Galactosemia (Galactokinase Deficiency): *GALK1* Gene Deletion/Duplication

**Test Code:** JA  
**Turnaround time:** 2 weeks  
**CPT Codes:** 81228 x1

### Condition Description
Galactokinase (GALK) deficiency is one of the three known forms of galactosemia, along with galactose-1-phosphate uridylyltransferase (GALT) deficiency (classic galactosemia) and UDP-galactose-4'-epimerase (GALE) deficiency [1-2]. GALK deficiency is an autosomal recessive disorder characterized by an elevation of blood galactose concentration and diminished galactose-1-phosphate concentration, leading to production of alternative metabolic products such as galactitol [3]. Galactokinase-deficiency may present in the neonatal period with cataracts; no other clinical complications have been consistently associated with GALK-deficiency [2].

GALK deficiency should be considered in individuals with cataracts, elevated red cell galactose, galactosuria, or elevated urinary galactitol and normal GALT enzyme activity. GALK activity is used to rule-out variant galactosemia due to galactokinase deficiency which should not be confused with classical galactosemia secondary to GALT deficiency, or epimerase-deficiency galactosemia secondary to GALE deficiency. The vast majority of patients with biochemical diagnosis of GALK deficiency have mutations in the GALK1 (17q25) gene [4-6]. Gene sequence analysis is available to test for mutations in the GALK1 gene in patients with a biochemical diagnosis of GALK deficiency (IQ).


Click [here](http://www.ThinkGenetic.com) for the GeneReviews summary on this condition. Also, refer to the Comprehensive Galactosemia Panel for a disease overview.

### References:
4. Sangiuliano et al. Biochemical Characterization of Two GALK1 Mutations in Patients with Galactokinase Deficiency Hum Mutat 2004 Apr;23(4):396-403
5. Park et al. Molecular and biochemical characterization of the *GALK1* gene in Korean patients with galactokinase deficiency. Mol Genet Metab. 2007. 91:234-8

### Genes

**GALK1**

### Indications
This test is indicated for:
- Individuals with elevated blood galactose but with normal GALT and GALE enzyme activities.
- Carrier testing for individuals with a family history of GALK deficiency.

### Methodology
Detection is limited to duplications and deletions. Array CGH will not detect point mutations or intronic mutations.

Please note that a "backbone" of probes across the entire genome are included on the array for analytical and quality control purposes. Rarely, off-target copy number variants causative of disease may be identified that may or may not be related to the patient's phenotype. Only known pathogenic off-target copy number variants will be reported. Off-target copy number variants of unknown clinical significance will not be reported.

### Detection
The vast majority of patients with clinical and biochemical diagnosis will have an abnormal DNA test.
Clinical Sensitivity: 26/26 mutations identified in 13 patients [8], 4/4 mutations identified in 2 patients [9].
Analytical Sensitivity: ~99%

### Specimen Requirements
Submit only 1 of the following specimen types

* Preferred specimen type: Whole Blood

### Type: Whole Blood
Specimen Requirements:

Disclaimer: This information is confidential and subject to change without notice. It may not be reproduced in whole or part unless authorized in writing by an authorized EGL representative.
In EDTA (purple top) or ACD (yellow top) tube:
Infants (2 years): 3-5 ml
Older Children & Adults: 5-10 ml

Specimen Collection and Shipping: Refrigerate until time of shipment. Ship sample within 5 days of collection at room temperature with overnight delivery.

**Type: Saliva**

Specimen Requirements:

Oragene™ Saliva Collection kit (available through EGL) used according to manufacturer instructions.

Specimen Collection and Shipping: Store sample at room temperature. Ship sample within 5 days of collection at room temperature with overnight delivery.

**Special Instructions**

Submit copies of diagnostic biochemical test results with the sample. Sequence analysis is required before deletion/duplication analysis by targeted CGH array. If sequencing is performed outside of Emory Genetics Laboratory, please submit a copy of the sequencing report with the test requisition.

**Related Tests**

- **GALT and GALE Gene Sequencing** for transferase deficient and epimerase deficient galactosemia
- **Comprehensive Galactosemia Panel** includes: GALT enzyme activity, isoenzyme pattern, gal-1-P concentration
- **Urine Galactitol Concentration**
- **Custom Diagnostic Mutation Analysis (KM)** is available to family members if mutations are identified by sequencing.
- **Prenatal testing** is available for known familial mutations only. Please call the Laboratory Genetic Counselor before collecting a fetal sample.