ABSTRACT

Objective: The objective of this study is to assess the effect of pregabalin in the management of post-operative pain and the quality of life (QOL) of osteoarthritis (OA) patients after total knee arthroplasty.

Methods: This prospective observational study was conducted in the Department of Orthopedics. A total of 96 patients were divided into two groups. Group A consists of 50 patients with the treatment of nonsteroidal anti-inflammatory drugs (NSAIDs) and Group B consists of 46 patients with the treatment of pregabalin with NSAIDs. The study subjects were followed once in 60 days for 6 months, and they were asked to answer the visual analog scale (VAS) and knee injury and OA outcome score (KOOS) questionnaire. The effect of the treatment was assessed by comparing the baseline score with follow-up score.

Results: Our study result showed that the pain score of Group B in VAS at 2nd follow-up was 2.56±0.34 and KOOS pain score was 92.73±3.45 (p<0.01). The QOL score was improved significantly to 81.56±5.29 (p<0.01) as compared to Group A.

Conclusion: The study concluded that pregabalin with NSAIDs group patients showed a better improvement in pain, symptoms, and QOL within short duration as compared to NSAIDs alone used group.

Keywords: Osteoarthritis, Total knee arthroplasty, Pregabalin, Nonsteroidal anti-inflammatory drugs, Visual analog scale.

INTRODUCTION

Osteoarthritis (OA) is a common, progressive disorder affecting primarily weight-bearing diarthrodial joints [1]. Worldwide, OA is estimated to be the fourth leading cause of disability. Recent reports, over 40% of the Indian population in the age group of 70 or above suffer from OA. Good pain control after surgery is important to prevent negative outcomes such as tachycardia, hypertension, and myocardial ischemia and decrease in alveolar ventilation, poor wound healing, and Insomnia [2].

Total knee arthroplasty (TKA) is an effective treatment for end-stage knee OA, but the optimal management of post-operative pain remains controversial. TKA is often cited as one of the most painful known procedures. Approximately 20% of the patients with OA of the knee do not want to undergo TKA because of the expectancy of high levels of pain. After TKA, the pain may also inhibit early rehabilitation to mobilize the knee joint [3].

Nonsteroidal anti-inflammatory drugs (NSAIDs) are often considered to be the preferred first-line treatment for OA. The use of non-opioid drugs during the pre-operative period can reduce excess intraoperative opioid use. The use of pre-operative NSAIDs and cyclooxygenase-2 (COX-2) inhibitors also has a significant effect on opioid requirements following surgery. This has been referred to as an "opioid-sparing effect." The clinical significance of this may be the reduction of opioid-related side effects, improved analgesia, and better patient satisfaction [4].

In addition to NSAIDs and COX-2 inhibitors, anti-neuropathic drugs are playing a role in the treatment of post-operative pain. Drugs such as gabapentin and pregabalin, intended for seizures and neuropathic pain syndromes, can inhibit central neuronal sensitization. Pregabalin was also found to have a synergistic effect with COX-2 inhibitors in clinical studies involving patient undergoing spinal fusion. The combination of the two drugs reduced post-operative pain, morphine consumption at 24 h, and opioid-induced side effects. Improvements in outcome were greater when these drugs were used in combination than either of the drugs used alone [5].

METHODS

A prospective observational study was carried out in the Department of Orthopedics in a Tertiary Care Hospital. The study was approved by the Institutional Ethical Committee. The duration of study was 1 year from November 2016 to October 2017. The patient’s history was collected from patient case records, designed data entry form and pain questionnaire, medication charts, laboratory reports, and patient/patient caretaker’s interview. The effect of the treatment was assessed using visual analog scale (VAS) [6] and knee injury and OA outcome score (KOOS) questionnaire [7].

Patients were eligible for the study if patients with 40–75 years of age, both male and female patients, patients with the treatment of NSAIDs or pregabalin with NSAIDs, and patients who underwent TKA. Patients were excluded if patients have an allergy or intolerance to any of the study drugs, severe liver failure, and heart or renal failure patients, patients with a history of opioid medication usage, and patients with bleeding disorder.

Randomly selected patients were assigned to receive study medication. A total of 96 patients are divided into two groups. Group A consists of 50 patients with the treatment of NSAIDs paracetamol 325 mg + aceclofenac 100 mg and Group B consists of 46 patients with the treatment of pregabalin 75 mg + NSAIDs (paracetamol 325 mg + aceclofenac 100 mg). The patient demographics data were collected using the data entry form. The baseline score of pain,
Table 1: Patient demographics

| Age (years) | Number of patient (%) |
|-------------|-----------------------|
| 40–49       | 9 (9.40)              |
| 50–59       | 23 (24.00)            |
| 60–69       | 46 (48.00)            |
| 70–79       | 18 (18.75)            |

Gender

| Male        | 40 (41.66) |
|-------------|------------|
| Female      | 56 (58.34) |

BMI

| Normalweight | 63 (65.62) |
|--------------|------------|
| Overweight   | 33 (34.38) |

BMI-Body mass index

Table 2: Comparison of VAS score

| VAS score | Baseline | Follow-up-1 | Follow-up-2 |
|-----------|----------|-------------|-------------|
| Group A   | 5.7±0.58 | 5.18±0.59   | 4.42±0.86** |
| Group B   | 5.6±0.93 | 3.97±0.46   | 2.56±0.34** |

VAS-Visual analog scale, **p<0.01

Table 3: Assessment of Group A using KOOS questionnaire

| Parameters | Baseline | Follow-up-1 | Follow-up-2 |
|------------|----------|-------------|-------------|
| Pain       | 62.78±7.92 | 70.8±6.70 | 79.0±6.68** |
| Symptoms   | 61.2±11.72 | 68.36±11.84| 78.56±9.80**|
| ADL        | 76.4±5.46  | 82.4±5.41  | 88.1±5.55** |
| SRF        | 91.3±2.52  | 92.7±3.41  | 93.1±3.28** |
| QOL        | 57.7±9.63  | 59.3±8.80  | 64.9±6.00** |

Table 4: Assessment of Group B using KOOS questionnaire

| Parameters | Baseline | Follow-up-1 | Follow-up-2 |
|------------|----------|-------------|-------------|
| Pain       | 67.19±5.73 | 84.9±6.02 | 92.7±3.45** |
| Symptoms   | 67.8±8.35  | 83.5±6.23  | 92.9±3.59** |
| Activity of daily living | 80.2±5.40 | 89.5±2.14 | 94.9±2.71** |
| SRF        | 92.3±5.55  | 93.5±4.03  | 92.6±2.52  |
| QOL        | 57.5±8.18  | 71.0±3.49  | 81.5±5.29** |

TKA: Total knee arthroplasty, KOOS: Knee injury and osteoarthritis outcome score, SRF: Sport and recreation function, QOL: Quality of life, **p<0.01

ASSESSMENT OF GROUP A USING KOOS QUESTIONNAIRE

KOOS mean pain score at baseline was 62.78±7.92 and at 2nd follow-up was 79.0±6.68. The mean score of symptoms at follow-up-2 was significantly high as compared to Group A (Table 2).

ASSESSMENT OF GROUP B USING KOOS QUESTIONNAIRE

KOOS mean pain score at baseline was 67.19±5.73 and at 2nd follow-up was 92.7±3.45. The mean score of QOL at follow-up-2 was significantly improved as compared with other parameters 81.5±5.29. The mean difference was statistically significant from baseline (p<0.01) (Table 3).

Comparison of Effect of Treatment

The Group B treatment was more effective in reduction of pain and symptoms as compared with Group A. The mean difference of pain and symptoms score between Group A and Group B was statistically significant (p<0.01). The improvement of QOL was high in Group B as compared to Group A (Table 5).

Discussion

Among 96 patients, of which 46 patients were in the age group of 60–69 years. The mean age of the study population was 61.6±7.56 years. A study conducted by Ohtori et al. showed that the mean age of patients was 70±8 years [8,9]. As per the study results, female patients (58.3%) were more suffered by OA than male (41.66%). The majority of patients were in our study were normal weights. Earlier study was reported that the majority of patients were having over body weight [10].

As per assessment, by VAS, Group B was found to have a significant reduction in pain intensity than Group A patients. Mathiesen et al. have reported that pregabalin administration could reduce the VAS score at 24 h compared with placebo postoperatively [11].

In this study, oral administration of pregabalin with NSAIDs was more effective in reducing pain as compared with NSAIDs alone in post-operative cases. Fassoulaki et al. said that the administration of gabapentin to the woman undergoing total abdominal hysterectomy did not reduce acute post-operative pain. However, there was a decrease in pain at 1 month postoperatively [12].

Group B was also effective in impaired active and passive range of motion (ROM) after TKA as compared with Group A patients. The mean difference of KOOS symptoms score was statistically significant (p<0.01) and result indicated that the symptom was decreased and knee flexion was improved as compared with Group A. Chiu et al. reported that oral pre-operative administration of pregabalin improved knee flexion and ROM after TKA [13,14].

The mean difference of activities of daily living (ADL) and QOL after TKA between Group A and B was statistically significant at final follow-up. The result indicates that the ADL and QOL were improved in Group B patients as compared with Group A patients. Based on the study findings, Group B patients had reduced pain and improved QOL of TKA as compared with Group A patients.
CONCLUSION

The present study showed that pregabalin with NSAIDs group was more effective in reducing pain after TKA as compared with NSAIDs group. As per the study, pregabalin with NSAID group results revealed that a better improvement in OA symptoms, ADL and QOL within a short duration. Regular follow-up and good patient compliance may improve the QOL of patients in the day-to-day activities. Long-term treatment of pregabalin may reduce the dependence of analgesics and decrease OA-related neuropathic pain incidence.

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CONFLICT OF INTEREST

The author has declared no conflict of interest.

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