

Immunological Tests.

## **ANA**

### **Antinuclear Antibody**

Directed against a variety of nuclear antigens and may be induced by drugs such as Hydralazine. Can be of any Ig class.

Different staining patterns

Homogenous staining; suggests lupus

Speckled Staining: Suggests mixed connective tissue disease

Centromere Staining; suggests CREST syndrome

Found in

Drug induced lupus (100%)

SLE (99%)

Scleroderma (97%)

Sjogrens syndrome (96%)

Mixed connective tissue disease (93%)

Polymyositis(78%)

### **Extractable Nuclear Antigens**

These are usually associated with positive ANA

Anti Ro Sjogrens syndrome, Congenital Heart Block, ANA negative SLE

Anti La primary Sjogrens syndrome

Anti Sm SLE (20%)

Anti RNP MCTD (100%)

Anti Jo1 Polymyositis

Anti SCL 70 progressive systemic sclerosis (20%)

Anticentromere CREST Syndrome

**Antineutrophil cytoplasmic antibody (ANCA)** has been shown to be a serological marker for a spectrum of diseases characterised by systemic necrotising vasculitis and crescentic vasculitis.

The presence of ANCA characterises a group of vasculitides, including Wegener's granulomatosis (WG),

Microscopic polyangiitis (MPA),

Churg–Strauss syndrome (CSS)

Indications for ANCA testing:

- patients suspected of Wegener's granulomatosis, microscopic polyangiitis, Churg–Strauss syndrome or idiopathic necrotising glomerulonephritis
- chronic destructive disease of the upper airways Pulmonary nodules (not obviously malignant)
- subglottic stenosis of the trachea
- pulmonary–renal syndrome
- glomerulonephritis
- vasculitis of the skin with evidence of systemic disease
- mononeuritis multiplex
- retro-orbital mass
- any other condition resembling systemic vasculitis
- 

ANCA patterns and antibody specificity ANCA describes a number of circulating autoantibodies specifically directed against the cytoplasmic constituents of neutrophils and monocytes

- two ANCA patterns were originally identified by indirect immunofluorescence (IIF): the cytoplasmic (C-ANCA) and the perinuclear (P-ANCA) patterns
  - these apparent morphological differences are purely artefactual and based on the fixative used to preserve the neutrophil substrate
    - 'classical' C-ANCA is associated with antibodies reacting with the 29–30 kDa elastinolytic enzyme, serine proteinase 3 (PR3)
      - composed of 229 amino acids and found in the azurophilic granules of neutrophils and monocytes
    - 'classical' P-ANCA pattern is associated with antibodies to myeloperoxidase (MPO), a 140 kDa heterodimeric enzyme also associated with the antimicrobial properties of neutrophils

#### **Presence of C-ANCA (1)**

- Wegener's granulomatosis usually
- microscopic polyangiitis sometimes
- idiopathic glomerulonephritis sometimes
- Churg-Strauss syndrome sometimes
- ulcerative colitis sometimes

Other positive c-ANCA test results have been reported in patients with tuberculosis, Hodgkin's lymphoma, human immunodeficiency virus infection, nasal septal perforation, monoclonal gammopathies, and drug-induced Wegener-like disease (2).

P-ANCA pattern is associated with antibodies to myeloperoxidase (MPO), a 140 kDa heterodimeric enzyme also associated with the antimicrobial properties of neutrophils if IIF and ELISA results are combined, the presence of P-ANCA and anti-MPO has 99% specificity for the diagnosis of primary systemic vasculitis, as does the combination of C-ANCA and anti- PR3 P-ANCA and anti-MPO are more often seen in microscopic polyangiitis (MPA), Churg-Strauss Syndrome (CSS) and idiopathic necrotising glomerulonephritis

- in the context of vasculitis, C-ANCA in more than 90% of cases is directed against PR3, whereas in 80–90% of cases P-ANCA reacts with MPO

#### Auto- antibodies in GI and Liver disease

Anti-mitochondrial antibody – primary biliary cirrhosis (96%)

Anti – smooth muscle antibody - auto-immune hepatitis, cryptogenic cirrhosis

Gastric parietal cell antibodies - pernicious anaemia (90%) gastric atrophy (40%)

Intrinsic Factor Antibodies – pernicious anaemia (70%)

Anti-gliadin, anti –endomesial antibody – coeliac disease (continued positive anti-gliadin tests suggest inadequate dietary

#### Autoantibodies in thyroid disease

Antithyroglobulin antibody high titre in autoimmune thyroiditis (90%)

Antimicrosomal antibodies low titre in Graves Disease (35%) and in adenocarcinoma (10%)

Anti glomerular basement membrane goodpastures syndrome

Acetylcholine receptor antibody – myaesthesia gravis (87%)