Management of lateral epicondylitis (tennis elbow) by local infiltration of platelet rich plasma an outcome study

Pradeep Kumar Kumawat1*, Rajesh Goel2, Urmila Kumhar3, Rajesh Sharma4, Rahul Parmar4, Bharat Sharma4

INTRODUCTION
Lateral epicondylitis is a common cause of lateral elbow pain that was first described by Runge. It is thought to represent an inflammation of the common extensor origin of the forearm and have a prevalence of 1.3% among those between 30 and 64 years of age, peaking between 45 and 54. Mainly dominant upper limb is affected and associated with repeated forceful activity of limb.

Lateral epicondylitis, commonly known as “tennis elbow,” occurs secondary to tendinosis of the extensor carpi radialis brevis origin immediately distal to the lateral epicondyle. The name is misleading because the lateral epicondyle is not the site of involvement, inflammation is not present, and most patients are not active tennis players.

Activities that require repeated contraction of the wrist extensors are responsible for tennis elbow like as lifting pots and pans or gripping a container of milk, wringing of washing clothes etc. Mainly extensor carpi radialis brevis
(ECRB) tendon involved. Over time, the pain can become very severe and may compromise with routine activities. Lateral epicondylitis evolves through several stages, beginning with degenerative angiogenesis and ends with fibrosis and calcification. Lateral epicondyle demonstrates angio fibroblastic hyperplasia at the extensor origin of the forearm.3

Various nonsurgical methods have been advocated for treating elbow tendinosis, including rest, nonsteroidal anti-inflammatory medication, bracing, physical therapy, iontophoresis, extracorporeal shock wave therapy, botulinum toxin and corticosteroid injection.

Platelet-rich plasma (PRP) has been recently the emerging biological therapy in which a large pool of signals released from platelets producing an instructional biological microenvironment for local and migrating cells for tissue regeneration. PRP modulate inflammation and angiogenesis largely because of their ability to secrete high levels of growth factors and chemokines. 4 PRP injection for musculoskeletal injuries have advantage over other method of treatment in relation to have no risk of an anaphylactic reaction, no blood-borne transmitted infection, low cost, safety, shorter recovery time compared to surgical management and availability for outpatient preparation and delivery.

The objectives of present study were; to determine demographic distribution of patient with lateral epicondylitis (tennis elbow), to study effect and functional outcome of lateral epicondylitis treated with PRP Infiltration at local site.

METHODS

This prospective randomized observational study was conducted in the department of orthopaedic, Govt. Medical College and associated group of hospitals, Kota between December 2015 to November 2017 after approval by ethical committee. 100 consecutive patients with lateral epicondylitis of the elbow were selected for the study

Inclusion criteria

A minimum age of 18 years and positive findings from two of the following clinical tests: cozen, mill, Gardner and Maudsley, tendinopathy present in lateral elbow, patients who had failed conservative treatment, symptoms lasting at least 3 months or longer, commitment to comply with all study procedures and the patient must give written informed consent were included in this study.

Exclusion criteria

Presence of full tendon tear with history of trauma, systemic autoimmune rheumatologic disease (connective tissue diseases and systemic necrotizing vasculitis), diabetes mellitus patient, blood disorders (thrombopathy, thrombocytopenia, anemia with Hb <9), patient receiving immunosuppressive treatment, received local steroid injection within 3 months of randomization, received non-steroidal anti-inflammatory, opioids, or oral corticosteroids within 15 days before inclusion in the study, severe heart disease, patients unable to comply with scheduled visits, patients with active cancer or cancer diagnosed, patient of hepatitis B, C, or HIV infection, pregnant or lactating women, other causes of elbow pain such as osteochondritis dissecans of capitellum, Varus instability, radial head arthritis, posterior interosseous nerve syndrome, synovitis of radio humeral joint, cervical radiculopathy, osteoarthritides of elbow patients were excluded from this study.

Method of PRP preparation and infiltration

PRP is derived from the centrifugation of autologous blood. 10 ml autologous peripheral venous blood was collected from cubital vein of unaffected upper limb of patient and 9ml transferred into 1 ml CDP-A (citric acid, sodium citrate, dextrose, monobasic sodium phosphate, adenine) containing sterile test tube. Rest 1 ml was sent for complete blood count. Sample was processed through two staged centrifugations (first with 1600 rpm for 10 minutes for separation of RBC and next with 3200 rpm for 7 minutes in order to concentrate platelets). Final product was 1-1.5 ml PRP containing WBCs. The PRP quantification was performed using Automated Symex analyzer and if approved, the injection was proceeded.

PRP infiltration

With patient in supine posture, the bony anatomical landmarks were identified. The elbow was flexed to 90° with the palm facing down. With proper aseptic precautions 1-1.5 ml PRP infiltrated using peppering technique just anterior and below the lateral epicondyle humerus.

Outcome measures

The clinical outcome was rated on the basis of VAS scale and PRTEE scoring system at every follow up. VAS pain scale: no pain (0-4 mm), mild pain (5-44 mm) - pain present occasionally while at work, moderate pain (45-74 mm) - pain present but can continue with work, severe pain (75-100 mm) - pain forces discontinuation of the work but can be resumed after rest.5

PRTEE (patient-rated tennis elbow evaluation)

Scoring system for both pain and functional disability. total score=pain subscale + function subscale best score=0 worst score=100.6

RESULTS

Mean age of patients was 43.07±6.73 (30-50) years. Mostly were house workers (57%), laboure’s (18%), farmers (10%) and others (15%) including tailors,
teachers, and drummers. Other patient’s characteristics are shown in Table 1. In this study the mean VAS score and mean PRTEE score before injection were 75 (SD-5.99) and 78.62 (SD-4.68) respectively.

Table 1: Demographic characteristics.

<table>
<thead>
<tr>
<th>Demographic characteristics</th>
<th>Sex</th>
<th>Mean age</th>
<th>Side of involvement</th>
<th>Dominance</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Male 35 (35%)</td>
<td>43.07 years±6.73</td>
<td>Left 15 (15%)</td>
<td>Dominant side involved 90 (90%)</td>
</tr>
<tr>
<td></td>
<td>Female 65 (65%)</td>
<td></td>
<td>Right 85 (85%)</td>
<td>Non-dominant side involved 10 (10%)</td>
</tr>
</tbody>
</table>

At the end of 6 months, 61 patients (61%) were completely relieved of pain and functional disability. 34 patients (34%) have mild pain mean VAS and PRTEE score were 6.05 and 5.63 respectively. 5 patients (34%) have mild pain mean VAS and PRTEE score were 19.25 (SD-9.43), showed statistically significant decrease in VAS score and PRTEE. 15 patients (15%) were completely free from pain VAS score was 0. PRTEE score was 0 in 10 (10%) patients.

At 2 weeks follow up, statistically significant difference in VAS scoring and PRTEE was seen. Mean VAS score was 55.25 (SD-7.50) and mean PRTEE score was 58.92 (SD-8.52), showed statistically significant decrease in VAS score and PRTEE.

At 4th week follow-up, there was statistically significant decrease in VAS score and PRTEE score. Mean VAS score was 40.25 (SD-8.31) and mean PRTEE score was 32.77 (SD-7.99).

At 3 months follow-up, mean VAS score was 19.25 (SD-13.47) and mean PRTEE score was 16.35 (SD-9.43), showed statistically significant decrease in VAS score and PRTEE. 15 patients (15%) were completely free from pain VAS score was 0. PRTEE score was 0 in 10 (10%) patients.

Table 2: Mean VAS scoring.

<table>
<thead>
<tr>
<th></th>
<th>Pre-injection VAS</th>
<th>2 weeks VAS</th>
<th>4th weeks VAS</th>
<th>3rd months VAS</th>
<th>6th months VAS</th>
</tr>
</thead>
<tbody>
<tr>
<td>Mean VAS</td>
<td>75</td>
<td>55.25</td>
<td>40.25</td>
<td>19.25</td>
<td>6.05</td>
</tr>
<tr>
<td>SD</td>
<td>5.99</td>
<td>7.50</td>
<td>8.31</td>
<td>13.47</td>
<td>7.89</td>
</tr>
<tr>
<td>P value</td>
<td>&lt;0.0001</td>
<td>&lt;0.0001</td>
<td>&lt;0.0001</td>
<td>&lt;0.0001</td>
<td>&lt;0.0001</td>
</tr>
</tbody>
</table>

Table 3: Mean PRTEE score.

<table>
<thead>
<tr>
<th></th>
<th>Pre-injection PRTEE</th>
<th>2 weeks PRTEE</th>
<th>4th weeks PRTEE</th>
<th>3rd months PRTEE</th>
<th>6th months PRTEE</th>
</tr>
</thead>
<tbody>
<tr>
<td>Mean PRTEE score</td>
<td>78.62</td>
<td>58.92</td>
<td>32.77</td>
<td>16.35</td>
<td>5.63</td>
</tr>
<tr>
<td>SD</td>
<td>4.68</td>
<td>8.52</td>
<td>7.99</td>
<td>9.43</td>
<td>7.51</td>
</tr>
<tr>
<td>P value</td>
<td>&lt;0.0001</td>
<td>&lt;0.0001</td>
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The combined action of all these growth factors is complex, and each may have a different effect on a particular tissue. Growth factors may also interact with each other, activating different sets of signaling pathways. Different isoforms of growth factors have varying effects that may enhance or inhibit osseous and soft-tissue repair, depending on the mode of release of the factor and the dynamics of the wound environment.9

At the end of 6 months, 61 patients (61%) were completely relieved of pain and functional disability. 34 patients (34%) have mild pain mean VAS and PRTEE score were 6.05 and 5.63 respectively. 5 patients follow-up were lost at 6 months. No recurrence of pain and functional disability was seen. It was seen that there was a significant increase in post intervention pain for few days, 70 participants (70%) complained of increase of pain after local infiltration that was treated with rest and ice fomentation for few days.

In this study mean platelet concentration in CBC was 241.25×10⁷/µl and in PRP was 808.03×10⁷/µl that was approximate 3-4 times to mean platelet concentration in CBC. It was seen that 80 patients (80%) received 3 PRP injection, 2 injections were given in 20 patients (20%). Maximum benefit was also seen after 3 injections. single injection was not given in any patient.

DISCUSSION

Platelet-rich plasma (PRP) is a volume of plasma fraction of autologous blood having platelet concentrations above baseline. The platelet α granules are rich in growth factors that play an essential role in tissue healing, such as transforming growth factor-β, vascular endothelial growth factor, and platelet-derived growth factor.

The properties of PRP are based on the production and release of multiple growth and differentiation factors when the platelets are activated. Platelets begin actively secreting these proteins within ten minutes of clotting, with more than 95% of the pre-synthesized growth factors secreted within one hour.7 After the initial burst of growth factors, the platelets synthesize and secrete additional such factors for the remaining several days of their life span.8

The combined action of all these growth factors is complex, and each may have a different effect on a particular tissue. Growth factors may also interact with each other, activating different sets of signaling pathways. Different isoforms of growth factors have varying effects that may enhance or inhibit osseous and soft-tissue repair, depending on the mode of release of the factor and the dynamics of the wound environment.9
Medical literature has sufficient studies to prove definitive role of PRP (platelet rich plasma) in healing of injured tissue. Cellular response to injury progresses through four general stages: hemostasis, inflammation, proliferation and finally remodeling. Each phase is characterized by enhanced cellular or Molecular activity, all of which involve platelets. Activated platelets and leukocytes mediate inflammation while various growth factors derived from platelets alfa granules influence tissue regeneration. Specifically, angiogenic and mitogenic growth factor concentrations are believed to aid tissue regeneration.

In this current study, the mean age encountered was 43 years (Range; 30 to 56 years); the peak incidence was seen from 30 to 50 years. Similar observation (mean age 45±5.9 years) was seen in Raeissadat et al.

In this current study, out of the 100 participants, 65 (65%) were female patients and 35 (35%) were male patients. Near similar observation (male 37% and female 63%) seen in study Omar et al and (male 40% and female 60%) in Raeissadat et al. More number of female patients in this current study may be due to that, females were more involved with household work which causes repetitive stress at the extensor carpi radialis brevis origin causing micro trauma, a relevant etiology for the initiation of the disease.

In this current study, out of the 100 participants, 85 (85%) participants had their right-side elbow affected and 15 (15%) had their left side affected. Out of the 100 participants, 90 (90%) participants had their Dominant elbow affected and 10 (10%) had their Non dominant elbow affected. In other two studies, one had 86% of the patients with their dominant elbow affected, while in another 80% of the patients with their dominant side affected. Parameters like age, sex, side of elbow involved, dominance of upper limb involved were comparable.

In this study the mean VAS score and PRTEE score before injection were comparable. Mean VAS score was 75 (SD-5.99) and mean PRTEE score was 79.52 (SD-5.56) Raeissadat et al. At 2 weeks follow-up, statistically significant (p value <0.0001) difference in VAS scoring and PRTEE was seen. Mean VAS score was 55.25 (SD-7.50) which was comparable with a study (Gautam et al.). Mean PRTEE score was 58.92 (SD-8.52).

At 4th week follow-up, there was statistically significant (p value <0.0001) decrease in VAS score and PRTEE score. Mean VAS score was 40.25 (SD-8.31) which was comparable (41.7±2.2) with a study Raeissadat et al. Mean PRTEE score was 32.77 (SD-7.99).

At 3 months follow-up, mean VAS score was 19.25 (SD-13.47) which was comparable (18±6) with a study (VK Gautam et al.). Mean VAS was 16 in Yadav et al and mean PRTEE score was 16.35 (SD-9.43) comparable (mean PRTEE-13) with Palacio et al, showed statistically significant (p value <0.0001) decrease in VAS score and PRTEE. 15 patients (15%) were completely free from pain, VAS score was 0. PRTEE score was 0 in 10 (10%) patients. At the end of 6 months there was no recurrence.

At the end of 6 months, 61 patients (61%) were completely relieved of pain and functional disability. 34 patients (34%) have mild pain. Mean VAS and PRTEE score were 6.05 and 5.63 respectively showed statistically significant (p value <0.0001). 5 patients follow-up were lost at 6 months. It was seen that there was a significant increase in post intervention pain for few days, 70 participants (70%) complained of increase of pain after local infiltration. This was also seen in other PRP studies Peerboom et al, Krogh et al.

In this current study it was seen that mean platelet concentration in whole blood was about 241.25±10^5/µl (SD-58.01) and mean platelet concentration in PRP was about 808.03±10^5/µl (SD-138.82) that is average 3-4 folds increase in platelet concentration over baseline. Another study had shown that clinical efficacy can be expected with a minimum increase in platelet concentration of 4 to 6 folds from whole blood baseline (200±10^5 platelets/µl). In this study the number of mean PRP injection was 2.8 (approximate 3).

CONCLUSION

Platelet rich plasma injection technique for lateral epicondylitis offers a better treatment with these advantages; (1) its application is minimally traumatic, (2) it has a reduced risk for immune mediated reactions anaphylaxis, devoid of potential complications such as hypoglycemia, skin atrophy, tendon tears associated with corticosteroid injection, (3) it is simple to acquire and prepare, easy to carry out as outpatient procedure and (4) it is inexpensive (5) better relief of pain, (6) low recurrence rate.

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Conflict of interest: None declared
Ethical approval: The study was approved by the institutional ethics committee

REFERENCES
