

IGH-BCL2 Fusion, t(14;18) by FISH

Indications for Ordering

Diagnosis of follicular lymphoma (FL) in conjunction with clinical, morphologic, and flow cytometric data

Test Methodology

- Fluorescence in situ hybridization (FISH)
- Dual color, dual fusion probes detect t(14;18) or *IGH-BCL2* gene rearrangement

Tests to Consider

Typical Testing Strategy

FL can often be diagnosed using the combination of morphology, immunohistochemistry (IHC), immunophenotyping, and clinical presentation

- Lymph node biopsy with morphologic and IHC evaluation
- Leukemia/lymphoma phenotyping by flow cytometry
- CD5 negative and CD10 positive suggests FL
- Bone marrow (BM) evaluation for staging – marrow involvement is common

Chromosome analysis by FISH – may be useful when the above combination does not yield a diagnosis

- Presence of *IGH-BCL2* fusion, t(14;18) supports a diagnosis of FL

Primary Test

[IGH-BCL2 Fusion, t\(14;18\) by FISH 3001298](#)

- Most sensitive method to detect *IGH-BCL2* fusion in formalin-fixed, paraffin-embedded (FFPE) tissue specimens

Related Tests

[Leukemia/Lymphoma Phenotyping Evaluation by Flow Cytometry 3001780](#)

- Aids in diagnosis of hematopoietic neoplasms

[Chromosome FISH, Interphase 2002298](#)

- Specific FISH probe for t(14;18) must be requested

[Chromosome Analysis, Bone Marrow 2002292](#)

- Diagnosis, prognosis, and monitoring of lymphoma in BM

[Chromosome Analysis, Solid Tumor 2002296](#)

- May identify additional, useful cytogenetic abnormalities in tissues that are not targeted by FISH assays

Disease Overview

Incidence – most common non-Hodgkin B-cell lymphoma

Signs/symptoms

- Early – usually asymptomatic
- Advanced
 - Peripheral and central lymphadenopathy
 - Splenomegaly
 - BM involvement
 - B symptoms – fever, night sweats, weight loss

Diagnostic Criteria for Follicular Lymphoma

- Morphology
 - Lymph node – neoplastic mixture of centrocytes and centroblasts recapitulating germinal centers of secondary lymphoid follicles
 - BM – paratrabecular lymphoid aggregates
- Flow cytometry
 - CD5 negative; CD10 positive monoclonal B cells

Genetics

Gene – *IGH-BCL2*

Structure/Function

- Translocation juxtaposes the immunoglobulin enhancer region (*IGH*, 14q32) with the *BCL-2* oncogene (18q21)
- Translocation causes overexpression of BCL-2, an anti-apoptotic protein, which prevents normal cell death

Test Interpretation

Results

- Positive – presence of the t(14;18) translocation substantiates a diagnosis of FL
 - 20% or more of the cells examined were abnormal
- Negative – absence of t(14;18) translocation

Limitations

- *IGH-BCL2* fusion, t(14;18) by FISH has not been validated for tissue fixed in alcohol-based or non-formalin fixatives
- Negative result does not exclude the possibility of translocations involving other partners nor rule out FL