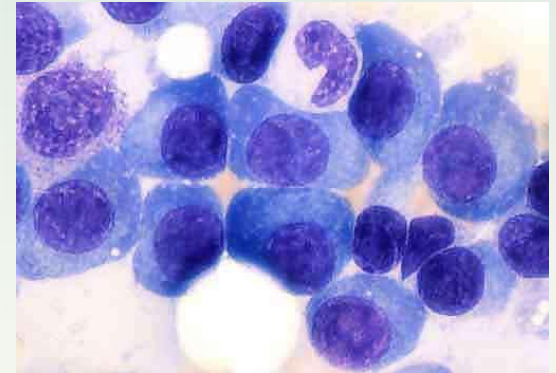
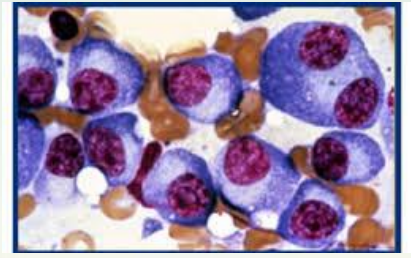


# MYELOM VE LENFOMALARDA ALGORİTMALAR

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04.06.2017



# Multiple Myelom Tanı Anında



- Öykü ve Fizik İnceleme
- Tam kan sayımı, trombosit sayısı
- Serum BUN, kreatinin, albumin, Ca
- Serum LDH ve  $\beta$ -2 mikroglobulin
- Serum Ig ler, Serum protein elektroforezi
- 24 saat idrar protein
- Serum hafif zincir assay
- İskelet surveyi
- Kİ biyopsisi (Kİ İHK, Kİ flow sitometri)
- Kİ sitogenetik inceleme
- Plazma hücre FISH

# MM Genetik Test

## Kİ Sitogenetik İnceleme

$t(4;14)$

kötü prognostik belirteç (relaps)

kötü prognostik belirteç (bir defa)

$t(14;16)$

KRAS sık  $t(11;14)$

kötü prognostik belirteç (tekrarlanabilir)

$del17p13$

1q21 amplifikasyon

tekrarlanabilir

kötü prognostik belirteç (tekrarlanabilir)

$del13$

# Özetle;

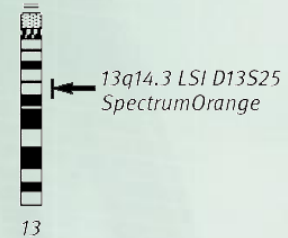
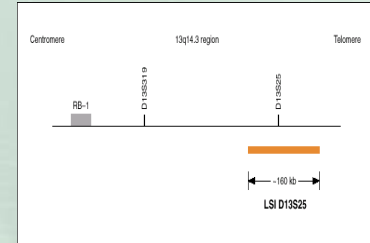
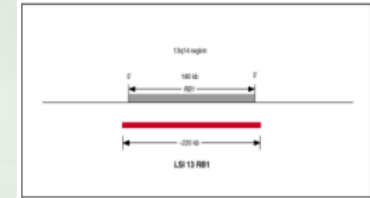
- 14q32 (*IGH* geni)' yi içeren yeniden düzenlenmeler kötü prognostik belirteç
- del13q FISH ile sık görülen bir anomali ancak yalnızca metafazda görünür olduğunda kötü prognostik belirteç
- del17p kötü prognostik belirteç

- Birçok merkezde genetik tarama prognostik danışma, hasta seçimi ve tedavi planlamada rutin olarak kullanılmakta
- NCCN myelom panelinde en az **t(4;14)(p16;q32)** **t(14;16)(q32;q23)** **17p13 delesyonu** ve **kromozom1 amplifikasyonu** çalışılmalı diye öneriliyor
- Güncel kılavuzda GEP(gen ekspresyon profilleri) 15gen, 70gen, 92 gen şeklinde paneller yer alıyor. Rutin kullanımı henüz önerilmiyor ancak seçilmiş hastalarda bilgi verici olabileceği vurgulanıyor

# Tartışma



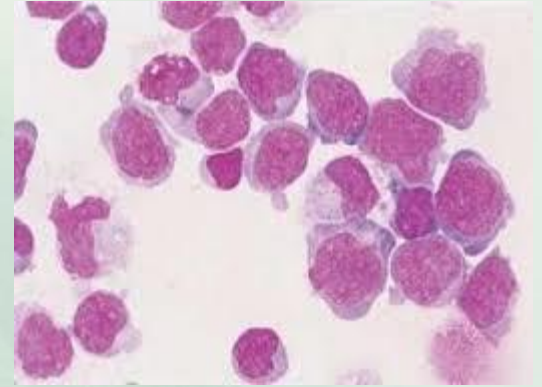
- Konvansiyonel yöntemle anomalileri saptayamama nedenleri??
- 13q delesyonu farklı bölgelere özgü problarla değerlendirmeli mi??  
FISH ile 13q del taraması D13S319 (13q14.3) ve D13S25 (13q14.3) ve RB-1 probları
- 13q RB1 rutin taramada kullanılabilir mi??





# Lenfoma

- Farklı ve deęişken klinik, patolojik ve **genetik** özellikler gösteren lenfoid doku neoplazilerdir
- Lenfoma sınıflaması epidemiyolojik, morfolojik, immünofenotipik ve genetik özelliklere göre yapılır



# Lenfomalar

## Classification

**Table 1**

### WHO Classification of the Mature B-Cell, T-Cell, and NK-Cell Neoplasms (2017)

#### Mature B-Cell Neoplasms

- Chronic lymphocytic leukemia/small lymphocytic lymphoma
- Monoclonal B-cell lymphocytosis
- B-cell prolymphocytic leukemia
- Splenic marginal zone lymphoma
- Hairy cell leukemia
- *Splenic lymphoma/leukemia, unclassifiable\**
  - ▶ *Splenic diffuse red pulp small B-cell lymphoma\**
  - ▶ *Hairy cell leukemia-variant\**
- Lymphoplasmacytic lymphoma
  - ▶ Waldenström's macroglobulinemia
- Monoclonal gammopathy of undetermined significance (MGUS), IgM
- Mu heavy chain disease
- Gamma heavy chain disease
- Alpha heavy chain disease
- Monoclonal gammopathy of undetermined significance (MGUS), IgG/A
- Plasma cell myeloma
- Solitary plasmacytoma of bone
- Extraoesophageal plasmacytoma
- Monoclonal immunoglobulin deposition diseases
- Extranodal marginal zone lymphoma of mucosa-associated lymphoid tissue (MALT type)
- Nodal marginal zone lymphoma
  - ▶ *Pediatric nodal marginal zone lymphoma\**
- Follicular lymphoma
  - ▶ In situ follicular neoplasia
  - ▶ Duodenal-type follicular lymphoma
  - ▶ Pediatric-type follicular lymphoma
- *Large B-cell lymphoma with IRF4 rearrangement*
- Primary cutaneous follicle center lymphoma
- Mantle cell lymphoma
  - ▶ In situ mantle cell neoplasia
- Diffuse large B-cell lymphoma (DLBCL), NOS
  - ▶ Germinal center B-cell type
  - ▶ Activated B-cell type
- T-cell/histiocyte-rich large B-cell lymphoma
- Primary DLBCL of the central nervous system (CNS)
- Primary cutaneous DLBCL, leg type
- EBV-positive DLBCL, NOS
- *EBV-positive mucocutaneous ulcer\**
- DLBCL associated with chronic inflammation
- Lymphomatoid granulomatosis
- Primary mediastinal (thymic) large B-cell lymphoma
- Intravascular large B-cell lymphoma
- ALK-positive large B-cell lymphoma
- Plasmablastic lymphoma
- Primary effusion lymphoma
- *HHV8-positive DLBCL, NOS\**
- Burkitt lymphoma
- *Burkitt-like lymphoma with 11q aberration\**
- High-grade B-cell lymphoma, with *MYC* and *BCL2* and/or *BCL6* rearrangements
- High-grade B-cell lymphoma, NOS
- B-cell lymphoma, unclassifiable, with features intermediate between DLBCL and classical Hodgkin lymphoma

\*Provisional entities are listed in italics.

[Continued on next page](#)



## Classification

*Table 1 continued*

### WHO Classification of the Mature B-Cell, T-Cell, and NK-Cell Neoplasms (2017)

#### Mature T-Cell and NK-Cell Neoplasms

- T-cell prolymphocytic leukemia
- T-cell large granular lymphocytic leukemia
- *Chronic lymphoproliferative disorder of NK-cells\**
- Aggressive NK-cell leukemia
- Systemic EBV-positive T-cell lymphoma of childhood
- Hydroa vacciniforme–like lymphoproliferative disorder
- Adult T-cell leukemia/lymphoma
- Extranodal NK/T-cell lymphoma, nasal type
- Enteropathy-associated T-cell lymphoma
- Monomorphic epitheliotropic intestinal T-cell lymphoma\*
- *Indolent T-cell lymphoproliferative disorder of the GI tract\**
- Hepatosplenic T-cell lymphoma
- Subcutaneous panniculitis-like T-cell lymphoma
- Mycosis fungoides
- Sézary syndrome
- Primary cutaneous CD30-positive T-cell lymphoproliferative disorders
  - ▶ Lymphomatoid papulosis
  - ▶ Primary cutaneous anaplastic large cell lymphoma
- Primary cutaneous gamma-delta T-cell lymphoma
- *Primary cutaneous CD8-positive aggressive epidermotropic cytotoxic T-cell lymphoma\**
- *Primary cutaneous acral CD8-positive T-cell lymphoma\**
- *Primary cutaneous CD4-positive small/medium T-cell lymphoproliferative disorder\**
- Peripheral T-cell lymphoma, NOS
- Angioimmunoblastic T-cell lymphoma
- *Follicular T-cell lymphoma\**
- *Nodal peripheral T-cell lymphoma with TFH phenotype\**
- Anaplastic large-cell lymphoma, ALK positive
- Anaplastic large-cell lymphoma, ALK negative
- *Breast implant–associated anaplastic large-cell lymphoma\**

#### Hodgkin Lymphoma

- Nodular lymphocyte-predominant Hodgkin lymphoma
- Classical Hodgkin lymphoma
  - ▶ Nodular sclerosis classical Hodgkin lymphoma
  - ▶ Lymphocyte-rich classical Hodgkin lymphoma
  - ▶ Mixed cellularity classical Hodgkin lymphoma
  - ▶ Lymphocyte-depleted classical Hodgkin lymphoma

#### Posttransplant Lymphoproliferative Disorders (PTLD)

- Plasmacytic hyperplasia PTLD
- Infectious mononucleosis-like PTLD
- Florid follicular hyperplasia PTLD
- Polymorphic PTLD
- Monomorphic PTLD (B- and T/NK-cell types)
- Classical Hodgkin lymphoma PTLD

#### Histiocytic and dendritic cell neoplasms

- Histiocytic sarcoma
- Langerhans cell histiocytosis
- Langerhans cell sarcoma
- Indeterminate dendritic cell tumor
- Interdigitating dendritic cell sarcoma
- Follicular dendritic cell sarcoma
- Fibroblastic reticular cell tumor
- Disseminated juvenile xanthogranuloma
- Erdheim-Chester disease

\*Provisional entities are listed in italics.

Swerdlow SH, Campo E, Pileri SA, Harris NL, Stein H, Siebert R, Advani R, Ghielmini M, Salles GA, Zelenetz AD, Jaffe ES. The 2017 revision of the World Health Organization classification of lymphoid neoplasms. *Blood* 2017;127:2375-2390.

# Diffuz Büyük B Hücreli Lenfoma (DBBHL)

- Tüm NHL' ların %30-58' i
- Tanı: Eksizyonel LN biyopsi

National Comprehensive Cancer Network® **NCCN Guidelines Version 3.2017** Diffuse Large B-Cell Lymphoma [NCCN Guidelines Index](#) [Table of Contents](#) [Discussion](#)

**DIAGNOSIS<sup>a,b</sup>**

**ESSENTIAL:**

- Hematopathology review of all slides with at least one paraffin block representative of the tumor. Rebiopsy if consult material is nondiagnostic.
- An FNA or core needle biopsy alone is not generally suitable for the initial diagnosis of lymphoma. In certain circumstances, when a lymph node is not easily accessible for excisional or incisional biopsy, a combination of core biopsy and FNA biopsies in conjunction with appropriate ancillary techniques for the differential diagnosis (immunohistochemistry, flow cytometry, PCR for IgH and TCR gene rearrangements, and FISH for major translocations) may be sufficient for diagnosis.
- Adequate immunophenotyping to establish diagnosis and GCB versus non-GCB origin<sup>c,d</sup>
  - ▶ IHC panel: CD20, CD3, CD5, CD10, CD45, BCL2, BCL6, Ki-67, IRF4/MUM1, MYC with or without
  - ▶ Cell surface marker analysis by flow cytometry: kappa/lambda, CD45, CD3, CD5, CD19, CD10, CD20

**USEFUL UNDER CERTAIN CIRCUMSTANCES:**  
Additional immunohistochemical studies to establish lymphoma subtype

- ▶ IHC panel: Cyclin D1, kappa/lambda, CD30, CD138, Epstein-Barr virus in situ hybridization (EBER-ISH), ALK, HHV8, SOX11
- **Revisyotipe or FISH: MYC, BCL2, BCL6 rearrangements<sup>e</sup>**

**SUBTYPES**

- Subtypes included:
  - ▶ DLBCL, NOS<sup>f</sup>
  - ▶ DLBCL coexistent with follicular lymphoma of any grade
  - ▶ DLBCL coexistent with gastric MALT lymphoma
  - ▶ DLBCL coexistent with nongastric MALT lymphoma
  - ▶ Follicular lymphoma grade 3<sup>g</sup>
  - ▶ Intravascular large B-cell lymphoma
  - ▶ DLBCL associated with chronic inflammation
  - ▶ ALK-positive DLBCL<sup>h</sup>
  - ▶ EBV-positive DLBCL of the elderly
  - ▶ T-cell-/histiocyte-rich large B-cell lymphoma
- Subtypes not included:
  - ▶ Primary cutaneous B-cell lymphomas (See [NCCN Guidelines for Primary Cutaneous B-Cell Lymphomas](#))
  - ▶ Primary DLBCL of the CNS (See [NCCN Guidelines for CNS](#))

→ [See Workup \(BCEL-2\)](#)

Primary Mediastinal Large B-Cell Lymphoma (PMBL), see [BCEL-B 1 of 4](#).  
Grey Zone Lymphoma, see [BCEL-B 2 of 4](#).  
Double Hit Lymphomas, see [BCEL-B 3 of 4](#).  
Primary Cutaneous B-cell Lymphomas, Leg type, see [BCEL-B 4 of 4](#).

<sup>d</sup>See [Use of Immunophenotyping/Genetic Testing in Differential Diagnosis of Mature B-Cell and NK/T-Cell Neoplasms \(NHODG-A\)](#).  
<sup>e</sup>Cases with double expression of MYC and either BCL2 or BCL6 by IHC having a GCB-like immunophenotype should undergo FISH testing for MYC, BCL2, and BCL6 rearrangement.  
<sup>f</sup>Germinal center (or follicle center) phenotype is not equivalent to follicular lymphoma and can occur in DLBCL and Burkitt lymphoma. Morphology is required to establish diagnosis.  
<sup>g</sup>Controversy exists regarding management of FL grade 3. Some may treat FL grade 3a as follicular lymphoma and others may treat it as DLBCL.  
<sup>h</sup>These are most often CD20 negative and rituximab is not necessary.

<sup>a</sup>Burkitt lymphoma intermediate histology or DLBCL CD10+ tumors with very high proliferation >90% with or without Burkitt lymphoma-like features might be considered for more aggressive treatment as per [BURK-A](#).  
<sup>b</sup>See [International Prognostic Index \(BCEL-A\)](#).  
<sup>c</sup>Typical immunophenotype: CD20+, CD45+, CD3-; other markers used for subclassification.

# Diffüz Büyük B Hücreli Lenfoma Tanı

## Tanı Anında Gerekli

Histopatolojik inceleme

Ayırıcı Tanı için İHK, flow sitometri, IGH ve TCR için PCR, majör translokasyonlar için FISH)

İmmunfenotiplendirme (CD20, CD3, CD5, CD10, CD45, BCL2, BCL6, Ki67, IRF4/MUM1, MYC)

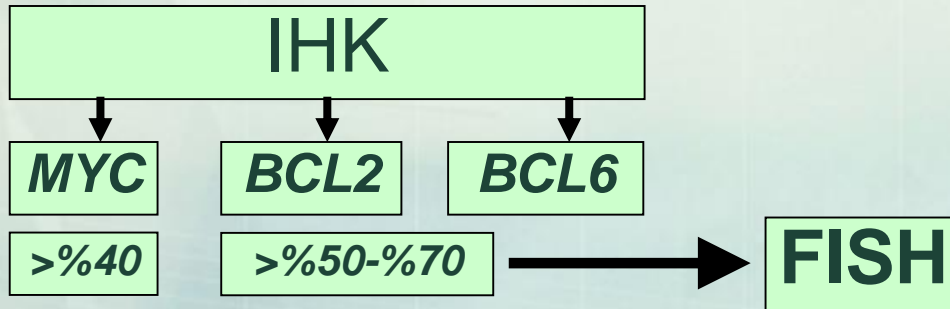
## Bazı Koşullarda Faydalı Olabilecek

İHK paneli: Siklin D1, kappa/lambda CD30, CD138, ALK, HHV8, SOX11

Karyotip ve FISH: MYC, BCL2, BCL6 yeniden düzenlenmeleri

# Diffuz Büyük B Hücreli Lenfoma (DBBHL)

- *MYC* ve *BCL2* yeniden düzenlenme varlığı 'double-hit'
- *MYC*, *BCL2* ve *BCL6* yeniden düzenlenme birlikteliği 'triple-hit'
- Kötü prognostik, agresif tedavi



- FISH ile doğrulanmıyorsa 'double expressor lenfoma'
- Double-hit lenfoma kadar olmasa da kötü prognoz

# Foliküler Lenfoma

- NHL grubunda 2. en sık görülen lenfoma
- Normal germinal merkez B hücrelerinin malign formu
- Tümör mikroçevresi makrofaj, follikül dendritik hücreler, fibroblastlar, endotel hücrelerinden oluşur
- Tanı: eksizyonel lenf nodu biyopsisi
- Evreleme: Ann Arbor sistemi



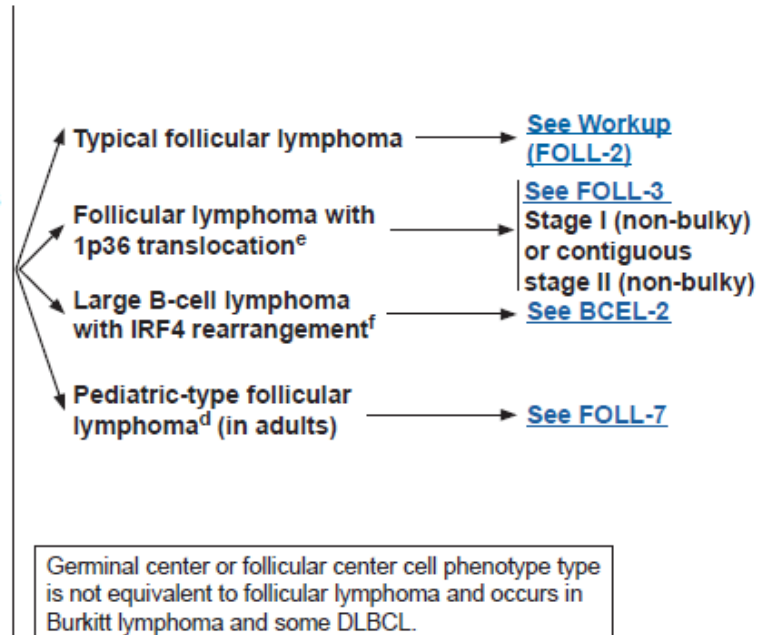
### DIAGNOSIS

#### ESSENTIAL:

- Hematopathology review of all slides with at least one paraffin block representative of the tumor. Rebiopsy if consult material is nondiagnostic.
- An FNA or core needle biopsy alone is not generally suitable for the initial diagnosis of lymphoma. In certain circumstances, when a lymph node is not easily accessible for excisional or incisional biopsy, a combination of core biopsy and FNA biopsies in conjunction with appropriate ancillary techniques for the differential diagnosis (immunohistochemistry, flow cytometry, PCR for IGHV and TCR gene rearrangements, and FISH for major translocations) may be sufficient for diagnosis. Histologic grading cannot be performed on an FNA.
- Adequate immunophenotyping to establish diagnosis<sup>b,c</sup>
  - ▶ IHC panel: CD20, CD3, CD5, CD10, BCL2,<sup>d</sup> BCL6, CD21, or CD23, with or without
  - ▶ Cell surface marker analysis by flow cytometry: kappa/lambda, CD19, CD20, CD5, CD23, CD10

#### USEFUL UNDER CERTAIN CIRCUMSTANCES:

- Molecular analysis to detect: antigen receptor gene rearrangements; *BCL2* rearrangements<sup>d</sup>
- Karyotype or FISH:<sup>e,f</sup> t(14;18); *BCL6*, 1p36, IRF4/MUM1 rearrangements<sup>d</sup>
- IHC panel: Ki-67;<sup>g</sup> IRF4/MUM1 for FL grade 3, cyclin D1



<sup>a</sup>Follicular lymphoma (FL), grade 1-2. FL, grade 3 is an area of controversy. The distinction between follicular grade 3a and 3b has not been shown to have clinical significance to date. However, controversy exists regarding management of FL grade 3. Some may treat FL grade 3a as FL and others may treat it as diffuse large B-cell lymphoma (DLBCL). FL, grade 3b is commonly treated according to the [NCCN Diffuse Large B-Cell Lymphoma Guideline \(BCEL-1\)](#). Any area of DLBCL in a FL of any grade should be diagnosed and treated as a DLBCL.

<sup>b</sup>Typical immunophenotype: CD10+, BCL2+, CD23+/-, CD43-, CD5-, CD20+, BCL6+. Rare cases of follicular lymphoma may be CD10- or BCL2-.

<sup>c</sup>See [Use of Immunophenotyping/Genetic Testing in Differential Diagnosis of Mature B-Cell and NK/T-Cell Neoplasms \(NHODG-A\)](#).

<sup>d</sup>In young patients with localized disease that lacks BCL2 expression or t(14;18), consider entity of pediatric-type FL. Analysis of BCL6 rearrangement may be useful for evaluating the diagnosis of pediatric-type FL.

<sup>e</sup>FL with 1p36 deletions have a predominant diffuse pattern in inguinal nodes, large localized mass, CD23+, typically grade 1-2 and have a good prognosis.

<sup>f</sup>Lymphomas with IRF4 translocations are usually DLBCL but occasionally are purely FL grade 3b and often DLBCL with FL grade 3b. Patients typically present with Waldeyer's ring involvement and are often children/young adults. The tumor is locally aggressive but responds well to chemotherapy +/- RT. These lymphomas do not have a BCL2 rearrangement and should not be treated as low-grade FL.

<sup>g</sup>There are reports showing that Ki-67 proliferation fraction of >30 % may be associated with a more aggressive clinical behavior, but there is no evidence that this should guide treatment decisions.



# Foliküler Lenfoma Tanı

## Tanı Anında Gerekli

eksizyonel lenf nodu biyopsisi

Ayırıcı Tanı için İHK, flow sitometri, *IGVH* ve *TCR* için PCR, majör translokasyonlar için FISH)

İmmunfenotiplendirme

## Bazı Koşullarda Faydalı Olabilecek

Antijen reseptör gen yeniden düzenlenmelerinin, *bcl-2* mutasyonunun gösterilmesi (pediyatrik tip FL)

Karyotip ve FISH; *t(14;18)*, *bcl-6*, **1p36**, *IRF4/MUM1* yeniden düzenlenmeleri

IHK paneli: Ki-67, *IRF4/MUM1*, Siklin D1

# Mantle Hücreli Lenfoma

- Tüm lenfomaların %6-9'u
- Ekstranodal tutulum sık (Kİ, KC, dalak, Waldeyer halkası ve GİS kanal tutulumu)
- Tanı: eksizyonel lenf nodu biyopsisi
- Preanalitik aşamada dikkat edilmesi gerekenler doku kendi hacminin 8-10 katı **%10 nötral tamponlu formalin** içinde 18-24 saat fikse edilmeli. 48 saat formalin içinde beklemiş dokularda İHK ve moleküler testlerde yanlış negatiflikler olabilmektedir

### DIAGNOSIS

#### ESSENTIAL:

- Hematopathology review of all slides with at least one paraffin block representative of the tumor. Rebiopsy if consult material is nondiagnostic.
- An FNA or core needle biopsy alone is not generally suitable for the initial diagnosis of lymphoma. In certain circumstances, when a lymph node is not easily accessible for excisional or incisional biopsy, a combination of core biopsy and FNA biopsies in conjunction with appropriate ancillary techniques for the differential diagnosis (immunohistochemistry, flow cytometry, PCR for IgH and TCR gene rearrangements, and FISH for major translocations) may be sufficient for diagnosis.
- Adequate immunophenotyping to establish diagnosis<sup>a,b</sup>
  - ▶ IHC panel: CD20, CD3, CD5, cyclin D1, CD10, CD21, CD23, BCL2, BCL6, Ki-67<sup>c</sup> with or without
  - ▶ Cell surface marker analysis by flow cytometry: kappa/lambda, CD19, CD20, CD5, CD23, CD10

#### USEFUL UNDER CERTAIN CIRCUMSTANCES:

- IHC: LEF1 may help distinguish from variant CLL; SOX11 or IGHV sequencing may be useful for determination of indolent MCL; may also help in diagnosis of CCND1- MCL.
- Karyotype or FISH: t(11;14), t(14;18), CLL panel
- Cell surface marker analysis by flow cytometry: CD200

<sup>a</sup>Typical immunophenotype: CD5+, CD20+, CD43+, CD23+/-, cyclin D1+, CD10+/-.  
Note: Some cases of MCL may be CD5- or CD23+. If the diagnosis is suspected, cyclin D1 staining or FISH for t(11;14) should be done. There are rare cases of CCND1- MCL (<5%) with an otherwise typical immunophenotype.

<sup>b</sup>See [Use of Immunophenotyping/Genetic Testing in Differential Diagnosis of Mature B-Cell and NK/T-Cell Neoplasms \(NHODG-A\)](#).

<sup>c</sup>Ki-67 proliferation fraction of <30% in lymph nodes is associated with a more favorable prognosis. However, it is not used to guide treatment.

### WORKUP

#### ESSENTIAL:

- Physical exam: Attention to node-bearing areas, including Waldeyer's ring, and to size of liver and spleen
- Performance status
- B symptoms
- CBC, differential, platelets
- Comprehensive metabolic panel
- LDH
- Bone marrow biopsy ± aspirate
- C/A/P CT with contrast of diagnostic quality and/or whole-body PET/CT scan
- Hepatitis B testing<sup>d</sup> if rituximab contemplated
- Echocardiogram or MUGA scan if anthracycline or anthracenedione-based regimen is indicated
- Pregnancy testing in women of child-bearing age (if chemotherapy or RT planned)

#### USEFUL UNDER CERTAIN CIRCUMSTANCES:

- Endoscopy/colonoscopy<sup>e</sup>
- Neck CT with contrast
- Uric acid
- Discussion of fertility issues and sperm banking
- Lumbar puncture (for blastic variant or CNS symptoms)
- Beta-2-microglobulin

→ [See Induction Therapy \(MANT-2\)](#)

<sup>d</sup>Hepatitis B testing is indicated because of the risk of reactivation with immunotherapy + chemotherapy. Tests include hepatitis B surface antigen and core antibody for a patient with no risk factors. For patients with risk factors or previous history of hepatitis B, add e-antigen. If positive, check viral load and consult with gastroenterologist.

<sup>e</sup>Essential for confirmation of stage I-II disease. See Discussion for details.

Note: All recommendations are category 2A unless otherwise indicated.

Clinical Trials: NCCN believes that the best management of any patient with cancer is in a clinical trial. Participation in clinical trials is especially encouraged.

# Mantle Hücreli Lenfoma Tanı

## Tanı Anında Gerekli

eksizyonel lenf nodu biyopsisi

Ayırıcı Tanı için İHK, flow sitometri, *IGVH* ve *TCR* için PCR, majör translokasyonlar için FISH)

İmmunfenotiplendirme

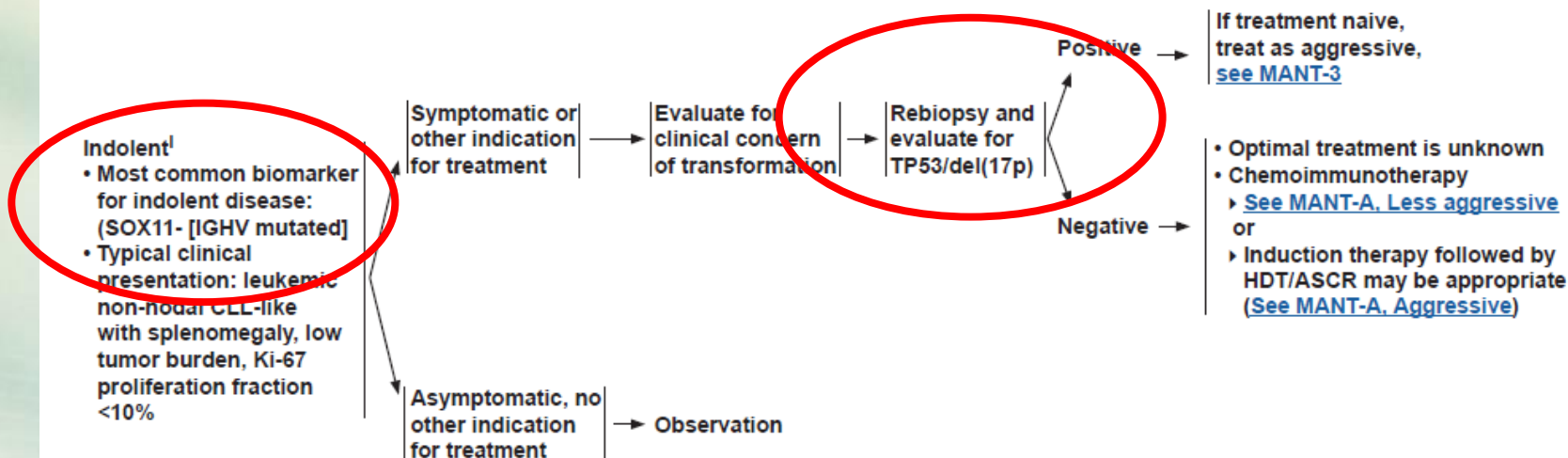
## Bazı Koşullarda Faydalı Olabilecek

İHK: LEF1 (KLL' den ayırmada)

İHK: *SOX11* ve *IGVH* indolent MCL'yı tanımda önemli(CCND1-olanlarda)

Karyotip ve FISH: *t(11;14)*, *t(14;18)*, KLL paneli

Flow sitometri CD200



Consider prophylaxis for tumor lysis syndrome ([See NHODG-B](#))

See monoclonal antibody and viral reactivation ([NHODG-B](#))

<sup>1</sup>The description represents the most common indolent presentation; however, there are some patients with GI or blood/bone marrow involvement only, which may express SOX11 and have an indolent course.

Note: All recommendations are category 2A unless otherwise indicated.

Clinical Trials: NCCN believes that the best management of any patient with cancer is in a clinical trial. Participation in clinical trials is especially encouraged.

# Hodgkin Lenfoma

- %20 Ig gen lokusu içeren genetik değişiklikler
- del1p, dup1q, del6q, del7q
- NFkappa/beta
- JAK/STAT yolağı



# TESTİÇİN ENDİKASYON Lenfositöz BK yüksekliği veya PY anormalliği

Reaktif Lenfositöz Şüphesi → Nedeni tedavi et ve BK tekrar değerlendir  
Açıklanamayan yükseklik

Flow sitometri ile lösemi/lenfoma fenotipleme

Anormal fenotipleme

Evet

Hayır

Atipik T-hücre veya NK fenotipi

CD19+ ve/veya CD20+  
Hafif zincir restriksiyonu

Reaktif lenfositöz düşün

T- veya NK lenfoproliferatif hastalıklar

B hücreli lenfoproliferatif hastalıklar

Enfeksiyöz etyoloji:  
Viral (EBV, CMV, HIV, HTLV, hepatit) Veya (tuberoskleroz, riketsiya, brucella, toxoplazma)

En az 1 pozitiflik:  
CD56, CD57 veya CD16

CD57-, CD56-,  
ve CD16-

Evet

Hayır

Geniş granüler lenfositik lenfoma (LGLL)

CD10+  
CD4+

Anji-immunoblastik B hücreli lenfoma

CD4+/-  
CD8+/-

Periferik T hücreli lenfoma düşün

CD3+

T-cell LGLL

CD30+

Anaplastik büyük hücreli lenfoma düşün

CD3-

NK-cell LGLL

CD4+  
CD7-  
CD26-

Sezary sendrom düşün

Tedavi et ve BK tekrar bak

CBC tekrar Otoimmün tiroidit, timoma düşün

Post-transplant

CD5-  
CD10+

CD5-  
CD10-

CD5+  
CD10-

DÜŞÜN EBV

DÜŞÜN  
Foliküler lenfoma  
Burkitt lenfoma  
Bazı diffüz büyük B hücreli lenfomalar  
Gastrik MALToma

CD11c+  
CD25+  
CD103+

Hairy cell lösemi

MYD88 L265P mutasyonu

Lenfoblastik lenfoma

Belirsiz CD20 CD23+  
Düşük hafif zincir yoğunluğu

KLL/SLL

PROGNOSTİK/TERAPOTİK  
Dizi analizi ile IGHV  
Mutasyon analizi  
Kromozom FISH,  
KLL Paneli

Belirgin CD20, CD43,  
CD23-, Cyclin D+  
Yüksek hafif zincir yoğunluğu

Mantle hücreli lenfoma

DİYAGNOSTİK  
Kromozom FISH, taze dokuda  
IGH-CCND1 Füzyonu, t(11;14)

DİYAGNOSTİK  
Flow sitometri ile T-hücre klonalitesi

DİYAGNOSTİK/PROGNOSTİK  
Kromozom FISH, interfaz IGH-BCL2 füzyonu, t(14;18)  
FISH ile BCL6 (3q27) Gen yeniden düzenlenmesi  
FISH ile t(8;14) IGH/MYC füzyonu  
MYC (8q24) Gen yeniden düzenlenmesi  
FISH ile lenfoma paneli

İSTENECEK  
Hairy cell lösemide Real Time PCR ile BRAF V600E Mutasyon taraması

# Kaynaklar

- National Comprehensive Cancer Network (NCCN Guidelines)
- European Society of Medical Oncology (ESMO) Guidelines
- International Myeloma Working Group (IMWG)
- Türk Hematoloji Derneđi (THD) Tanı ve Tedavi Kılavuzları

# MYELOM VE LENFOMALARDA ALGORİTMALAR

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