

High Performance Exome Built on Advanced and Proven Technology

Key Advantages

- Advanced technology that you can rely on
- Superior performance for confident results and efficient sequencing
- Comprehensive content for more complete coverage
- Native automation support for increased productivity
- Seamless integration with cloud-based informatics solutions for faster time-to-answers

Introduction

The SureSelect Human All Exon V8 provides up-to-date content and complete coverage* of protein coding regions from RefSeq, CCDS, and GENCODE. It also covers the *TERT* promoter and hard-to-capture exons that are omitted by other exomes on the market. Powered by machine learning-based probe design and a new production process, the SureSelect Human All Exon V8 spans a 35.1 Mb target region of the human genome with an efficient end-to-end design size of only 41.6 Mb. The panel delivers excellent enrichment performance, as well as efficient and cost-effective exome sequencing. In addition, the panel is manufactured in large scale to provide consistent results for many years.

The SureSelect Human All Exon V8 is compatible with the legacy SureSelectXT target enrichment system, as well as the streamlined SureSelectXT HS2, SureSelectXT HS, SureSelectXT Low Input, and SureSelectQXT library preparation and target enrichment systems, which feature a fast, 90-minute hybridization protocol and a single-day workflow. The V8 exome workflow is natively supported by the Bravo Automated Liquid Handling platform for high-throughput sample preparation, and the Magnis NGS Prep system for complete, walkaway automation. The sequencing data can be readily analyzed using Alissa Interpret for efficient variant interpretation and reporting, providing an end-to-end exome sequencing solution.

Built on More Than 10 Years of Exome Technology



Figure 1. Superior by design. The new Human All Exon V8 is built on more than ten years of SureSelect technology and is now powered by machine learning for improved coverage and efficiency.

Excellent Target Coverage and Uniform Distribution for Confident Results

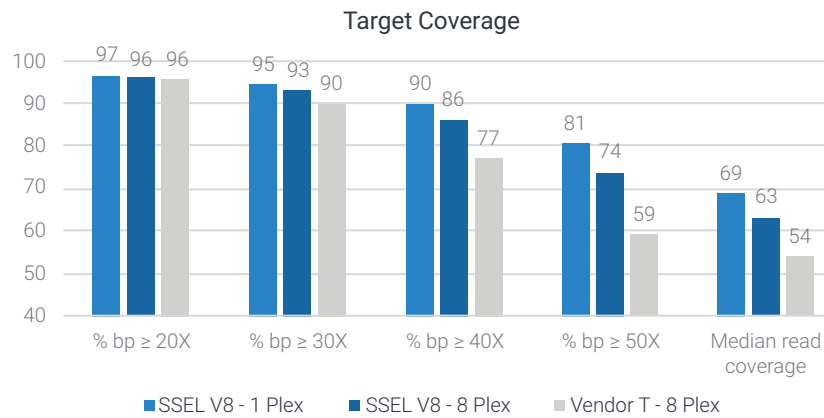


Figure 2. The SureSelect Human All Exon V8 delivers broader target coverage across relevant depths and deeper median coverage for more confident variant detection. Exome-enriched libraries were generated following vendor protocols** and sequenced on an Illumina HiSeq 4000 instrument. All samples were downsampled to 5 Gb for analysis.

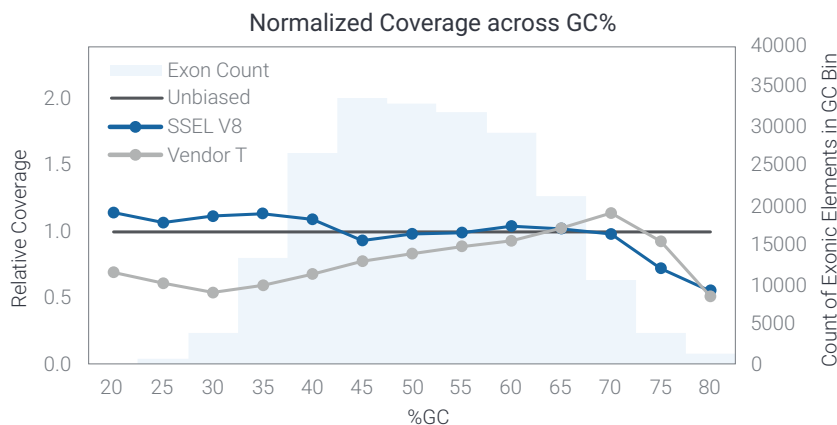


Figure 3. The SureSelect Human All Exon V8 provides more uniform distribution of coverage across the GC spectrum. Variants in GC-rich and GC-poor regions are better represented and less likely to be missed due to low coverage. Exome-enriched libraries were generated from 8-plex captures following vendor protocols** and sequenced on an Illumina HiSeq 4000 instrument. All samples were downsampled to 5 Gb for analysis.

Efficient Sequencing with High-Performance Enrichment

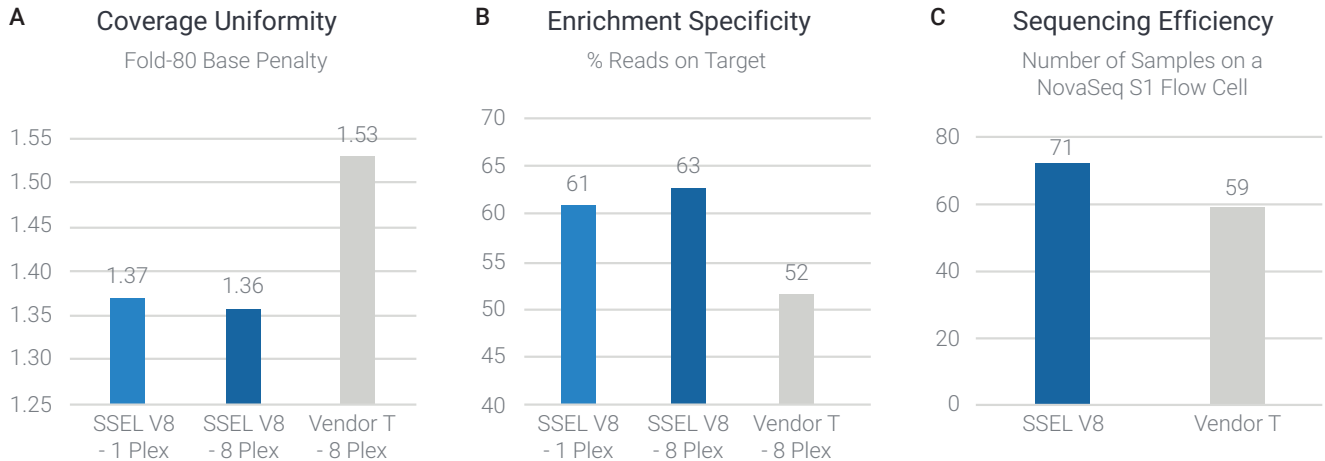


Figure 4. SureSelect Human All Exon V8 provides superior enrichment performance, resulting in lower sequencing costs. A) The SureSelect Human All Exon V8 delivers excellent coverage uniformity as measured by a lower fold-80 base penalty (lower is better). B) It also provides higher on-target rate, indicative of a highly specific enrichment of exonic sequences. C) SureSelect Human All Exon V8's high-performance exome enrichment, combined with an efficient design, reduces sequencing costs, allowing more samples to be sequenced together on a sequencing run. Exome-enriched libraries were generated from 8-plex captures following vendor protocols**, and the number of reads required to achieve a 30X coverage of 90% was calculated to determine the number of samples that can be sequenced together on a NovaSeq S1 flow cell.

Superior Coverage of the Exome

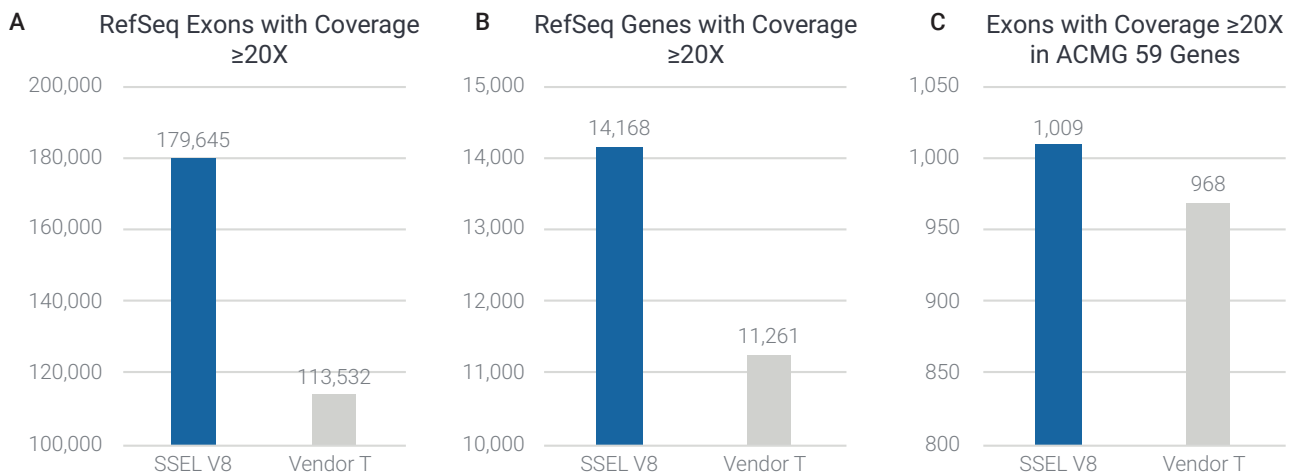


Figure 5. The SureSelect Human All Exon V8 panel achieves better coverage of the exome. The SureSelect HumanAll Exon V8 panel provides 58% more RefSeq exons (A) and 26% more RefSeq genes (B) covered at a minimum depth of 20X. (C) In addition, within the clinically actionable ACMG 59 genes, the SureSelect Human All Exon V8 panel also provides more exons with a minimum depth of 20X. Exome-enriched libraries were generated from 8-plex captures following vendor protocols** and sequenced on an Illumina HiSeq 4000 instrument using 2x100bp reads. All samples were downsampled to 5 Gb for analysis.

More Complete Coverage of Genes Associated with Rare Diseases

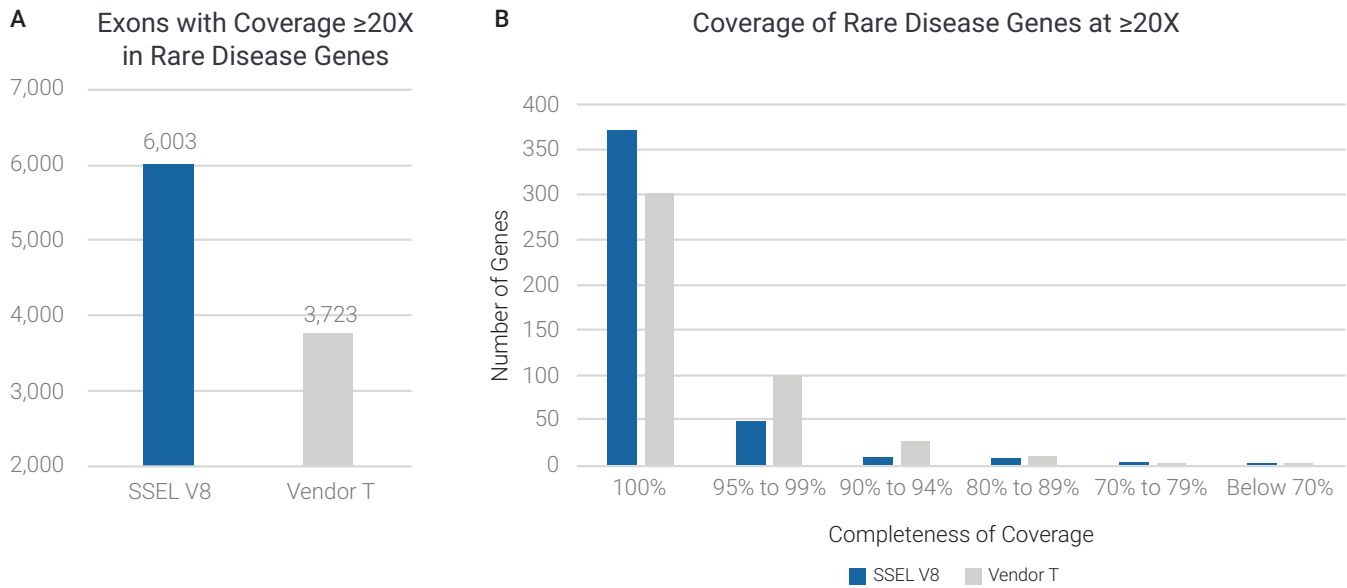


Figure 6. The SureSelect Human All Exon V8 panel provides more complete coverage of genes associated with rare diseases to ensure better detection of disease-causing variants. The coverage of a panel of 438 genes responsible for 448 childhood recessive diseases¹ is analyzed at both the exon and gene level. A) The SureSelect Human All Exon V8 panel delivers superior coverage at the exon level across the 448 genes, providing 60% more exons that have a minimum coverage of 20X. B) The SureSelect Human All Exon V8 panel also provides 24% more genes that are completely covered, such that 100% of bases have a coverage greater than 20X. Exome-enriched libraries were generated from 8-plex captures following vendor protocols** and sequenced on an Illumina HiSeq 4000 instrument using 2x100bp reads. All samples were downsampled to 5 Gb for analysis.

Consistent Performance Across Manual and Automated Workflows

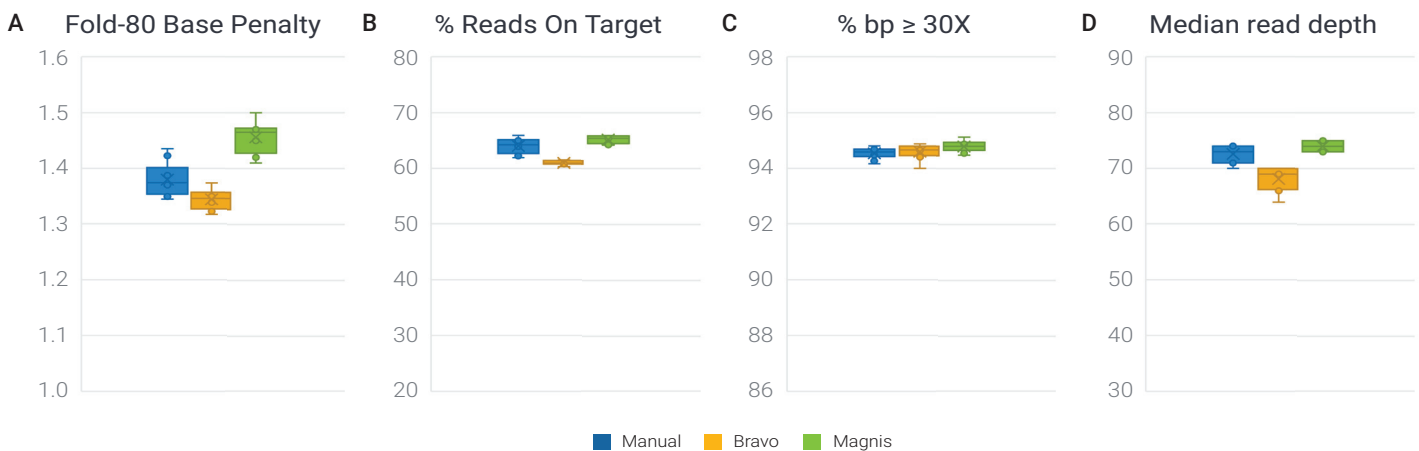


Figure 7. SureSelect Human All Exon V8 provides consistent enrichment performance using both manual and automated library preparation. Both the Bravo automated liquid handling platform and the Magnis NGS Prep system generated libraries with reproducible and comparable performance as those prepared using a manual workflow by experienced operators. For all workflows, 200 ng of HapMap genomic DNA from the Coriell Institute (Camden, New Jersey) was Covaris-sheared and used as input material. For manual (4 operators, total of 10 samples) and Bravo (8 samples) workflows, libraries were constructed using the SureSelectXT Low Input Reagent kit and enriched as 1-plex captures using a 90 min hybridization. For the Magnis (4 runs, total of 10 samples) workflow, libraries were generated using the Agilent Magnis SureSelectXT HS Human All Exon V8. Rev B Kit, which is supplied as pre-aliquoted reagents in plates. Samples were sequenced on an Illumina HiSeq 2500 instrument (manual and Bravo) or HiSeq 4000 instrument (Magnis) using 2x100bp reads and downsampled to 50 million reads or 5 Gb of data for analysis.

Increased Productivity with Automated Library Preparation and Variant Assessment

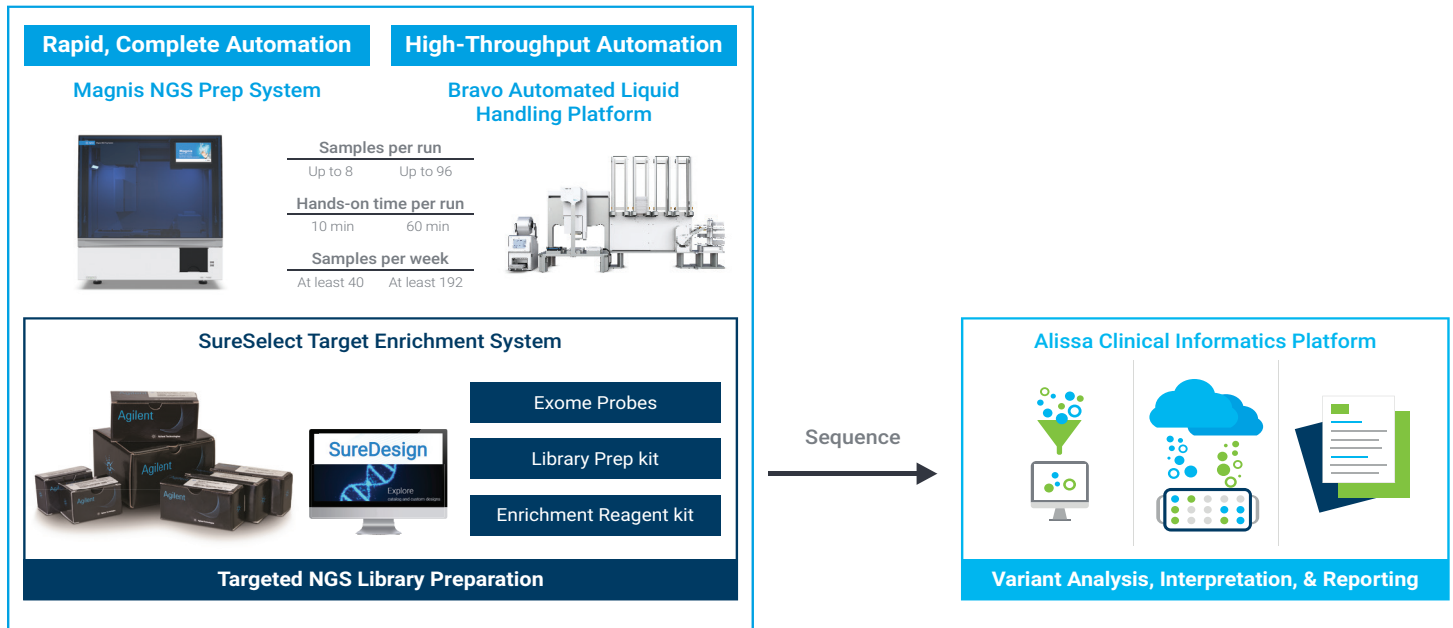


Figure 8. Agilent whole-exome sequencing solutions. The SureSelect Human All Exon V8 panel can be easily supplemented with additional targets required for exome sequencing using the Agilent SureDesign web-based application. The V8 exome panel can be paired with Bravo and Magnis automation and the Alissa Clinical Informatics platform to achieve greater operational efficiency. The exome is readily automated on the Bravo automated liquid handling platform for high-throughput applications, requiring only 60 min of hands-on time to process up to 96 samples. The Agilent Magnis SureSelect Human All Exon V8 kit is pre-allocated for use on the Magnis NGS Prep system. Only 10 minutes of hands-on time is needed to set up the instrument run and load reagents and consumables onto the instrument's deck. From there, the Magnis system delivers 8 exome-enriched libraries in less than 9 hours without further operator intervention. The Alissa Clinical Informatics platform allows clinical workflow to be automated from variant filtration and annotation, through curation and classification, enabling clinical genetics and molecular pathology labs to efficiently triage, curate, and report genomic variants.

* SureSelect Human All Exon V8 covers 100% of the protein coding sequences from CCDS release 22, GENCODE release 31, and RefSeq release 95, accessed in August 2019.

** For both vendors, the respective protocol was followed without modification using NA12878 genomic DNA from the Coriell Institute (Camden, New Jersey). The starting DNA amount was 200 ng for samples prepared with the SureSelect Human All Exon V8 and 50 ng for Vendor T, as specified in Vendor T's protocol. For samples prepared with the SureSelect Human All Exon V8, libraries were constructed from Covaris-sheared DNA using the SureSelectXT Low Input Reagent kit and enriched as 1-plex or 8-plex captures, using 90-min hybridization. For Vendor T, DNA was enzymatically fragmented and libraries were generated and captured in 8-plex, using overnight hybridization. All samples were sequenced on an Illumina HiSeq 4000 instrument using 2x100bp reads and downsampled to 50 million reads or 5 Gb of data. Analysis was carried out using vendor provided target BED files. Fold-80 base penalty was reported from analysis with Picard HsMetrics (Broad Institute). The percentage of on-target reads was calculated as the total number of reads with at least 50% overlap with any base of the target, divided by the number of uniquely mapped reads.

References

1. Kingsmore, S. Comprehensive Carrier Screening and Molecular Diagnostic Testing for Recessive Childhood Diseases. PLoS Curr. 2012, 4, e4f9877ab8ffa9.

Ordering Information

| Product Description | 16 Rxns | 96 Rxns | 96 Rxns Auto |
|--|---------------|-----------|--------------|
| SureSelectXT HS Human All Exon V8 | 5191-6873 | 5191-6874 | 5191-6875 |
| Sure SelectXT Human All Exon V8 | 5191-6879 | 5191-6891 | 5191-6892 |
| Product Description | 2 Hybs | 12 Hybs | 12 Hybs Auto |
| SureSelectXT HS PreCap Human All Exon V8 | 5191-6876 | 5191-6877 | 5191-6878 |
| Product Description | 32 Rxns | 96 Rxns | |
| Magnis SSEL XT HS Human All Exon V8. Rev B | G9772C | G9772D | |
| Product Description | | | |
| Alissa Interpret | Contact Sales | | |

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