Renpenning Syndrome 1: *PQBP1* Gene Deletion/Duplication

**Test Code:** YQ  
**Turnaround time:** 2 weeks  
**CPT Codes:** 81404 x1

### Condition Description

Mutations in the *PQBP1* gene (Xp11.23) cause X-linked recessive mental retardation that is often syndromic but can be nonsyndromic. *PQBP1* mutations have been associated with Renpenning syndrome, Sutherland-Hann syndrome, cerebropalatocardiac (Hamel) syndrome, and Golabi-Ito-Hall syndrome. Common features of X-linked mental retardation caused by *PQBP1* mutation seem to be mental retardation, microcephaly, and short stature. Considerable phenotypic variability is observed between families with different mutations and even between families with the same mutation.

In addition to mental retardation, microcephaly, and short stature, other reported features include small testes, ocular colobomas, cardiac malformations, cleft palate, spastic diplegia, and anal anomalies. Facial characteristics include narrow and tall craniofacies with upslanting palpebral fissures, abnormal nasal configuration, cupped ears, and short philtrum. The nose may appear long or bulbous, with overhanging columella.

Historically, Renpenning syndrome has been associated with mental retardation with short stature, moderate microcephaly, but no remarkable facies and no other neurologic abnormalities. Sutherland-Haan syndrome has been associated with mental retardation, short stature, microcephaly, brachycephaly, spastic diplegia, small testes, and possibly intrauterine growth retardation. Cerebropalatocardiac (Hamel) syndrome has been associated with severe mental retardation with congenital heart defects, microcephaly, spasticity, short stature, cleft or highly arched palate, and other craniofacial abnormalities. Golabi-Ito-Hall syndrome has been associated with mental retardation, microcephaly, postnatal growth deficiency, and other anomalies, including atrial septal defect.

**Click here** for the OMIM summary on this condition.

### Genes

*PQBP1*

### Indications

This test is indicated for:
- Confirmation of a clinical/biochemical diagnosis of Renpenning syndrome in individuals who have tested negative for sequence analysis
- Carrier testing in adult females with a family history of Renpenning syndrome who have tested negative for sequence analysis

### Methodology

DNA isolated from peripheral blood is hybridized to a CGH array to detect deletions and duplications. The targeted CGH array has overlapping probes which cover the entire genomic region. 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

Detection is limited to duplications and deletions. The CGH array will not detect point or intronic mutations. Results of molecular analysis must be interpreted in the context of the patient's clinical and/or biochemical phenotype.

### Specimen Requirements

Submit only 1 of the following specimen types

- preferred specimen type: Whole Blood

**Type:** Whole Blood
Specimen Requirements:

In EDTA (purple 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, if appropriate. Contact the laboratory if further information is needed.

Sequence analysis is required before deletion/duplication analysis by targeted CGH array. If sequencing is performed outside of EGL Genetics, please submit a copy of the sequencing report with the test requisition.

Related Tests

- Sequencing analysis of the PQBP1 gene is available (YP) and is required before deletion/duplication analysis.
- A CGH array-based test for deletion/duplication analysis of 64 different X-linked intellectual disability genes is available (OL).
- Prenatal testing is available to adult females who are confirmed carriers of mutations. Please contact the laboratory genetic counselor to discuss appropriate testing prior to collecting a prenatal specimen.