Introduction

Exome sequencing offers a cost-effective approach to interrogate the protein coding portion of the human genome, which contains approximately 85% of known disease-causing mutations. The Agilent SureSelect Human All Exon V8 features a slim and efficient design, powered by machine learning-based probe selection and placement. The panel provides comprehensive and updated coverage of human protein-coding regions from major public databases, such as RefSeq, CCDS, and GENCODE. It also covers the TERT promoter and hard-to-capture exons that are omitted by other exome sequencing panels on the market. Most importantly, the panel delivers excellent enrichment performance and sequencing economy, making it ideal for routine exome sequencing.

The SureSelect Human All Exon V8 exome panel is fully customizable. For applications that require augmentation or enhancement of the exome, Agilent provides a suite of predesigned content that can be leveraged and combined with the core exome content offered by SureSelect Human All Exon V8 to quickly create a customized exome panel. Examples of predesigned content include designs such as untranslated regions (UTRs), non-coding clinical variants (NCVs), copy number variations (CNVs) backbone, pan-cancer biomarkers, and the mitochondrial genome. Custom content can also be easily created using the Agilent SureDesign web-based custom design portal, or with the help of the Agilent SureSelect panel design team.

The SureSelect Human All Exon V8 exome enrichment workflow is natively supported on the Agilent Bravo Automated Liquid Handling platform for high-throughput sample preparation and the Agilent Magnis NGS Prep system for complete, walkaway automation. The sequencing data can be readily analyzed using the Agilent Alissa Interpret software for efficient variant interpretation and reporting, providing an end-to-end exome sequencing solution.
Easily Customizable with Custom or Predesigned Content

Routine Exome Sequencing
SureSelect Human All Exon V8
- 35.1 Mb | 41.6 Mb
  - Protein coding regions from RefSeq, CCDS, and GENCODE
  - TERT promoter region

Exome V8
Core Exome
NCV Non-coding Clinical Variants
UTR Untranslated Regions
OneSeq CNV Backbone
Pan-Cancer Cancer Biomarkers
Mito Mitochondrial Genome
NCV Non-coding Clinical Variants

Clinical Research
SureSelect Human All Exon V8+NCV
- All ClinVar variants, including pathogenic (P) & likely pathogenic (LP) variants
- All HGMD variants, including disease-causing mutations (DM)
- All pathogenic (P) and likely pathogenic (LP) variants in American College of Medical Genetics (ACMG) 73 genes for secondary findings
* Both coding and non-coding variants

SureSelect Human All Exon V8+UTR
- Untranslated regions from GENCODE

SureSelect Human All Exon V8+Pan-Cancer
- 35.2 MB
- 34.1 Mb
- 35.8 Mb
- 36.5 Mb

SureSelect Human All Exon V8+NCV
- 38.0 Mb | 49.0 Mb

SureSelect Human All Exon V8+UTR
- 78.0 Mb | 85.0 Mb

SureSelect OneSeq CNV Backbones
- 1 Mb CNV backbone probes
- 300 kb CNV backbone probes

SureSelect Mitochondrial Panel
- 37 genes of the mitochondrial genome
- 15.7 kb

SureSelect Pan-Cancer
- 1.60 Mb | 2.15 Mb
- 519 genes plus MSI and TMB

Detection of Mitochondrial DNA Mutations
SureSelect Mitochondrial Panel
- Detection of Mitochondrial DNA Mutations
- 37 genes of the mitochondrial genome

Detection of CNVs and Chromosomal Aneuploidies
SureSelect OneSeq CNV Backbones
- 1 Mb CNV backbone probes
- 300 kb CNV backbone probes

Biomarker Discovery and Translational Research
SureSelect Human All Exon V8+UTR
- Untranslated regions from GENCODE

Oncology Research
SureSelect Pan-Cancer
- 1.60 Mb | 2.15 Mb
- 519 genes plus MSI and TMB

Figure 1. Customizable exome content. The SureSelect Human All Exon V8 exome panel can be rapidly customized via the wide selection of predesign content offered by Agilent to meet specific needs in exome sequencing. User-defined custom content can also be easily created using the SureDesign web-based portal, or with the help of the SureSelect panel design team.

Up-to-Date and Clinically Relevant Content

Table 1. Clinically relevant exome content for any application. The SureSelect Human All Exon V8 exome panels target up-to-date content from relevant databases, such as coding sequences from CCDS, RefSeq, and GENCODE, pathogenic variants in ACMG, ClinVar, and HGMD, untranslated regions (UTR) defined by GENCODE, and genes recommended by the National Comprehensive Cancer Network (NCCN). The table summarizes design coverage compared to other commercially available exome panels. The ACMG variants are pathogenic (P) and likely pathogenic (LP) variants in the 73 genes that ACMG recommends reporting secondary findings for in clinical exome sequencing. SureSelect Human All Exon V8+Pan-Cancer provides boosted coverage of 143 NCCN genes from the 13 most prevalent cancers according to the National Cancer Institute.

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Coding Content
- CCDS Release 22: 100%db
- RefSeq Release 95: 100%db
- GENCODE V31: 100%db

Non-coding Content
- ACMG variants: 43.2%db
- ClinVar P/LP variants: 12.4%db
- HGMD DM variants: 8.7%db

UTR Content
- GENCODE V33: 0%db | 0.31% | 100% | 3.0% | 0% | 0.33% | 0.88%

NCCN Guidelines
- Enhanced coverage of NCCN genes: No | No | No | Yes, 143 NCCN genes | No | No | No
Figure 2. Comprehensive coverage of both coding and non-coding content for clinical research. The SureSelect Human All Exon V8+NCV exome panel expands coverage beyond the human protein-coding regions to include non-coding clinical variants from major databases. The 49 Mb panel provides excellent (>30X) coverage across pathogenic (P) and likely pathogenic (LP) variants in the ACMG 73 genes and the ClinVar database, as well as the disease-causing mutations (DM) in the HGMD database. Exome-enriched libraries were generated using the Agilent SureSelect XT Low Input Reagent kit with 200 ng of mechanically-sheared HapMap genomic DNA from the Coriell Institute and enriched as 1-plex captures using a 90 min hybridization. Samples were sequenced using 2x150bp reads and downsampled to 9 gigabases (Gb) for analysis.

Figure 3. Robust coverage of untranslated regions for translational research and biomarker discovery. The SureSelect Human All Exon V8+UTR exome panel expands coverage beyond the human protein-coding regions to include untranslated regions (UTRs) defined by GENOCODE V33. The 85 Mb panel provides excellent (>30X) coverage across the protein-coding regions and UTRs. Exome-enriched libraries were generated using the SureSelect XT Low Input Reagent kit with 200 ng of mechanically-sheared HapMap genomic DNA from the Coriell Institute and enriched as 1-plex captures using a 90 min hybridization. Samples were sequenced using 2x100bp and 2x150bp reads and downsampled to 12 Gb and 14.4 Gb, respectively, for analysis.

Figure 4. Enhanced coverage of cancer biomarkers for oncology research. The SureSelect Human All Exon V8+Pan-Cancer exome panel augmented with a custom SureSelect pan-cancer panel provides improved coverage of important cancer biomarkers. This enhanced exome panel allows confident calling of low-frequency somatic variants in tumor sequencing. The pan-cancer panel targets somatic variants including single nucleotide variants (SNVs), insertions and deletions (indels), CNVs, and translocations in 519 genes, as well as genomic loci for determining tumor mutational burden (TMB) and microsatellite instability (MSI) status. The blend ratio of the V8 exome panel and the pan-cancer panel can be customized to provide the desired target coverage. Exome-enriched libraries were generated using the SureSelect XT Low Input Reagent kit with 10 ng of HD799 moderately formalin-compromised DNA from Horizon Discovery and enriched as 1-plex captures using a 90 min hybridization. The capture probes consisted of just the SureSelect Human All Exon V8 exome panel (V8 only), or with the Pan-Cancer panel as a spike-in at different ratios (1:2, 1:4, and 1:8). Samples were sequenced using 2x150bp reads and downsampled to 15 Gb for analysis.
References


Ordering Information

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Product Description

- SureSelect Human All Exon V8+Pan-Cancer: Contact Sales
- SureSelect Human All Exon V8+Mitochondrial Genome: Contact Sales
- SureSelect Human All Exon V8+OneSeq CNV Backbone: Contact Sales
- Alissa Interpret: Contact Sales

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For Research Use Only. Not for use in diagnostic procedures.

Alissa Interpret is a USA Class I Exempt Medical Device, Europe CE IVD, Canada and Australia Class I IVD Device.

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