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Role of intra articular injection of autologous platelet rich plasma in osteoarthrosis of the knee joint

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Abstract

Osteoarthrosis (OA) represents a failure of the diarthrodial, synovial lined joint. Among the elderly, knee Osteoarthrosis is the most frequent cause of chronic disability ^[1]. Because of the increased life expectancy, the prevalence of osteoarthrosis is on the rise in the Indian population. Platelet rich plasma (PRP) is defined as a volume of plasma with a platelet concentration higher than average in peripheral blood. Many basic, preclinical and even clinical case studies and trails report PRP's ability to improve musculoskeletal conditions including osteoarthrosis. Our study of platelet rich plasma has thrown up an exciting choice of treatment modality in knee osteoarthrosis, and it has proved efficient in observation period of one year.

Keywords: anti-inflammatory intra-articular therapies, clinical evidence, knee osteoarthrosis, platelet rich plasma

Introduction

Osteoarthrosis is age-related, affecting more than 80% of people over the age of 55 OA in weight-bearing joints is strongly linked to BMI (Body Mass Index). OA has a preference for knees, hips, shoulders, and the spine and phalanges. Occurrence in an atypical joint, such as an elbow, can usually be before the trauma, congenital joint abnormality, underlying systemic disease, or a chronic crystalline arthropathy. The heterogeneity of OA can be due to many factors that can contribute to cartilage damage.

Symptomatic OA of the knee, which is having pain during most days of a month along with other evidence (e.g., radiological) of arthritis, has a prevalence of 30% to 40% in India^[3, 4].

Osteoarthrosis is a chronic disorder of synovial lined joints leading to progressive softening and disintegration of articular cartilage accompanied by the new growth of bone at the joint margins, cyst formation, and sclerosis at subchondral regions of bone, mild synovitis, and capsular fibrosis.

Aim of the study

Treatment of osteoarthrosis with NSAIDs is associated with Osteoarthrosis is a clinically heterogenous degenerative condition characterized by the destruction of articular cartilage due to the uncoupling of a balance between cartilage degeneration and regeneration.

The management of osteoarthrosis ^[15] has varied from conventional therapy with physical education, NSAIDS, intraarticular steroid injection increased the risk of GI disturbances along with the alarming rise in NSAID induced complications. Arthroplasty is usually reserved for advanced stages of OA.

The main aim of this study is to evaluate the effectiveness of Platelet Rich plasma in decreasing pain, stiffness, and improving function in patients with early osteoarthrosis knee, as Platelet Rich Plasma provides a cocktail of growth factors into the joint cavity. Platelet Rich Plasma is postulated to modify the disease process, unlike other methods of nonsurgical treatment which provide symptomatic relief. Platelet Rich Plasma is a cost-effective tool that could prevent the need for Total Joint Arthroplasty. Or at least reduce the number of surgeries.

Review of literature

Khosbin *et al.*, in a systematic review with the quantitative synthesis in 2013, concluded that intraarticular PRP injections might have benefits in the treatment of adult patients with mild to moderate osteoarthrosis. The study also reported an increased incidence of non-specific adverse events among patients treated with Platelet-rich plasma.

Kalbkhani *et al.* in 2014 studied the effect of PRP in experimentally induced OA in rabbits knee joint concluded that the PRP group had near-normal joint structure at 16-week postop interval, and hence PRP could potentially be used for the treatment of osteoarthrosis.

Giuseppe Filardo, in 2010, studied platelet-rich plasma intraarticular knee injections for the treatment of degenerative cartilage lesions, and osteoarthrosis concluded that treatment with PRP could reduce pain and improve knee function and quality of life with short term efficacy.

Kon *et al.* Trial - the effectiveness of PRP injections were compared to hyaluronic acid (HA) intra-articular injection therapy. The primary outcomes of pain reduction function improvements were measured through the International Knee Document Committee (IKDC) and Visual Analogue Scale (EQ-VAS) scoring system.

Evaluation of the IKDC score in the PRP group showed a steady increase from

(Baseline) to 62.7 and 64.0 at 2 and 6 months follow up, respectively.

Objectives

To evaluate the role of Autologous Platelet Rich Plasma (PRP) in treating patients presenting with early osteoarthrosis knee and analyze whether it could be a cost-effective disease-modifying the measure.

Study Design

A prospective study was done in 150 patients in which standardized injection protocol was given and was assessed on variables such as pain, physical function using the WOMAC scale, and for depression using a visual analog scale at preinjection, 6 weeks post-injection, 3 months and 6 months and 1yr post-injection.

Materials and Methods

Patients attending the outpatient department of Orthopaedics at NRI Medical College with complaints of bilateral knee pain and stiffness were screened, and those diagnosed as bilateral Knee Osteoarthrosis were chosen for the study.

The Patients, classified either grade 0 to 4 on the Kellgren-Lawrence grading scale or grade 1 to 4 on the Ahlback scale, were included in the study after prior well informed written consent.

150 Patients were chosen and were given an intraarticular injection of protein- rich plasma

Inclusion Criteria

- 1. Kellgren–Lawrence Grade 1, Grade 2, Grade 3A
- 2. Age of more than 35 years.
- 3. A patient who gave consent for the study.
- 4. Platelet count (minimum 2 lakhs per microliter)
- 5. Random blood sugar <160 mg/dl
- 6. haemoglobin >10 gm%

Exclusion Criteria

- 1. Platelet dysfunction syndrome.
- 2. Critical thrombocytopenia (<105 /mL).
- 3. Hypofibrinogenemia.
- 4. Septicemia.
- 5. Coagulopathies.
- 6. Presence of tumors or metastasis.
- 7. Active infection.
- 8. Pregnancy or breast-feeding.
- 9. Immune deficiencies.
- 10. Patients with vascular injuries.

Kellgren Lawrence grading of osteoarthritis10

Grade 0: no radiographic features of OA are present

Grade 1: doubtful narrowing of joint space with possible osteophytes Lipping

Grade 2: definite osteophytes, definite narrowing of joint space Grade 3: moderate multiple osteophytes, definite narrowing of joints space, some sclerosis and possible deformity of bone contour

Grade 4: large osteophytes, marked narrowing of joint space, severe sclerosis and definite deformity of bone contour

Ahlback radiological criteria1

Grade 1: Joint Space Narrowing (Less Than 3 Mm)

Grade 2: Joint Space Obliteration

Grade 3: Minor Bone Attrition (0-5 Mm)

Grade 4: Moderate Bone Attrition (5-10 Mm)

Grade 5: Severe Bone Attrition (More Than 10 Mm)

Autologous Platelet Rich Plasma^[7] was prepared in the NRI medical college orthopedics department.

The process of separating Platelet Rich Plasma was done under strict aseptic conditions.

The Patients baseline platelet count and leukocyte count were determined, and Platelet Rich Plasma was quantified as having eight to ten times the baseline valve of platelets. The Concentration of Platelets in the final product was corroborated by the Department of the pathology of NRI medical college on a periodic basis.

We in this study did not use leucocyte filter, and the final Platelet Rich Plasma contained minute traces of leucocytes.

Injection protocol

The Injection procedure was performed in the emergency operation theatre. The patient was placed supine on the operation table. Parts scrubbed, painted, and draped. Under sterile aseptic precautions, 3ml of Platelet Rich Plasma was injected into the joint cavity from a medial approach sterile bandaging given. Cycling of the knee is done 10 times for the distribution of PRP in the joint cavity. The Patient has advised analgesics and ice fomentation for pain after the procedure.

The Patients were advised to carry on with their regular work from Day 1.

Outcome analysis

The patients were advised to follow up at 6 weeks, 3 months, and 6 months and 12 months. Outcome analysis was done for the reduction in pain, decrease in stiffness, and improvement in physical function using the WOMAC scale. The Patients were also assessed for reduction in pain using Visual analog scale both at pre-injection and at 12 months post-injection.



Fig 1: Procedure



Fig 2: The process of separating Platelet Rich Plasma was done under strict aseptic conditions

Womac (western ontario and mcmaster universities) index of osteoarthritis 5, 17

The WOMAC index is used to assess patients with osteoarthrosis of the hip or knee using 24 parameters. It can be used to monitor the course of the disease or to determine the effectiveness of Therapy

Circle one number for each activity Pain

Scale of difficulty: 0 = None,

- 1 = Slight,
- 2 = Moderate,
- 3 = Very,
- 4 = Extremely

1.	Walking	0 1 2 34
2	Climbing stairs	01234
3.	Nocturnal	01234
4.	Rest	01234
5.	Weight bearing	01234
	Stiffness	
1.	Morning stiffness	01234
2.	Stiffness occurring later in the day	01234

Physical function

1. Descending stairs	01234
2. Ascending stairs	01234
3. Rising from sitting	01234
4. Standing	01234
5. Bending to floor	01234
6. Walking on flat surface	01234
7. Getting in / out of car	01234
8. Going shopping	01234
9. Putting on socks	01234
10. Lying in bed	01234
11. Taking off socks	01234
12. Rising from bed	01234
13. Getting in and out of bath	01234
14. Sitting	01234
15. Getting on and off toilet	01234
16. Heavy domestic duties	01234
17. Light domestic duties	01234
5	

Total Score: ____/ 96 = ___%

Visual analog scale for pain

The Pain, the Patient, perceives is graded on a visual scale and the score calculated.



Ask patients about their:

- 1. Intensity
- 2. Location
- 3. Duration
- 4. Onset
- 5. Variation
- 6. Quality



Fig 3: Age

Statistical analysis Gender distribution



Fig 4: Sex distribution in PRP

Weight distribution



Fig 5: Weight distribution of PRP

WOMAC Score



Fig 6: Mean WOMAC score

Pain Score



Fig 7: Mean Pain score

Stiffness score Physical function score





Fig 9: Mean Physical function score

Visual Analog Score



Fig 10: Mean visual analog score

Discussion

Osteoarthrosis is a synovial joints disorder caused mainly by the uncoupling of balance between cartilage regeneration and degeneration due to focal loss of hyaline cartilage leading to proliferation of cells and the formation of new bone and remodeling of joint surfaces, Osteoarthrosis is a dynamic repair process of synovial joints that may be triggered by a variety of causes.

The use of biological agents, including PRP and mesenchyal stem cells (MSCs) in orthopedics, has increased exponentially over the last few years due to its autologous nature, lack of side-effects, and supposed effectiveness.

Platelet-rich plasma is an autologous blood product with platelet concentrations above baseline.

Tissue repair is a complex process comprising chemotaxis, angiogenesis, cell proliferation, and matrix formation. Platelets are involved in all of these functions by the release of growth factors.

High concentrations of proteins such as platelet-derived growth factor (PDGF), vascular endothelial growth factor (VEGF), endothelial cell growth factor, and the fibroblast growth factor have led to suggestions that PRP may be useful in conditions requiring tissue healing. Conversely, the other protein in PRP, transforming growth factor (TGF- β 1), has an inhibitory effect and can lead to non-predictable results.

Preparation of PRP can result in four products

- 1. Pure PRP (P-PRP) with a low content of leucocytes. This can be injected as a liquid or a gel.
- 2. Leucocyte-rich PRP (L-PRP) has a higher concentration of platelets than P- PRP. Similarly to P-PRP, it can be used as an activated gel or in a liquid form to be injected intra-articular.
- 3. Pure platelet-rich fibrin (P-PRF). This is obtained by double-spinning centrifugation without adding anticoagulants. The end product is a platelet-rich fibrin scaffold, which is stiffer than the conventional PRP and takes the form of a gel and can be used for the healing of wounds.
- 4. Leucocyte- and platelet-rich fibrin (L-PRF), which is a leucocyte-rich gel which is non-injectable and is applied locally.

Platelets are regarded as the primary mediators of hemostasis. They contain alpha granules enriched with growth factors. Platelets also have anti-bacterial and fungicidal agents, which provoke the synthesis of interleukins and chemokine's. When platelets get activated, this causes the release of growth factors. Among them, important ones include transforming growth factor (PDGF), insulin-like growth factor (IGF), and fibroblast growth factor (FGF), etc. In the presence of calcium chloride, the platelet concentrate is activated, which causes the release of these growth factors, eventually promoting healing.

PRP has also been found to have anti-inflammatory actions. The inflammatory cascade generated by cyclooxygenase family can be inhibited by anti-inflammatory mediators present in PRP.

PRP has an influence on all structures of joint. Chemotactic assays have revealed that the PRP stimulated the differentiation of type-II collagen cells and the production of prostaglandins along with the migration of corticospongious bone cells

We, in our study, had 150 patients with classic findings of Osteoarthritis. They were comparable to baseline characteristics of age, weight, and pre-injection, and post- injection WOMAC score. All the patients received intraarticular injections.

The Efficacy of Platelet Rich Plasma (PRP) in decreasing pain, stiffness, physical function were assessed and scored on the WOMAC index.

Age distribution revealed a mean age in to be 49.92.

Gender distributions were comparable in both groups, with 66.6 % being male 33.3% being female.

Thus the study of platelet-rich plasma ensured that all patients were comparable to baseline.

The Global WOMAC score showed a mean of 74.1 at the preinjection period, which decreased to 62.45 at 6 weeks follow up and 47.68 at 3 months and declining to 33.40 at 6 months and 33.20 at one year.

Individual variables such as pain, physical function, etc. were assessed. The mean score for pain showed a decrease from 16.57 to 11.31 at 6 weeks post-injection. At the end of 1 year follow up, the mean was 5.34.

The mean of Physical function decreased from a pre-injection score of 51.11 to 24.62 at 6 months follow up and 24.50 at one year.

Conclusion

The Epidemic of Modernization, along with adequate health care delivery, has led to an expanded lifespan of human beings. The focus of health care providers is undergoing a drift towards noncommunicable and degenerative disorders.

Osteoarthrosis represents a failure of the diarthrodial joint, characterized by degenerative changes in articular cartilage of joints. The management of Osteoarthrosis has undergone a sea change during the last century.

Our study of platelet-rich plasma relied on injecting a highly concentrated mix of platelets into the joint cavity and observing the patients for the decrease in symptoms of pain, stiffness, and improvement in physical function. Our study has revealed a consistent reduction in pain and stiffness and a definite improvement in lifestyle of the patients.

Our study of platelet-rich plasma has thrown up an exciting choice of treatment modality using Platelet Rich Plasma in the treatment of Knee Osteoarthrosis, and it has proved efficient in the observation period of one year.

Complications

There were no significant complications of infection in our patients.

Limitations

Long term follows up needed with magnetic resonance imaging to assess the

Regeneration of cartilage.

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