ASSESSMENT OF CERVICAL LYMPHADENOPATHY BY ULTRASONOGRAPHY AND COLOUR DOPPLER

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BACKGROUND

Grey scale ultrasonography can be used as a first line imaging modality along with histopathology for evaluation of cervical lymphadenopathy. Morphologic features like size, shape, hilum and necrosis within the lymph node and vascularity are analysed. We wanted to identify the role of Grey scale ultrasonography and colour Doppler in characterisation of enlarged cervical lymph nodes, evaluate findings in benign and malignant cervical lymphadenopathy and assess the sensitivity, specificity, predictive value, likelihood ratios and accuracy in malignant and benign cervical lymph node enlargement.

METHODS

This was a cross sectional study with diagnostic test evaluation. All suspected cases of cervical lymphadenopathy referred for sonological evaluation to the Department of Radiodiagnosis of Vinayaka Mission's Medical College were included in the study. About 75 patients included in the study, underwent Grey scale and colour Doppler sonography of cervical lymphadenopathy.

RESULTS

When three criteria were positive for malignancy, the sensitivity and specificity of grey scale ultrasonography and colour Doppler were 100 % and 73.80 %, and when four criteria were positive, the sensitivity and specificity were 72.73 %, and 90.48 % respectively. When we analysed individual variables, status of hilum (absence / presence) has reasonably good sensitivity and specificity, positive predictive value, negative predictive value and accuracy (90.09 %, 92.90 %, 90.09 %, 92.90 % and 92.0 % respectively). In our study the important criteria positive for malignant lymph nodes were size > / 11 mm in short axis diameter, round shape, absent hilum and peripheral and central + peripheral type of vascularity of cervical lymph nodes. For benign lymph nodes, oval shape, presence of hilum and central vascularity were the important criteria noted.

CONCLUSIONS

Grey scale ultrasonography can be complemented by colour Doppler ultrasonography (USG) in patients presenting with cervical lymphadenopathy to differentiate between benign and malignant lesions by reducing unnecessary biopsies by more accurate discrimination.

Introduction

Lymphadenopathy is an abnormality in the size or character of lymph nodes caused by the invasion or propagation of inflammatory or neoplastic cells to the lymph nodes. It usually presents as enlargement of lymph nodes and are very common in developing countries like India. The lymph nodes affected are abnormal in size, consistency or number. It may be

generalised or localised involving specific groups of lymph nodes. Causes for cervical lymphadenopathy may vary. They include primary or secondary malignant neoplasms of lymph node, bacterial, viral, fungal and protozoal infections, auto immune conditions or drugs. Majority of the patients may have benign lymphoid hyperplasia or reactive lymphadenitis. Lymphadenopathy may be the only clinical finding or one of the several nonspecific findings in some patients. Early detection of malignancy is of great clinical significance. It improves the patient survival by proper surgical and radiation treatment. Pretreatment staging and therapeutic planning in patients with primary malignant tumours is possible with sonographical evaluation. Biopsy and other pathological tests are invasive and time consuming.

But, Ultrasonography is easily available, cost effective, non-invasive, radiation free and safe investigation.1 For assessing cervical lymph node, sonography is a useful imaging tool. It is a non-invasive, non-ionising imaging technique. This Technique with the help of ultrasound waves provides valuable diagnostic information with a high diagnostic accuracy. The conventional ultra sound uses a grey scale to detect the various anatomical structures of body and study their morphological characteristics. Even small blood vessels in lymph nodes can be identified with ultrasonography. Grey scale sonography is widely used in the evaluation of number, size, site, shape, borders, matting, adjacent soft tissue oedema and internal architecture of cervical lymph node. The differentiation of benign from malignant lymphadenopathy is of critical importance in both treatment planning and prognosis.2 The use of higher frequency sonic wave's results in an improvement in spatial and contrast resolution and also shows the internal architecture of lymph nodes.3 This readily identifies normal, inflammatory and malignant lymph node.4,5 Colour Doppler can show flow in all the lymph nodes including benign and malignant lesions of lymph node. The analysis of patterns of nodal vascularity helps to differentiate benign from malignant lesions.

Transducer and the equipment used determines the ability to detect a colour flow pattern within a small lymph node. High frequency transducer wave has the capacity to detect superficial low velocity signals. Vessels in inflamed lymph nodes are dilated. But vessels in lymph nodes affected with metastasis may be compressed by tumour cells. This can be easily detected with the help of colour Doppler. Presence of extra hilar blood vessels is another sign of malignancy in the sonographic diagnosis of lymph node. Presence of calcification and necrosis is also more common in malignant lymph nodes. So, in our study, size, shape, border, echogenicity, hilum, necrosis, calcification and matted / discrete nature of lymph nodes are studied for distinguishing benign and malignant lymph nodes by sonological methods.6,7 This study has been conducted to evaluate the efficacy of ultrasonography and colour Doppler to differentiate benign from malignant cervical lymphadenopathy.

Materials and Methods

Our study is a cross sectional study with diagnostic test evaluation. It was conducted at the Department of Radio diagnosis, Vinayaka Mission's Medical College. Sampling Technique Patients with suspected cervical lymphadenopathy from other departments (Surgery, General Medicine) referred to the Department of Radiodiagnosis, for sonographic evaluation of neck were included for the study.

Cervical lymph nodes measuring 5 mm or more than 5 mm in long axis diameter by ultrasonography were included for the study in patients from 5 to 80 years. Patients not giving consent and in whom follow up lost were excluded from the study.8,9 By using the formula, $N = (Z\alpha)2 \times pq / d2$ one would get the total number of positive cases (n) which is required for the study. Where $Z\alpha$ is 1.96 at 5 % α . P is expected sensitivity, q is (100 - p), d is precision, which can be any value between 5 to 20 % of p (taking as 10 % of P in this study). Hence the total number of positive cases, $N = (1.96)2 \times 87.5 \times 12.5 / 122 = 29.37$ approximately 30 cases. From the cases referred to the Department of Radio diagnosis, On examining records, an average of 40 % prevalence was noted. Hence the total sample size (N) would be equal to total number of positive cases (N) required for the study divided by prevalence (P).10 Sample size $N = n / P = 30 \times 100 / 40 = 75$. C

ervical lymph nodes are characterised on grey scale by the shape, size, border, hilum and calcification of the lymph nodes and their vascular pattern on Colour Doppler.11,12 All lymph nodes were evaluated by histopathology to differentiate malignant or benign.13,14 Data was collected from a total of 75 cases referred for the ultrasonography of neck. All sonographic examinations were done in GE Logic F Series ultra-sonographic and Colour Doppler equipment with a linear array high frequency (7 - 13 MHz) transducer. Data from all histopathological evaluations were collected.15,16 Data was entered in Microsoft excel.

Statistical Analysis

Analysis of the data was done in SPSS version 16.0 and DAG_Stat (Diagnostic & Agreement Statistics analysis package). Frequencies of variables, sensitivity, specificity, positive predictive value, negative predictive values were calculated.

Results

Grey Scale Ultra Sonographic Findings

Analysis of Cervical Lymph Node by Size Test positive criteria: size of the cervical lymph node (short axis diameter) > / = 10 mm. Out of 75 lymph nodes, 30 nodes (40 %) were > 10 mm in short axis diameter and 45 (60 %) < 10 mm in short axis diameter. Out of the 33 (100 %) malignant lymph nodes, 27 (81.20 %) were > / = 10 mm in short axis diameter and 6 (18.80 %) were < 10 mm in short axis diameter. Out of 42 (100 %) benign lymph nodes, 39 (92.85 %) lymph nodes were < 10 mm in short axis diameter and 3 lymph nodes (7.15 %) were > / = 10 mm. Statistical indices showing comparison of size of lymph nodes with histopathology (Malignant Vs Benign) showed the following results. By comparing the size of the lymph node, the sensitivity to differentiate benign and malignant and benign lymph nodes were 81.8. Specificity was 92.9. False Negative was 18.2 and false positive was 7.1. Positive predictive value was 90. Negative Predictive Value was 86.7. Positive Likelihood ratio was 11.5. Negative Likelihood ratio was 0.2. Accuracy of the test was 88. Analysis of Cervical Lymph Node by Shape Shape of the cervical lymph node correlated with histopathology (Malignant vs. Benign) using test positive criteria as round shape. Out of the 33 malignant lymph nodes, 28 (84.84 %) were round in shape and 5 (15.12 %) were oval in shape. Out of the 42 benign lymph nodes, 37 (90.1 %) were oval in shape and 5 (11.90 %) were round in shape. Statistical indices showing comparison of shape of lymph node with histopathology (Malignant vs. Benign) were done. Sensitivity of the test was 84.8. Specificity

was 88.1. False negative was 15.2. False positive was 11.9. Positive Predictive value was 84.8. Negative Predictive value was 88.1. Positive Likelihood ratio was 7.1. Negative Likelihood ratio was 0.2. Accuracy of the test was 86.7.

Analysis of Cervical Lymph Node by Border

Test positive criteria for analysis of cervical lymph node was irregular border of the lymph node. Out of 33 malignant nodes, 14 cases (42.42 %) were with irregular borders and 19 cases (57.58 %) were with regular borders. Out of 42 benign nodes 40 (95.40 %) were with regular borders and 2 cases (4.60 %) were with irregular borders. Positive Predictive value was 84.8. Negative Predictive value was 88.1. Accuracy was 86.7.

Analysis of Cervical Lymph Node by Hilum

Out of 33 malignant lymph nodes, 30 cases (90.90 %) showed absent hilum and 3 cases (9.1 %) showed preserved hilum. Out of 42 benign nodes 39 Cases (92.90 %) showed presence of hilum and 3 cases (7.1 %) showed absence of hilum. Test positive criteria is absent hilum of lymph node. Sensitivity was 90.9. Specificity was 92.9. False Negative was 9.1. False positive was 7.1. Positive predictive value was 90.9. Negative predictive value was 92.9. Positive Likelihood ratio was 12.7. Negative Likelihood ratio was 0.1 Accuracy was 92. Analysis of Cervical Lymph Node by Necrosis Test positive criteria was presence of necrosis in lymph node. Out of 33 cases of malignant cervical lymph nodes 14 cases (42.42 %) were with necrosis and 19 (57.58 %) were without necrosis. Out of 42 lymph nodes, only 4 cases (9.58 %) showed presence of necrosis. 38 cases (90.42 %) were without necrosis. Sensitivity was 42.4. Specificity was 90.5. False Negative was 57.6. False positive was 9.5. Positive predictive value was 77.8 negative predictive value was 66.7. Positive Likelihood ratio 4.5. Negative Likelihood ratio was 0.6. Accuracy was 69.3.

Analysis of Cervical Lymph Node by Hypoechogenecity

Test positive criteria was hypoechogenecity of lymph node. Out of 33 malignant cases, 30 (90.9 %) cases were hypoechoic and 3 (9.1 %) cases were hetero-hypoechoic. Out of 42 benign cases, 41 (97.38 %) cases were hypoechoic and 1 (2.62 %) cases were hetero-hypoechoic. Sensitivity of the test was 90.9. Specificity was 2.4. False negative was 9.1. False positive was 97.6. Positive predictive value was 42.3. Negative predictive value was 25. Positive likelihood ratio was 0.9. Negative likelihood ratio was 3.8. Accuracy was 41.3.

Analysis of Cervical Lymph Nodes by Colour Doppler

Sonographic Findings Among 33 malignant lymph nodes 14 cases (42.42 %) were with peripheral vascularity and 19 cases (48.58 %) with peripheral and central vascularity. Only 3 cases (9.09 %) showed central vascularity. Among 42 benign nodes 34 cases (80.95 %) showed central vascularity and 8 (19.04 %) cases showed peripheral and central vascularity. Test positive criteria was peripheral / Central + peripheral vascularity of lymph node. Sensitivity was 90.9. Specificity was 81. False negative was 9.1. False positive was 19. Positive predictive value was 78.9. Negative predictive value was 91.9. Positive likely ratio was 4.8. Negative likely ratio was 0.1. Accuracy was 85.3.

CONCLUSIONS

Grey scale ultrasonography and colour Doppler have a sensitivity and specificity of 100 % and 73.80 % respectively, in detecting malignancy in cervical lymphadenopathy when three

criteria are positive. As the number of criteria increases the specificity increases, but sensitivity decreases. When analysing individual variables, absence of hilum has got a good sensitivity and specificity, positive predictive value, negative predictive value and accuracy. In our study, the most important criteria positive for malignant lymph nodes were size of lymph node > / 11 mm in short axis diameter, round shape, absent hilum and peripheral and central+ peripheral type of vascularity of cervical lymph nodes.

References

- 1. Baatenburg De Jong RJ, Rongen RJ, Lameris JS, Harthoorn M, Verwoerd CD, Knegt P. Metastatic neck disease. Palpation vs ultrasound examination. Arch Otolaryngol Head Neck Surg 1989; 115:689–690. Back to cited text no. 1
- 2. Zhao RN, Zhang B, Jiang Y ×. Sonographic evaluation of metastatic cervical lymph nodes. Zhongguo Yi Xue Ke Xue Yuan Xue Bao 2012; 34:633–639. Back to cited text no. 2
- 3. Gupta A, Rahman K, Shahid M. Sonographic assessment of cervical lymphadenopathy: role of high resolution and color Doppler imaging. Head Neck 2011; 33:297–302. Back to cited text no.
- 4. Lyshchik A, Higashi T, Asato R, Tanaka S, Ito J, Hiraoka M. Cervical lymph node metastases: diagnosis at sonoelastography initial. Radiology 2007; 243:258–267. Back to cited text no. 4
- 5. Ying M, Pang BSF. Three dimensional ultrasound measurement of cervical lymph node volume. Br J Radiol 2009; 82:617–625. Back to cited text no. 5
- 6. Ying M, Ahuja A, Yuen HY. Grey-scale and power Doppler sonography of unusual cervical lymphadenopathy. Radiol Source 2004; 30:449–454. Back to cited text no. 6
- 7. Sakai F, Kiyono K, Sone S, Kondo Y, Oguchi M, Watanabe T, et al. Ultrasonic evaluation of cervical metastatic lymphadenopathy. J Ultrasound Med 1988; 7:305–310. Back to cited text no. 7
- 8. Kaddah AM, Hegazy R. Value of gray-scale and color Doppler sonography in the evaluation of cervical lymphadenopathy. Med J Cairo Univ 2011; 79:59–65. Back to cited text no. 8
 - Sathyanarayan V, Siva Bharani K. Enlarged lymph nodes in head and neck cancer: analysis with triplex ultrasonography. Ann Maxillofac Surg 2013; 3:35–39. Back to cited text no.
- 9. Ahuja AT, Ying M, Ho SY. Ultrasound of malignant cervical lymph nodes. Cancer Imaging 2008; 8:48–56. Back to cited text no. 10
- 10. Moritz D, Ludwig A, Oestmann W. Color Doppler sonography for differential diagnosis of enlarged cervical lymph nodes. Am J Roentgenol 2000; 174:1279–1284. Back to cited text no. 11
- 11. Chan JM, Shin LK, Jeffrey RB. Ultrasonography of abnormal neck lymph nodes. Ultrasound Q 2007; 23:47–54. Back to cited text no. 12
- 12. Rosario PW, de Faria S, Bicalho L. Ultrasonographic differentiation between metastatic and benign lymph nodes in patients with papillary thyroid carcinoma. J Ultrasound Med 2005; 24:1385–1389. Back to cited text no. 13

- 13. Ahuja AT, Ying M. Sonographic evaluation of cervical lymph nodes. Am J Roentgenol 2005; 184:1691–1699. Back to cited text no. 14
- 14. Gor DM, Langer JE, Loevner LA. Imaging of cervical lymph nodes in head and neck cancer: the basics. Radiol Clin North Am 2006; 44:101–110. Back to cited text no. 15