XLMR 58: TSPAN7 Gene Deletion/Duplication

Test Code: DTSPA  
Turnaround time: 2 weeks  
CPT Codes: 81228 x1

**Condition Description**

Intellectual disability (ID) is a nonprogressive cognitive impairment affecting 1-3% of the Western population. It is estimated that up to 50% of moderate-severe cases have genetic causes and approximately 10% are due to X-linked intellectual disability disorders (XLID). XLID can be syndromic or nonsyndromic and is observed in all ethnic groups. More than 100 XLID syndromes have been described in the literature to date. Fragile X is the most common XLID syndrome (~1 in 4000 males) while others can be quite rare with only a few patients reported in the literature. Males can have moderate to severe intellectual disability depending on the syndrome, and carrier females can also be affected, but typically have milder clinical symptoms.

Zemni et al. (2000) described a female individual with mild intellectual disability and minor autistic features. She had a balanced translocation involving the X chromosome [46,X,t(X;2)(p11.2;p21.3)]. The TSPAN7 gene, formerly known as the TM4SF2 gene, is located at the breakpoint on the X chromosome. TSPAN7 transcripts from this individual were barely detectable when compared with TSPAN7 transcripts from controls. The TSPAN7 gene is expressed in both the fetal and adult brain.

Zemni et al. identified two other mutations in the TSPAN7 gene in two families with XLID. Both mutations were not found in 100 control chromosomes. Males and females can have mild to moderate intellectual disability.

**References:**
- OMIM #300096: TSPAN7 gene

**Genes**

TSPAN7

**Indications**

This test is indicated for:
- Confirmation of a clinical/biochemical diagnosis of XLMR 58 in individuals who have tested negative for sequence analysis.
- Carrier testing in adults with a family history of XLMR 58 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**

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 TSPAN7 gene is available and is required before deletion/duplication analysis.
- 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.
- X-Linked Intellectual Disability panels are available for 30, 60, and 90 genes.