

Phosphatidyl-choline, -ethanolamine, -glycerin, -inositol, -serine IgA, IgG, IgM  
 see Phospholipid-Antibodies, Serum

Platelet, free antibodies (serum), bound antibodies (EDTA-blood)  
 see Platelet Antibodies (free, bound)

RNP-U1 see Ribonucleoprotein U1-snRNP Antibody

Proteinase-3 (PR3) see Antineutrophil Cytoplasmatic Antibody (ANCA)

Scl-70 see Scl-70 Antibody

Scleroderma Antibody see Scl-70 Antibody

Sm Protein (Smith) see Smith (SM) Antibody

Smooth muscle (ASMA) see Smooth Muscle Antibodies (SMA)

Smith (SM) Antibody

Soluble liver antigen see Soluble Liver Antigen (SLA)-Antibody (Anti-SLA)

Sjogren Antibodies see SS-A/Ro and SS-B/La Antibodies

SS-A (Ro) see SS-A/Ro and SS-B/La Antibodies

SS-B (La, Ha) see SS-A/Ro and SS-B/La Antibodies

Thyroperoxidase (MAK)

Thyroid (thyroglobulin, microsomal, TSH-Receptor) see Thyroglobulin Antibody

Thyroperoxidase Autoantibody

Thyrotropin Receptor Antibody, Serum

Topoisomerase I Antibody see Scl-70 Antibody

**Sampling:** for each test 1 mL of serum

**B- and T Lymphocytes** see Lymphocyte Immunophenotyping

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## **Bartonella henselae, Serology**

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**Synonyms:** Bacillary Angiomatosis Serology  
 Cat Scratch Disease Serology

**Background:** Bartonella species includes *B. quintana* and *B. henselae*, responsible for cutaneous bacillary angiomatosis, bacillary peliosis of the liver and spleen, fever and bacteriemia (*Bartonella* bacteremic syndrome), endocarditis (so called culture negative endocarditis). Associated only with *B. quintana*: trench fever and with *B. henselae* only: Cat scratch disease.

Patients presenting with cat scratch disease are in >80% younger than 20 years, an inoculation papule is seen in 50 % followed by local lymphadenopathy and cats bites or scratches in 75 %. *B. henselae* has been isolated from blood and tissue. In the normal population, approx. 15% of the people showed IgG titres > or = 1:128. In patients with cat scratch disease, 85 % display titres > or = 1:128. Cross reactivity within other Bartonella species occur, e.g. *Bartonella quintana*, associated with trench fever. Seroconversion may take up to 3 weeks post infection; a late specimen may be helpful.

**Sampling:** 1 mL serum early and a specimen 3-6 weeks post infection.

**Reference Interval:** Negative: titer < 1:64  
A threefold increase in titre is considered significant.

**Bence-Jones Protein see** Free Light Chains Structure, Urine

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**Beta-2-Microglobulin, Serum or Urine see** Microglobulin  $\beta$ -2-, Serum or Urine

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## Beta-Crosslaps, Serum

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**Related Information:** Pyridinolines  
Vitamin D, Serum

**Background:** Collagen  $\alpha_1$  or  $\alpha_2$  chains are linked at the carboxy or aminoterminal end with the helix of adjunct collagen by hydroxypyridiniumderivates. During proteolytic breakdown of collagen type I the hydroxypyridinium interlinked  $\alpha_1$  or  $\alpha_2$  chains are released into serum and urine and are known as N-terminal crosslaps (NTX), C-terminal crosslaps (CTX) and collagen type I C-terminal telopeptide. All peptides are released during the turnover of skin and bone tissue, however, the major amount in serum or urine of CTX and NTX are products of bone resorption.

Alpha-CTX is the non-isomerized form of aspartic acid; the beta form which is the isomerized form, forms spontaneously but delayed posttranslational. It indicates tissues degradation composed of mature collagen such as mature bone, but not recently formed bone tissue.

**Sampling:** All types of collagen resorption products have a circadian rhythm peaking during the night and with a low during daytime (plus/minus 30%)

Standard procedure:

Schedule sampling between 7:30 -8:30 in the morning after a strict 12h fasting period (only pure water allowed!). Sample into EDTA tube. EDTA whole blood is stable for 8h at room temperature; EDTA plasma is stable for 24h room temperature, 1 month at -20°C.

**Reference Interval:**

Male		< 0.60 ng/mL
Female	< 45 years	< 0.60 ng/mL
	> 45 years	< 1.00 ng/mL

Levels increases after menopause in serum and urine by 50-70%

During anti-bone-resorptive therapy, levels are expected to decrease after 6 month by 55-70%

## Bile Acids, Serum or Feces

**Related Information:** Bilirubin, Fractionated, Serum

Gamma Glutamyl Transferase (Gamma-GT), Serum

**Synonyms:** Cholylglycine; Bile Acid Conjugates; Cholic and Chenodeoxycholic Acid

**Background:** Bile acids are produced by hepatocytes from cholesterol. The two primary bile acids are cholic and chenodeoxycholic acids. In bile they solubilize cholesterol, thus being essential for cholesterol elimination and conjugate with glycine, taurine, amino acids. These primary bile acids undergo bacterial degeneration to form deoxycholic, lithocholic acids and oxidation products (ursodeoxycholic acids). 95% of bile acids are reabsorbed in the small intestine and undergo enterohepatic circulation. Normally < 1% of the bile acid pool is found in serum, but is elevated in cirrhosis, obstructive jaundice, hepatitis due to decreased clearance, but with normal ratio secondary to primary bile acids. In cirrhosis, a disproportionate decrease in cholic acid occurs with a reduced ratio of primary to secondary bile acids. In cholestasis, no secondary bile acids are formed.

Determination of bile acids in serum is one of the most sensitive methods to evaluate liver function.

**Sampling:** Serum: 1 mL serum. Overnight fasting sample preferred.

Feces: Collect feces for 24h in a sterile container without preservatives.

Keep refrigerated during collection period. Note total weight on the request sheet. Before aliquoting approx. 2 g of stool, mix well.

**Reference Interval:** Serum: < 7.0  $\mu\text{mol/L}$

Feces: 410 - 1210  $\mu\text{mol} / 24\text{h}$

## Bilharzia (Schistosomiasis) Serology

**Background:** For definite diagnosis of schistosomiasis identification of ova of *S. haematobium*, *S. mansoni*, *S. japonicum* in rectal or bladder biopsy in feces or urine is necessary.

Antibodies persist following therapy. Differentiation between recent or chronic infection is not possible; Positive results indicate chronic active or inactive schistosomiasis. No distinction between the sites of infection or the various forms of the fluke by serology.

**Sampling:** 1 mL serum, avoid hemolysis, avoid highly lipemic serum

**Reference Interval:** Serum antibody titer negative: < 1:16

## Bilirubin Fractionated, Serum

<b>Related Information:</b>	Acetaminophen, Serum
	Alanine Aminotransferase (AST), Serum
	Alkaline Phosphatase, Serum
	Amylase, Isoenzymes, Serum
	Amylase, Total, Serum
	Amylase, Total, Urine
	Aspartate Aminotransferase (AST), Serum
	Ethanol, Blood, Urine
	Gamma-Glutamyl Transferase (Gamma-GT), Serum
	Hepatitis A Antibodies, IgG and IgM (IgG anti-HAV, IgM anti-HAV)
	Hepatitis B (HBV), Serology and Antigen Detection
	Hepatitis B Virus DNA Detection (HBV-DNA)
	Hepatitis C Antibody (Anti-HCV)
	Hepatitis C Genotyping
	Hepatitis C Virus RNA Quantification (HCV-RNA)
	Hepatitis D Antibody (Anti-Delta Serology)
	Hepatitis E Antibody (Anti-HEV)
	Leucine Aminopeptidase (LAP), Serum
	Lipase, Serum
	Prothrombin Time
<b>Test includes:</b>	Total, conjugated and unconjugated Bilirubin
<b>Synonyms:</b>	Total, direct, indirect Bilirubin

**Background:** Produced in the breakdown of heme and in the reduction of biliverdin, circulating in the plasma and is conjugated, called the direct form, by the liver to bilirubin diglucuronide, a water soluble pigment excreted in the bile.

Conjugated, direct bilirubin in serum is the most sensitive test for liver function and occurs in biliary diseases, intra- and extrahepatic lesions, hepatitis, cirrhosis, neoplasms, cholestatic drug reactions and in faulty excretory function of hepatocytes such as Dubin Johnson syndrome, Rotor's syndrome.

A total bilirubin increase is associated with:

Hemolytic diseases, hepatocellular dysfunction, diseases of hepatic ducts or common bile ducts. Also Gilbert's syndrome, an asymptomatic hereditary jaundice with increasing hyperbilirubinemia caused by Glucuronyl transferase deficiency. Dubin Johnson syndrome, anorexia, prolonged fasting, pulmonary embolism or infarction, congestive heart failure.

Limitations:

Drugs can interfere with the diazo method used for the test as well as causing jaundice in vivo. Such drugs are: diphenylhydantoin, azathioprine, phenothiazine, erythromycin, penicillin, sulfonamides, contraceptives, anabolic androgenic steroids, halothane, aminosalicic acid, isoniazid, methyl dopa, indomethacin, pyrazinamide.

False positive reactions with the diazo method are often seen during cephotetan, pansporin, cefuroxime therapy.

**Sampling:** 1 mL serum or capillary plasma (heparin or EDTA), avoid hemolysis, protect from light. Transport to laboratory within 4h or separate serum/plasma within 8 h and refrigerate thereafter.

**Reference Interval:**

Neonates:	Bilirubin, total (mg/dL), maximum		
	Age	premature	full term
	<1 day	8	6
	24-48h	12	10
	3-5 days	15	12
	6-7days	15	10
Children and adults:	Total :		< 1.2 mg/dL
	Conjugated or direct bilirubin:		< 0.3 mg/dL
	Unconjugated or indirect bilirubin:		< 0.9 mg/dL

Critical value:

Newborn: Total bilirubin more than 15 mg/dL in term infants,  
10-15 mg/dL in prematures.  
Increase of more than 1 mg/dL per hour for more than 6 hours.

## Biotin (Vitamin H), Serum

**Background:** Tissue biotin is a cofactor for carboxylation of pyruvate, acetyl-coenzyme A (CoA), propionyl CoA, and beta-methylcrotonyl CoA .

Deficiency presents as severe exfoliative dermatitis and alopecia, similar to zinc deficiency. Secretion in the urine as intact biotin, and to lesser amounts as bis-norbiotin and biotin sulfoxide. Sources for biotin are organs, egg yolk, milk, fish, and nuts. Biotin is stable to cooking. Daily recommended intake approx. 30 µg, the bacterial intestinal flora contributes in part to the supply. Therapeutic large doses (5-10 mg) are applied to infants with seborrhea or genetic alteration of biotin dependent enzymes, no toxicity has been reported so far.

**Sampling:** 2 mL serum

**Reference Interval:** > 200 pg/mL  
< 100 pg/mL interpreted as a biotin deficiency

## Bordetella pertussis, Culture

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**Related information:** Bordetella pertussis, Serology

**Background:** B. pertussis causes whooping cough. The disease begins with mild respiratory infection and develops within 2 weeks into the distinctive whooping cough. Pertussis like symptoms are also caused by Chlamydia trachomatis, adenoviruses, and respiratory syncytial virus. Diagnosis can be made by B. pertussis cultivation from nasopharyngeal specimens. Despite vaccination programs, pertussis is still one of the most common causes of death from infectious diseases worldwide. Pertussis is highly contagious, immunized persons can be transiently colonized and spread the organism.

**Sampling:** Perform a nasopharyngeal swap near septum and the floor of the nose under rotating. Inoculate at bedside immediately. Since routine agar does not support the growth of the organism, either the swap should be plated at beside on Bordet-Gengou agar plate, derived from the laboratory or a special transport medium (to order from the laboratory) has to be used. Successful cultivation even under optimal conditions is poor about 50%.

**Reference Interval:** Report on diagnostic finding  
Growth or no growth of Bordetella pertussis or B. parapertussis

## Bordetella pertussis, Serology

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**Related information:** Bordetella pertussis, Culture

**Background:** The value of the antibody detection is limited by the time required for seroconversion. There is a lack of association between antibody levels and protective immunity. Patients with acute infection develop IgG and IgM and IgA antibody responses, vaccinated individuals show increase of IgM and IgG, IgA antibodies are lacking.

**Sampling:** 1 mL serum

**Reference Interval:** Differentiation of immunoglobulin class

IgA antibody negative:	< 9 RE/mL
borderline:	9-11 RE/mL
positive:	>11 RE/mL
IgG antibody negative:	< 9 RE/mL
borderline:	9-11 RE/mL
positive:	>11 RE/mL
IgM antibody negative:	< 9 RE/mL
borderline:	9-11 RE/mL
positive:	>11 RE/mL

## Borrelia, Serology

**Synonyms:** Borreliosis; Lyme Disease

**Test includes:** B. burgdorferi, -afzelii, -garinii

**Background:** *Borrelia* spp. are irregular coiled spirochetes, Giemsa and silver stain positive, culturable from the tick vector but not from patient's specimens in serum containing media and transmitted by arthropods. Diseases caused are Lyme disease and relapsing fever.

*Borrelia burgdorferi* is the cause of Lyme disease transmission by *Ixodes dammini*, or *I. pacificus*. The reservoirs are mammals (white footed mouse) and large mammals. Feeding time of the ticks must exceed 24 h to transmit a sufficient dose for infection. In the US, the most common tick borne diseases are Lyme disease, Rocky Mountain spotted fever, ehrlichiosis, relapsing fever, and tularemia.

*B. burgdorferi* is disseminated into heart, joints, CNS. The disease occurs stage wise: Erythema chronicum migrans, a non-pruritic red rash with a clear center at the site of the bite in 75% of the cases and later usually accompanied by flu-like transient symptoms (arthralgias, headache, fever chills, fatigue). The second stage with heart and CNS involvement (neuroborreliosis) occurs after weeks to month (myocarditis, aseptic meningitis, neuropathies with Bell's palsy). The third phase is characterized by arthritis of the large joints and CNS symptoms.

*B. recurrentis* is transmitted by the human body louse and *B. hermsii* by soft ticks (*Ornithodoros*) with rodents as the reservoirs.

Antibodies of the IgM class are detectable after 2-3 weeks post infection, peaking at 3-6 weeks.

Treatment: Doxycycline, amoxicillin, penicillin G, ceftriaxone.

Prevention: Two Lyme disease vaccines have been developed: LYMErix™ by SmithKline Beecham Pharmaceuticals and ImuLyme™ by Pasteur Merieux Connaught, however the vaccine is not effective in all individuals.

**Sampling:** 1 mL serum

**Reference Interval:** Screening by EIA method. Validation by Western blot assay. Differentiation of immunoglobulin class for *B. burgdorferi*, -afzelii, -garinii antibody.

IgG antibody negative:	< 17 RE/mL
borderline:	17-20 RE/mL
positive:	> 20 RE/mL
IgM antibody negative:	< 15 RE/mL
borderline:	15-20 RE/mL
positive:	> 20 RE/mL

## Brain Natriuretic Peptide, Serum

**Related Information:** Albumin, Serum  
Creatinine, Serum or Plasma  
Digoxin, Serum  
Magnesium (Mg), Serum  
Osmolality, Serum  
Renin Activity, Plasma

**Synonyms:** B-Type Natriuretic Peptide; BNP; Natriuretic Peptide, Brain

**Background:** BNP, is a polypeptide, which is produced by the ventricular myocardium under stimulation of volume expansion and overload. The hormone displays natriuretic and vasodilatory effects, suppressing the renin angiotensin system.

BNP is a useful parameter in diagnosis and treatment of congestive heart disease (CHF) since correlating with severity and prognosis. Also helpful in the diagnosis of cardiac versus non-cardiac dyspnea.

Limitation: In the very early stage of acute CHF presenting with dyspnea or edema, BNP may be normal. Elevated levels may occur in pulmonary embolism, right heart failure, and pulmonary hypertension.

**Sampling:** 2 mL serum, place on ice, if transit time is less than 6h, otherwise freeze.

**Reference Interval:**

In healthy individuals < 125 pg/mL

Varies with age and sex.

Values correlate with degree of congestive heart failure.

	BNP pg/mL 5th-95th percentile approx	median approx (pg/mL)
Patients with cardiac disease, but without symptoms	15-499	95
Patients with limited ability to perform exercise	10-1100	220
Patients with limitations in ordinary physical activity	40-1300	450
Patients not able to perform exercise	150-1300	1000

## Brucella, Serology

**Background:** Brucella species organisms are small gram negative rods. The major human pathogens are *B. melitensis* (reservoir goat and sheep) and *B. abortus* (cattle).

Transmission is via unpasteurized milk or cheese from infected animals of the domestic livestock or by skin contact.

The organism localizes in the reticuloendothelial cells (liver, spleen, bone marrow, lymph nodes) and replicates intracellularly. After an incubation period of 1-3 weeks, fever, chills, fatigue, malaise and weight loss occurs. The undulating fever occurs in a minority of patients. Liver, spleen and lymph nodes are often enlarged.

Drug of choice: Tetracyclines plus rifampin

**Sampling:** 1 mL serum

**Reference Interval:** Antibody agglutination test: negative

## Bunyaviruses, Serology

**Background:** Bunyaviruses are enveloped viruses, with helical capsid symmetry, 100 nm in size, circular, single stranded with negative polarity RNA. The virus was first isolated in Africa (Bunjamwera), causing encephalitis. Now 4 distinct Hantaviruses are known in N. America and more than 20 serotypes worldwide. As a member of the Bunyavirus family, Hantaviruses cause Korean hemorrhagic fever in Europe and Asia with headache, petechial hemorrhages, shock, and renal failure. Mortality is about 10%. Hantaviruses are also classified as rodent-borne viruses, since they are transmitted from rodents directly without an arthropod vector in opposition to arboviruses (arthropod-borne). In the early nineties, first in New Mexico an outbreak (hantavirus pulmonary syndrome) occurred clinically presenting influenza-like syndromes (fever, myalgia), followed in 3-6 days by progressive cough and shortness of breath with pleural effusions and in severe cases by respiratory failure, tachypnea, tachycardia, hypotension. Laboratory chemistry revealed hemoconcentration, thrombocytopenia, increased partial thromboplastin time, leucocytosis, increased serum lactate dehydrogenase and aspartate aminotransferase levels were observed. Antibodies may be present at the time of clinical symptoms. The virus is transmitted by inhalation of aerosols created from urine or feces from deer and mice (*Peromyscus*). Transmission from person to person was observed in an outbreak in Argentina. Mortality in Hantavirus pulmonary syndrome is close to 40%. No vaccine, no effective antiviral drug available.

**Sampling:** 1 mL serum at the acute phase and follow up sera 2-3 weeks later.

**Reference Interval:** Differentiation of antibodies of immunoglobulin classes IgG and IgM for Hantaviruses (Hantaan, Puumala, Dobrava, Seoul) and Sandfly fever virus (Toscana virus)  
(Method: Immunoblot)