Exome: Sanger Confirmation and Interpretation Only

Test Code: EXSAN
Turnaround time: 11 weeks
CPT Codes: S1417 x1

Condition Description

What is the Medical EmExome?
The human exome is the complete coding (exonic) region of the genome. It is estimated to encompass approximately 1-2% of the genome, yet contains approximately 85% of disease-causing pathogenic variants. Current off-the-shelf exome kits used for clinical exome sequencing cover 92% of the exome. Traditionally, gene discovery has been done in research laboratories; however, now with the ability to sequence nearly the entire coding region of the human genome, it is possible for clinical laboratories to use this information to identify a previously unrecognized cause of disease. The Medical EmExome is the next level in clinical exome sequencing offered by EGL Genetics (EGL). The exome sequencing design provides >97% coverage of 22,000 genes, with a mean read depth of 100X. Of the ~4600 disease-associated genes analyzed, 3000 have 100% coverage (720X) of all exons; twice the number of genes with complete coverage offered by competitors, making it the most comprehensive exome sequencing test available. This is also the highest coverage offered by any clinical exome sequencing performed in a CLIA-/CAP-certified laboratory.

The Medical EmExome also features the EmExome Boost Option (Medical EmExome Trios only), which allows clinicians to choose an EGL gene panel relevant to the patient’s phenotype to ensure coverage of ALL exons, at no additional cost. EGL is the first clinical laboratory to offer near complete coverage of all disease-associated genes with an exome boost option. A research protocol is also available for extended exome and genome testing for the discovery of novel disease genes.

What gene coverage levels can be expected?
The Medical EmExome is performed on genomic DNA, using the Agilent V5 Plus designed to target the exome with greater coverage of known disease-associated genes. These targeted regions are then sequenced using the Illumina HiSeq 2500 sequencing system, with 100 basepair (bp) paired-end reads (similar to bidirectional Sanger sequencing) and an average coverage of 100X in the target region. The target region includes the exon and 10 bp of flanking intronic region. The DNA sequence is mapped to, and analyzed in, comparison with the published human genome build UCSC hg19 reference sequence. The targeted coding exons and splice junctions of the known protein-coding RefSeq genes are assessed for the depth of coverage and data quality threshold values. The Medical EmExome bioinformatics analysis pipeline is used to compare sequence changes in the individual being tested to the reference sequence. All potential positive sequence variants in the proband are confirmed by conventional di-deoxy DNA sequence analysis (Sanger sequencing) using a separate DNA isolation.

The Medical EmExome also features the EmExome Boost Option (Medical EmExome Trios only), which allows clinicians to choose an EGL gene panel relevant to the patient’s phenotype to ensure coverage of ALL exons, at no additional cost. Targeted sequencing of parental samples for the proband only option will be completed at no additional charge for exome sequencing only if needed (such as a variant of uncertain clinical significance).

The interpretation service is offered for exomes sequenced outside Emory, in either clinical or core laboratories. The exome data is mapped to, and analyzed in, comparison with the published human genome build UCSC hg19 reference sequence. The targeted coding exons and splice junctions of the known protein-coding RefSeq genes are assessed for the depth of coverage and data quality threshold values. The Medical EmExome bioinformatics analysis pipeline is used to compare sequence changes in the individual being tested to the reference sequence.

Indications

This service is offered for exomes sequenced outside Emory in either clinical or core research laboratories.

Methodology

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The Medical EmExome also features the EmExome Boost Option, which allows clinicians to choose an EGL gene panel relevant to the patient’s phenotype to ensure coverage of ALL exons, at no additional cost.

Targeted sequencing of parental samples for the proband only option will be completed at no additional charge for exome sequencing only if needed (such as a variant of uncertain clinical significance).

### Specimen Requirements

**Additional Specimen Collection/Handling Instructions Required for this Test**

Please send parental samples with the sample of the affected individual. If parental samples are not available, please notify the laboratory. No extra charge is applied for processing parental samples.

Please contact the lab before submitting any other specimen type.

**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.