# **PacBi**

PacBi●

# **Revio System** Reveal more with accurate long-read sequencing at scale



Scale

1,300 human HiFi genomes per year



### Ease of use

Simplified consumables and flexible run setup

#### Compute power

Google DeepConsensus and more on board



### Affordability

\$1,000 per human HiFi genome

### This is your moment for a more complete view of biology



#### Comprehensive variant calling with phasing + 5mC

Access high accuracy for all variant types — SNVs, indels, structural variants, tandem repeat expansions, and methylation — including in challenging regions.<sup>1</sup>



#### Truly complete assembly of complex genomes

Accurately construct the full sequence of chromosomes, including telomeres and centromeres.<sup>2</sup>



## Targeted sequencing to study genes at scale

Focus the power of HiFi variant calling by enriching for regions of interest using hybrid capture or PCR amplification.



## Single-cell transcriptome sequencing

Catalog RNA isoforms at single-cell resolution, moving beyond gene counting to catalog full-length transcripts.<sup>3</sup>

### See it all with HiFi sequencing on the Revio<sup>™</sup> system

#### Long reads

HiFi reads are tens of kilobases long, providing the ability to resolve large variants — like structural variants and tandem repeat expansions — and map to difficult, repetitive regions of the genome.

#### Direct methylation detection in native molecules

HiFi sequencing identifies base pair-level DNA methylation in all runs, without bisulfite treatment.

#### **Exceptional accuracy**

HiFi sequencing provides reliable answers through exceptional accuracy, with 90% of bases  $\geq$ Q30 and median read accuracy  $\geq$ Q30.

#### Uniform coverage across sequence contexts

With an optimized polymerase and no amplification, HiFi sequencing provides uniform coverage even for high-[GC] regions and tandem repeats.



HiFi reads for the sample HG002 reveal accurate sequence of both maternal and paternal alleles, providing phasing that spans over 84 kb of the cystic fibrosis gene, *CFTR*.

### Sequence with HiFi reads at high throughput

	Sequel <sup>®</sup> lle system	Revio system	Increase
Higher density	8 million ZMWs	25 million ZMWs	З×
Independent stages	1	4	4×
Shorter run times	30 hours	24 hours	1.25×
30× HiFi human genomes/year	88	1,300	15× overall

### Enjoy a fully automated sequencing workflow



### The easy-to-use Revio system fits your schedule



The Revio system offers configurable sequencing run times to tailor to your experiments.



The new flowcell design of the Revio SMRT<sup>®</sup> Cell eliminates the need for a nitrogen supply, simplifying laboratory requirements.



The ability to load consumables while sequencing is in progress makes it easy to maximize instrument utilization without off-hours work.





### Set up runs instantly and use less plastic with smart consumables

- The Revio system has only three workdeck consumables — a sequencing plate, a SMRT Cell tray, and pipette tips.
- Sequencing plates are linked automatically to run designs through an NFC tag, enabling rapid run setup on instrument.
- The Revio sequencing plate combines what was previously four parts, eliminating extraneous plastics and making runs easier to manage. The single standard 96-well plate includes sample libraries, reagents, and space for mixing and waste.





### Get the most accurate reads – fast

- The Revio system features cutting-edge NVIDIA GPUs with >20× more compute power than the Sequel<sup>®</sup> Ile system.
- The GPUs provide rapid turnaround time for basecalling and HiFi read generation, keeping pace with the sequencing throughput of the Revio system.
- With Google DeepConsensus<sup>4</sup> on board, Revio is the most accurate PacBio long-read system to date. DeepConsensus uses advanced deep learning techniques to further extend the accuracy and yield of HiFi sequencing.



### Reduce your compute and storage costs

- · The Revio system is capable of much more than just basecalling. Every run also measures DNA methylation status, calculated with a deep learning algorithm that processes polymerase kinetics.
- All fundamental processing steps are performed on instrument, including barcode demultiplexing and conversion to the standard BAM format.
- A more efficient data representation reduces file size by 50% per base  $-20 \times$  smaller than for other long-read technologies. Base quality scores are grouped into seven bins, and similar reads are sorted together for more effective compression.



	On instrument	Off instrument
Revio system	<ul> <li>Base calling</li> <li>HiFi read generation with DeepConsensus</li> <li>Methylation calling</li> <li>Barcode demultiplexing</li> <li>BAM file generation</li> </ul>	<ul><li>Variant calling</li><li>Genome assembly</li></ul>
Other long-read technology	<ul><li>Signal collection</li><li>Initial base calling</li></ul>	<ul> <li>Additional base calling</li> <li>Methylation calling</li> <li>Barcode demultiplexing</li> <li>BAM file generation</li> <li>Variant calling</li> <li>Genome assembly</li> </ul>

### Add to what you love about HiFi sequencing with the Revio system

#### Keep what's great about HiFi sequencing

- · Long, accurate reads of native DNA molecules
- Easy sample and library prep with the Nanobind® CBB kit and SMRTbell® prep kit 3.0
- On-instrument generation of HiFi reads + methylation calls, with demultiplexing
- Standard BAM file format compatible with downstream analysis tools

#### ...and get even more on the Revio system

- High-throughput up to 1,300 HiFi genomes per year
- Onboard Google DeepConsensus for the most accurate HiFi reads yet
- A simplified user experience, including adaptive loading, fewer consumables, rapid run setup, and real-time performance run previews
- 50% smaller output files to reduce storage costs



The Revio system produces the same – or better – read length, quality, and variant calling performance that have made HiFi sequencing so celebrated. Data shown is for a single Revio SMRT Cell for HG002/GM24385.<sup>5</sup>

### System specifications

The Revio system utilizes nanofabricated Revio SMRT<sup>®</sup> Cells, each of which has 25 million zero-mode waveguide wells. Revio has four independent stages, allowing sequencing of multiple SMRT Cells in parallel. The onboard compute provides accurate basecalling with Google DeepConsensus, plus methylation calling in every run.

Library	Run time*	Q30+ bases	HiFi yield per SMRT Cell $^{\dagger}$	Methylation
1–5 kb	12 hours	95%	6 million reads	
5–10 kb	24 hours	90%	30 Gb	5mC at CpG sites for native DNA
10-15 kb			60 Gb	
15-20 kb			90 Gb	
20-25 kb	30 hours	85%	90 Gb	

\* Run time refers to the data collection step, which determines the time between processing SMRT Cells.

+HiFi yield is dependent on library quality and sequencing preparation procedures. Specified yield is based on high-quality samples prepared following best practices.

### Key applications and sample throughput

The Revio system supports a variety of applications that benefit from accurate long HiFi reads. Its four independent stages allow different samples and applications to be sequenced in parallel. A subset of key applications is below.

Library	Sample	Per Revio SMRT Cell	Per year*	Expected coverage <sup>‡</sup>
1-5 kb	Amplicon	>1,000	>2.6 million	50×
5–10 kb	Microbial genome	96	124,800	50×
15-20 kb	Human genome	1	1,300	30×
15–20 kb	Transcriptome with Kinnex™ full-length RNA kit	4	5,200	10 million reads

\* Annual throughput is based on 2,600 Revio SMRT Cells for 12 hour runs; 1,300 for 24 hour runs; and 1,050 for 30 hour runs. Adaptive loading – which increases the consistency of yield per SMRT Cell – adds about 4 hours to run time, affecting the maximum number of SMRT Cells per year.

†Expected coverages are estimates.



#### $\square$

We have offices in countries around the world. Visit **pacb.com/contact** for contact info.

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### What will you discover with Revio?



Learn more about the Revio system: **pacb.com/revio** 

#### **Ordering information**

Part number	Product
102-090-600	Revio system
102-301-900	Nanobind CBB kit
102-182-700	SMRTbell prep kit 3.0
102-817-600	Revio polymerase kit
102-202-200	Revio SMRT Cell tray
102-587-400	Revio sequencing plate
Instrument operating	g environment
Temperature	19–25°C (66–77°F)
Humidity	20–80% relative humidity, non-condensing
Ventilation	16,400 BTU/hr (4,800 W)
W × D × H	36.5 × 35.5 × 68.7 in (92.7 × 90.2 × 174.5 cm)
Weight	449 kg (989 lb)
Power	200–240 VAC at 50–60 Hz, 30A
Network	1 GbE or 10 GbE, copper

#### **KEY REFERENCES**

1. Lincoln, S. E., et al. (2021). One in seven pathogenic variants can be challenging to detect by NGS: an analysis of 450,000 patients with implications for clinical sensitivity and genetic test implementation. *Genetics in Medicine*, 23(9), 1673–1680.

- 2. Nurk, S., et al. (2022). The complete sequence of a human genome. Science, 376(6588), 44-53.
- 3. Al'Khafaji, A. M., et al. (2021). High-throughput RNA isoform sequencing using programmable cDNA concatenation. bioRxiv, doi:10.1101/2021.10.01.462818.
- 4. Baid, G., et al. (2022). DeepConsensus improves the accuracy of sequences with a gap-aware sequence transformer. Nature Biotechnology, 1–7.
- 5. PacBio (2022). HG002 sequence data from Revio system.

\$1,000 per human HiFi genome assumes US list price of \$995 for sequencing reagents for one Revio SMRT Cell, which has an expected yield of 90 Gb, equivalent to a 30x human genome.

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