

Liver Cytosolic Antigen Type 1 (LC-1) Antibody

Indications for Ordering

- Differential evaluation of autoimmune liver disease (ALD) of unknown etiology, especially autoimmune hepatitis (AIH) of childhood onset
- Consider
 - o If other serologic tests in ALD panel are negative, **or** o In combination with specific tests
 - Anti-LKM1
 - ANA
 - F-actin
 - SMA

Test Description

Liver-Kidney Microsome-1 Antibody, IgG

 Semiquantitative enzyme-linked immunosorbent assay (ELISA)

Liver Cytosolic Antigen Type 1 (LC-1) Antibody, IgG

• Qualitative immunoblot

Tests to Consider

Primary tests

Liver-Kidney Microsome - 1 Antibody, IgG 0055241

- Use in combination with Liver Cytosolic Antigen Type 1 (LC-1) Antibody, IgG
- More likely to be positive than LC-1

Liver Cytosolic Antigen Type 1 (LC-1) Antibody, IgG 2010711

 Use in combination with Liver-Kidney Microsome - 1 Antibody, IgG

Related tests

<u>Autoimmune Liver Disease Evaluation with Reflex to Smooth Muscle Antibody (SMA), IgG by IFA 2007210</u>

Recommended first-line panel for evaluation of ALD

 Negative results do not rule out disease

ANCA-Associated Vasculitis Profile (ANCA/MPO/PR-3) with Reflex to ANCA Titer 2006480

 Initial test in conjunction with Autoimmune Liver Disease Evaluation with Reflex to SMA, IgG by IFA for evaluation of ALD

Disease Overview

Incidence

AIH – 0.85-1.9/100,000 per year for white adults of northern European ancestry

Prevalence

- ALD
 - o 5% of all liver diseases
- AIH type 1
 - o Most common type of AIH
- AIH type 2
 - o Rare 4% of AIH individuals in the U.S.

Age

- AIH type 1 bimodal peaks (10-30 years, 40-50 years)
- AIH type 2 childhood

Physiology

AIH

- Etiology antibodies directed against the liver
- Chronic course with slow progression may resemble other chronic liver diseases (eg, alcoholic cirrhosis, chronic viral hepatitis)

Clinical presentation

- Clinical features vary widely, ~25% asymptomatic
 Arthralgias
 - o Anorexia, fatigue, lethargy, malaise
 - o Hepatomegaly, nausea, upper abdominal pain, jaundice
 - Progression to cirrhosis and liver failure possible
- Antibody-negative disease
 - Same presentation and histology as antibody-positive AIH

Diagnostic/prognostic issues

- May be difficult to clinically distinguish between AIH types 1 and 2
- Important to distinguish AIH type 1 from type 2 (prognostically)
 - Higher risk of fulminant course and progression to cirrhosis in type 2
 - o Cirrhosis often present in children at time of diagnosis
- Antibody testing may be helpful for diagnosis
 - o Antibodies to liver-kidney microsome-1 (LKM1) and/or LC-1 are diagnostic for AIH type 2
 - LKM1 antibodies should be analyzed by measuring antibodies to cytochrome P4502D6
- Diagnosis of exclusion
 - No other etiology found for liver disease/cirrhosis key to this diagnosis
- Liver biopsy may be appropriate in certain individuals
 Should be performed when diagnosis is still unclear
 Considered "gold standard"

Test Interpretation

Typical antibody pattern in AIH-1

- pANCA atypical staining
- SMA, F-actin positive
- ANA homogeneous pattern most common
- LC-1 negative
- SLA variably positive, more common in children

Results

Typical antibody pattern in AIH-2

- pANCA positive (rare) or negative
- LKM-1 positive
- LC-1 positive
- ANA, SMA, F-actin, M2, SLA negative

Limitations

- Negative antibody testing does not rule out ALD
- All interpretation of antibody patterns must be done in conjunction with clinical presentation
 - o Overlap may occur between diseases and antibodies
- Neither LKM-1 nor LC-1 has absolute diagnostic sensitivity for AIH type 2