A TWO YEARS STUDY OF SHOWCASING THE ROLE OF BONE MARROW ASPIRATION IN DIAGNOSING THE SPECTRUM OF HAEMATOLOGICAL DISORDERS IN A TERTIARY CARE HOSPITAL

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ABSTRACT

BACKGROUND
Evaluation of bone marrow has widespread use in clinical practice. It is a powerful diagnostic tool for many haematological and non-haematological disorders. Bone marrow aspiration provides definitive information regarding bone marrow cellularity, its architecture and the stage of maturation of different blood cells. Bone marrow aspiration is a safe invasive procedure done routinely in the hospitals for the diagnosis and management of haematological disorders. It is also essential for follow-up of patients undergoing chemotherapy, bone marrow transplantation and other modalities of medical treatment.

The objectives of this study were:
1. To evaluate the aetiological spectrum of haematological disorder as diagnosed on bone marrow aspiration examination.
2. To know the age incidence and gender distribution in various haematological disorders.

MATERIALS AND METHODS
This was a prospective and retrospective study done in the Department of Pathology, Yenepoya Medical College, Mangalore, Karnataka, between 2015 and 2016. A total of 164 patients underwent bone marrow evaluation for diagnostic purposes. Bone marrow was collected by bone marrow aspiration needle from posterior iliac spine of each selected patient after giving local anaesthesia by 2% Lidocaine hydrochloride under aseptic condition. Bone marrow aspiration slides were stained with Leishman stain and special stains like PAS (Periodic Acid Schiff), Perl and Sudan B whenever needed.

RESULTS
Out of 164 cases males were 91 cases (55%), whereas females were 73 cases (45%) with a male-to-female ratio of 1.24; 27 cases (16.5%) of haematological disorders who underwent bone marrow aspiration were in the age group of 40-49 years and < 10 years’ age group was consisting of 10 (6%) patients; 33 (21.24%) cases of haematological malignancies were found. Out of 33 cases 14 cases were of Acute Myeloid Leukaemia (AML), which is the commonest malignancy in our study. In this study, among non-malignant haematological disorder of bone marrow study, megaloblastic anaemia, 13 cases (7.92%) was the most common diagnosis followed by iron deficiency 10 (6%).

CONCLUSION
Bone marrow aspiration study is a powerful diagnostic tool for many haematological and non-haematological disorders. Histopathological evaluation of bone marrow biopsy is gold standard.

KEYWORDS
Aspiration, Marrow, Haematological, Leukaemia, Anaemia.


BACKGROUND
Evaluation of bone marrow has widespread use in clinical practice. It is a powerful diagnostic tool for many haematological and non-haematological disorders. Bone marrow aspiration is carried out mainly for the cytological assessment of bone marrow cells. Bone marrow biopsy is not necessarily a marrow substitute by aspiration and smear, but is an advantageous complementary procedure. Here larger amounts of marrow can be examined, cellularity readily assessed, architectural patterns analysed and structures other than haematopoietic cells can also be examined. Bone marrow aspiration provides definitive information regarding bone marrow cellularity, its architecture and the stage of maturation of different blood cells. It also come up with detailed information regarding the presence of haemoparasites within the bone marrow, the presence of infiltrates and storage diseases. It also assists in the diagnosis and staging of haematological malignancies, especially leukaemias.

It is also essential for follow-up of patients undergoing chemotherapy, bone marrow transplantation and other modalities of medical treatment. It is very useful in diagnosing unsuspected conditions. When other test results turn out to be non-contributory or inconclusive during the evaluation process. Bone marrow examination also gives explanation for unexplained cytopenias and leukaemia. It gives a more complete picture of the reaction of the haemopoietic tissue to anaemia than can be gained from Peripheral Blood Smear (PBS) alone.
Bone marrow aspiration is a safe invasive procedure done routinely in the hospitals for the diagnosis and management of haematological disorders. The risk associated with it is 0.08%. The most common complications of this procedure are bleeding, infection and pain at the biopsy site.

Aims and Objectives
1. To evaluate the aetiopathological spectrum of haematological disorders as diagnosed on bone marrow aspiration examination.
2. To know the age incidence and gender distribution in various haematological disorders.

MATERIALS AND METHODS
This was a prospective and retrospective study done in the Department of Pathology, Yenepoya Medical College, Mangalore, Karnataka, between 2015 and 2016. A total of 164 patients underwent bone marrow evaluation for diagnostic purposes.

Detailed clinical history was taken, physical examination to look specifically for the presence of anaemia, lymphadenopathy and hepatosplenomegaly. Complete blood count including haemoglobin, total and differential leucocyte count, total platelet count and blood indices were performed using automated haematology analyser (Sysmex XS1000i). Peripheral blood smear examination was done after Leishman and retic stain.

Bone marrow was collected by bone marrow aspiration needle from posterior iliac spine of each selected patient after giving local anaesthesia by 2% Lidocaine hydrochloride under aseptic condition. After the procedure, patients were observed to make sure that their vitals remained stable (especially pulse, blood pressure and temperature). Biopsy site was observed for infection and bleeding.

Bone marrow aspiration slides were stained with Leishman stain and special stains like PAS (Periodic Acid Schiff), Perl and Sudan B whenever needed.

RESULTS
Results from a total 164 patients who underwent BMA procedure were tabulated. The age range was 1 year to 85 years with a mean age of 41.2 years. Majority of them were males 91 cases (55%), whereas females were 73 cases (45%) with a male-to-female ratio of 1.24 (Table 1) and respectively. Table 3 shows frequency of haematological conditions in the study population.

<table>
<thead>
<tr>
<th>Age in Years</th>
<th>No. of Patients</th>
<th>Percentage</th>
</tr>
</thead>
<tbody>
<tr>
<td>&lt; 10</td>
<td>10</td>
<td>6</td>
</tr>
<tr>
<td>10-19</td>
<td>18</td>
<td>11</td>
</tr>
<tr>
<td>20-29</td>
<td>21</td>
<td>12.8</td>
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<tr>
<td>30-39</td>
<td>26</td>
<td>15.86</td>
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<tr>
<td>40-49</td>
<td>27</td>
<td>16.5</td>
</tr>
<tr>
<td>50-59</td>
<td>25</td>
<td>15.24</td>
</tr>
<tr>
<td>60-69</td>
<td>24</td>
<td>14.64</td>
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<tr>
<td>&gt; 70</td>
<td>13</td>
<td>7.9</td>
</tr>
<tr>
<td>Total</td>
<td>164</td>
<td>100</td>
</tr>
</tbody>
</table>

Table 2. Age Distribution

Out of 164 cases, 14 cases (8.53%) were diagnosed as Acute Myeloblastic Leukaemia (AML), Chronic Myeloid Leukaemia (CML) was seen in 8 cases (4.87%), Acute Lymphoblastic Leukaemia (ALL) in 3 cases.

Out of 164 cases nutritional deficiency anaemias were noted in 31 cases (18.79%), out of which megaloblastic anaemia was seen in 13 patients, whereas iron deficiency and combined nutritional anaemia was found in 10 and 8 cases respectively.

In iron deficiency anaemia, bone marrow cellularity is mildly increased due to erythroid hyperplasia. Erythropoiesis shows micronormoblastic type of maturation. (Figure A) Perls stain was done for deficient iron stores in the marrow.

The bone marrow aspiration of megaloblastic anaemia showed numerous megaloblasts with large cells with an increased nuclear/cytoplasmic ratio, sieve-like chromatin and giant metamyelocytes (Figure B).

In acute myeloid leukaemia, aspiration revealed abundant myeloblasts with large cells, increased nuclear/cytoplasmic ratio, fine chromatin with 2-4 variably prominent nucleoli and Auer rods (Figure C).

Aspirates from multiple myeloma show numerous atypical plasmablasts with binucleate and trinucleate atypical plasma cells with high nuclear/cytoplasmic ratio with prominent nucleoli (Figure D).

<table>
<thead>
<tr>
<th>Diseases/Findings</th>
<th>No. of Patients</th>
<th>Percentage</th>
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</thead>
<tbody>
<tr>
<td>Acute myeloblastic leukaemia</td>
<td>14</td>
<td>8.53</td>
</tr>
<tr>
<td>Chronic myeloid leukaemia</td>
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<td>4.87</td>
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<tr>
<td>Iron deficiency anaemia</td>
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<td>6</td>
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<tr>
<td>Megaloblastic anaemia</td>
<td>13</td>
<td>7.92</td>
</tr>
<tr>
<td>Combined nutritional anaemia</td>
<td>8</td>
<td>4.87</td>
</tr>
<tr>
<td>Multiple myeloma</td>
<td>6</td>
<td>3.65</td>
</tr>
<tr>
<td>Acute splenic sequestration</td>
<td>6</td>
<td>3.65</td>
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<tr>
<td>Anaemia of chronic disease</td>
<td>5</td>
<td>3</td>
</tr>
<tr>
<td>Idiopathic thrombocytopenic purpura</td>
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<td>2.43</td>
</tr>
<tr>
<td>Acute lymphoblastic leukaemia</td>
<td>3</td>
<td>1.82</td>
</tr>
<tr>
<td>Myelodysplastic syndrome</td>
<td>4</td>
<td>2.43</td>
</tr>
<tr>
<td>Bone secondaries</td>
<td>3</td>
<td>1.82</td>
</tr>
<tr>
<td>Dry tap</td>
<td>3</td>
<td>1.82</td>
</tr>
<tr>
<td>Eosinophilia</td>
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<td>1.21</td>
</tr>
<tr>
<td>Myeloproliferative disorders</td>
<td>2</td>
<td>1.21</td>
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Table 1. Gender Distribution

Majority of the patients, i.e. 27 cases (16.5%) of haematological disorders who underwent bone marrow aspiration were in the age group of 40-49 years and <10 years’ age group was consisting of 10 (6%) patients. Table 2 shows age distribution of different age groups of patients.
Chronic lymphocytic leukaemia 1 0.6
Chronic myelomonocytic leukaemia 1 0.6
Hemophagocytic syndrome 1 0.6
Lymphoma 1 0.6
NHL 1 0.6
Storage disorder 1 0.6
Secondary polycythemia 1 0.6
Remission 3 1.82
Reactive marrow 27 15.51
Unsatisfactory 20 12.20
No definitive opinion 3 1.82
Normal study 13 7.92
Total 164 100

Table 3. Frequency of Haematological Conditions in the Study Population

DISCUSSION

Bone marrow examination is an important investigation performed in the routine practice for the diagnosis of various haematological and non-haematological disorders. It is one of the most common and safe procedures in medical practice. Rarely infection, excessive bleeding or embolism have been reported after bone marrow biopsy.\textsuperscript{13}

In this study out of 164 study population, maximum 27 (16.5%) were in 40-49 years’ age group and lowest 10 (6%) were in < 10 years’ age group (Table 2). Out of 164 cases 91 (55%) were male and 73 (45%) were female, which is comparative to study done by Pudasaini et al.\textsuperscript{11}

In our study out of 164, 33 (21.24%) cases of haematological malignancies were found. Out of 33 cases, 14 cases were of Acute Myeloid Leukæmia (AML) which is the commonest malignancy in our study, which is similar to the study done by Rahim et al.\textsuperscript{2}

The incidence of ALL (3 cases) in our study is lower as compared to Kuperan et al.\textsuperscript{13} The decrease in number of ALL in the present study might be due to the fact that we have included more adult population than the children in the present study.\textsuperscript{13}

Out of 164 cases, 8 cases were of Chronic Myeloid Leukæmia (CML), which is similar to the study done by Anjum et al\textsuperscript{8} and one case of Chronic Lymphocytic Leukæmia (CLL) was found.

**Figure 1.** Bone Marrow Aspiration showing (A) Erythroid Hyperplasia in Iron Deficiency Anaemia, (B) Showing Megaloblasts in Megaloblastic Anaemia (Leishman stain, 400x)

**Figure 2.** Bone Marrow Aspiration showing (C) Myeloblasts in Acute Myeloid Leukæmia (AML) and (D) Atypical Plasma Cells in Multiple Myeloma
Other malignancies in this study were multiple myeloma 6 cases (3.65%) and Myelodysplastic Syndrome (MDS) 4 cases (2.42%), which is comparable to the study done by Afzal Khan et al. Other series of studies showed the occurrence of multiple myeloma ranging from 0.94% to 4.1%,1,11,15,16,17 whereas myelodysplastic syndrome showed 2 - 7.9%.18

In this study among non-malignant haematological disorder of bone marrow study, megaloblastic anaemia23 cases (7.92%) was most common diagnosis followed by iron deficiency anaemia in 8 cases (4.87%). Similar situation was observed in the study done by Rahim et al, because due to the fact that mostly iron deficiency anaemia is treated on an outpatient basis and bone marrow examination is not routinely done to confirm its diagnosis.8

There were three cases of metastatic deposits, which is comparatively more than study done by Shastry et al.10 There was one case of storage disorder noted, which is similar to the study done by Chang et al.19

Out of 164 cases, ITP was found in 4 cases (2.43%) which is significantly lower than the study conducted by Ahmed and Khodke et al.20,21

In our study out of 164 cases dry tap was observed in 3 cases (1.82%), which is remarkably lower than study done by Novone et al, Engeset et al and Humphries JE et al.22,23

CONCLUSION

Bone marrow aspiration study is a powerful diagnostic tool for many haematological and non-haematological disorders. Although, there is a dependence on newer methodologies and ancillary assays including immunochemistry, cyto genetic analysis, flow cytometry and molecular assays, which may have augmented and refined the diagnostic criteria of previously obtained by light microscopy, the traditional and crucial role of bone marrow aspiration and histopathological evaluation of bone marrow biopsy remains unchanged.

REFERENCES