Astana Medical University Department of Internal Disease №1

# SIW

Topic: chronic lymphocytic leucosis

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# Chronic lymphocytic leucosis

 The chronic lymphocytic leucosis is a widespread kind of a blood cancer at which emergence of the pathological mature neoplastic lymphocytes having abnormally long life expectancy is observed.



## Etiology

 More than 2/3 all patients passed an age threshold in 60 years.



- ОМонЛ острый моноцитарный лейкоз
- ОЛЛ острый лимфобластный лейкоз
- СКВ системная красная волчанка
- ЛКМЗ лимфома из клеток мантийной зоны
- ХЛЛ хронический лимфоцитарный лейкоз

Exogenic factors	Endogenic factors
The ionizing and magnetic radiations	Metabolic disturbances in an organism
Chemicals (benzene, pistetsids)	Immunodeficiency
Medicinal preparations	Chromosomal mutations
Viruses	

#### Pathogenesis

- There is a mutation of cells of predecessors of lymphocytes, DNA is programmed on a proliferation of abnormal lymphocytes.
- At first the damaged lymphocytes gather in lymph nodes. After achievement of a certain quantity they with current of a lymph migrate in a lien and a liver, thus, promoting augmentation of the sizes of the above-named organs. In process of the attack of marrow, malignant lymphocytes replace normal cells, provoking, anemic processes and reduction of quantity of healthy formulated elements of a blood. In parallel to these processes note decrease of the activity of antibodies.

# Clinics

- Often at early stages the chronic lymphocytic leukosis doesn't prove in any way. If symptoms nevertheless appeared, belong to them:
- The lymphadenitis which isn't followed by pain
- Fatigue
- Temperature increase
- Pain in the top left part of a stomach which can be caused by lien augmentation
- Night sweating
- Loss of weight
- Frequent infections







- The generalized lymphadenitis which merge in huge soft or dense packages becomes perceptible.
- The lien reaches the appreciable sizes, its mass is enlarged

#### Classification

Stage 0	Lymphocytosis of a peripheral blood and marrow (more than 10000 cells in 1 мкл bloods and not less than 40% of lymphocytes among formulated elements of marrow)
Stage the I	Lymphocytosis and the enlarged lymph nodes
The stage the II	Exept of signs of stages 0 and I, is available gepato-and a splenomegaly
Stage of III	Besides signs stages of 0, I and II, HB 110 g/I

- Group A (forecast good, more than 10-year survival) HB> 100 g/l; the quantity of thrombocytes > 100x10/l is struck less than 3 organs
- Group B (the forecast intermediate) the Maintenance of HB and thrombocytes the same, as in group A; 3 organs and more are struck
- Group C (forecast bad, less than 2-year survival) HB <100 g/l; quantity of thrombocytes <100x10/l</li>

## Diagnosis

- Blood test. By quantity of blood cells and their look under a microscope it is possible to suspect a leucosis. Most of patients with a chronic leucosis has an increased quantity of leucocytes and I, depression of number of erythrocytes and lymphocytes d thrombocytes. The maintenance of cells of a leykolizis is enlarged (Botkin's Gumprekht cells)
- Biochemical blood test helps to specify function of kidneys and structure of a blood.
- The research of marrow gives the chance to establish the diagnosis of a leukosis and to estimate efficiency of treatment. Hyperplasia of lymphocytic elements.

- For the purpose of specification like a leukosis special methods of a research are used: cytochemistry, flowing cytometry, immunocytochemistry, cytogenetics and molecular and genetic research.
- X-ray inspections of a thorax and bones allow to tap a lesion of lymph nodes of a mediastinum, bones and joints.
- The Computer Tomography (CT) gives the chance to find a lesion of lymph nodes in a thoracal cavity and a stomach.

• The Magnetic Resonance Imaging (MRI) is especially shown at a research of a head and spinal cord.

 Ultrasonography (US) allows to distinguish tumoral and cystic educations, to tap a lesion of kidneys, liver and lien, lymph nodes.

#### Treatment

- Patients are younger than 70 years and without serious associated diseases of Hemoimmunoterapiya;
- Fludarabin + Cyclophosphamide + Rituximab (FCR); Fludarabin + Rituximab (FR);
- Pentostatin + Cyclophosphamide + Rituximab (PCR);
- Bendamustin + Rituximab (BR);
- Obinutuzumab + Hlorambutsil.

- Patients are more senior than 70 years, or with serious associated diseases Obinutuzumab + Hlorambutsil;
  Rituximab + Hlorambutsil;
- Bendamustin (70 mg/sq.m in 1 cycle with rising to 90 mg/sq.m) + Rituximab (BR);

Cyclophosphamide + Prednizolon± Rituximab; Rituximab;

- Флударабин±Ритуксимаб;
- Kladribin;

Hlorambutsil.

# Treatment of a recurrence and refractory options of HLL

• Choice drug at treatment of a recurrence and refractory options of HLL is Ibrutinib. Ibrutinib in a dose of 420 mg is applied (3 x 140-mg in capsules).

Indications for treatment ibrutiniby:

- • ECOG status 0-1.
- The diagnosis of HLL, is established according to criteria of the mezhunanarodny working group on studying of HLL, 2008;
- • Existence of indications by the beginning of therapy.
- To the patient должнен to be conducted at least one course of therapy of HLL with including of purine analogs or is taped

#### Complications

- Frequent infectious diseases. The people suffering from a chronic lymphocytic leukosis often have infectious diseases. In most cases it is infections of the top and lower respiratory tracts. In certain cases there can be more serious infectious diseases.
- Formation of more aggressive form of cancer. A small amount of the people suffering from a chronic lymphocytic leukosis can have more aggressive form of cancer, a so-called diffuse V-macrocellular lymphoma. Sometimes such degeneration is called Richter's syndrome.



- Augmentation of risk of emergence of other forms of malignant neoplasms. At the people suffering from a chronic lymphocytic leukosis the risk of formation of other types of cancer, such as melanoma, cancer of a lung and cancer of digestive tract is increased.
- Disturbances from immune system. At a small share of patients with a chronic lymphocytic leukosis the autoimmune disease at which the cells of immune system designed to protect an organism from an infection by mistake attack erythrocytes or thrombocytes develops

