

Gastrointestinal lymphomas of mucosa associated lymphoid tissue (MALT) – a report of 7 Sri Lankan cases

M V D de Silva *, M S Fernando **, N Ratnatunga ***

Journal of the Ceylon College of Physicians, 1998, 31; 1 & 2, 58-61

Abstract

Introduction: Lymphoma of mucosa associated lymphoid tissue (MALT) is a distinct recently described subtype with a good prognosis. It is often confused with other variants of Non Hodgkin's lymphoma (NHL) with a poor prognosis.

Objective: 1. To determine the proportion of MALT type lymphoma in gastrointestinal lymphomas previously diagnosed as NHL. 2. To determine if low grade MALT lymphomas are associated with high grade lymphomas.

Method: Histology slides of nine NHL of the stomach, ileum and colon, diagnosed between 1987-1995 at the University Department of Pathology, Peradeniya were reviewed for diagnostic criteria of MALT type lymphoma.

Results: Seven tumours were subtyped as low grade MALT lymphomas. Four of these showed coexistent high grade large cell lymphoma.

Conclusion: 77.7% of gastrointestinal lymphomas were of MALT type. The coexistence of high grade lymphoma in 4 cases indicates that MALT lymphomas can transform into large cell lymphomas.

Introduction

Lymphomas of mucosa associated lymphoid tissue (MALT) were first described by Isaacson and Wright in 1983¹ as a distinctive type of B-cell Non Hodgkin's Lymphoma (NHL) involving the gastrointestinal tract (GIT). However all GIT lymphomas

are not of MALT type. They could be large cell or large cell immunoblastic lymphomas, mantle cell lymphomas, small lymphocytic lymphomas or T cell lymphomas². Recognition of MALT lymphomas as a separate subtype of NHL is important because they remain localized at the site of origin for a long time, tend to relapse in the same or other mucosal sites and if localized may be cured^{3,4}.

Most pathologists in Sri Lanka use the international working formulation to classify NHL⁵. In this classification there is no category named MALT lymphoma. Thus the objective of this retrospective study was to determine the proportion of MALT type lymphoma in GIT lymphomas previously diagnosed as other categories of NHL. We also wanted to determine if low grade MALT lymphomas were associated with high grade lymphomas.

Method

A total of nine gastrointestinal lymphomas had been reported at the Department of Pathology, University of Peradeniya during a 9 year period between 1987-1995. The slides which had been stained with haematoxylin and eosin were reviewed for features of low grade MALT lymphoma.

Results

Seven tumours were subtyped as low grade MALT lymphomas. Four of these showed coexistent high grade large cell lymphoma. Two cases were high grade NHL without evidence of coexistent MALT lymphoma. The clinical features of the 7 cases of MALT lymphoma are shown in Table 1. Follow up data of patients is not known as this was a retrospective study.

All cases showed diagnostic criteria for low grade MALT lymphoma which included the presence of lymphoepithelial lesions (Figure 1), reactive follicles and a mixed diffuse population (Figure 2) of centrocyte like cells, small round lymphocytes,

* Senior Lecturer

** Lecturer, Department of Pathology, Faculty of Medicine, Colombo.

*** Associate Professor, Department of Pathology, Faculty of Medicine, Peradeniya.

Table 1
Clinical Features of 7 patients with MALT lymphoma

Case No:	Age (years)	Sex	Site	Mesenteric nodes	Microscopy	Depth of penetration
1	56	Female	Terminal ileum	Enlarged	High + low grade MALT lymphoma	Confined to bowel wall,
2	57	Female	Ascending colon	Enlarged	High + low grade MALT lymphoma	Extended to mesenteric fat
3	55	Male	Pylorus of stomach	Enlarged	High + low grade MALT lymphoma	Extended to mesenteric fat
4	33	Male	Ileum	Not enlarged	High + low grade MALT lymphoma	Penetrated the muscularis propria and extended to the serosal surface
5	50	Male	Stomach	Not enlarged	low grade MALT lymphoma	Penetrated the muscularis propria and extended to the serosal surface
6	Not known	Not known	Ileum	Not enlarged	low grade MALT lymphoma	Penetrated the muscularis propria and extended to the serosal surface
7	10	Male	Colon	Not enlarged	low grade MALT lymphoma	Confined to bowel wall

monocytoid cells and plasma cells^{2,6}. In the 4 cases where there was co-existent high grade lymphoma, the low grade MALT lymphoma component was situated at the periphery with the high grade lymphoma occupying the central part of the tumour mass or ulcer. There was marked destruction of glands in the high grade areas and lymphoepithelial lesions were absent. The high grade component was composed of lymphocytes with larger nuclei, clumped chromatin and frequent mitotic figures (Figure 3). In all cases where the mesenteric lymph nodes were enlarged they were effaced by a high grade large cell lymphoma similar in appearance to that in the GIT. Low grade MALT lymphomas without a high grade component were not associated with enlarged mesenteric lymph nodes.

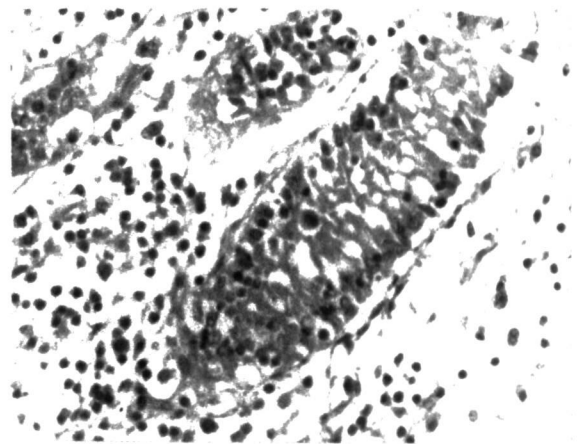


Figure 1. Lymphoepithelial lesion; infiltration of mucosal glands by lymphocytes (haematoxylin and eosin $\times 400$).

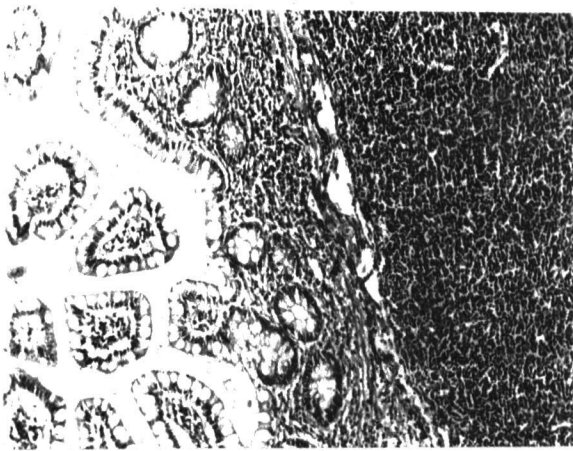


Figure 2. Low grade MALT lymphoma; diffuse infiltration of mucosa and submucosa by lymphocytes (haematoxylin and eosin $\times 100$).

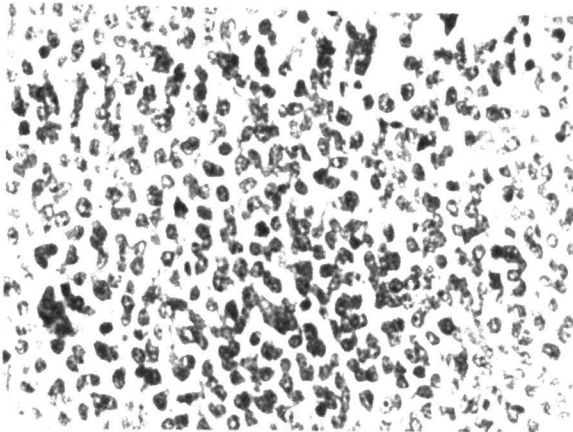


Figure 3. High grade area in MALT lymphoma (haematoxylin and eosin $\times 400$).

Discussion

The clinical, behavioural and pathological features of MALT lymphomas are sufficiently distinctive for them to be considered as a specific sub type of NHL. The concept of lymphomas of MALT has been currently expanded to include several extranodal sites^{4,7}. Most extranodal MALT lymphomas arise in sites without normal MALT such as the stomach, salivary glands and thyroid^{6,8}. This is explained by the proposal that they arise in a setting of "acquired MALT". This nonindigenous lymphoid tissue is acquired secondary to an infection such as *Helicobacter pylori* in the stomach⁹ or an

autoimmune disease such as Sjogren's syndrome in salivary glands or Hashimoto's disease in the thyroid⁶. The growth pattern and immunophenotype of the centrocyte like cells in MALT lymphoma suggest that they may represent neoplastic marginal zone cells⁸. The tendency of MALT lymphomas to evolve slowly, remain localized and to involve other mucosal sites when they do spread, may be due to specific homing patterns of MALT derived lymphocytes⁴. MALT lymphomas are consistently CD 10 negative and fail to show rearrangement of the *bcl-2* proto-oncogene both of which are characteristic features of follicular center cell lymphomas³. Similarly MALT lymphomas are CD 5 negative and fail to show *bcl-1* gene rearrangement which helps to distinguish them from mantle cell lymphomas^{3,10}.

The association of high grade lymphoma and low grade MALT lymphoma as seen in 4 of our cases has been described previously by Chan and co-workers who found 10 cases with this association in a series of 48 gastric lymphomas¹¹. They suggested that the likelihood of finding this association may increase with the number of sections examined. In their series the immunophenotype of the low and high grade components were similar providing strong evidence that the two components evolve from the same clone. Both high and low grade MALT lymphomas show a favourable behaviour when compared with nodal lymphomas^{8,12}.

Early gastric MALT lymphomas are known to regress with treatment for *Helicobacter pylori*¹³. Local surgery has been adopted as a potentially curative therapeutic modality for MALT lymphoma. However local relapse has been reported often after a long disease free interval, even in patients where the initial excision seemed to be complete¹⁴. Therefore patients should be followed up regularly.

References

1. Isaacson PG, Wright DH. Malignant lymphoma of mucosa-associated lymphoid tissue. A distinctive type of B-cell lymphoma. *Cancer* 1983; **52**: 1410-1416.
2. Rosai J. *Ackerman's Surgical Pathology*. Volume 1. 8th edition. St Louis: Mosby 1996; 693-695.

3. Isaacson PG. Lymphomas of mucosa-associated lymphoid tissue (MALT). *Histopathology* 1990; **16**: 617-619.
4. Isaacson PG, Wright DH. Extranodal malignant lymphoma arising from mucosa associated lymphoid tissue. *Cancer* 1984; **53**: 2515-2524.
5. Non-Hodgkin's Lymphoma Pathologic Classification Project: National Cancer Institute sponsored study of classification of Non-Hodgkin's lymphoma. Summary and description of a working formulation for clinical usage. *Cancer* 1982; **49**: 2112-2135.
6. Burke JS. Extranodal lymphomas and lymphoid hyperplasias. In: Jaffe ES, ed. *Surgical pathology of the lymph nodes and related organs*. 5th edition. Philadelphia: WB Saunders 1995; 479-482.
7. Pelstring RJ, Essell JH, Kurtin PJ, Cohen AR, Banks PM. Diversity of organ site involvement among malignant lymphomas of mucosa-associated tissues. *American Journal of Clinical Pathology* 1991; **96**: 738-745.
8. Isaacson PG, Spencer J. Malignant lymphoma of mucosa-associated lymphoid tissue. *Histopathology* 1987; **11**: 445-462.
9. Wotherspoon AC, Oritz-Hidalgo C, Falzon M, Isaacson PG. Helicobacter pylori associated gastritis and primary B-cell gastric lymphoma. *Lancet* 1991; **338**: 1175-1176.
10. Wotherspoon AC, Pan L, Diss TC, Isaacson PG. A genotypic study of low grade B-cell lymphomas, including lymphomas of mucosa-associated lymphoid tissue (MALT). *Journal of Pathology* 1990; **162**: 135-140.
11. Chan KC, Ng CS, Isaacson PG. Relationship between high grade lymphoma and low grade B-cell mucosa-associated lymphoid tissue lymphoma (MALToma) of the stomach. *American Journal of Pathology* 1990; **136**: 1153-1164.
12. Isaacson PG, Spencer J, Wright DH. Classifying primary gut lymphomas. *Lancet* 1988; 1148-1149.
13. Wotherspoon AC, Doglioni C, Diss TC, Pan L, Moschini A, de Boni M, Isaacson PG. Regression of primary low grade B-cell gastric lymphoma of mucosa-associated lymphoid tissue type after eradication of Helicobacter pylori. *Lancet* 1993; **1343**: 575-577.
14. Wotherspoon AC, Doglioni C, Isaacson PG. Low grade gastric B-cell lymphoma of mucosa-associated lymphoid tissue (MALT): a multifocal disease. *Histopathology* 1992; **20**: 29-34.