including sex, age,outcomes of disease. Haemorrhoidal symptoms (pain, heaviness, bleeding, pruritus and anal discharge) and in the proctoscopic appearance of the haemorrhoids comparing baseline visit.

Statistical Analysis

Results were analysed using chi-square test, Bar chart, Paired T test

RESULTS

52 cases of varicose veins and 66 cases of hemorrhoids were monitored for two years at a tertiary care hospital. The age and sex distribution of hemorrhoids and varicose veins are displayed in charts 1, 2, 6, and 7 Respectively. Out of the 66 patients, 41 had improved symptoms

(62%), and 24 (36.4%) needed surgery in the end.In one instance, rectal prolapse occurred as shown in chart 3,4,5.0f the 52 patients with varicose veins, 42 (80.7%) had improved symptoms, while 9 (17.3%) needed surgery in the end.One (1.9%) developed necrotizing fasciitis as shown in chart 8and 9.

DISCUSSION

General practitioners frequently treat diseases including varicose veins and hemorrhoids. The doctor can select from a variety of therapy options for both problems. Mechanical compression stockings are used as a treatment for varicose veins. For hemorrhoids, there are numerous over-the-counter topical treatments available. Dietary modifications, lifestyle adjustments, and hydrotherapy are examples of conservative treatments for both illnesses that depend heavily on patient compliance to be successful.

CONCLUSION

When comparing the results of the initial visit and the last appointment, which took place eight weeks after starting diosmin hesperidin treatment, the symptoms of hemorrhoids pain, heaviness, bleeding, pruritus, and anal dischargeas well as their proctoscopic appearance have decreased for early grades of hemorrhoids. Following two months of treatment with diosmin and hesperidin, venous symptoms decreased. Diosmin plus hesperidin for the conservative management of hemorrhoids and varicose veins has great efficacy

Age Distribution (hemorroids Cases):chart 1

| 9 | | |
|------------------|--------------|----------------|
| AGE DISTRIBUTION | NO. OF CASES | PERCENTAGE (%) |
| 21-30 | 11 | 16.7% |
| 31-40 | 10 | 15.2% |
| 41-50 | 15 | 22.7% |
| 51-60 | 12 | 18.2% |
| 61-70 | 12 | 18.2% |
| 71-80 | 6 | 9.1% |
| TOTAL | 66 | 100% |

Sex Distribution (hemorroids Cases) Chart 2

| | , | |
|------------------|--------------|----------------|
| SEX DISTRIBUTION | NO. OF CASES | PERCENTAGE (%) |
| MALE | 45 | 68.2% |
| FEMALE | 21 | 31.8% |
| TOTAL | 66 | 100% |

Follow up chart 3

| FOLLOW UP | NO. OF | PERCENTAGE |
|------------------------|--------|------------|
| | CASES | (%) |
| SYMPTOMATICALLY BETTER | 41 | 62.1% |
| SURGICAL INTERVENTION | 24 | 36.4% |
| RECTAL PROLAPSE | 1 | 1.5% |
| TOTAL | 66 | 100% |

Chart 4

| PROCTOSCOPY BEFORE TREATMENT * PROCTOSCOPY | | | | | | |
|--|-----------|-------------------|------|------|-------|------|
| AFTER TREATMENT | | | | | | |
| | | PROCTOSCOPY AFTER | | | | Tota |
| | | TREATMENT | | | | 1 |
| | | NIL | MODE | GOOD | EXCE | |
| | | | RATE | | LLENT | |
| PROCTOSC | GRADE I | 0 | 5 | 18 | 4 | 27 |
| OPY | HEMORROID | | | | | |
| BEFORE | GRADE II | 2 | 7 | 7 | 0 | 16 |
| TREATME | HEMORROID | | | | | |
| NT | GRADE III | 12 | 1 | 0 | 0 | 13 |
| | HEMORROID | | | | | |
| | GRADE IV | 9 | 1 | 0 | 0 | 10 |
| | HEMORROID | | | | | |
| Total | | 23 | 14 | 25 | 4 | 66 |

Chi-Square = 58.926 P = < 0.05

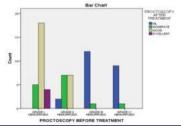


Chart 5

Age Distribution(varicose Veins) Chart 6

| AGE DISTRIBUTION | Count | Percent |
|------------------|-------|---------|
| 21-30 | 2 | 3.8% |
| 31-40 | 6 | 11.5% |
| 41-50 | 9 | 17.3% |
| 51-60 | 13 | 25.0% |
| 61-70 | 15 | 28.8% |
| 71-80 | 3 | 5.8% |
| 81-90 | 4 | 7.7% |
| TOTAL | 52 | 100% |

Sex Distribution (varicose Veins) Chart 7

| ` | , | |
|------------------|--------------|----------------|
| SEX DISTRIBUTION | NO. OF CASES | PERCENTAGE (%) |
| MALE | 33 | 63.5% |
| FEMALE | 19 | 36.5% |
| TOTAL | 52 | 100% |

Follow Up:chart 8

| 1 ono ii epicharto | | | | |
|------------------------|--------|------------|--|--|
| FOLLOW UP | NO. OF | PERCENTAGE | | |
| | CASES | (%) | | |
| SYMPTOMATICALLY BETTER | 42 | 80.7% | | |
| SURGICAL INTERVENTION | 9 | 17.3% | | |
| NECROTIZING FASCITIS | 1 | 1.9% | | |
| TOTAL | 52 | 100% | | |

Paired T Test: Chart 9

| | MEAN | N | STD. | t | p |
|-------------|---------|----|-----------|-------|--------|
| | | | DEVIATION | VALUE | VALUE |
| VCSS BEFORE | 12.3269 | 52 | 1.18357 | 7.945 | < 0.05 |
| TREATMENT | | | | | |
| VCSS AFTER | 10.8077 | 52 | 1.89983 | | |
| TREATMENT | | | | | |

REFERENCES

- Feldo M., Wójciak-Kosior M., Sowa I., Kocki J., Bogucki J., Zubilewi cz T., Kęsik J., Bogucka-Kocka A. Effect of Diosmin Administration in Patients with Chronic Venous Disorders on Selected Factors Affecting Angiogenesis. *Molecules*. 2019;24:3316. doi: 10.3390/molecules24183316. [PMC free article] [PubMed] [CrossRef] [Google Scholar]
- Ramelet A.A., Boisseau M.R., Allegra C., Nicolaides A., Jaeger K., Carpentier P., Cappelli R., Forconi S. Veno-active drugs in the management of chronic venous disease. An international consensus statement: Current medical position, prospective views and final resolution. Clin. Hemorheol. Microcirc. 2005;33:309–319. [PubMed] [Google Scholar]
- Perumal S. Effect of diosmin on apoptotic signaling molecules in Nnitrosodiethylamine-induced hepatocellular carcinoma in experimental rats. *Mol. Cell. Biochem.* 2018;449:27–37. doi: 10.1007/s11010-018-3339-3. [PubMed] [CrossRef] [Google Scholar]
- Katsenis K. Micronized purified flavonoid fraction (MPFF): A review of its pharmacological effects, therapeutic efficacy and benefits in the management of chronic venous insufficiency. Curr. Vasc. Pharmacol. 2005;3:1–9. doi: 10.2174/ 1570161052773870. [PubMed] [CrossRef] [Google Scholar
- Meshikhes A.W.N. Dafton for haemorrhoids: A prospective, multi-centre observational study. Surgeon. 2004;2 doi: 10.1016/S1479-666X(04)80032-5. [PubMed] [CrossRef] [Google Scholar]
- Cypriani B., Limasset B., Carrié M.L., Le Doucen C., Roussie M., de Paulet A.C., Damon M. Antioxidant activity of micronized diosmin on oxygen species from stimulated human neutrophils. *Biochem. Pharmacol.* 1993;45:1531–1535. doi: 10.1016/0006-2952(93)90056-3. [PubMed] [CrossRef] [Google Scholar]
- Oh-hora M., Komatsu N., Pishyareh M., Feske S., Hori S., Taniguchi M., Rao A., Takayanagi H. Agonist-Selected T Cell Development Requires Strong T Cell Receptor Signaling and Store-Operated Calcium Entry. *Immunity*. 2013;38:881–895. doi: 10.1016/j.immuni.2013.02.008. [PMC free article] [PubMed] [CrossRef] [Google Schelber
- Tsouderos Y. Venous tone: Are the phlebotonic properties predictive of a therapeutic benefit? A comprehensive view of our experience with Daflon 500 mg. Z. Fur Kardiol. 1991-80((Supul) 57):95-101 [PubMed] [Google Scholar]
- 1991;80((Suppl. S7)):95–101. [PubMed] [Google Scholar]
 Behar A., Nathan P., Lavieuville M., Allaert F.A. Effect of veinotonyl 75 on the capillary permeability test using technetium albumin in cyclic orthostatic edemas. *Phlebologie*. 1993;46:721–731. [PubMed] [Google Scholar]
- 10. Sardinha T.C., Corman M.L. Hemorrhoids. Surg. Clin. North Am. 2002;82:1153–1167.
- [PubMed] [Google Scholar
 11. Lohsiriwat V. Hemorrhoids. From basic pathophysiology to clinical management.
 World J. Gastrointest. Oncol. 2017;18:2009–2017. [PMC free article] [PubMed] [Google Scholar]
- Gurel E., et al. Herbal haemorrhoidal cream for haemorrhoids. Chin. J. Physiol. 2013;56:253–262. [PubMed] [Google Scholar]