PERIPHERAL ENTHESOPATHY: AN OSTEOLOGICAL STUDY

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ABSTRACT

Introduction: Enthesis is the site of attachment of tendon or ligament to the bone. Enthesitis is characterized by erosive, inflammatory lesions that may undergo ossification. Ossified entheses are commonly described as enthesophytes. Enthesitis is a common clinical finding in spondyloarthropathies (SpA). Diffuse idiopathic skeletal hyperostosis (DISH) is another condition which is characterized by ossification of entheses, Entheseal ossifications are usually observed during our bone demonstrations to Medical students.

Objective: To describe the sites and frequency of ossification of entheses in the appendicular skeleton.

Materials and Methods: This is an observational study conducted by examining matured and intact bones of upper and lower limbs procured from the osteology section of the Department of Anatomy, VSS Institute of medical sciences and research to find out if there is any ossification of ligaments or tendons at their attachment sites.

Results: Entheseal ossifications were more common in bones of the lower limb in comparison to that of the upper limb. Distal parts of the lower limb were more involved by enthesial ossification compared to the proximal parts. Ligamentum patellii (33%), achillis tendon(32%) & plantar aponeurosis (27.5%) were the most common structures ossified and presented as bony spurs at their attachment sites to the calcaneum and tibial tuberosity. In the upper limb, most of the enthesial ossification sites were around elbow joint, like medial epicondylo, lateral epicondylo, radial tuberosity, olecranon process, and articular margin of elbow joint.

Conclusion: In addition to the usual sites of enthesial ossification described in previous imaging studies, many new sites of involvement were detected on gross examination of bones. This study can guide the clinicians and radiologists to focus on these unusual sites to correlate with their findings.

KEYWORDS: Enthesis, Enthesophytes, Spondyloarthropathy, DISH, Ossification, Spur.

INTRODUCTION

Enthesis is the site of attachment of tendon or ligament to the bone. There are two types of the enthesis, fibrous and fibrocartilaginous. Fibrous entheses are usually found at the site of attachment of massive muscle bodies to long bones which are mostly large areas of the diaphyses or metaphyses thus dissipating stress from tendons over bone tissue. Fibrocartilaginous entheses have more sophisticated architecture, attached to epiphyses, often in the vicinity of synovial joints. In addition to the dispersion of contractile force, they allow precise limb movement about the joints. Most of
the entheses in the body are fibrocartilaginous. These may incorporate uncalcified as well as calcified fibrocartilage, fat pads, bursae, and synovium in addition to the tendon, composed of dense fibrous connective tissue and adjacent bone [1,2].

Due to their location at the interface of ligaments and bones, entheses are more prone to mechanical forces. Both mechanical and inflammatory factors can trigger enthesitis[3,4].

Inflammation of enthesis or enthesitis is a common clinical finding in spondyloarthropathies (SpA). These are a group of chronic inflammatory disorders characterized by asymmetrical peripheral arthritis predominantly of lower limbs and axial skeletal inflammation. Enthesitis is characterized by erosive, inflammatory lesions that may undergo ossification [5,6].

Ossified entheses are commonly described as enthesophytes. Diffuse idiopathic skeletal hyperostosis (DISH) is another condition that is characterized by ossification of entheses, especially in the axial skeleton but also in peripheral entheses [7]. DISH is a systemic, relatively common condition, with an average prevalence of approximately 10% in people >50 years of age [7,9].

Most commonly enthesal ossifications are localized around the spine. Peripheral enthesopathic abnormalities are more frequently reported from the lower limb[8,9].

Capsules of synovial joints, ligamentum patellae, tendoachilis, and plantar fascia are described to be involved frequently[10]. These are detected by X-ray, Ultrasound, CT scan, or MRI in patients suffering from various types of spondyloarthropathies and DISH.

Enthesal ossifications were usually observed during our bone demonstrations to Medical students. In addition to the commonly described areas, ossification of entheses were observed by us in some unusual structures like interosseous membrane, intermuscular septa, transverse scapular ligament, costoclavicular ligament, etc. Though upper limb enthesophytes are rarely reported, those were also observed in many parts of upper limb bones. The bones we studied for enthesal ossifications were of an unknown source, so the etiology of enthesal ossifications were not clear. Therefore the present study was conducted to describe only the sites and frequency of bony proliferation at enthesal sites in the appendicular skeleton. As only imaging studies are available on this topic, an osteological study on dry bones may add some useful information to the existing knowledge of enthesopathy.

MATERIALS AND METHODS

All the intact matured bones of upper and lower limbs preserved in the osteology section of the Department of Anatomy, VSS Institute of medical sciences, and research were included in our study. This is an observational study conducted to find out the sites of attachments of ligaments and tendons where they have been ossified. We also calculated the percentage of involvement of these structures to describe their frequency.

RESULTS

Among lower limb bones, enthesophytes were found in all long bones including other bones like patella and calcaneum. Out of 152 femurs, ossification was found in 20 enthesal sites highest being the capsular ligament of the knee joint (5.2%) at the articular margin of condyles. 33% of ligamentum patellae at tibial tuberosity showed enthesal ossification among 77 tibias. Single fibula out of 58 had enthesophytes at malleolar fossa due to posterior talofibular ligament ossification. Patella showed enthesophytes at its base and apex due to the ossification of quadriceps tendon (12.5%) and ligamentum patellae (2.7%). Among short bones, only calcaneum had bony spurs at the attachment sites of tendoachilis (32%) and plantar aponeurosis (27.5%) [Table1]. Among upper limb bones, suprascapular ligaments (6%), costoclavicular ligaments (13.5%), capsule ligament of the shoulder (4%) and elbow joint (1%), common flexor (2%) and extensor tendons (1%) were found to be ossified at their attachment sites. Ossification of the interosseous membrane (11%) was observed at its radial attachment. In radius and ulna tendons of biceps and triceps were ossified at their sites of insertion respectively [Table 2].
**Fig. 1:** Enthesophytes at Articular margin of condyles of femur (a,b), Spiral line and medial lip of linea aspera(c), Tibial tuberosity (d), Soleal line(e).

**Fig. 2:** Enthesophytes at Malleolar fossa of fibula (a), Proximal surface of calcaneum(b), Medial process on the plantar surface of calcaneum (c), Apex of patella(d), base of patella (e).

**Fig. 3:** Enthesophytes at Inferior surface of the medial end of the clavicle(a), Suprascapular notch(b), Anatomical neck of humerus (c), Articular margin of lower end of the humerus(d, e), Lateral supracondylar ridge of the humerus(f), medial epicondyle of the humerus(g), Lateral epicondyle of the humerus(h), Radial tuberosity of radius (i), Interosseous border of the radius(j), Olecranon process of the ulna(k).

**Table 1:** Ossification of enthesis in lower limb bones.

<table>
<thead>
<tr>
<th>Name of the bone</th>
<th>Number of bones</th>
<th>Enthesis involved</th>
<th>Sites of attachment to the bone</th>
<th>No. of ossified enthesis</th>
<th>% of ossified enthesis</th>
</tr>
</thead>
<tbody>
<tr>
<td>Femur</td>
<td>152</td>
<td>Capsular ligament of hip joint</td>
<td>Intertrochanteric line</td>
<td>5</td>
<td>3.28</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Capsular ligament of knee joint</td>
<td>Articular margin of condyles</td>
<td>8</td>
<td>5.2</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Insertion of gluteus maximus</td>
<td>Gluteal tuberosity</td>
<td>2</td>
<td>1.3</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Medial intermuscular septum &amp; aponeurotic origin of vastus medialis</td>
<td>Spiral line and medial lip of linea aspera</td>
<td>5</td>
<td>3.2</td>
</tr>
<tr>
<td>Patella</td>
<td>72</td>
<td>Quadriceps tendon</td>
<td>Base of patella</td>
<td>7</td>
<td>12.5</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Ligamentum patellae</td>
<td>Apex of patella</td>
<td>2</td>
<td>2.7</td>
</tr>
<tr>
<td>Tibia</td>
<td>77</td>
<td>Ligamentum patellae</td>
<td>Tibial tuberosity</td>
<td>26</td>
<td>33</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Insertion of tendon of sartorius, gracilis &amp; semitendinosus</td>
<td>Upper end of medial surface of tibia</td>
<td>1</td>
<td>1.3</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Tendinous origin of soleus</td>
<td>Soleal line</td>
<td>4</td>
<td>5.4</td>
</tr>
<tr>
<td>Fibula</td>
<td>58</td>
<td>Posterior talofibular ligament</td>
<td>Malleolar fossa</td>
<td>1</td>
<td>1.7</td>
</tr>
<tr>
<td>Calcaneum</td>
<td>87</td>
<td>Tendoachillis</td>
<td>Proximal surface of calcaneum</td>
<td>28</td>
<td>32</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Plantar aponeurosis</td>
<td>Medial process on the plantar surface</td>
<td>24</td>
<td>27.5</td>
</tr>
</tbody>
</table>
Table 2: Ossification of enthesis in upper limb bones.

<table>
<thead>
<tr>
<th>Name of the bone</th>
<th>Total number of bones studied</th>
<th>Observation of ossified enthesis</th>
<th>Sites of attachment to the bone</th>
<th>No. of ossified enthesis</th>
<th>% of ossified enthesis</th>
</tr>
</thead>
<tbody>
<tr>
<td>Scapula</td>
<td>100</td>
<td>Transverse scapular ligament</td>
<td>Suprascapular notch</td>
<td>6</td>
<td>6</td>
</tr>
<tr>
<td>Clavicle</td>
<td>74</td>
<td>Costoclavicular ligament</td>
<td>Inferior surface of medial end of clavicle</td>
<td>10</td>
<td>13.5</td>
</tr>
<tr>
<td>Humerus</td>
<td>100</td>
<td>Capsular ligament of shoulder joint</td>
<td>Anatomical neck</td>
<td>4</td>
<td>4</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Capsular ligament of elbow joint</td>
<td>Articular margin</td>
<td>1</td>
<td>1</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Lateral intermuscular septum</td>
<td>Lateral supracondylar ridge</td>
<td>1</td>
<td>1</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Common extensor tendon</td>
<td>Lateral epicondyle</td>
<td>1</td>
<td>1</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Common flexor tendon</td>
<td>Medial epicondyle</td>
<td>2</td>
<td>2</td>
</tr>
<tr>
<td>Radius</td>
<td>90</td>
<td>Interosseous membrane</td>
<td>Interosseous border</td>
<td>10</td>
<td>11</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Insertion of tendon of biceps</td>
<td>Radial tuberosity</td>
<td>3</td>
<td>3.33</td>
</tr>
<tr>
<td>Ulna</td>
<td>80</td>
<td>Insertion of Triceps</td>
<td>Olecranon process</td>
<td>5</td>
<td>6.25</td>
</tr>
</tbody>
</table>

DISCUSSION

It is well known that plane radiography, ultrasound and MRI demonstrate soft tissue thickening, bone erosion, new bone proliferation, and bone structure alterations at inflamed entheses. But this osteological study focused only on the sites and frequency of bony proliferation at enthesal sites in the appendicular skeleton. Mechanical factors, physiological and anatomical characteristics of the enthesis, play an important role in the localization of enthesitis. In the course of normal joint function, strain levels at insertion sites maybe 4 times those in the tendon midsubstance [11]. So microtrauma and degenerative changes mostly affect the enthesis which is the principle of enthesitis development in spondyloarthopathy. This also explains the finding that most of the clinically relevant enthesitis sites are present in the lower limbs, particularly at the heel (Achilles enthesitis and Planter fasciitis).

Peggy Jacques et al have proved that enthesitis and new bone formation are driven by mechanical strain [12]. According to their study on animal model, the entheses are subjected to repetitive biomechanical stressing forces that are applied during normal muscle, ligament, and tendon action. In SpA patients, new bone formation often occurs in close relationship with the entheses. Development of spinal syndesmophytes along the anterior intervertebral ligaments, and bony spurs at the achilles tendon and plantar fascia are the results of this process. Subsequent bone erosion after inflammation appeared in anatomically distinct sites as compared with new bone formation at the achilles’ enthesis. Erosions preferentially developed in regions undergoing compression, whereas spur formation occurred in regions prone to tensile forces. Baraka et al [13] in their study on ultrasonographic evaluation of lower limb enthesopathy in ankylosing spondylitis and Behçet’s disease have reported that achillis tendon was most commonly affected followed by plantar aponeurosis. In both studied groups, the most commonly affected enthesal site was the achilles tendon insertion which was found to be affected in (30.5%) sites in group I and (41.5%) in group II, followed by the plantar aponeurosis attachment which was found to be affected in (27.78%) sites of group I and 23/65 (35.38%) in group II where the group I & II represent cases of ankylosing spondylitis and Behcet’s disease respectively. Our osteological study is consistent with the findings of this study with 32% in achillis tendon and 27.5% in plantar aponeurosis involvement.

Tibial tuberosity enthesophyte was the least frequent to be detected in their study. According to them several anatomical factors such as bone widening and sharp changes in the fiber orientation can lead to an anisotropic misleading artifact. But in the present study, tibial tuberosity enthesophyte was found in 33% cases which were higher as compared to other sites.

P V Balint et al [14] in an Ultrasonographic study of lower limb enthesis in SpA patients have reported that enthesophyte formation (3-30%) was the most common USG bony abnormality of
heel, patella were mainly involved. Enthesopathy had an increased predictive value in diagnosing DISH [21,22]. Common sites of hyperostotic spurs were at the olecranon, lateral and medial epicondyle. The formation of these spurs is influenced by mechanical factors like physical activities, handedness, and sex. [23]. In the present study also hyperostotic spurs were mostly present around the elbow.

Common flexor tendon, common extensor tendon, tendon of biceps, tendon of triceps, and capsular ligament of elbow joint were ossified at their attachment sites.

Though we found bony spurs in olecranon (6.25%), lateral (1%) and medial (2%) epicondyles they were less frequent as compared to costoclavicular ligaments (13.5%), interosseous membrane (11%) and suprascapular ligaments (6%). Interestingly enthesophytic changes in the above structures have not been reported by earlier authors.

In the present study, we observed significant osteophytic changes along the medial lip of linea aspera in the femur and soleal line of the tibia which are the site of attachments of vastus medialis and soleus respectively. As described earlier, these types of entheses come under a different category called fibrous entheses which are usually associated with massive muscle bodies that attach directly to large areas of the diaphyses or metaphyses of long bones and dissipate stress from tendons over bone tissue [1]. We did not find a report on enthesopathy in these areas in any other literature.

Compared to lower limb enthesopathy less number of studies and reports are found on upper limb enthesopathy. Wes Cormic has mentioned about supraspinatus tendon enthesopathy, lateral epicondylar enthesopathy [18]. Resnick et al. [19] showed that the majority of the patients with DISH also showed clinical and radiographic abnormalities at locations other than the classical osteoarthritic sites including elbows, wrists, and shoulders. Littlejohn JO et al [20] have reported about hyperostosis on olecranon though the tibial spine, entheseal sites while bony erosion was seen at 1–13% of sites. They observed ossification of entheses in 8/70 quadriceps, 7/69 proximal patellar ligament, 2/69 distal patellar ligament, 21/70 achillis tendon, 4/70 plantar aponeurosis. In contrast, our study showed more number of enthesophytes formation of the distal patellar ligament (26/77) and plantar aponeurosis (24/87).

Antonio Spadaro et al [15] in an imaging study of peripheral enthesitis in ankylosing spondylitis have reported that the distal portion of the lower limbs is more frequently affected by enthesopathic abnormalities compared with the proximal part of the lower limb. Similar results were reported by Lenthinen et al [16], Kiris et al. [17] and Wes Cormic [18]. All the above studies show that patellar tendon, achillis tendon, and plantar fascia are more commonly affected compared to the attachment of adductor muscles or gluteal muscles, which is consistent with our result. The reasons for the predilection of the distal part of lower limbs by the enthusiastic process is unknown, but anatomic and physiological factors, such as the major length of the tendon, may play a role [15,19].

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