

Leukemias

Learning Objective:

1-To figure the type of leukemia.

2-What is clinical feature of leukemia.

3-How do you differentiate from other disease.

4-The aim of treatment

-Induce a clinical and hematologic remission

-To maintain remission by chemotherapy and prophylactic CNS therapy

-To treat the complications of therapy and of the disease).

5-How to deal with family and child with cancer

-Understanding of the importance of sensitive early diagnostic counseling.

-Understanding of the psychological impact of cancer on children and their families.

Type of Leukemia

1-Acute leukemias represent a clonal expansion and arrest at a specific stage of normal myeloid or lymphoid hematopoiesis, types of acute leukemia:

1. Acute lymphoblastic leukemia 75% .
2. Acute myeloblastic leukemia 20%.

2-Chronic leukemia

Chronic myeloid leukemias 3%.

1. Philadelphia chromosome +ve
2. Juvenile myelomonocytic leukemia

Acute lymphoblastic leukemia

Incidence

- Peak incidence between 2 and 5 years of age.
- Account for 25-30% of all childhood cancer.

Etiology :

- 1- Ionizing radiation.
- 2- Chemicals.
- 3- Drugs .
- 4- Genetic:
 - Identical twins.
 - Chromosomal abnormalities (down syndrome , Bloom, fanconie anemia).
- 5- Increase incidence:
 - Ataxia teleneglectasia .
 - Diamond blackfan

6-Breast feeding of more than (6) months duration-decreasing risk of leukaemia to (11%).

7-conjugate Hb vaccination at the ages of (3)month and (2)years-(28%)lower incidence of leukaemia

Clinical features of Acute lymphoblastic leukemia.

1. General system effects.
 1. Fever (60%)
 2. Lassitude (50%)
 3. Pallor (40%)
2. Hematologic effects arising from Marrow invasion
 - 1- Anemia
 - 2- Neutropenia
 - 3- Thrombocytopenic
 - 4- One -2% pancytopenia
3. Clinical manifestations arising from lymphoid system invasion.
 - 1- Lymphadenopathy → sometimes mediastinal lymphadenopathy.
 - 2- Splenomegaly
 - 3- Hepatomegaly.
4. Clinical manifestations of extramedullary invasion:
 - Central nervous system involvement.
 - ↑ intracranial pressure.
 - Focal neurological signs .
 - Hypothalamic syndrome (polyphagia with excessive wt gain) .
 - Diabetes insipidus.
 - Chloromas of the spinal cord in ALL) .
 - CNS-Hg.
5. Genitourinary system involvement .
 - Testicular involvement

- Renal involvement present with hematuria hypotension renal failure
- 6. Gastrointestinal involvement. leukemia infiltrate in the GI--silent until terminal stages most common site is the ceccum.
- 7. Bone and joint involvement :
 - Bone pain is one of the initial symptoms in 25%.
 - Direct leukemic infiltration of the periosteim
 - Bone infarction
 - Expansion of marrow cavity by leukemic cells.
- 8. Skin involvement occurs in neonatal leukemia.
- 9. Cardiac involvement due to leukemia infiltration .
- 10.Lung involvement

Diagnosis

- 1- C.B.P(Hb may be low or normal, W.B.C high or low, Plt low.
Blood film show normochromic, normocytic and blast)
- 2- Bone marrow send for:
 - Histochemistry : by special stain(periodic acid Schiff,
 - Immunophentypes :by immunophenotyping of blast cell by flow cytometric analysis to detect intracellular antigens.
 - Cytogentic:by cytogenetic analysis of leukemia blasts and there are two major classes of cytogenetic aberration (visible loss or gain chromosomal,a balanced exchange without apparent loss or gain of DNA)
- 3- Chest radiograph for mediastanial mass
- 4- Blood chemistry (Bu ,Uric acid ,Ca , PH, LDH,K,SGOT,SGPT.
- 5- CSF for the diagnosis of CNS leukemia.
 - Present of more than 5WBC/mm³
 - Identification of blast cells on cytocentrifuge examination.

CNS involvement in leukemia

-CNS 1 : < 5 WBC/mm³ No. blast

-CNS 2: < 5 WBC / mm³ blast

-CNS 3: > 5 WBC /mm³ blast.

D.D

1-Aplastic anemia

2-AML

3-Mylofibrosis

4-Involvement of bone marrow by other malignancy like lymphoma, neuroblastoma.

Poor Prognosis

- 1) Age under one year and greater than 10 years, poor prognosis .
- 2) $WBC \geq 50.000$. plt. < 20.000 .
- 3) Immunophenotype T-cell, mature B- cell .
- 4) DNA index < 1.16 hypodiploidy(no of chromosomal less than 45).
- 5) Cytogenetics philadelphic chromosome + (9.22) or infantile leukemia(4.11).
- 6) CNS disease at diagnosis CNS 2 or 3.
- 7) mediastinal mass

- 8) Early response to induction therapy patients: who are not in remission at the end of induction therapy(M2 or M3) this because the blast cell contain ACTH receptor if increase in receptor more response to steroid and no. of blast cell in 8 days less than one 1000 in CBP .

Classification of bone marrow remission :

M1 → blast in B.M : $< 5\%$

M2 → blast in B.M : $5 \leq 25\%$

M3 → blast in B.M : $> 25\%$

Treatment

- 1- General care
- 2- Hydration and hydroxyurea
- 3- Plt. and blood transfusions.
- 4- Antibiotics
- 5- Chemotherapy

Aim of treatment:

- 1- Induce a clinical and hematologic remission.
- 2- To maintain remission by chemotherapy and prophylactic CNS therapy
- 3- To treat the complications of therapy and of the disease.
- 4- To prevent relapse (B.M,CNS,Testicular).