# B-Cell Lymphoma (80%

- B-Cells help make antibodies, which are proteins that attach to and help destroy antigens
- Lymphomas are caused when a mutation arises during the B-cell life cycle
- Various different lymphomas can occur dur several different stages of the cycle
  - Follicular lymphoma, which is a type of B-cell lymphoma is caused by a gene translocation which results in an over expressed gene called BCL-2, which blocks apoptosis.

# T-Cell Lymphoma (15%

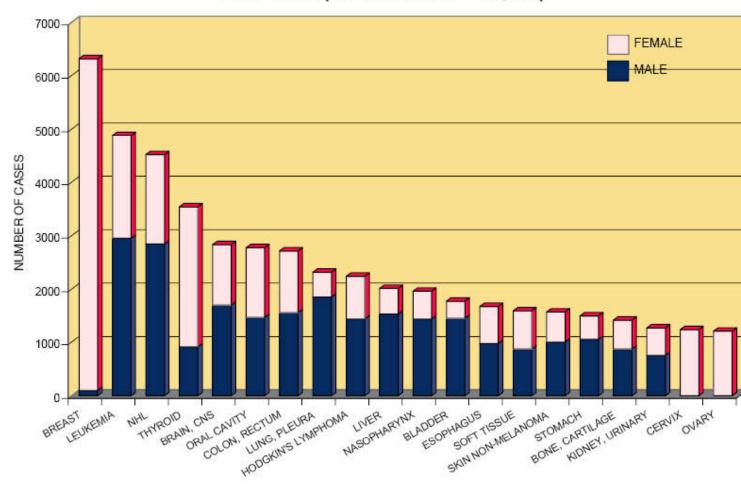
- The T-cells are born from stem cells, similar to that of B-cells, but mature in the thymus.
- They help the immune system work in coordinated fashion.
  - These types of lymphomas are categorian by how the cell is affected
    - Anaplastic Large cell Lymphoma, t-cell lymphoma caused by a gene translocation i chromosome 5

## Mechanisms of lymphomagen

- Genetic alterations
- Infection
- Antigen stimulation
- Immunosuppression

### **Epidemiology of lymphoma**

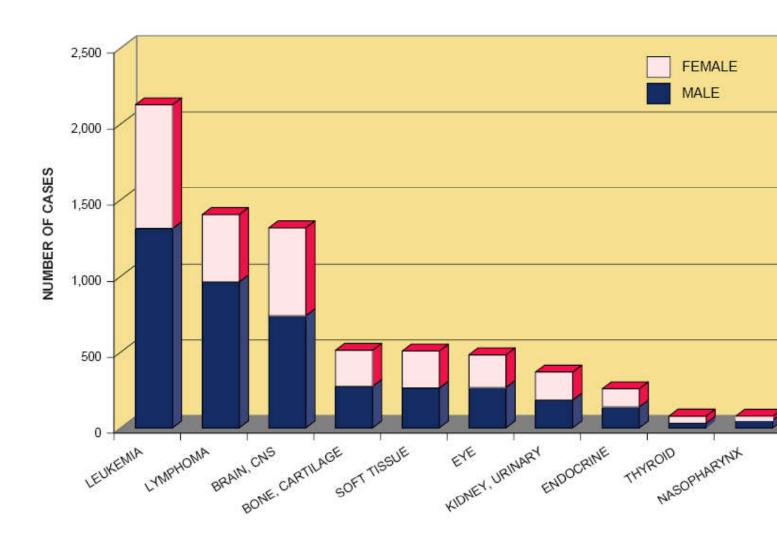
- males > females
- incidence
  - NHL increasing
  - Hodgkin lymphoma stable
- in NHL: 3<sup>rd</sup> most frequently diagnosed cancer in males and 4<sup>th</sup> in females
- in HL: 5<sup>th</sup> most frequently diagnosed cancer in males and 10<sup>th</sup> in females



#### DISTRIBUTION OF 20 MOST COMMON MALIGNANCIES 1975 - 2005 (TOTAL CASES = 57,255)

#### FIGURE 6

#### DISTRIBUTION OF 10 MOST COMMON PEDIATRIC MALIGNANCIES 1975 - 2005 (TOTAL CASS = 7,548)



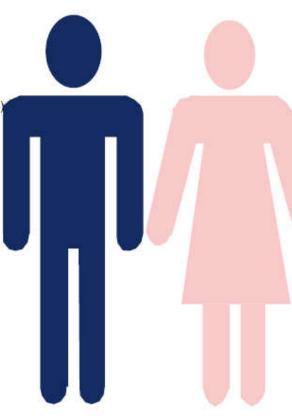
#### COMPARATIVE DATA - KFSH&RC vs NCR vs USA (% to TOTAL CANCER CASES)

SITE	KFSH&RC 2005 Analytics	NCR 2003 Saudis	USA 2005 Estimates
BREAST	14.9%	10.6%	15.5%
THYROID	7.9%	6.6%	1.9%
COLON, RECTUM	7.1%	8.7%	10.9%
NON-HODGKIN'S LYMPHOMA	6.8%	8.0%	4.1%
LEUKEMIA	6.2%	6.5%	2.5%
BRAIN, CNS	5.1%	3.2%	1.3%
HODGKIN'S LYMPHOMA	4.6%	3.7%	0.5%
LUNG, PLEURA	3.5%	3.7%	12.7%
BONE, CARTILAGE	2.9%	1.2%	0.2%
SOFT TISSUE	2.2%	1.8%	0.7%
SKIN MELANOMA	0.2%	0.3%	4.3%
PROSTATE (% to MALES)	2.6%	2.7%	32.7%

#### DISTRIBUTION OF 20 MOST COMMON MALIGNANCIES 2005 ANALYTIC CASES (TOTAL CASES = 2,336)

MALE

LEUKEMIA 92 (8.5%) COLON, RECTUM 90 (8.3%) NHL 83 (7.7%) BRAIN, CNS 67 (6.2%) HODGKIN'S LYMPHOMA 63 (5.8%) BLADDER 62 (5.7%) PROSTATE 61 (5.6%) ORAL CAVITY 58 (5.4%) NASOPHARYNX 58 (5.4%) LUNG, PLEURA 55 (5.1%) THYROID 44 (4.1%) BONE, CARTILAGE 43 (4.0%) STOMACH 37 (3.4%) LARYNX 31 (2.9%) SOFT TISSUE 31 (2.9%) SKIN NON-MELANOMA 31 (2.9%) KIDNEY, URINARY 26 (2.4%) LIVER 20 (1.8%) TESTIS, GENITAL 19 (1.8%) PANCREAS 16 (1.5%) ESOPHAGUS 16 (1.5%)



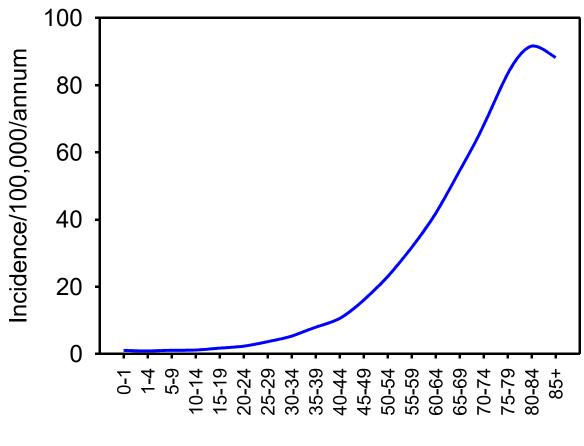
#### FEMALE

BREAST 344 (27.4%) THYROID 141 (11.2%) COLON, RECTUM 77 (6.1%) NHL 76 (6.1%) UTERUS, GENITAL 58 (4.6%) ORAL CAVITY 53 (4.2%) LEUKEMIA 53 (4.2%) BRAIN, CNS 52 (4.1%) CERVIX 44 (3.5%) HODGKIN'S LYMPHOMA 44 (3.5 OVARY 40 (3.2%) KIDNEY, URINARY 28 (2.2%) LUNG, PLEURA 26 (2.1%) BONE, CARTILAGE 25 (2.0%) ESOPHAGUS 24 (1.9%) SKIN NON-MELANOMA 23 (1.8% NASOPHARYNX 21 (1.7%) SOFT TISSUE 21 (1.7%) STOMACH 15 (1.2%) LIVER 12 (1.0%) OTHER G.I. 12 (1.0%)

#### ANALYTIC CASES SEEN AT KFSH&RC BY SITE\* AND AGE 2005

SITE	0-4	5-9	10-14	15-19	20-24	25-29	30-34	35-39	40-44	45-49	50-54	55-59	60-64	65-69	70-74	75-79
Oral Cavity	0	3	0	1	3	3	4	2	7	10	13	14	7	15	11	5
Nasopharynx	0	0	1	1	3	2	6	10	6	10	13	8	5	8	6	0
Esophagus	0	0	0	0	0	0	2	0	1	3	3	1	5	5	7	6
Stomach	0	1	0	0	1	0	4	3	3	1	3	5	8	5	8	5
Colon, Rectum	0	0	0	0	1	5	6	15	8	24	23	20	15	17	15	11
Liver	3	2	0	1	0	1	1	0	3	2	1	3	5	6	1	3
Pancreas	0	0	0	0	0	0	1	1	2	0	1	3	4	4	5	0
Other G.I.	0	0	0	0	0	0	2	0	3	5	0	1	6	1	2	3
Larynx	0	0	0	0	0	1	0	1	1	2	4	6	6	3	6	2
Lung, Pleura	0	0	0	0	0	0	1	4	13	6	12	9	9	7	9	4
Multiple Myeloma	0	0	0	0	1	0	1	0	2	2	1	2	3	1	1	2
Lymphoid Leukemia	25	13	13	11	5	3	1	1	1	0	2	0	1	0	0	0
Myeloid Leukemia	8	4	5	7	6	3	8	6	5	9	1	0	1	0	0	0
Other Leukemias	2	1	1	1	0	0	0	0	1	0	0	0	0	0	0	0
Reticuloendothelium	0	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0
Bone, Cartilage	1	7	20	23	7	1	0	1	1	1	1	1	1	1	1	1
Soft Tissue	5	4	7	5	4	2	1	5	4	3	5	3	2	1	0	1
Skin Melanoma	0	0	0	0	0	0	0	0	2	0	0	0	1	0	0	2
Skin Non-Melanoma	0	2	2	3	0	0	1	1	6	3	5	5	10	3	3	5
Breast	0	0	0	0	5	11	31	60	61	61	46	29	19	7	11	4
Uterus, Genital	0	0	1	0	4	1	3	1	6	4	6	5	7	7	9	3
Cervix	0	0	0	0	0	0	3	2	11	8	5	6	4	1	3	1
Ovary	0	2	2	1	1	3	0	2	4	6	6	1	2	4	2	2
Prostate	0	0	0	0	0	0	0	0	1	0	1	6	8	12	17	7
Testis, Genital	0	0	0	0	4	3	3	0	3	3	1	0	0	1	1	0
Bladder	1	0	0	1	0	2	0	2	3	4	8	11	13	8	8	4
Kidney, Urinary	10	3	1	0	1	0	1	3	6	3	3	4	7	5	3	3
Eye	10	1	0	0	0	0	0	1	0	0	0	0	0	0	0	0
Brain, CNS	24	15	15	3	10	4	2	9	7	6	4	6	8	2	3	1
Thyroid	0	1	2	9	18	22	23	22	22	15	17	11	9	5	4	2
Other Endocrine	15	2	2	2	0	0	1	0	0	0	0	1	0	0	0	0
NHL - Lymph Nodes	3	5	2	4	8	6	7	4	5	4	10	4	13	8	4	4
NHL - Extra-nodal	4	1	3	3	3	2	3	2	5	11	4	9	5	5	1	3
Hodgkin's Lymphoma-LNs	7	16	19	16	17	7	6	2	4	3	2	1	1	0	1	2
HL - Extra-nodal	0	0	0	0	0	0	0	0	2	0	0	0	0	0	0	0
Primary Unknown	0	0	0	0	1	0	1	5	0	1	2	4	5	6	4	0
All Other Sites	1	2	0	0	1	0	1	2	2	4	2	0	1	0	1	1
TOTAL	119	85	96	93	104	82	124	167	211	214	205	179	191	148	147	87

### **Age distribution of new NHL**



Age (years)

## **Risk factors for NHL**

- immunosuppression or immunodeficie
- connective tissue disease
- family history of lymphoma
- infectious agents
- ionizing radiation

# **Clinical manifestations**

- Variable
  - severity: asymptomatic to extremely ill
  - time course: evolution over weeks, months, years
- Systemic manifestations
  - fever, night sweats, weight loss, anorexia, p
- Local manifestations
  - lymphadenopathy, splenomegaly most com
  - any tissue potentially can be infiltrated

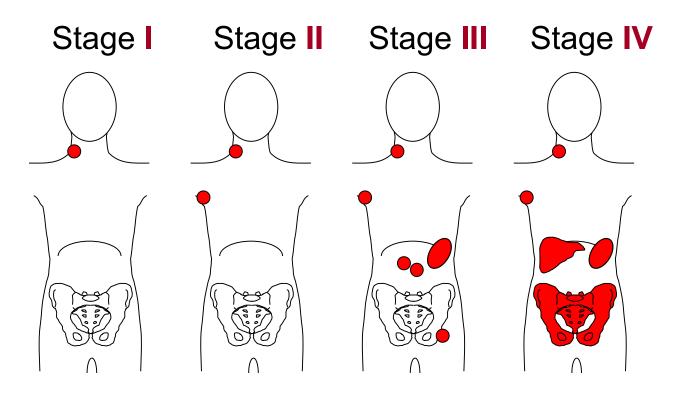
### **Other complications of lymphon**

- bone marrow failure (infiltration)
- CNS infiltration
- immune hemolysis or thrombocytope
- compression of structures (eg spinal cord, ureters)
- pleural/pericardial effusions, ascites

### Non-Hodgkin's Lymphoma Staging

- Stage is the term used to describe the extent of tumor that has spread through the body (I and II are localized where as III and IV are advanced.
- Each stage is then divided into categories
  A, B, and E
  - A: No systemic symptoms
  - B: Systemic Symptoms such as fever, night sweats and weight loss
  - E: Spreading of disease from lymph node to another organ

# **Staging of lymphoma**



- A: absence of B symptoms
- **B**: fever, night sweats, weight loss

# Staging

#### Stages of Non-Hodgkin's Lymphoma

Stage I (early disease): the cancer is found only in a single lymph n region OR one organ or area outside the lymph node.

Stage II (locally advanced disease): the cancer is found in two or n lymph node regions on one side of the diaphragm (the breathing n that separates the abdomen from the chest), OR the cancer is four one lymph node region plus a nearby area or organ.

Stage III (advanced disease): the disease involves lymph nodes bo above and below the diaphragm OR one nodal area and one organ opposite sides of the diaphragm.

Stage IV (widespread disease): the lymphoma is outside the lymph nodes and spleen AND has spread to one or more organs such as bone marrow, skin, or other organs.

## Symptoms

- Painful Swelling of lymph nodes locat in the neck, underarm and groin.
- Unexplained Fever
- Night Sweats
- Constant Fatigue
- Unexplained Weight loss
- Itchy Skin

Cancer Sourcebook



### **Causes and Risk Factors**

- The Exact causes are still unknown
  - Higher risk for individuals who:
    - Exposed to chemicals such as pesticides or solvents
    - Infected w/ Epstein-Barr Virus
    - Family history of NHL (although no heredita pattern has been established)
    - Infected w/ Human Immunodeficiency Virus (HIV)

Lymphoma.org

### **Diagnosis** Staging Studies

- Bone marrow aspiration and biopsy
- Radionuclide scans:
- GI x-rays
- Spinal fluid analysis
- CT scans
- Magnetic Resonance Imaging (MRI)
- Biopsy

#### Diagnosis requires an adequate biopsy

- Diagnosis should be <u>biopsy-proven</u> before treatment is initiated
- Need enough tissue to assess cells a architecture

- open bx vs core needle bx vs FNA

## Treatment

- Non-Hodgkin's Lymphoma is usually treated by a team of physicians including hematologists, medical oncologists and a radiation oncologist.
- In some cases such as for Indolent lymphomas, the Doctor may wait to start treatment until the patient starts showing symptoms, known as "watchful waiting"

## **Treatment Options**

- Chemotherapy
- Radiation
- Bone Marrow Transplantation
- •Surgery
- Immunotherapy

 Using the bodies own immune system combined wi material made in a lab.

### **Survival Rates**

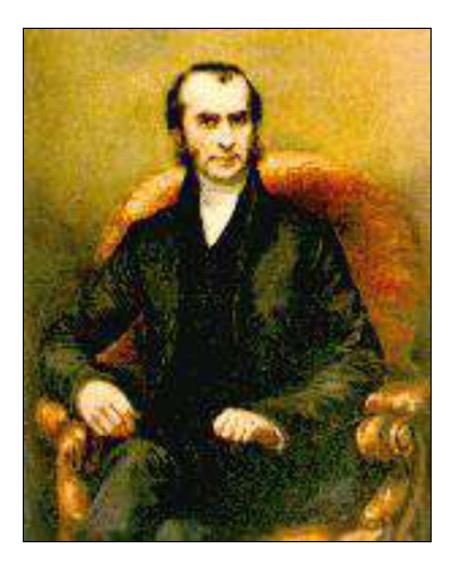
 Survival Rates vary widely by cell typ and staging.

– 1 Year Survival Rate: 77%

- 5 Year Survival Rate: 56%

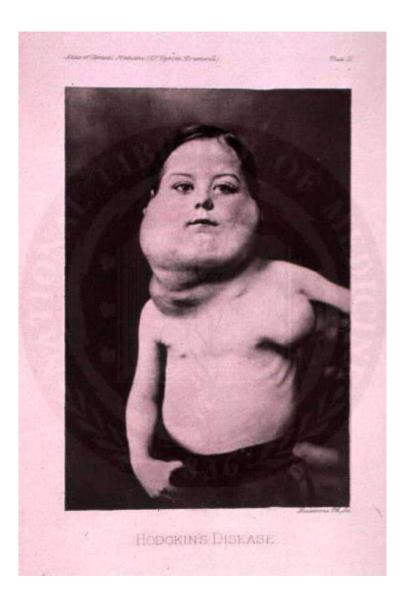
- 10 Year Survival Rate: 42% Cancer.org

## Hodgkin lymphoma



#### Thomas Hodg (1798-1866)

#### **Classical Hodgkin Lymphom**



# Hodgkin lymphoma

- cell of origin: germinal centre B-cell
- Reed-Sternberg cells (or RS variants in the affected tissues
- most cells in affected lymph node are polyclonal reactive lymphoid cells, no neoplastic cells

### **Reed-Sternberg cell**

