Silicone lymphadenopathy mimicking a lymphoma in a patient with a metatarsophalangeal joint prosthesis

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Abstract
With lymph node enlargement, the possibility of a malignant process such as metastatic carcinoma or lymphoma needs to be excluded. This report describes a 47 year old woman with inguinal lymph node enlargement initially suspicious for lymphoma. Fine needle aspiration findings favoured reactive hyperplasia, but a malignant process could not be excluded. The final histological diagnosis was a foreign body granulomatous inflammatory response as a result of regionally disseminated silicone particles from an overlooked metatarsophalangeal joint prosthesis. Because of the large number of joint prostheses worldwide, it should be kept in mind that migration of wear particles can create granulomatous inflammation and node enlargement.

Keywords: lymphadenopathy; silicone; prosthesis

Although complications associated with the use of joint prostheses are recognised in the orthopaedic literature, they are less well recognised by physicians in other specialties. The production of wear debris after arthroplasty has been widely reported and inflammatory responses to biomaterials are still described. Metal, silicone elastomers, and polyethylene are the biomaterials most commonly implicated. In tissues found adjacent to implants there is often a histiocytic reaction. Wear debris commonly disseminates to lymph nodes, whereas systemic dissemination with granulomatous reaction is less frequent. The same lymph node may harbour both granulomatous inflammation, as a result of wear particles, and metastatic carcinoma or lymphoma. Our present case draws attention to the fact that the possibility of lymph node enlargement a result of silicone wear particles from a 15 year old arthroplasty was not elicited in the initial evaluation for lymphadenopathy.

Case report
A 47 year old woman presented in October 1997 with left inguinal lymph node enlargement. She denied any fevers, chills, or other systemic symptoms. The only finding on physical examination was an enlarged, non-inflamed, mobile lymph node. Since 1992, she had been receiving gold salts and non-steroid anti-inflammatory drugs for rheumatoid arthritis. She underwent a fine needle aspiration of the enlarged lymph node and reactive hyperplasia was favoured, but a malignant process could not be excluded. One year later, she noted an increase in the size of the lymph node. The lymph node was surgically excised and submitted for pathological evaluation.

The specimen consisted of a lymph node, approximately 25 mm in diameter, with a light tan appearance and a homogeneous consistency on cut section. The specimen was separated into several portions. One segment was fixed in formalin, embedded in paraffin wax, and processed for light microscopy and immunohistochemistry. Another portion was frozen. Another segment was processed for cytogenetic and flow cytometry evaluation. The formalin fixed sections were stained with haematoxylin erythrosin safron (HES), Giemsa, periodic acid Schiff, Perls, and for CD20, CD3 and BCL-2 (all from Dako, Glostrup, Denmark) using the streptavidin–biotin method. The frozen sections were stained with oil red O. There was no abnormality on cytogenetic examination and no monoclonality detected in flow cytometry. The histological examination showed follicular hyperplasia and numerous multinucleated foreign body giant cells within sinuses and interfollicular spaces (fig 1). The immunohistochemistry study
showed normal compartmentalisation of B cells (mainly in the follicular centre) and T cells (in the interfollicular space). BCL-2 immunostaining showed negativity of the B cell population within the hyperplasic follicles. Many of the giant cells contained irregular particles of refractile material as well as asteroid bodies (fig 2). The particles showed a mean diameter of 24 µm; the largest measured particle was 40 µm in diameter. These particles were not doubly refractile under polarised light and special stains were negative. The histological appearance of these particles was suggestive of silicone, so we queried the patient about this possibility. She related that she had a left metatarsophalangeal joint prosthesis, which was placed 15 years ago for a bone deformity (hallux valgus). It was a Swanson silastic prosthesis, for which plain film radiographs did not reveal any evidence of damage.

**Discussion**

Silicone elastomer arthroplasties for joint diseases involving the small joints of the hands and feet have been used worldwide since the late 1960s. Medical silicones, or polydimethylsiloxane, can be formulated in different physical forms of silicone applied in various ways, resulting in materials of varying structures and consistencies, which can be individually tailored for different body parts. For example, elastomers are used for joint prostheses, gel for breast implants, and liquid for soft tissue injection. As in joint prostheses, a local granulomatous reaction can be encountered for the different physical forms of silicone applied in medicine. In fact, such a reaction is found in breast and testicular implants as well as in subcutaneous injections. In such cases, it is important to realise that silicone leakage can migrate to a location far from the site of application and can lead to—for example, regional lymphadenopathy, which might be clinically relevant. As in the few cases reported previously, in our present case, the adenopathy was initially thought to be caused by malignancy or infection. Further investigation disclosed that foreign material might be the cause.

The frequency of lymphadenopathy after this common procedure is still difficult to establish. In a study of 23 cases of silastic arthroplasty it was shown that silicone lymphadenopathy may be a frequent complication (13%). However, in a review of the literature regarding long term complications of 15 556 small joint implants, lymphadenopathy as a result of wear debris was reported in only 13 cases (0.08%).

A history of previous silastic arthroplasty might not always be available to the clinician (much less to the pathologist), but the histological appearance is sufficiently characteristic to prompt an appropriate inquiry. The chemical identity can be confirmed by infrared imaging spectroscopy or by electron microscopy coupled with energy dispersive x-ray analysis. This case emphasises the long latency period that can be associated with this clinical phenomenon, which may mimic other, more serious, diseases. Fine needle aspiration cytology can be useful in the diagnosis of such lymphadenopathy, if it shows a foreign body giant cell reaction to particulate matter morphologically compatible with silicone elastomer. In our case, retrospective review of the original cytology slides failed to identify any pronounced pathology. With currently increasing numbers of silastic joint arthroplasties performed for rheumatoid arthritis or bone deformity, silicone lymphadenopathy should always be considered in the differential diagnosis of node enlargement. Uncommonly, there may be an associated malignant process. A review of the reports of concurrent malignant tumour cells and exogenous silicone particles in the same lymph node has shown that lymphoma is usually found with particles from joint prostheses and carcinoma with particles from breast implants. All of the patients with both joint prosthesis related lymphadenopathy and non-Hodgkin’s lymphoma had longstanding rheumatoid arthritis. Patients with rheumatoid arthritis are at an increased risk of developing malignant lymphoma, compared with the general population. They form the population most likely to be treated with Swanson joint implants. Therefore, the presence of silicone joint implants in patients with rheumatoid arthritis and lymphoma might be a concomitant rather than a causal finding. Similarly, follicular hyperplasia, as initially seen in the present case, is more likely to be a manifestation of rheumatoid arthritis than a response to foreign material.

The long term outlook in terms of foreign body reaction is reported to be very good, but the first reported case of silicone lymphadenopathy associated with a finger joint implant had a recurrence in an ipsilateral node four years later. Therefore, it is recommended that prostheses should be replaced if they result in pronounced lymphadenopathy as a result of migrating wear debris.

Contents

Review

111 Demystified ... Monoclonal antibodies

P N Nelson, G M Reynolds, E E Waldron, E Ward, K Giannopoulos, P G Murray

Papers

118 Genetic relation of lobular carcinoma in situ, ductal carcinoma in situ, and associated invasive carcinoma of the breast

H Buerger, R Simon, K-L Schäfer, R Diallo, R Littmann, C Poremba, P J van Diest, B Dockhorn-Drews, W Böcker

122 Expression of π-class glutathione S-transferase: two populations of high grade prostatic intraepithelial neoplasia with different relations to carcinoma

R Montironi, R Mazzucchelli, D Stramazzotti, R Pomante, D Thompson, P H Bartels

129 Interphase argyrophilic nucleolar organiser regions and nucleolar counts in transitional cell bladder tumours

I A Korneyev, N N Mamaev, V V Kozlov, M G Rybakova, S H Al-Shukri

133 Telomerase activity in pancreatic endocrine tumours: a potential marker for malignancy

K Y Lam, C Y Lo, S T Fan, J M Luk

137 Role of chromosome 3p12–p21 tumour suppressor genes in clear cell renal cell carcinoma: analysis of VHL dependent and VHL independent pathways of tumorigenesis

A Martinez, P Fullwood, K Kondo, T Kishida, M Yao, E R Maher, F Latif

145 Expression of the matrix metalloproteinase 9 in Hodgkin’s disease is independent of EBV status

J R Flavell, K R N Baumforth, D M Williams, M Lukesova, J Madarova, V Noskova, J Prochazkova, D Lowe, Z Kolaj, P G Murray, P N Nelson

150 Detection of clonal T cell populations by high resolution PCR using fluorescently labelled nucleotides; evaluation using conventional LSS-SSCP

C L Wickham, C Lymans, S Ellard

155 Immunoblotting can help the diagnosis of ocular toxoplasmosis

D O Ho-Yen, D J Chapman, D Ashburn

Short report

159 Selective genetic analysis of p53 immunostain positive cells

M Phelps, B S Wilkins, D B Jones

Miscellanea

162 Correspondence • Book reviews • Notice