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# Investor & Analyst Day November 20, 2024



### **Disclaimer and Other Information**

#### **Forward Looking Statements**

This presentation includes "forward-looking statements" within the meaning of the "safe harbor" provisions of the United States Private Securities Litigation Reform Act of 1995. The actual results of the Company may differ from its expectations, estimates, and projections and, consequently, you should not rely on these forwardlooking statements as predictions of future events. Words such as "expect," "estimate," "project," "budget," "forecast," "anticipate," "intend," "plan," "may," "will," "could," "should," "believes," "predicts," "potential," "continue," and similar expressions (or the negative versions of such words or expressions) are intended to identify such forward-looking statements. These forward-looking statements include, without limitation, the Company's expectations with respect to future performance and development and commercialization of products and services, its anticipated cash runway and its financial guidance for the full year 2024. These forward-looking statements involve significant risks and uncertainties that could cause the actual results to differ materially from those discussed in the forwardlooking statements. Most of these factors are outside the Company's control and are difficult to predict. Factors that may cause such differences include, but are not limited to: the inability to maintain the listing of the Company's Class A common stock on The Nasdag Stock Market; the ability of the Company to grow and manage growth profitably and retain its key employees; the Company's ongoing leadership transitions; changes in applicable laws or regulations; the ability of the Company to raise financing in the future; the success, cost and timing of the Company's product development and commercialization activities; the commercialization and adoption of the Company's existing products and the success of any product the Company may offer in the future; the potential attributes and benefits of the Company's commercialized Platinum® protein sequencing instrument and kits and the Company's other products once commercialized; the Company's ability to obtain and maintain regulatory approval for its products, and any related restrictions and limitations of any approved product; the Company's ability to identify, inlicense or acquire additional technology; the Company's ability to maintain its existing lease, license, manufacture and supply agreements; the Company's ability to compete with other companies currently marketing or engaged in the development or commercialization of products and services that serve customers engaged in proteomic analysis, many of which have greater financial and marketing resources than the Company; the size and growth potential of the markets for the Company's products and services, and its ability to serve those markets once commercialized, either alone or in partnership with others; the Company's estimates regarding future expenses, future revenue, capital requirements and needs for additional financing; the Company's financial performance; and other risks and uncertainties described under "Risk Factors" in the Company's most recent Annual Report on Form 10-K and Quarterly Reports on Form 10-Q and in the Company's other filings with the SEC. The Company cautions that the foregoing list of factors is not exclusive. The Company cautions readers not to place undue reliance upon any forward-looking statements, which speak only as of the date made. The Company does not undertake or accept any obligation or undertaking to release publicly any updates or revisions to any forward-looking statements to reflect any change in its expectations or any change in events, conditions, or circumstances on which any such statement is based.

### **Investor Day Agenda**

Management	Q&A Session	11:30 AM–Noon
Jeff Hawkins, CEO	The Proteomics Lab of the Future	11:20–11:30 AM
John Vieceli, CPO	Platform Roadmap	11:00–11:20 AM
Brian Reed, PhD	Innovating Discovery Applications in Proteomics	10:40–11:00 AM
Todd Rearick, CTO	Technology Architecture for the Future	10:20–10:40 AM
Jeff Hawkins, CEO	Proteomics Market: Current & Future Perspective	10:00–10:20 AM

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### Proteomics Market: Current & Future Perspective



### **Proteins are the Core of Biological Discoveries**

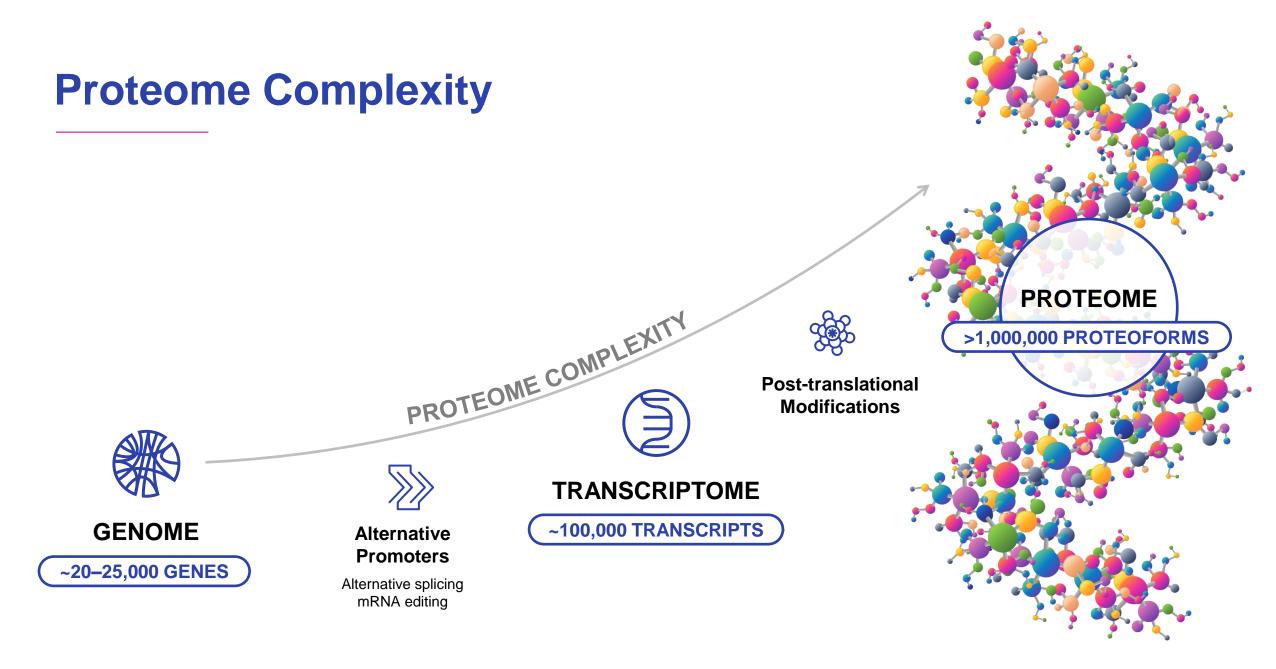


Disease Biomarkers Therapeutic Development

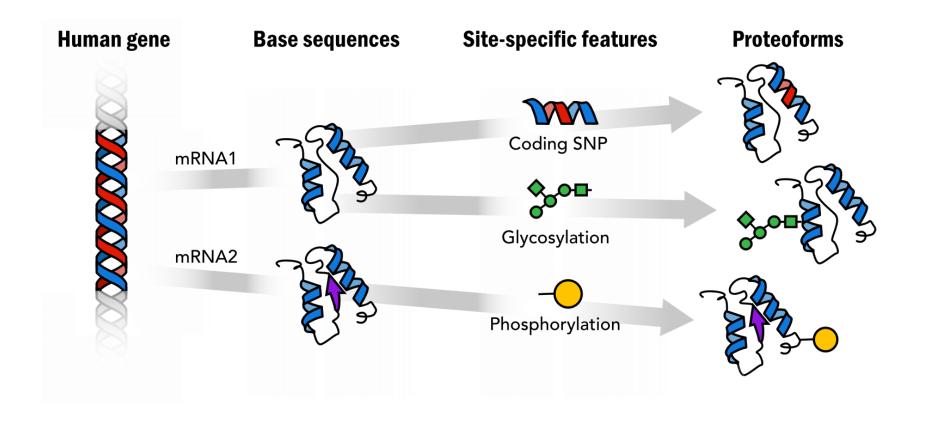
Biotech Innovation

Proteins are the vital engines of biological systems, playing crucial roles in both health and disease

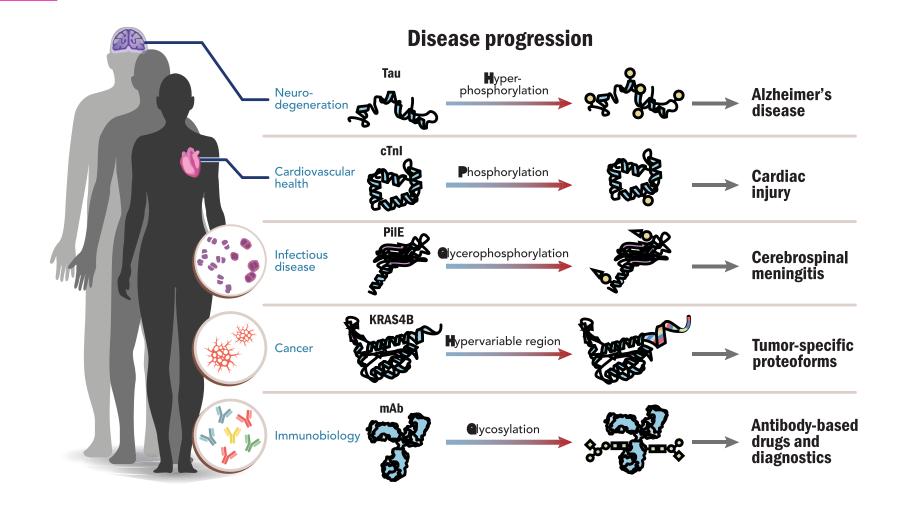




# **Transcriptomics Does Not Accurately Predict Protein Profiles**



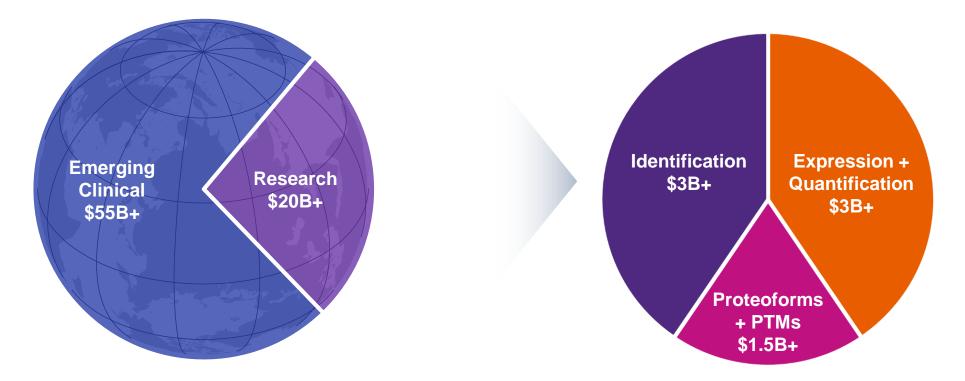
### **Disease Progression Goes Beyond the Protein Level**



# **Proteomics is a Large and Growing Market Opportunity**

**\$8B+ Initial Target Market**<sup>2</sup>

#### **\$75B+ Proteomics Market**<sup>1</sup>



#### 

SVB Leerink Research, "Proteomics: The Next Frontier in Life Science Tools and Diagnostics." September 28, 2021
 DeciBio Consulting Evaluation, June 2020

# **Platinum Use Cases Today**

#### **Identify Proteins Critical to Biology**



In-gel digest of bio samples



Characterize antibodies



Identify co-IP proteins

#### Uncover + Understand **Proteoforms**

Post-translational modifications

Amino acid variants



**RNA** isoforms

#### **Screen and Characterize Proteins with Barcodes**



Protein/antibody engineering



mRNA vaccine development

Lipid nanoparticle delivery

# **How QSI Customers Are Leveraging Platinum**



MRNA screening with protein barcodes for gene therapy



Studying citrullination PTMs



Characterizing far-flung extremophiles



Studying mRNA translation and PTMs



Studying disease isoforms



Mapping protein conformations using protein barcodes

# The Proteomics Market is Poised for Significant Growth

#### Large-scale screening studies designed to identify clinically relevant biomarkers are

#### Deep pr analysis

increasing

#### Deep proteoform-level

analysis will be needed to fully define and characterize the biomarkers with highest medical value



**Population-scale studies** will be needed to characterize what a "healthy" profile looks like





#### **Routine use of multiomics**

requires creating new data analysis tools — these tools will require large amounts of training data



#### Al-driven drug development

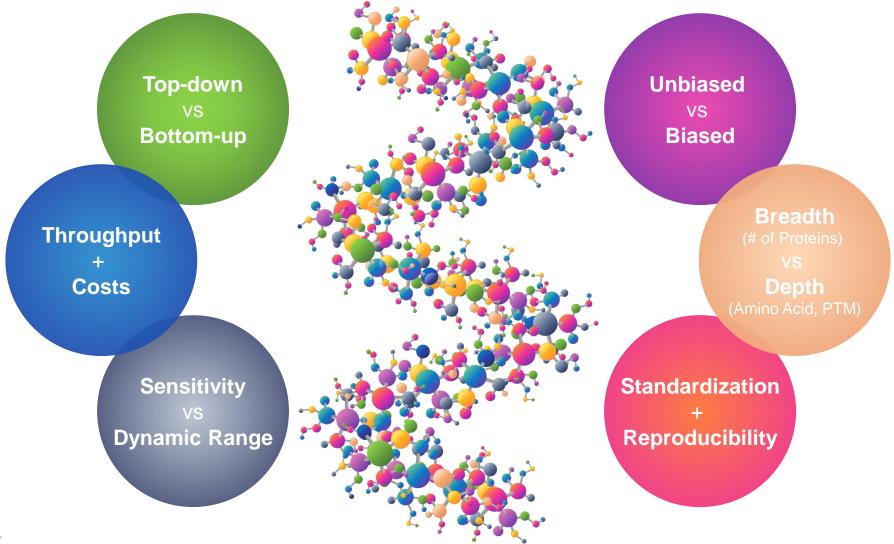
will drive the need for deeper proteomic data (amino acid level) to better inform and train the models



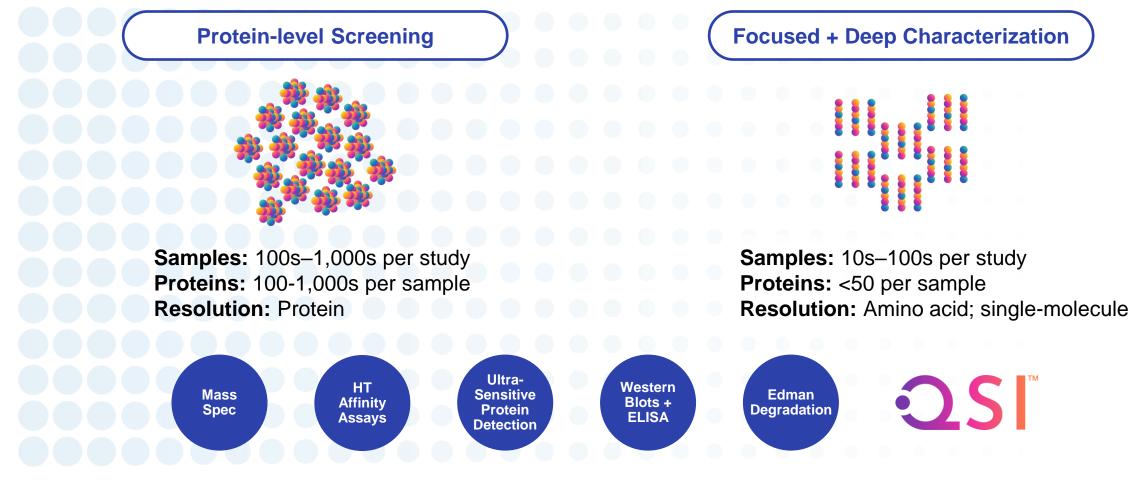
#### The proteome is dynamic

— longitudinal data will be needed (i.e., repeat testing)

### **Technical Challenges in Proteomics Today**



# Multiple Specialized Platforms Required to Fully Interrogate the Proteome



# QSI is Best Positioned to Usher in a New Paradigm in Proteomics



**Proteus**<sup>™</sup>

**Core technology** is the only commercially available tech that can enable single-molecule, top-down, and bottom-up proteomics methods



**New architecture** (Proteus<sup>™</sup>), combined with other ongoing technology development initiatives, creates clear path to *de novo* sequencing



**Ultrarapid sequencing chemistry** can enable significant increase in sample throughput per day and unlock time-sensitive applications (e.g., clinical diagnostics) in the future Platinum<sup>®</sup> Pro



# Distribution Agreement in Place to Scale Adoption Across the US + Canada



2	Sequencing Chip Rit Mode Kit	
Platinum' Pro	Sequencing Kit	Barcoding Kit
Quantum in	Library Prep Kit	Library Prep Kit



### **(** ) avantor<sup>™</sup>

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### **QUANTUM SI**

Technology Development Pipeline

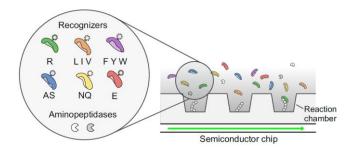
### **Quantum-Si Core Technologies**

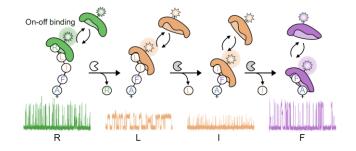
**Platinum<sup>®</sup> System 2M Chip** 





**Chemistry Biomolecules** 





**Algorithms Applications** 



ထိုး။ **Barcode Applications** 



**Variant Calling** 

**PTM Characterization** 



**Protein Identification** 

### **End-to-End Protein Analysis**

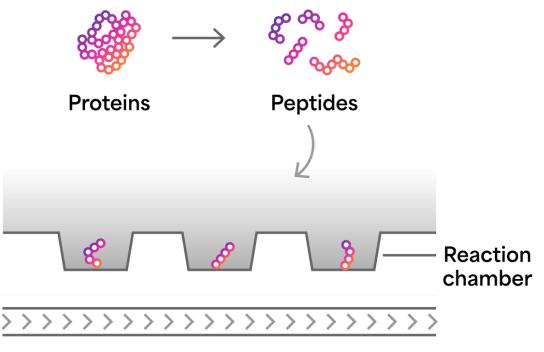


### **Prepare Proteins for Sequencing**

Proteins are digested into short fragments (peptides)



Peptides are immobilized at the bottom of reaction chambers on our chip



Semiconductor chip

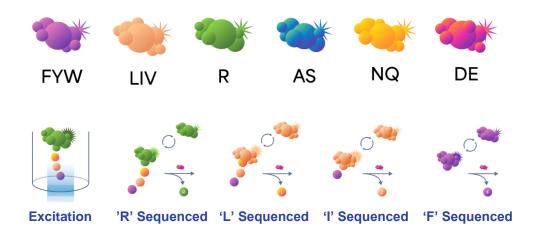
### **Kinetic Signatures Uniquely Identify Proteins + Proteoforms**

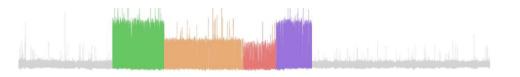


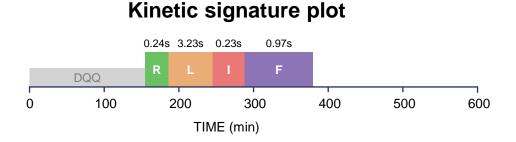
Recognizers bind amino acids in sequence



Recognition events produce kinetic signatures







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# **Rationale for New Technology Architecture**



Semiconductors require large R&D investment



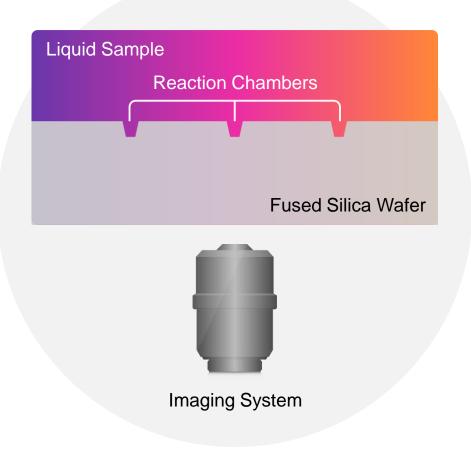
**Re-partitioning of system** allows for less expensive consumable



Leverage optical magnification to pack wells closer together and scale to billions of reads



Leverage high-performance, commerciallyavailable imaging components



### **QSI Core Technologies**



# **Chip + Surface Chemistry**

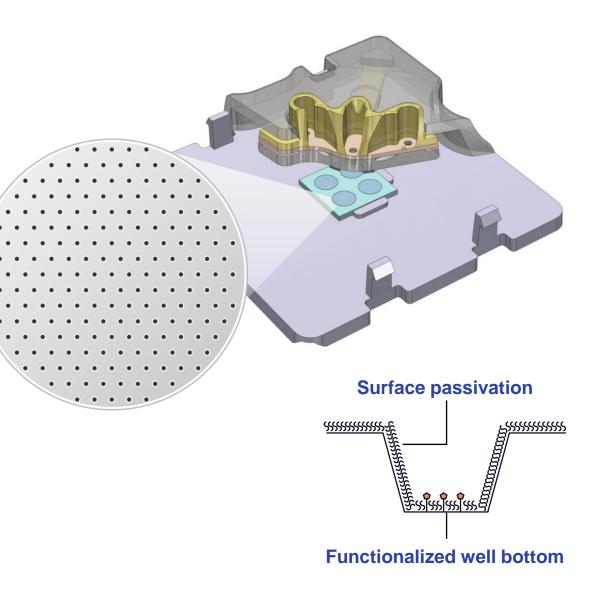
Simple passive device with approximately 20M wells (per flow cell) at initial launch



**Heavily de-risked** — leverages existing design, materials, and fabrication methods



**Compatible with** existing surface chemistry



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# **Proteus<sup>™</sup> Consumable Development**



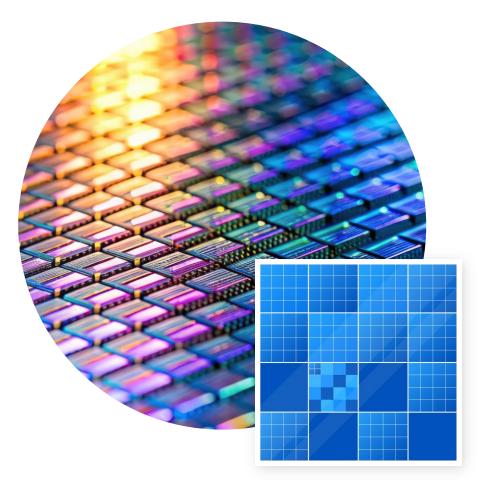
Wafer process flow developed in production foundry



**Prototype wafers** fabricated and tested



**Simple process** has low-risk path to high-volume production



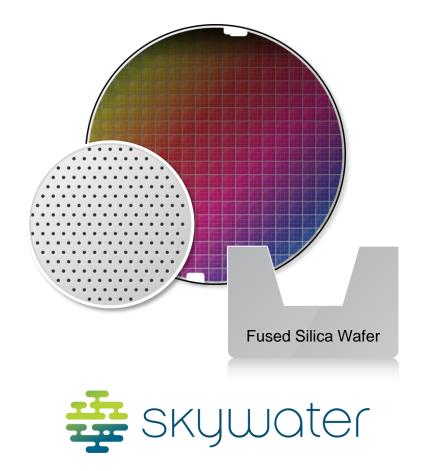
### **Wafer Fabrication Process Development**



### Foundry process modules work and produce the desired well structure



**Foundry partner** for development and production is in place



# **Proteus<sup>™</sup> Instrument Development**



Move imaging components to the system



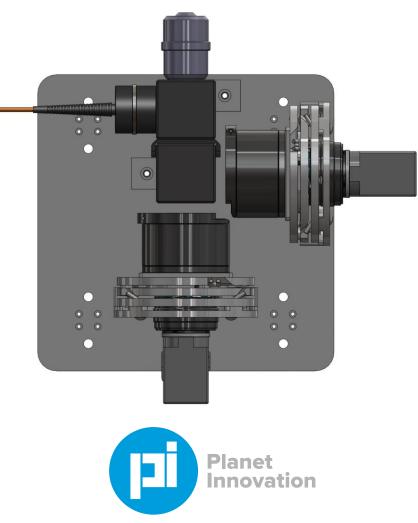
Increase workflow automation

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Leverage commercially available technologies for imaging and liquid handling



Takes advantage of significant investment in optics driven by NGS industry



# Library Prep + Sequencing Chemistry



Existing library prep and sequencing chemistry are completely portable



New system discriminates dyes with color rather than lifetime

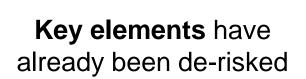


**Some new dye development** is necessary, but is underway and low risk



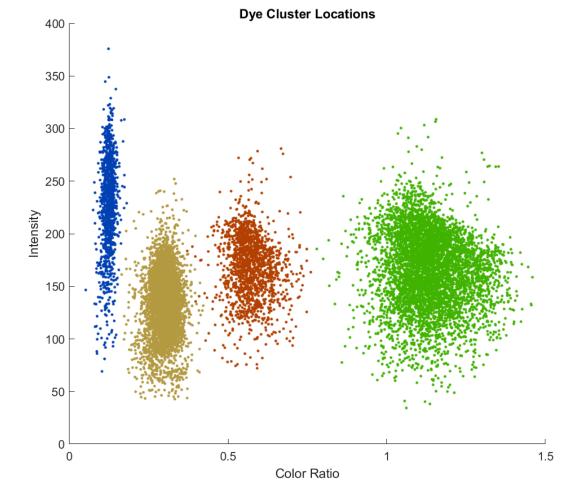
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## **Color Ratio is a Viable Alternative to Current Lifetime Detection**





Move to color means we can leverage off-the-shelf camera technology



# **Analysis Software**



# **Backend processing** is completely portable to new system

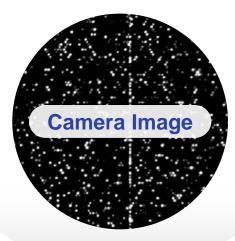
• Pulse detection, ROI calling, alignment, protein inference, and other applications



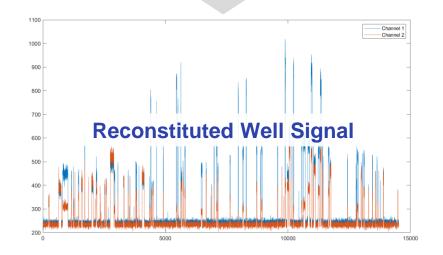
**Development required** for frontend image processing



Well within state-of-the-art capability



#### **Registration Deconvolution**



### **Instrument Roadmap**

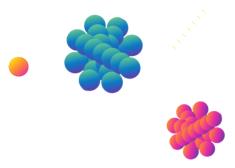


New architecture scales up to 10B reads per consumable



**Enables shotgun proteomics** Proteus<sup>™</sup> 2.0 of complex samples QUANTUM SI (**10B**) 0 Platinum<sup>®</sup> Pro Puts us on path to Proteus<sup>™</sup> 1.0 de novo sequencing (**50M+**) Platinum (2M)

### **QUANTUM SI**



### Innovation Toward the Most Advanced Set of Discovery Applications in Proteomics

**Brian Reed** 





### **Innovation at the Forefront of Proteomics**

- The path to complete proteome coverage
- **Ultrasensitive PTM detection for proteoforms**
- Deep, unbiased interrogation of high-complexity samples
- 4

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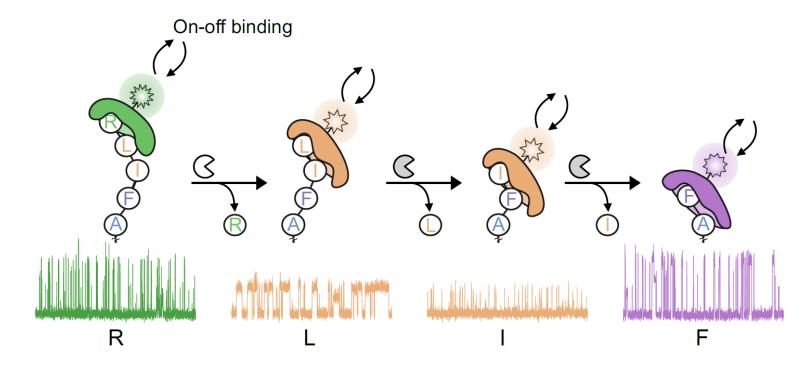
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Beyond sequencing: the first platform for top-down single-molecule proteomics



# Acceleration on the Path to Complete Proteome Coverage

### **Sequence Proteins on Platinum**

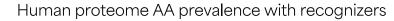


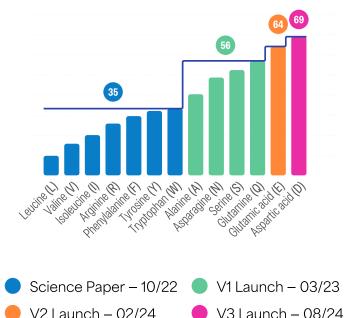
- Each recognizer binds 1-3 cognate N-terminal amino acids (NAAs)
- Rapid on-off binding generates a pulsing pattern detected by the chip
- Extremely information-rich data output: 10s-100s of pulsing events per amino acid

# A Rapid Path to Complete Amino Acid Coverage

Our team has mastered the engineering and evolution of amino acid recognizers

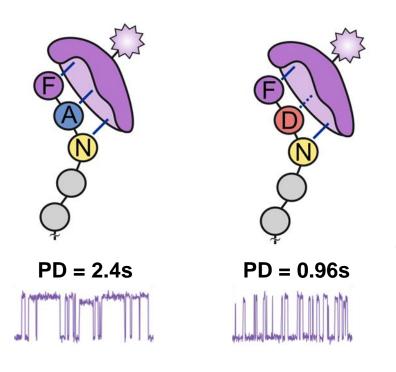
#### **Recognition Progress**





- As a result of our rich kinetic output, we have more data on our recognizers than possibly any other set of proteins in biotechnology
- Recognizers in the V3 kit recognize 13 of the 20 types of amino acids (69%)
- New recognizers have already been developed and are on track for release in our next kit update
- We are on track to enable complete reference-free sequencing: enables key applications like sequencing antibodies and cancer neoantigens de novo

### **Kinetic Signatures are Sensitive to Downstream Sequence**



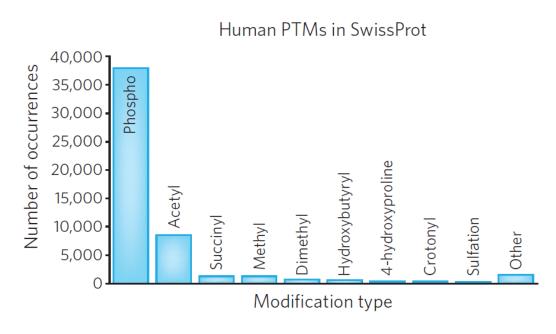
- Recognizers physically contact residues
  downstream of bound NAA
- Influence is encoded in the peptide's kinetic signature and is highly predictable
- Kinetic signatures are a unique and powerful feature of Quantum-Si's core technology
- The acquisition of single-molecule kinetic information gives us unprecedented insight into binding interactions

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# Ultrasensitive PTM Detection for Proteins and Proteoforms

### Phosphorylation is the Most Abundant PTM in the Human Proteome

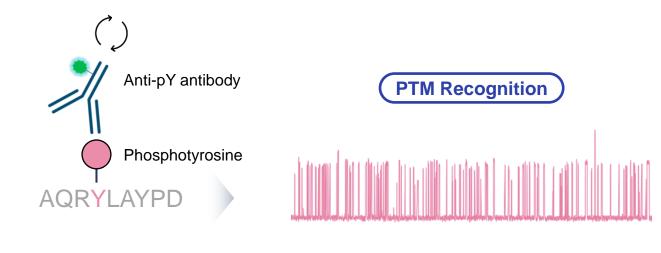


Nat Chem Biol 14, 206–214 (2018). https://doi.org/10.1038/nchembio.2576

- Post-translational modifications (PTMs) are central to protein function and implicated in human diseases
- There are more than 400 different types of PTMs; phosphorylation is the dominant type (~72% of all PTM sites)
- Phosphorylation has the largest disease association: 81% of all discovered PTMassociated diseases<sup>1</sup>

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### Affinity Reagents as Ultrasensitive PTM Recognizers

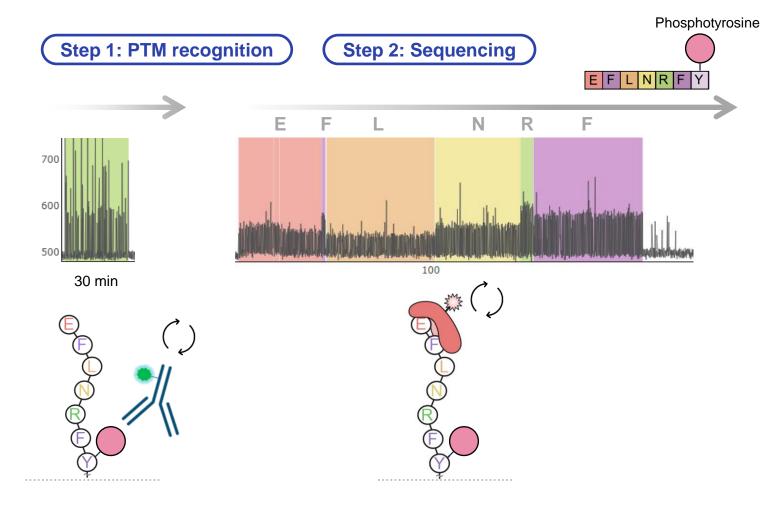


- Anti-PTM antibodies and other affinity reagents work on chip for ultrasensitive PTM detection
- Deliver the same real-time kinetic information as NAA recognizers
- Recognize PTMs anywhere in the peptide (not just at the N-terminus)

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AQRYLAYPE

# Affinity Reagents as Ultrasensitive PTM Recognizers

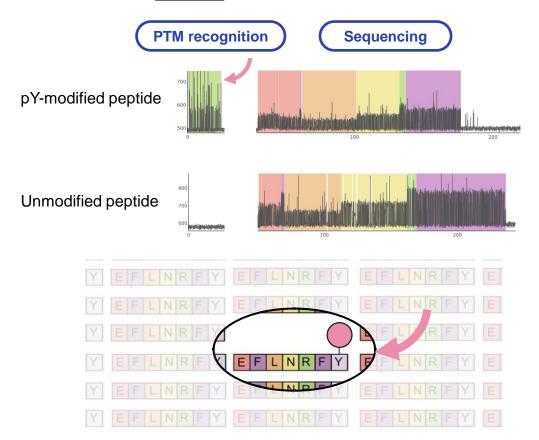


- **Step1:** PTM detection for 30 minutes with PTM recognizer
- Step 2: Normal protein sequencing with NAA recognizers
- PTM recogizers can be multiplexed and combined with kinetic signatures to pinpoint PTMs in multisite configurations

### **Ultrasensitive Phosphotyrosine Detection with CDNF**

#### CDNF\_HUMAN

..QEAGGRPGADCEVCK EFLNRFYK SLIDRGVNFSLDTIEK ELISFCLDTK..



Detection of less than 1 PTM-modified peptide in 1,000

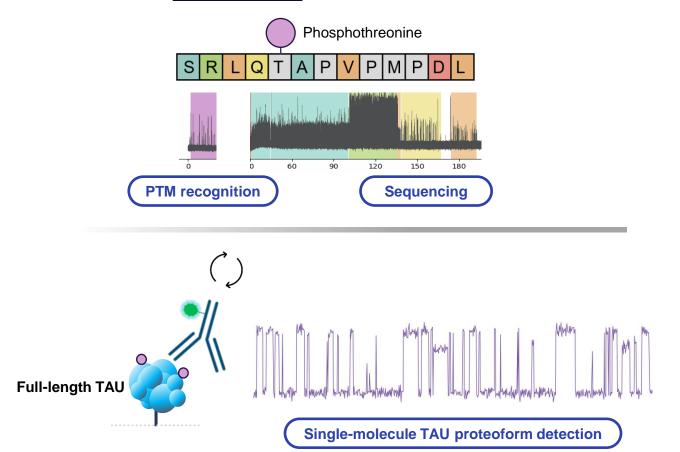
- Extreme sensitivity to PTM stoichiometry due to the clear pulsing pattern from PTM recognition
- Example: a CDNF peptide is detected at a ratio of less than 1 phosphorylated peptide in 1,000
- Method can be extended to other types of PTMs, e.g., ubiquitination, glycosylation; works with commercially available affinity reagents

#### QUANTUM SI

### **Recognition of Human TAU Proteoforms**

#### TAU\_HUMAN

..VAVVRTPPKSPSSAK **SRLQTAPVPMPDLK** NVK SK IGSTENLK HQPGGGK VQIINK K..



- Affinity reagents can be used in a bottom-up or top-down configuration
- **Example:** bottom-up recognition of pT\* on human TAU peptides, top-down detection of immobilized full-length TAU proteoforms
- Real-time approach enables proteoform detecting reagents to be run simultaneously
- First commercially available platform for detection and differentiation of full-length proteoforms

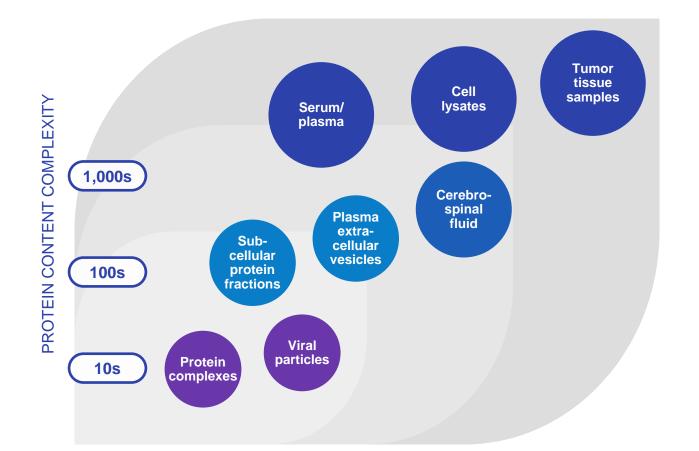
QUANTUM SI

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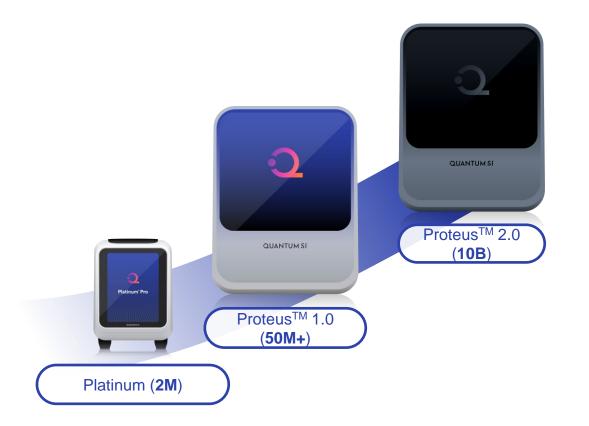
# Unbiased Interrogation of High-Complexity Samples with Quantum-Si's Core Technology

### Sequencing Complex Biological Samples Unlocks Broad Access to Proteomics



- **Biological samples like serum** contain hundreds to thousands of proteins with wide dynamic range of abundance
- Unbiased, consistent, accessible interrogation of these samples is a challenge in proteomics
- Sequencing is not limited to predefined content: enables discovery of changes in proteins and proteoforms that other methods are unable to access

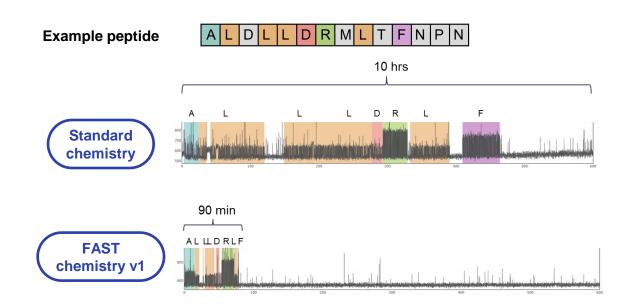
### **Unbiased Interrogation of High-Complexity Samples**



- New chip architectures and advances in sequencing chemistry will enable sequencing biological samples at ever-increasing depth
- Future versions of the platform will see shotgun sequencing with thousands of proteins identified
- **Barcoding approaches** and flowcell designs will enable sample multiplexing
- Innovative methods to fractionate proteins and to reduce sample complexity will be combined with these improvements

#### QUANTUM SI

### Fast Sequencing for Deep Coverage and Rapid Sampleto-Answer



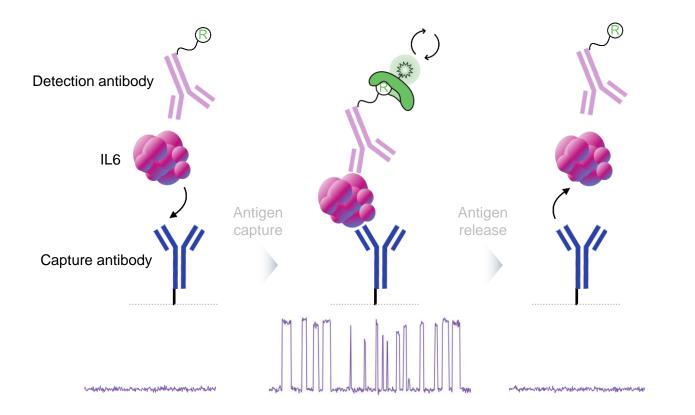
- We have developed new sequencing chemistry with a much faster rate of sequencing
- With FAST chemistry, we achieve equal performance to 10-hour runs in just 90 minutes (version 1)
- Path to runs <30 minutes for some applications with further development
- Enables deep sample coverage via iterative FAST sequencing and rapid sample-to-answer methods for clinical applications



Beyond Sequencing: the First Commercially Available Platform for Top-Down Single-Molecule Proteomics

### **Detecting Antibody Binding Events with the Power of Real-Time Kinetics**

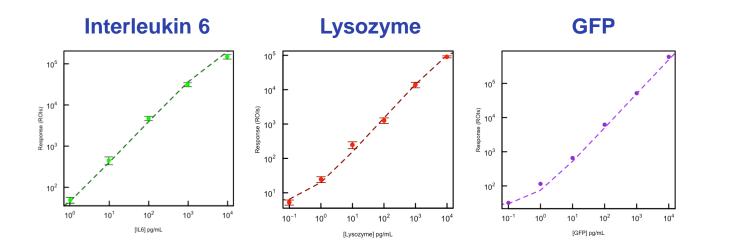
#### **Dye-cycling enables ultrasensitive real-time detection of biomarkers**



- Detection of fixed protein panels with high sensitivity is an increasingly important application in proteomics
- We developed a single-molecule sandwich assay that enables real-time detection of biomarkers
- Dye-cycling approach uses our existing kits to translate immune complex formation into a readily detected pulsing segment

### **Ultrasensitive Detection of Proteins in Serum**

#### Direct detection of proteins in serum with high sensitivity

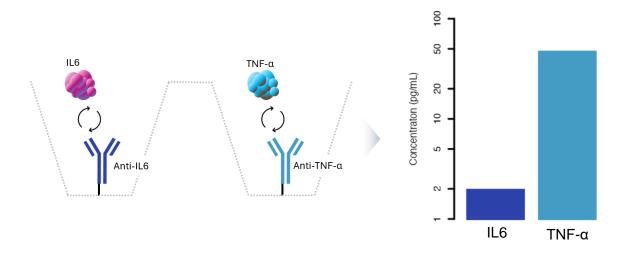


Spike-in titration experiments in serum demonstrate 0.1–1 pg/mL detection (path down to 10 fg/mL with further development)

#### **QUANTUM SI**

### **Multiplexed Ultrasensitive Protein Biomarker Detection**

#### Multiplexed detection of human IL6 and TNF- $\alpha$



- Affinity reagents against multiple biomarkers can be loaded on the chip
- Dye-cycling approach enables discrimination of biomarkers by fluorescence and kinetic properties in multiplexed assays, along with PTMs

#### **QUANTUM SI**

### **A Platform for Ultrasensitive Detection of Protein Panels**

#### **IL10** IL6 IL2 IL4 TNFα **IFNy IL12** Protein panel size scales with the platform <u>c</u> Proteus<sup>TM</sup> 2.0 (1,000+) Э Э Proteus<sup>™</sup> 1.0 (100's)Platinum (up to

#### Multiplexed biomarker detection directly in serum

- Sensitivity on Platinum is suitable for commercialization of panels with up to 10 proteins
- Panel size scales with the platform, as well as capacity to multiplex samples
- Proteins detected directly in serum on chip, eliminating complex sample prep
- Sample-to-answer in ~2 hours with one instrument

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### **QUANTUM SI**

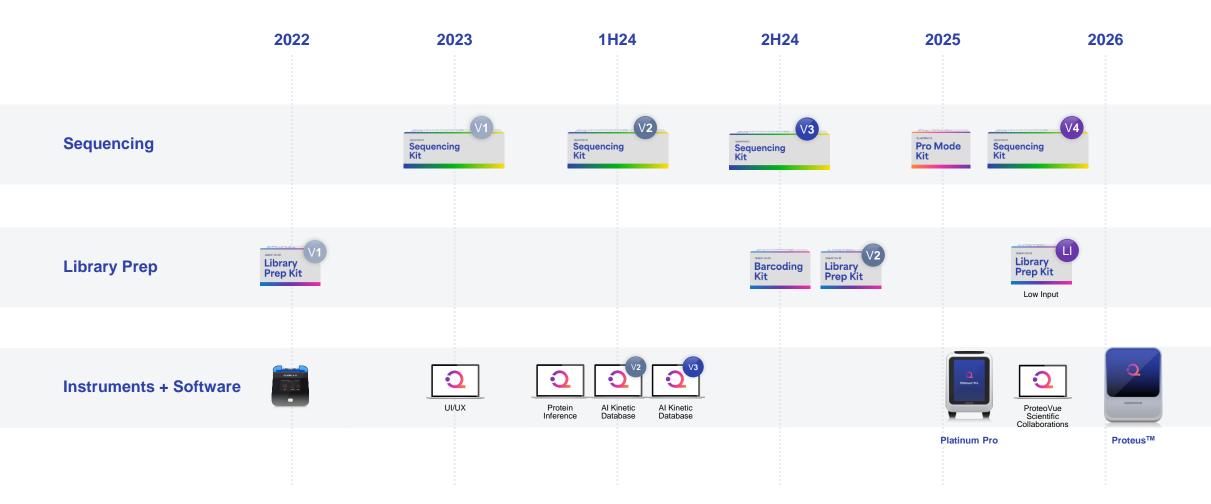
# **Platform Roadmap**

Nov 20, 2024

John Vieceli

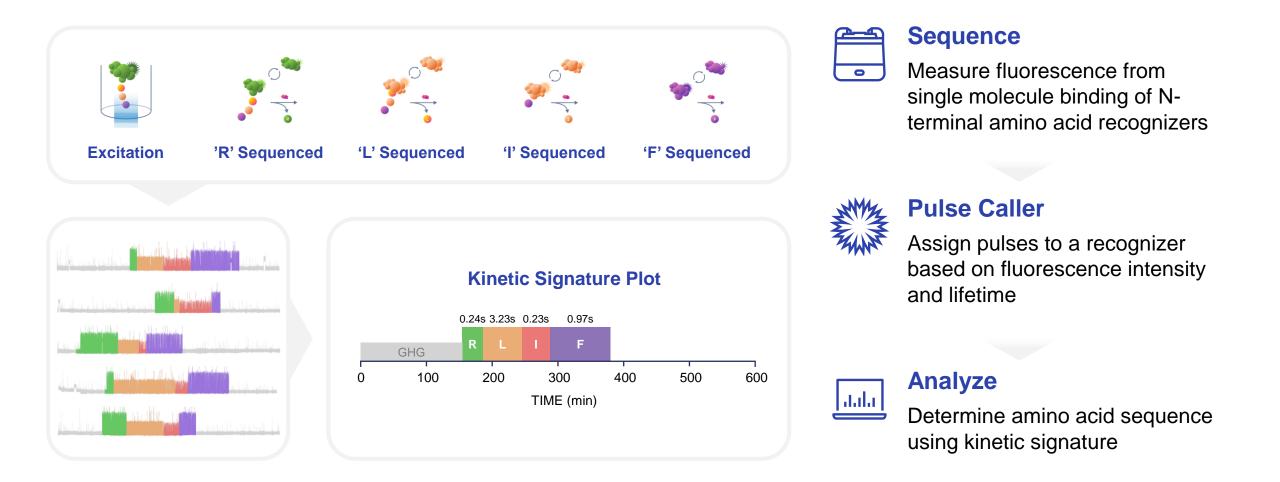


### **Innovation Pipeline is Robust and Accelerating**



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### **Sequencing Analysis Software**



### Software Workflows for Next-Gen Protein Sequencing™



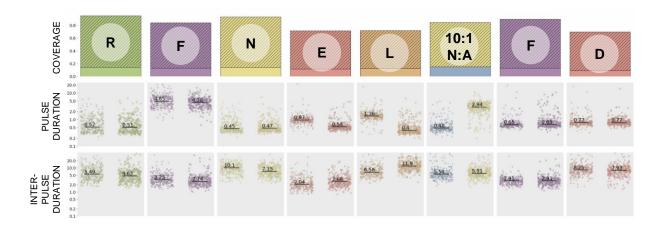
Protein Inference

Kinetic signature enables inference of sample protein from whole human proteome panel

Rank	Inferred Protein	Score	Likelihood	Mass (kDa)	Length
1 IL4	spIP04112IIL4_HUMAN	11.035496	99.99%	17	153
2	spIP06127ICD5_HUMAN	0.593929	44.78%	55	495
3	splQ15208ISTK38_HUMAN	0.582068	44.12%	54	465
4	splQ96LQ0IPPR36_HUMAN	0.506878	39.76%	49	422
5	splQ81WR1ITRI59_HUMAN	0.440162	35.6%	47	403
6	splQ9UMR3ITBX20_HUMAN	0.428596	34.85%	49	447
7	splQ96EU6IRRP36_HUMAN	0.403635	33.21%	30	259
8	splQ9H2F9ICCD68_HUMAN	0.371386	31.02%	39	335
9	splQ8IVI9INOSTN_HUMAN	0.365859	30.63%	58	506
10	splO9BZ81IMAGB5_HUMAN	0.310895	26.72%	32	275

#### **ProteoVue™ Variant Calling**

Kinetic signature enables differentiation of protein variations at the single amino acid level

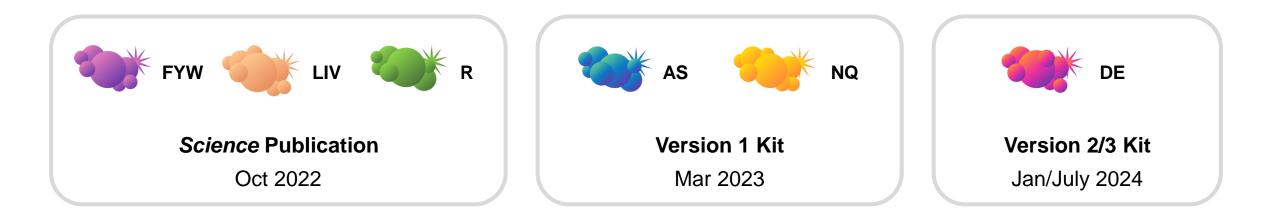


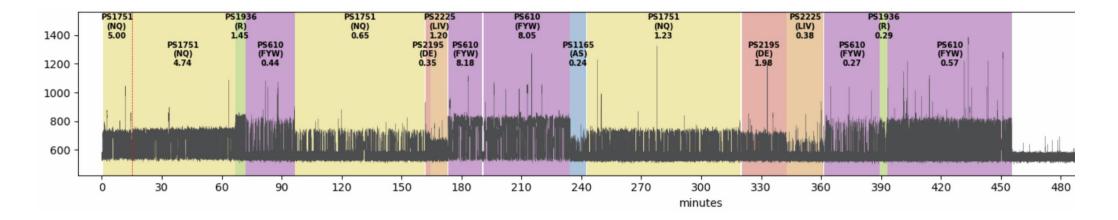
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### **QUANTUM SI**

# **Artificial Intelligence**

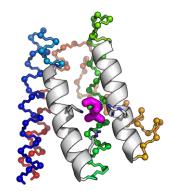
### **N-terminal Amino Acid Recognizer Development**

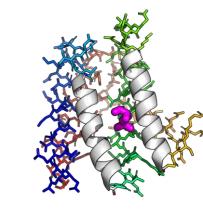


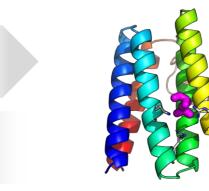


**QUANTUM SI** 

### **Recognizer Protein Design Al**







Amino acid recognizer **backbone design** 

Amino acid recognizer sequence design

Orthogonal verification of amino acid recognizer protein design



### **Pulse Width Prediction Using Artificial Intelligence**



### Platinum Sequencing

QSI is continually increasing the size of the training data with more proteins and/or new binders



Pulse Width Prediction Al

Currently predicts ~4.6 million pulse widths used in analysis



### Better Performance

Pulse width prediction Al trained with more platinum sequencing data improves protein detection performance

### **QUANTUM SI**

# Platinum<sup>®</sup> Pro

### **Platinum<sup>®</sup> Instrument**





**Customers identified opportunities** to improve workflow and UI/UX



Functionality limited to protein & peptide sequencing



Local analysis enabled by additional server

### Introducing Platinum<sup>®</sup> Pro





Streamlined workflow and reduced hands-on time



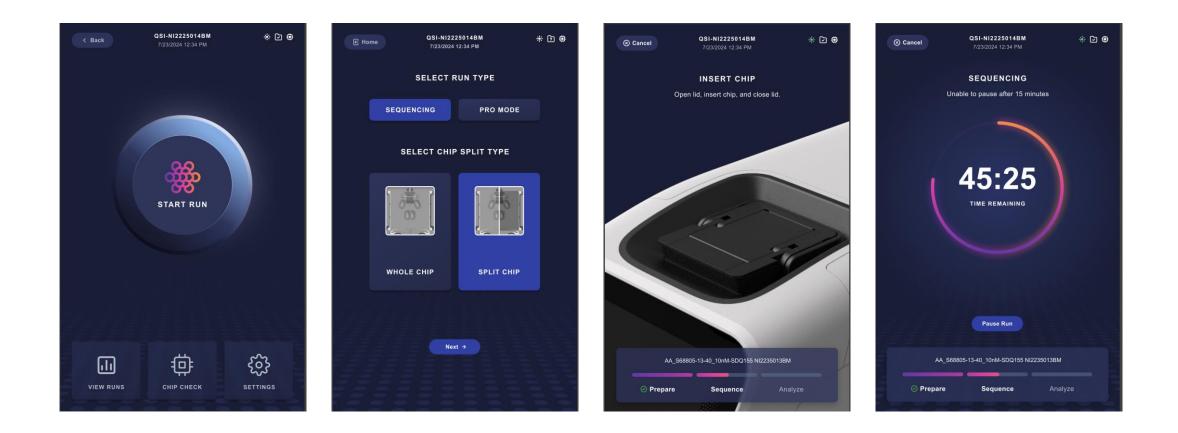
**Pro Mode** enables new applications

Onboard analysis or via the cloud

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**Enhanced user interface** 

### **Streamlined Workflow Improves Usability**



#### **QUANTUM SI**

### **Pro Mode Available only on Platinum Pro**







### Peptide Barcodes can be Used to Monitor Protein Expression Both *In Vivo* or *In Vitro*



Equimolar 5 barcode mix encoded as mRNA and packaged into LNPs



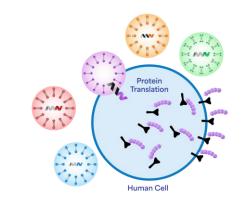
Inject into mouse model; harvest target tissues



Enrich for target protein, functionalize, cleave, and sequence barcodes

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### **V2 Library Preparation Kit Improvements**





Simplified workflow reduces need for buffer exchange



Improved performance with ~80% of proteins successfully inferred



Reduced protein input five-fold

### **QUANTUM SI**



### **Proteus<sup>™</sup> Increases Throughput + Automation**





Switch from semiconductor to optical architecture with patterned array for throughput scalability



Liquid handling automation simplifies workflow and reduces hands-on time



Up to an order of magnitude throughput increase per sample relative to Platinum at initial launch

### **Proteus<sup>™</sup> Increases Number of Samples**

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Run one or two samples simultaneously



Reagent cartridges with sequencing workflow automation



Run up to 8 samples in one sequencing run



### **QUANTUM SI**

# **The Proteomics Lab of** the Future



# QSI's Pipeline is Heavily De-risked Compared to Other Proteomics Companies



**Builds upon** QSI's existing commercially available technologies



**Industry-leading** protein and enzyme engineering program operating at scale and with high success rates



**Manufacturing infrastructure** in place and routinely producing and delivering product to customers today



Strategic Partnerships to Accelerate Commercial Adoption and Deliver on Innovation Roadmap









#### **QUANTUM SI**

### **QSI is Best Positioned to Usher in a New Paradigm in Proteomics**



**New platform architecture** designed so QSI will not be feature limited (can scale to billions of reads)



**QSI core technology** is the only commercially available tech that can enable single-molecule, top-down, and bottom-up proteomic analysis



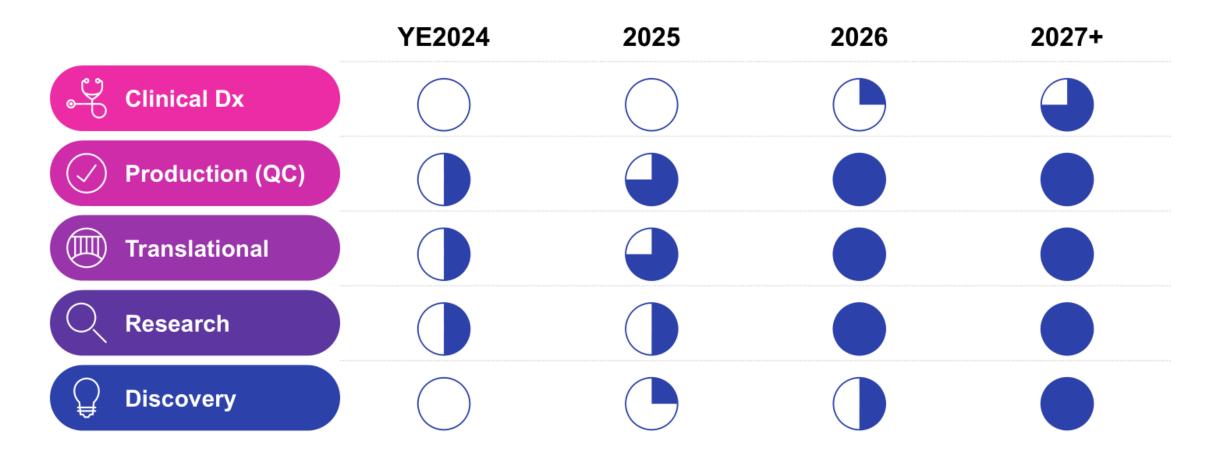
**QSI ultrarapid sequencing** can enable significant increase in sample throughput per day and unlock time-sensitive applications (e.g. clinical diagnostics) in the future



**QSI new architecture**, combined with other ongoing technology development initiatives, creates clear path to *de novo* sequencing

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### **QSI Near-term Pipeline Will Unlock Opportunities Across All Market Segments**



### **Proteomics Lab Today**



Many specialized platforms needed to fully interrogate the proteome



**Technical tradeoffs** when selecting between the breadth of protein coverage and depth of insights



High capital costs and manual workflows limit the number of laboratories capable of performing proteomics



### **QSI Will Power the Proteomics Lab of the Future**



**One platform** and core technology capable of addressing the broadest range of proteomics analysis methods



**Eliminate technical tradeoffs –** single-molecule, amino acids and PTMs, top-down or bottom-up, ultrasensitive, scalable throughput



Affordable and automated, allowing any lab — anywhere — to be a proteomics core lab



### **QUANTUM SI**<sup>®</sup>

Q&A

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