

Epstein-Barr Virus (EBV), Molecular Detection, PCR, Varies

### Overview

#### **Useful For**

Rapid qualitative detection of Epstein-Barr virus (EBV) DNA in specimens

Diagnosis of disease due to EBV

This test **should not be used** to screen asymptomatic patients.

#### Method Name

Real-Time Polymerase Chain Reaction (PCR)/DNA Probe Hybridization

#### NY State Available

Yes

### Specimen

Specimen Type Varies

### **Necessary Information**

1. Specimen source is required.

2. Source information must include main anatomical site of collection.

#### Specimen Required

Submit only 1 of the following specimens:

Specimen Type: Fluid
Sources: Spinal fluid, sterile body fluids (peritoneal fluid/ascites, pericardial fluid, pleural fluid/thoracentesis), amniotic, or ocular
Supplies: Sarstedt Aliquot Tube, 5 mL (T914)
Preferred: Sterile screwcap 5-mL plastic vial
Acceptable: Sterile container
Specimen Volume: 0.5 mL
Collection Instructions: Do not centrifuge.

Specimen Type: Fluid
 Sources: Respiratory; bronchial washing, bronchoalveolar lavage, nasopharyngeal aspirate or washing, sputum, or tracheal aspirate
 Supplies: Sarstedt Aliquot Tube, 5 mL (T914)



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**Container/Tube:** 

Preferred: Sterile screwcap 5-mL plastic vial Acceptable: Sterile container Specimen Volume: 1.5 mL

Specimen Type: Swab
Sources: Eye and upper respiratory (nasal, throat)
Supplies:
-Culturette (BBL Culture Swab) (T092)
-BD E-Swab (T853)
-M4-RT(T605)
Container/Tube: Multimicrobe media (M4-RT, M4, M5, Bartels, or Jiangsu) and E-Swab or Culturette
Collection Instructions: Place swab back into multimicrobe media.

Specimen Type: Bone marrow Container/Tube: Lavender top (EDTA) only Specimen Volume: 0.5 mL Additional Information: Clotted specimens will be rejected.

Specimen Type: Fresh tissue
Supplies: M4-RT (T605)
Container/Tube:
Preferred: Sterile container containing multimicrobe medium (M4-RT, M4, M5, Bartels, or Jiangsu)
Acceptable: Sterile container containing 1-2 mL of sterile saline
Specimen Volume: Entire collection
Collection Instructions: Submit only fresh tissue. Fixed tissue is not acceptable.

## Forms

If not ordering electronically, complete, print, and send a <u>Microbiology Test Request</u> (T244) with the specimen.

## Specimen Minimum Volume

Ocular Fluid, Spinal Fluid: 0.3 mL Sterile body fluids (peritoneal fluid/ascites, pericardial fluid, pleural fluid/thoracentesis): See Specimen Required Respiratory Specimens: 1 mL Tissue: 2 x 2-mm biopsy

## **Reject Due To**

Calcium	Reject
alginate-tipped	
swab	
Wood swab	
Transport	
swab	
containing gel	



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Formalin-fixed
and
paraffin-embe
dded tissues
Heat-inactivate
d specimens
Dry/flocked
ESwabs

### **Specimen Stability Information**

Specimen Type	Temperature	Time	Special Container
Varies	Refrigerated (preferred)	7 days	
	Frozen	7 days	

## Clinical & Interpretive

#### **Clinical Information**

Epstein-Barr virus (EBV) is the causative agent of infectious mononucleosis, Burkitt lymphoma, and in Southern China, nasopharyngeal carcinoma. EBV-associated central nervous system (CNS) disease is most frequently associated with primary CNS lymphoma in patients with AIDS. In addition, CNS infection associated with the detection of EBV DNA can be seen in immunocompetent patients.

#### **Reference Values**

Negative Reference values apply to all ages.

#### Interpretation

Detection of Epstein-Barr virus (EBV) DNA in cerebrospinal fluid (CSF) supports the clinical diagnosis of central nervous system (CNS) disease due to the virus. EBV DNA is not detected in CSF from patients without CNS disease caused by this virus.

#### Cautions

A negative result does not eliminate the possibility of Epstein-Barr virus (EBV) infection of the central nervous system.

This assay may detect viremia or viral shedding in asymptomatic individuals. However, this assay is only to be used for patients with a clinical history and symptoms consistent with EBV infection and must be interpreted in the context of the clinical picture.

#### Supportive Data

Thirty negative specimens of each matrix accepted for this assay were spiked with Epstein-Barr positive control plasmid at the approximate limit of detection (LOD; 10-20 targets/mcL). The 30 spiked specimens of each type were run in a blinded manner along with 30 negative (non-spiked) specimens; 93% to 100% of the spiked specimens were positive and



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100% of the non-spiked specimens were negative.

#### Analytical Sensitivity/LOD:

The 95% LOD for this assay is less than 10 targets per microliter using plasmid and whole virus spiked into matrix. The LOD was confirmed in each matrix type that is accepted for testing with this assay.

#### Analytical Specificity:

No <u>polymerase chain reaction</u> signal was obtained from extracts of 40 bacterial and viral isolates that could cause similar symptoms including herpes simplex virus 1 and 2; cytomegalovirus; varicella zoster virus; and human herpesvirus (HHV) 6, HHV 7, and HHV 8.

#### Precision:

Interassay precision was 100% and intraassay precision was 100%.

#### Reportable Range:

This is a qualitative assay and results are reported as either negative or positive for targeted Epstein-Barr virus DNA.

#### **Clinical Reference**

1. Tachikawa N, Goto M, Hoshino Y, et al. Detection of toxoplasma gondii, epstein-barr virus, and JC virus DNAs in the cerebrospinal fluid in acquired immunodeficiency syndrome patients with focal central nervous system complications. Intern Med. 1999;38(7):556-562. doi:10.2169/internalmedicine.38.556

 Antinori A, Cingolani A, De Luca A, et al. Epstein-Barr virus in monitoring the response to therapy of acquired immunodeficiency syndrome-related primary central nervous system lymphoma. Ann Neurol. 1999;45(2):259-261
 Cingolani A, De Luca A, Larocca LM, et al. Minimally invasive diagnosis of acquired immunodeficiency syndrome-related primary central nervous system lymphoma. J Natl Cancer Inst. 1998 ;90(5):364-369. doi:10.1093/jnci/90.5.364

 Niller HH, Wolf H, Minarovits J: Regulation and dysregulation of epstein-barr virus latency: implications for the development of autoimmune disease. Autoimmunity. 2008:41(4):298-328. doi:10.1080/08916930802024772
 Studahl M, Hagberg L, Rekvdar E, Bergstrom T. Herpesvirus DNA detection in cerebrospinal fluid: difference in clinical presentation between alph-, beta-, and gamma-herpes viruses. Scand J Infect Dis. 2000;32(3):237-248. doi:10.1080/00365540050165857

Lau AH, Soltys K, Sindhi RK, Bond G, Mazariegos GV, Green M. Chronic high epstein-barr viral load carriage in pediatric small bowel transplant recipients. Pediatr Transplant. 2010;14(4):549-553. doi:10.1111/j.1399-3046.2009.01283.x
 Fugl A, Andersen CL Epstein-barr virus and its association with disease - a review of relevance to general practice. BMC Fam Pract. 2019;20(1):62. doi:10.1186/s12875-019-0954-3

## Performance

#### **Method Description**

Viral nucleic acid is extracted by the MagNA Pure automated instrument (Roche Applied Science) from clinical specimens. Primers are directed to the target gene (latent membrane protein). The LightCycler instrument amplifies and monitors by fluorescence the development of target nucleic acid sequences after the annealing step during <u>polymerase</u>



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<u>chain reaction (PCR)</u> cycling. This is an automated PCR system that can rapidly detect (30-40 minutes) amplicon development through stringent air-controlled temperature cycling in capillary cuvettes. The detection of amplified products is based on the fluorescence resonance energy transfer (FRET) principle. For FRET product detection, a hybridization probe with a donor fluorophore, fluorescein, on the 3' end is excited by an external light source and emits light that is absorbed by a second hybridization probe with an acceptor fluorophore, LC-Red 640, at the 5' end. The acceptor fluorophore then emits a light of a different wavelength that can be measured with a signal that is proportional to the amount of specific PCR product. Melting curve analysis is performed following PCR amplification. Starting at 45 degrees C, the temperature in the thermal chamber is slowly raised to 80 degrees C and the fluorescence is measured at frequent intervals. Analysis of the PCR amplification and probe melting curves is accomplished through the use of LightCycler software.(Unpublished Mayo method)

PDF Report

No

Day(s) Performed Monday through Sunday

**Report Available** Same day/1 to 4 days

**Specimen Retention Time** 7 days

#### **Performing Laboratory Location** Mayo Clinic Laboratories - Rochester Main Campus

## Fees & Codes

#### Fees

- Authorized users can sign in to <u>Test Prices</u> for detailed fee information.
- Clients without access to Test Prices can contact <u>Customer Service</u> 24 hours a day, seven days a week.
- Prospective clients should contact their account representative. For assistance, contact <u>Customer Service</u>.

## **Test Classification**

This test was developed and its performance characteristics determined by Mayo Clinic in a manner consistent with CLIA requirements. It has not been cleared or approved by the US Food and Drug Administration.

#### **CPT Code Information**

87798

#### LOINC<sup>®</sup> Information

Test ID Test Order Name	Order LOINC <sup>®</sup> Value
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EBVPV	Epstein-Barr Virus, PCR, Varies	5005-4
Result ID	Test Result Name	Result LOINC <sup>®</sup> Value
EBVS	Specimen Source	31208-2