Outcome of Adult Acute Lymphoblastic Leukaemia with standard chemotherapy (remission induction & consolidation) in a tertiary hospital of Bangladesh

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## Background

Acute Lymphoblastic Leukaemia (ALL)

 A heterogeneous group of lymphoid neoplasm resulting from the proliferation of malignant lymphoid cells

## Background

 Complete remission rate with induction therapy in adult ALL 60-90%

5-year disease free survival 25-35%

Childhood ALL - complete remission rate & overall survival 80%

## Background

In adults- poor outcome associated with —

- older age
- male gender
- high white cell count
- Philadelphia chromosome positivity

## Aims & Objectives

General Objectives:

To describe the treatment outcome of adult ALL patients with standard chemotherapy (remission induction & consolidation)

## Aims & Objectives

Specific Objectives:

- To describe the clinical presentations of adult ALL patients
- To describe the clinical outcome of adult ALL patients
- To see the frequency of toxic effects of chemotherapy

Study design

Observational/ descriptive study with prospective record of outcome

Place of study

Dept. of Haematology & Medicine, SSMC & MH, Dhaka

Duration of study:

1<sup>st</sup> January, 2011 to 31<sup>st</sup>

December, 2011

Study population:

All Adult ALL patients admitted to Haematology & Medicine wards of SSMC & MH, Dhaka over study duration

Sample size: 20 cases

Inclusion Criteria

- Cases of ALL ≥ 15year age group
- ALL confirmed by bone marrow exam

Exclusion Criteria

- Patients with severe dysfunction of liver or kidney or with major ECG abnormality
- Patients unwilling to sign written informed consent
- History of treatment for ALL

- Cases- recruited both from Haematology & Medicine Department of SSMCH
- Data collection by-
- Detailed history taking
- Thorough clinical examination
- Recorded in a structured data collection sheet

- Pre-treatment investigation
- CBC with PBF
- Bone marrow examination
- Blood chemistry including liver and renal function tests
- Serum uric acid
- Serum electrolytes
- Only morphological evaluation
- Immunological & cytogenetic features not evaluated

 Medical Research Council United Kingdom ALL (MRC UKALL) X protocol

- remission induction therapy
- early intensification
- interim maintenance
- cranial irradiation

Remission induction therapy

- Inj. Daunorubicin 45mg/m² on day 1 & 2 iv
- Inj. Vincristine 1.5mg/m² on days 1, 8,15 & 22 iv
- Inj. L-asparaginase 6000mg/m² iv on days 10-18
- Prednisolone 45mg/m² PO on days 1 to 28
- Methotrexate 12.5mg IT 2 doses

Early intensification

- Inj. Vincristine 1.5mg/m<sup>2</sup> i.v. bolus on Day-1
- Inj. Methotrexate 12.5mg IT on Day 1
- Inj. Daunorubicin 45mg/m² i.v. Day 1 & 2
- Prednisolone 45mg/m<sup>2</sup> PO on days 1 to 5
- Inj. Cytarabine 100mg/m<sup>2</sup> i.v. 12hourly on Days 1 to 5
- Inj. Etoposide 100mg/m<sup>2</sup> i.v. on days 1 to 5
- Thioguanine 20mg/m<sup>2</sup> PO on days 1 to 5

- Interim maintenance
- Inj. Vincristine 1.5mg/m<sup>2</sup> iv on Day 1 (monthly)
- Prednisolone 45mg/m<sup>2</sup> PO on Days 1 to 5 (monthly)
- 6-mercaptopurine (80mg/m<sup>2</sup>) PO daily
- Methotrexate (20mg/m²) PO weekly

Cranial irradiation:

1800cGy in 10 fractions over 2 weeks

- During induction-
- Blood count on days 3,7,14 & 21
- At the end of each cycle (on D29)—
- BME & evaluation of marrow cellularity & marrow differential

## Follow-up

 Upto end of study period from completion of consolidation therapy

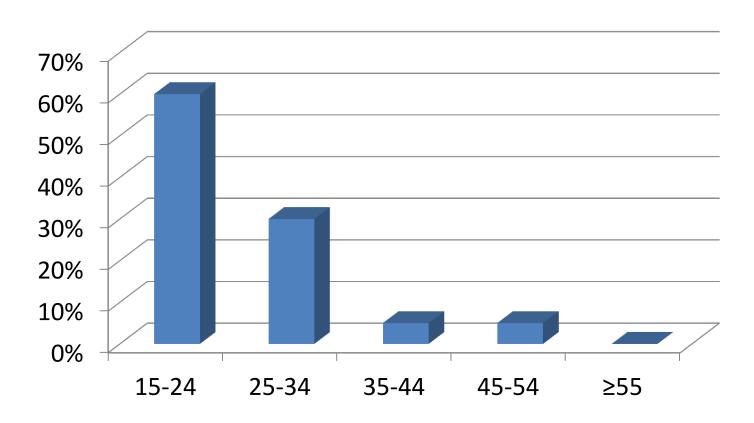


Fig 1: Age(in years) distribution of the patients(N=20)

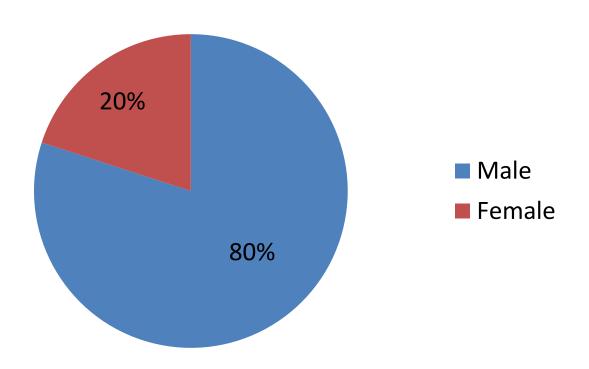


Fig 2: Sex distribution of the patients

Table 1: Distribution of patients by principal presentation(N=20)

| Presentation      | No. of patients | Percentage(%) |
|-------------------|-----------------|---------------|
| Fever             | 17              | 85            |
| Fatigue           | 18              | 90            |
| Bleeding          | 11              | 55            |
| Bone & joint pain | 10              | 50            |

Table 2: Distribution of patients by other clinical manifestation(N=20)

| Presentation                  | No. of patients | Percentage(%) |
|-------------------------------|-----------------|---------------|
| Sore throat                   | 05              | 25            |
| Cough                         | 04              | 20            |
| Anorexia                      | 08              | 40            |
| Abdominal pain                | 06              | 30            |
| Headache & blurring of vision | 01              | 05            |

Table 3: Distribution of patients by clinical findings (N=20)

| Clinical findings                  | No. of patients | Percentage(%) |
|------------------------------------|-----------------|---------------|
| Anaemia                            | 18              | 90            |
| Superficial lymphadenopathy        | 11              | 55            |
| Mediastinal lymphadenopathy on CXR | 02              | 10            |
| Splenomegaly                       | 14              | 70            |
| Hepatomegaly                       | 09              | 45            |

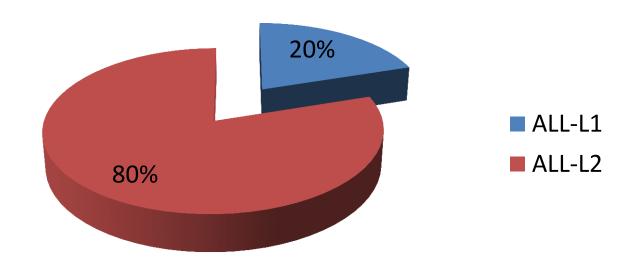


Fig 3: Distribution of patients by FAB subtype(N=20)

Table 4: Distribution of patients by laboratory parameter(N=20)

| WBC ×10 <sup>9</sup> /L | No. of patients | Percentage |
|-------------------------|-----------------|------------|
| ≤4                      | 04              | 20         |
| 4.1-10                  | 01              | 05         |
| 10.1-50                 | 06              | 30         |
| >50                     | 09              | 45         |
| Total                   | 20              | 100        |

Table 4: Distribution of patients by laboratory parameter(N=20)

| Hb(gm/dl)                 | No. of patients     | Percentage(%)    |
|---------------------------|---------------------|------------------|
| <6                        | 08                  | 40               |
| 6.1-10                    | 10                  | 50               |
| >10                       | 02                  | 10               |
| Total                     | 20                  | 100              |
|                           |                     |                  |
| Platelet ×10 <sup>9</sup> | No. of patients     | Percentage(%)    |
| Platelet ×10 <sup>9</sup> | No. of patients  12 | Percentage(%) 60 |
|                           |                     |                  |
| <30                       | 12                  | 60               |

Table 5: Overall toxicity of chemotherapy

| Co | mplications   | No. of patients | Percentage<br>(%) |
|----|---|-----------------|-------------------|
| A. | Haematological toxic effects during treatment         |                 |                   |
| 1. | Febrile neutropenia                                   | 09              | 45                |
| 2. | Severe thrombocytopenia TPC <10×10 <sup>9</sup> /L    | 05              | 25                |
| 3. | Severe pancytopenia(TC-0.5×10 <sup>9</sup> /L L-100%) | 02              | 10                |

Table 5: Overall toxicity of chemotherapy

| Complications                          | No. of patient | Percentage<br>(%) |
|--|----------------|-------------------|
| B. Other adverse events during Tx      |                |                   |
| 1. Nausea and vomiting                 | 18             | 90                |
| 2. Oral thrush                         | 10             | 50                |
| 3. Abdominal pain                      | 02             | 10                |
| 4. Diarrhoea                           | 02             | 10                |
| 5. Jaundice                            | 02             | 10                |
| 6. Paraesthesia, weakness, muscle pain | 08             | 40                |
| 7. Constipation                        | 05             | 25                |
| 8. Cushing's syndrome                  | 06             | 30                |
| 9. Hyperglycaemia                      | 01             | 05                |
| 10. Alopecia                           | 14             | 70                |

Table 6: Overall treatment outcome after remission induction & consolidation (n=20)

| Outcome                                     | No. of patients | Percentage(%) |
|---|-----------------|---------------|
| Complete remission & on maintenance therapy | 15              | 75            |
| No remission                                | 02              | 10            |
| Death                                       | 02              | 10            |
| Discontinued treatment                      | 01              | 05            |
| Relapse                                     | 00              | 00            |
|   |                 |               |

#### Limitation

 Small sample size – collected from one tertiary hospital only - limiting generalisability & representativeness

 Immunologic & cytogenetic features of ALL- not be evaluated

Very short follow-up - unequal for comparison or conclusion

### Conclusion

- Overall treatment outcome quite similar to international data
- Further large scale study with long term follow up is recommended
- to determine the overall & disease free survival of adult ALL patients treated with standard chemotherapy in Bangladeshi patients

## Acknowledgement

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# THANK YOU