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## Investor & Analyst Day November 20, 2024



# Disclaimer and Other Information

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## 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 forward-looking 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 forward-looking 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 Nasdaq 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, in-license 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

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

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## Proteomics Market: Current & Future Perspective



# Proteins are the Core of Biological Discoveries



**Disease  
Biomarkers**



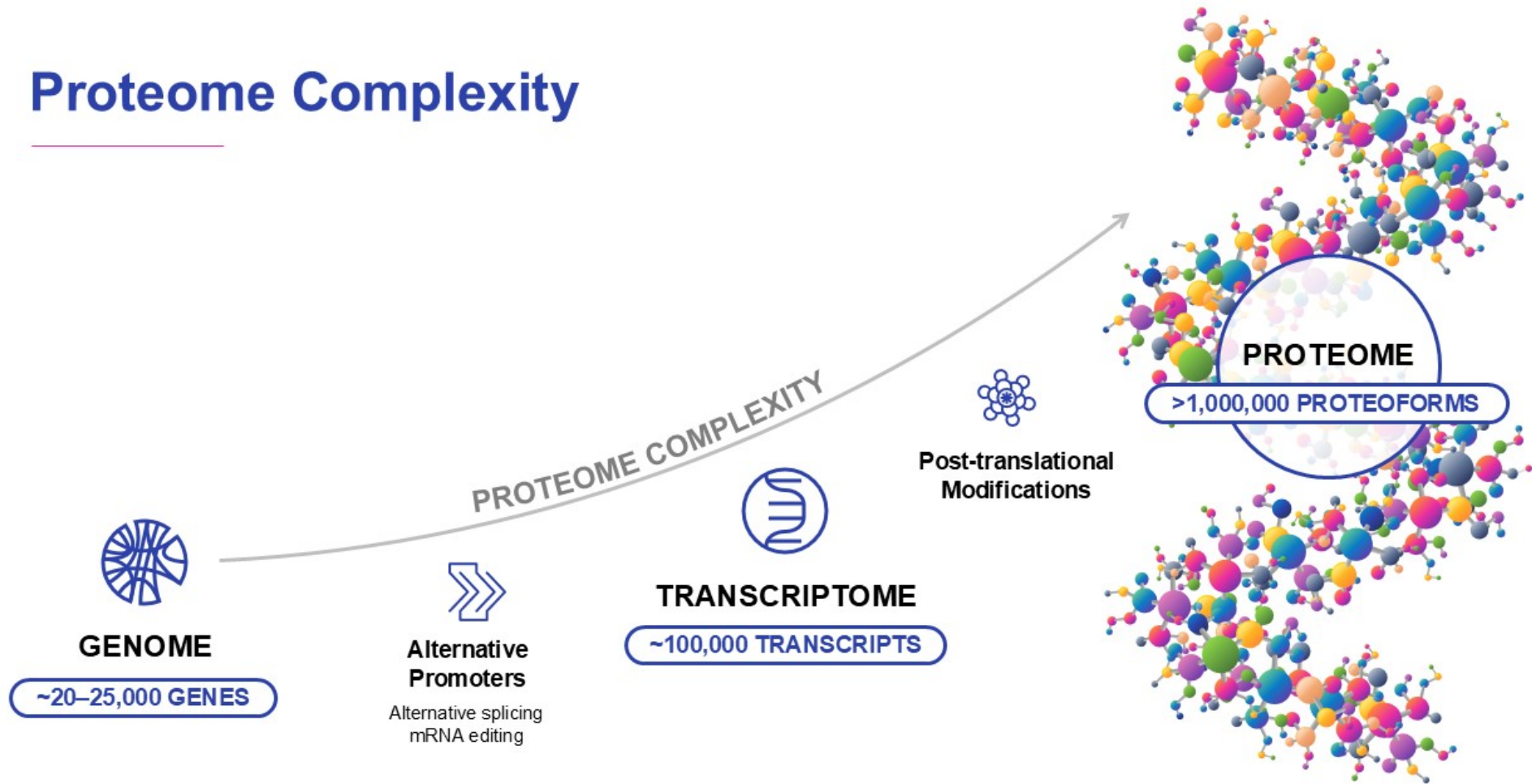
**Therapeutic  
Development**



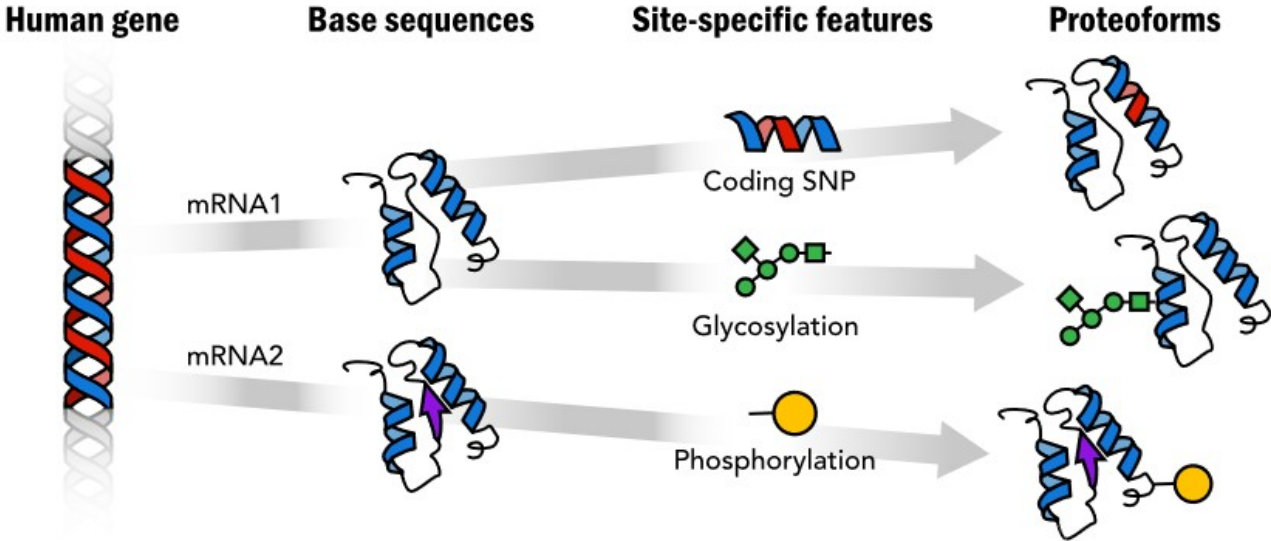
**Biotech  
Innovation**

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

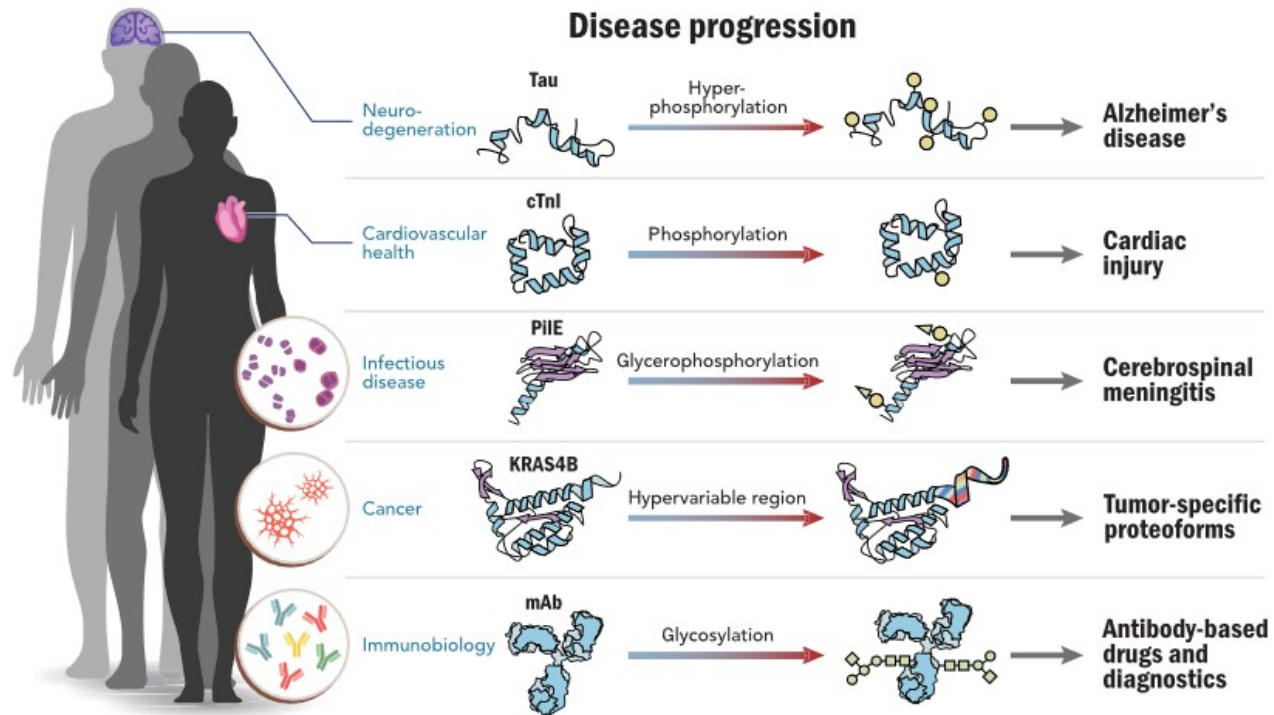
# Proteome Complexity



# Transcriptomics Does Not Accurately Predict Protein Profiles



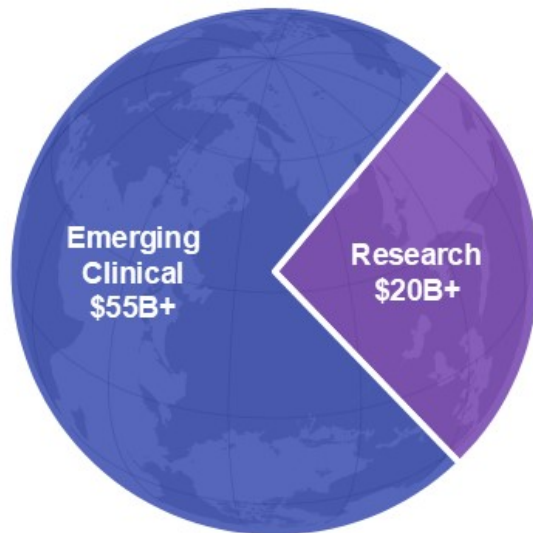
# Disease Progression Goes Beyond the Protein Level



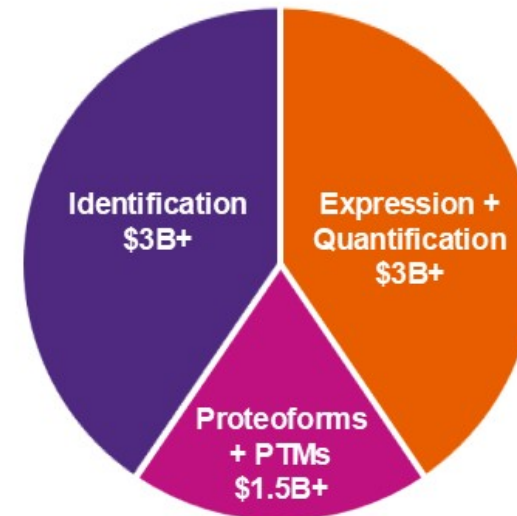


# Proteomics is a Large and Growing Market Opportunity

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



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



# Platinum Use Cases Today

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## 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

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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 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



**AI-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

## Protein-level Screening



**Samples:** 100s–1,000s per study  
**Proteins:** 100-1,000s per sample  
**Resolution:** Protein

Mass  
Spec

HT  
Affinity  
Assays

Ultra-  
Sensitive  
Protein  
Detection

Western  
Blots +  
ELISA

Edman  
Degradation

## Focused + Deep Characterization



**Samples:** 10s–100s per study  
**Proteins:** <50 per sample  
**Resolution:** Amino acid; single-molecule



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

Proteus™



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



**New architecture** (Proteus™), 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® Pro



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





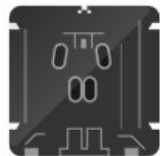
# Technology Development Pipeline

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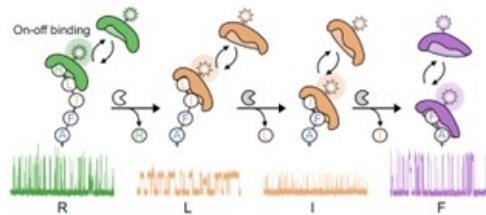
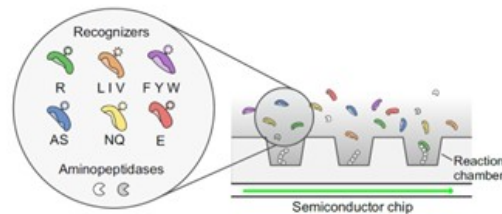
# Quantum-Si Core Technologies

## Platinum® System

### 2M Chip



## Chemistry Biomolecules



## Algorithms Applications

 Peptide Mapping

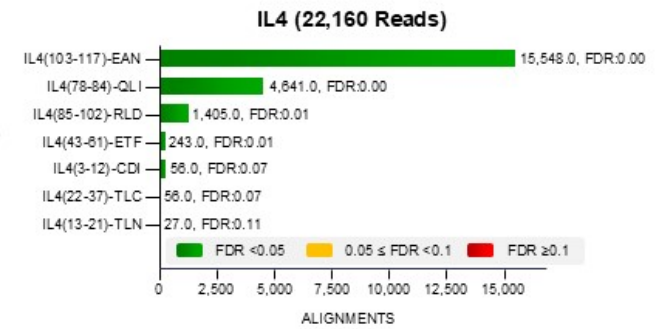
 Barcode Applications

 Variant Calling

 PTM Characterization

 Protein Identification

# End-to-End Protein Analysis



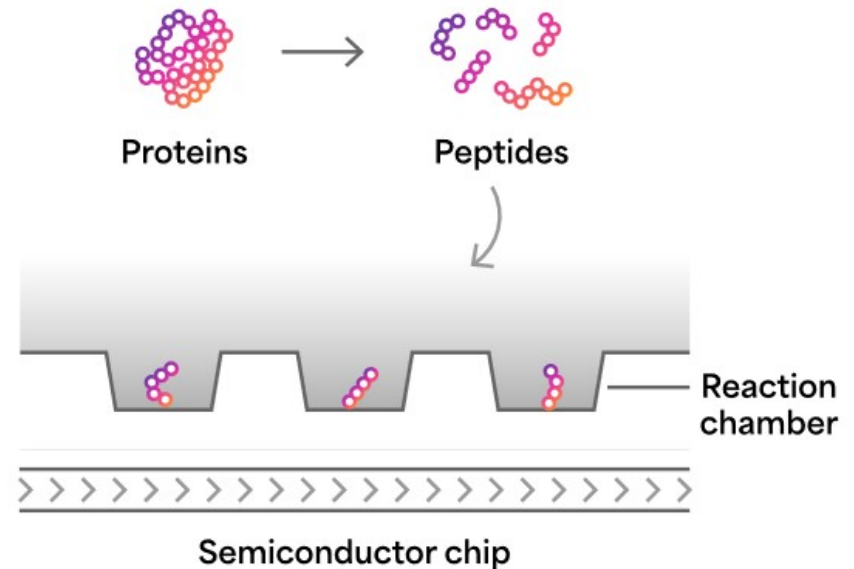
# Prepare Proteins for Sequencing



**Proteins are digested** into short fragments (peptides)



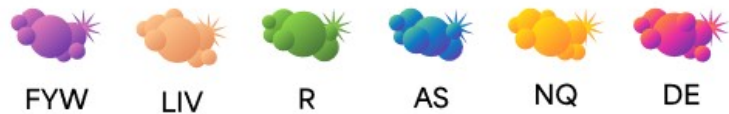
**Peptides are immobilized** at the bottom of reaction chambers on our chip



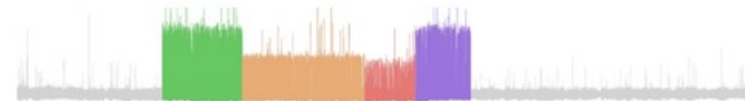
# Kinetic Signatures Uniquely Identify Proteins + Proteoforms



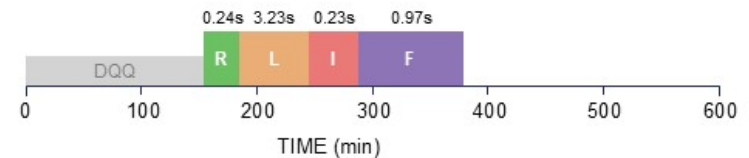
**Recognizers bind** amino acids in sequence



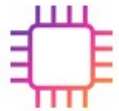
**Recognition events** produce kinetic signatures



**Kinetic signature plot**



# Rationale for New Technology Architecture



**Semiconductors** require large R&D investment



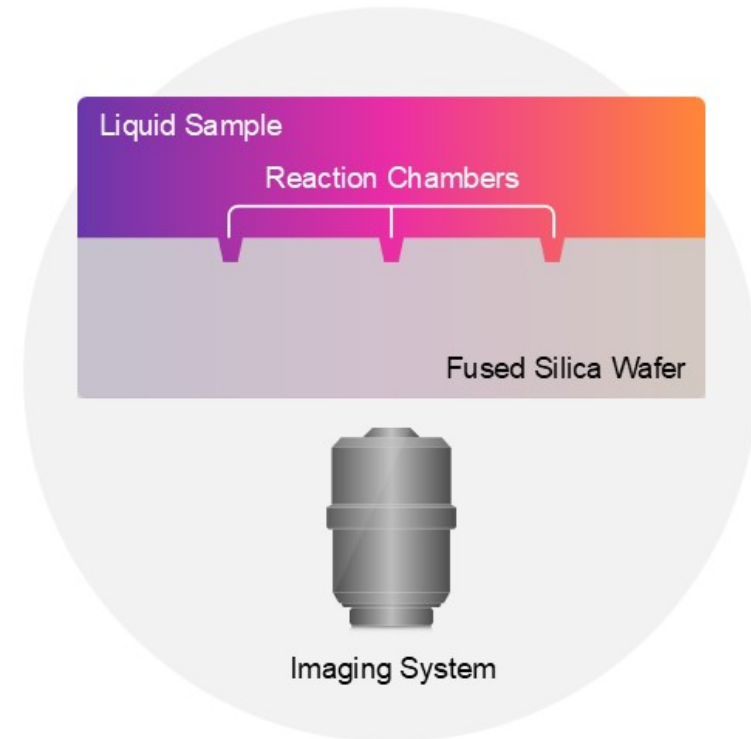
**Re-partitioning of system** allows for less expensive consumable



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



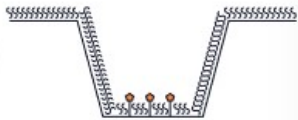
**Leverage high-performance**, commercially-available imaging components



# QSI Core Technologies



**Chip**



**Surface Chemistry**



**Instrument**



**Library Prep + Sequencing Reagents**



**Software**

# Chip + Surface Chemistry



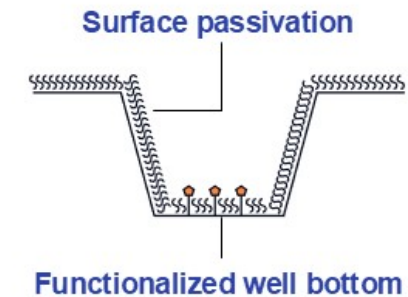
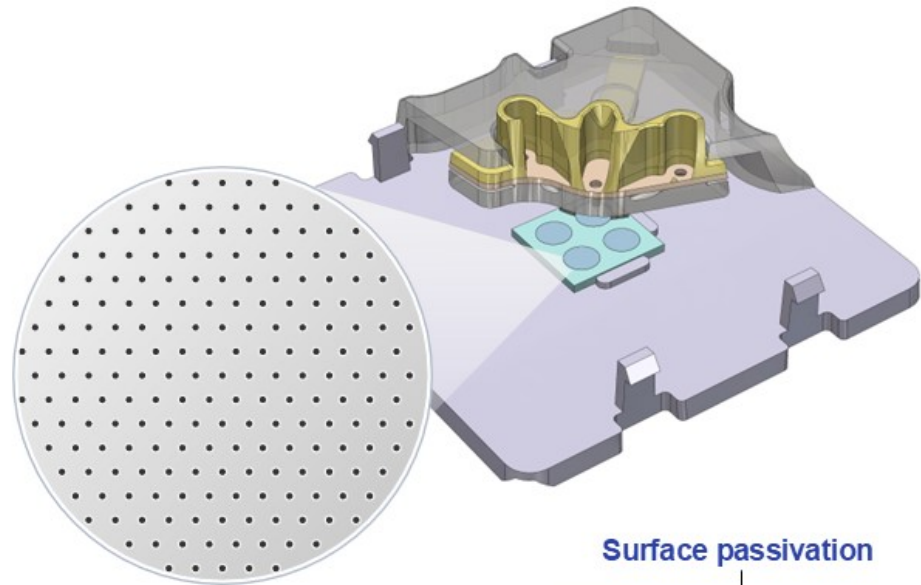
**Simple passive device** with approximately 20M wells (per flow cell) at initial launch



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



**Compatible with** existing surface chemistry





# Proteus™ Consumable Development



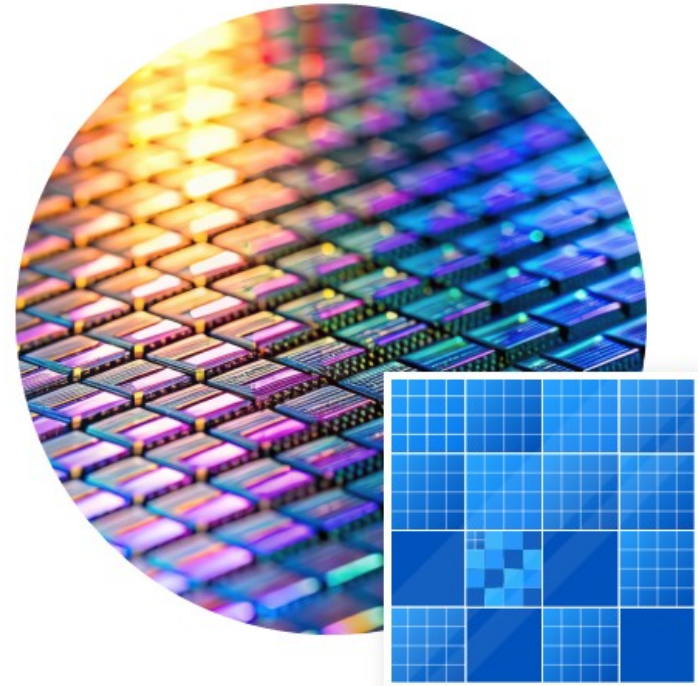
**Wafer process flow** developed in production foundry



**Prototype wafers** fabricated and tested



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



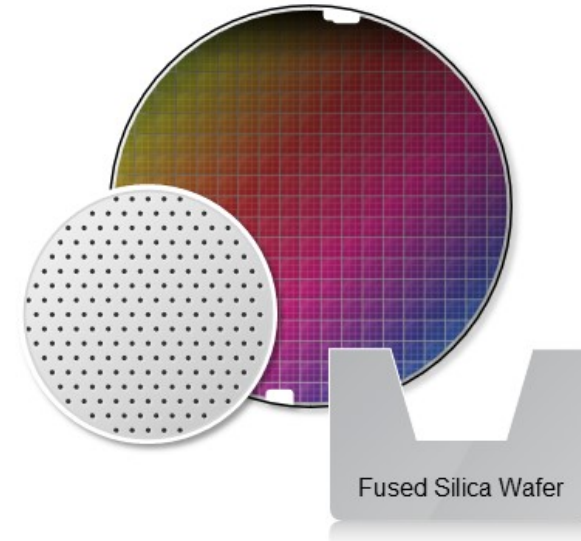
# Wafer Fabrication Process Development



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



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



# Proteus™ Instrument Development



**Move imaging components** to the system



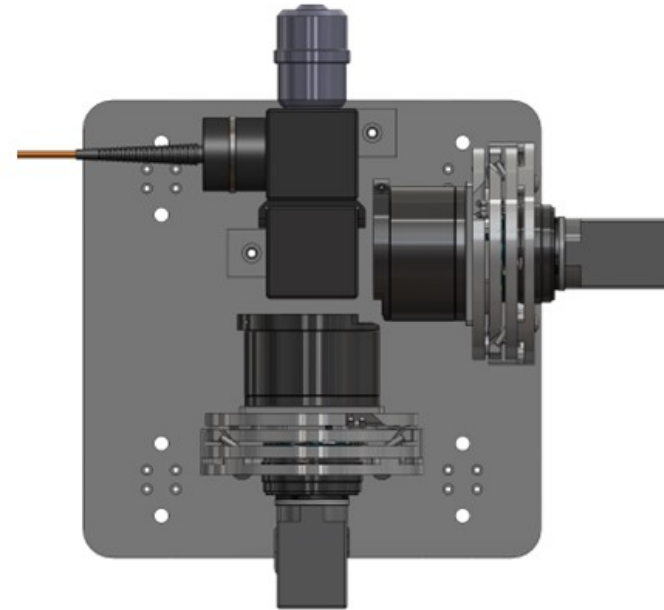
**Increase workflow automation**



**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



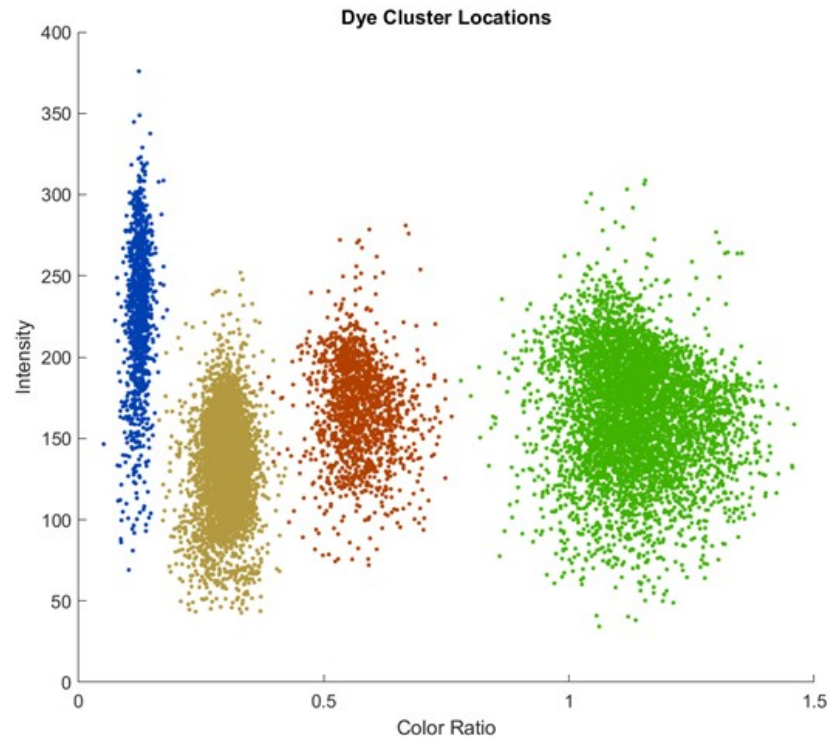
# Color Ratio is a Viable Alternative to Current Lifetime Detection



**Key elements** have already been de-risked



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



# Analysis Software



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

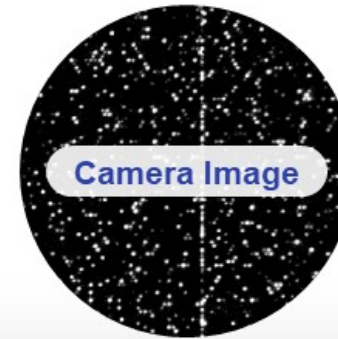
- Pulse detection, ROI calling, alignment, protein inference, and other applications



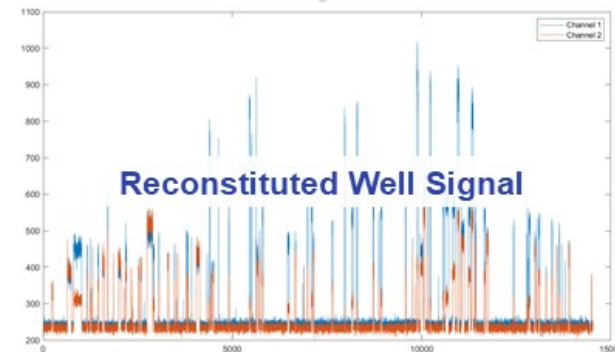
**Development required** for frontend image processing



**Well within** state-of-the-art capability



Registration Deconvolution



# Instrument Roadmap



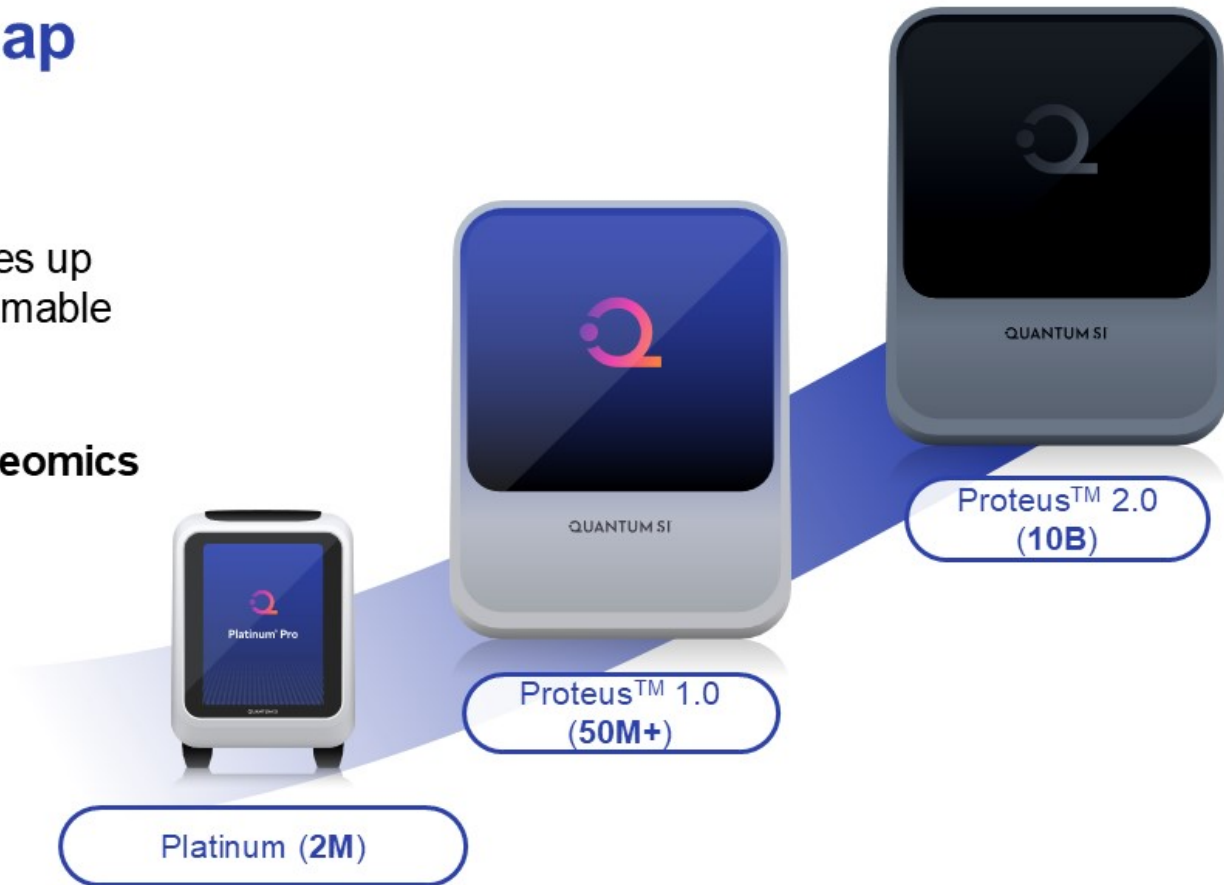
**New architecture** scales up to 10B reads per consumable



**Enables shotgun proteomics** of complex samples



**Puts us on path to** *de novo* sequencing



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

Brian Reed





# Agenda

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## Innovation at the Forefront of Proteomics

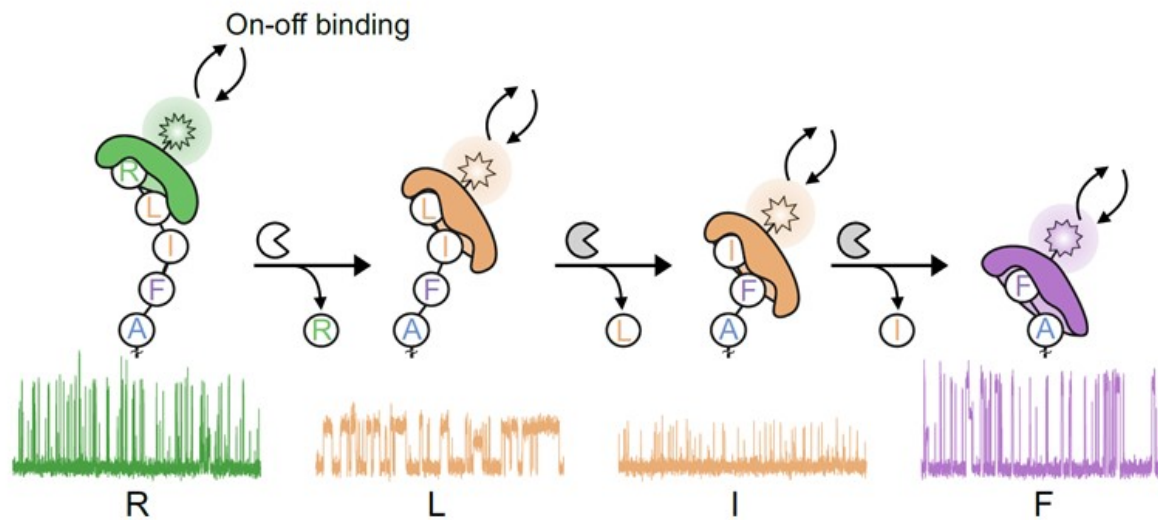
- 1 The path to complete proteome coverage
- 2 Ultrasensitive PTM detection for proteoforms
- 3 Deep, unbiased interrogation of high-complexity samples
- 4 Beyond sequencing: the first platform for top-down single-molecule proteomics

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## Acceleration on the Path to Complete Proteome Coverage

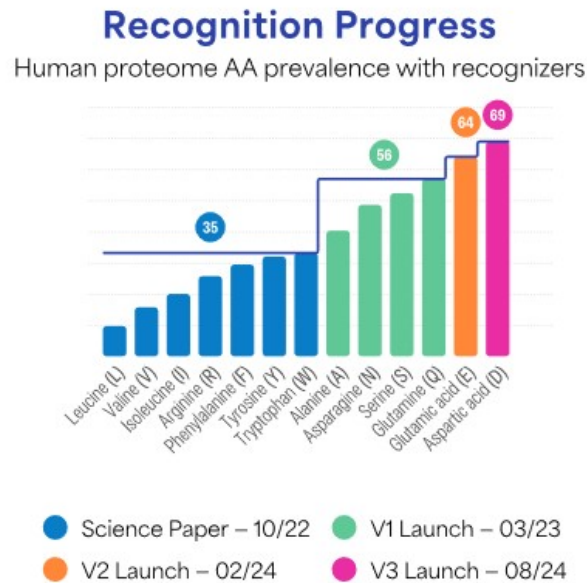
# Sequence Proteins on Platinum



- **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

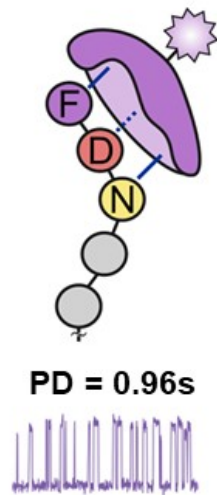
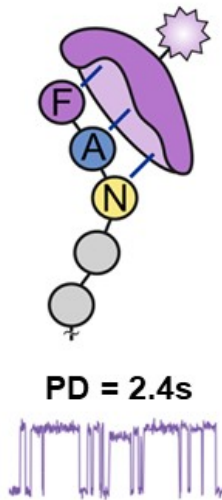
# A Rapid Path to Complete Amino Acid Coverage

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



- **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



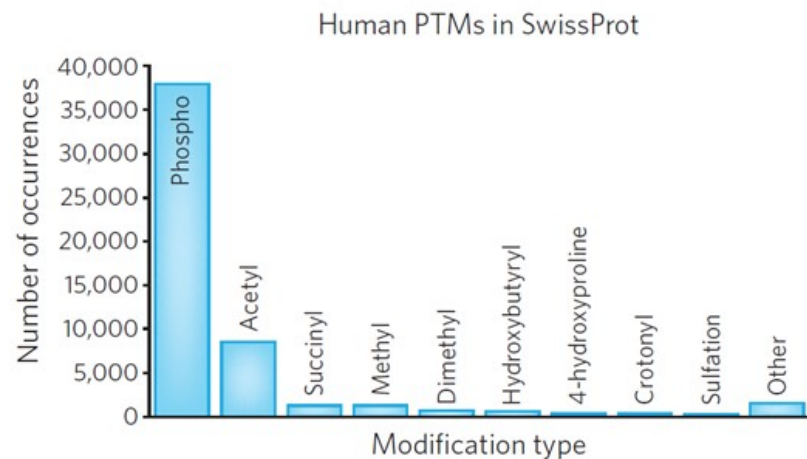
- **Recognizers physically contact residues downstream of bound NAA**
- **Influence is encoded in the peptide's kinetic signature** and is highly predictable
- **Kinetic signatures are a unique and powerful feature** of Quantum-Si's core technology
- **The acquisition of single-molecule kinetic information**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

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## Ultrasensitive PTM Detection for Proteins and Proteoforms

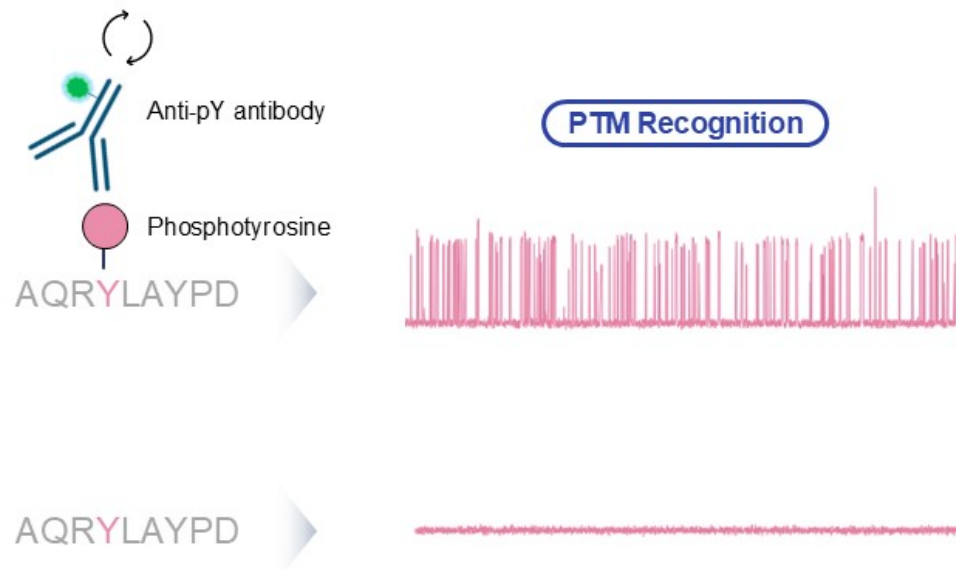
# Phosphorylation is the Most Abundant PTM in the Human Proteome



*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 PTM-associated diseases<sup>1</sup>

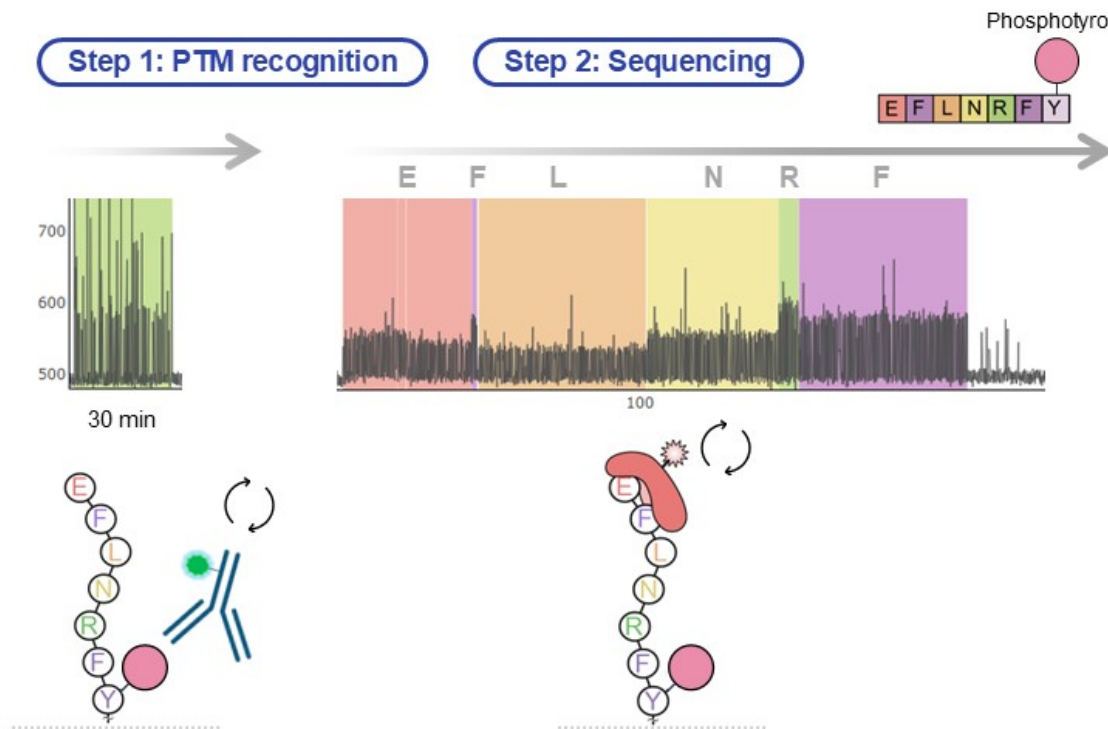
# Affinity Reagents as Ultrasensitive PTM Recognizers



- **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)



# Affinity Reagents as Ultrasensitive PTM Recognizers

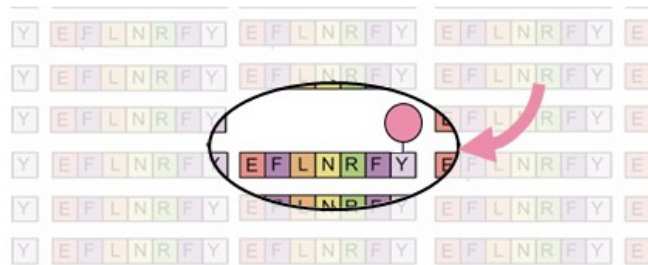
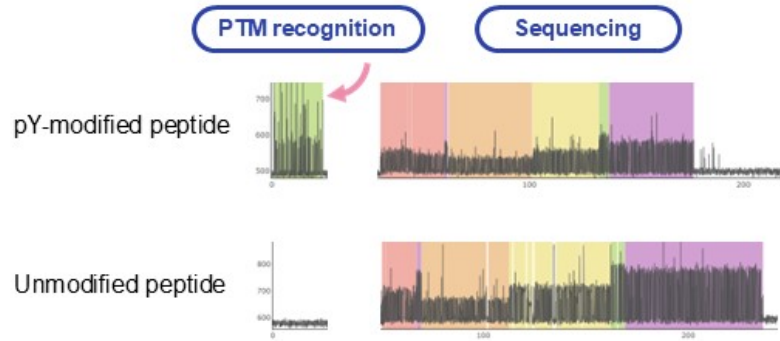


- **Step 1:** PTM detection for 30 minutes with PTM recognizer
- **Step 2:** Normal protein sequencing with NAA recognizers
- **PTM recognizers can be multiplexed and combined** with kinetic signatures to pinpoint PTMs in multisite configurations

# Ultrasensitive Phosphotyrosine Detection with CDNF

CDNF\_HUMAN

..QEAGGRPGADCEVCK[EFLNRFYK]SLIDRGVNFSLDTIEK ELISFCLDTK..



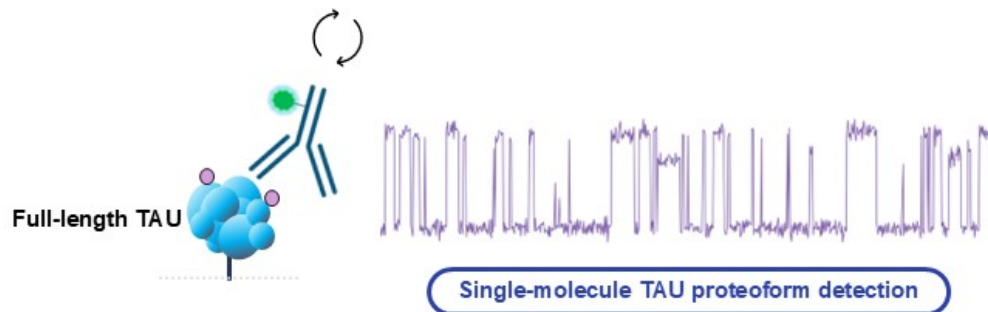
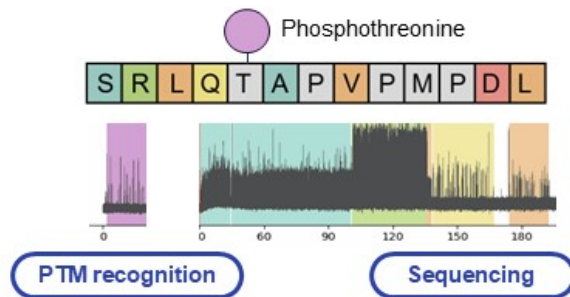
Detection of less than 1 PTM-modified peptide in 1,000

- **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

# Recognition of Human TAU Proteoforms

TAU\_HUMAN

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



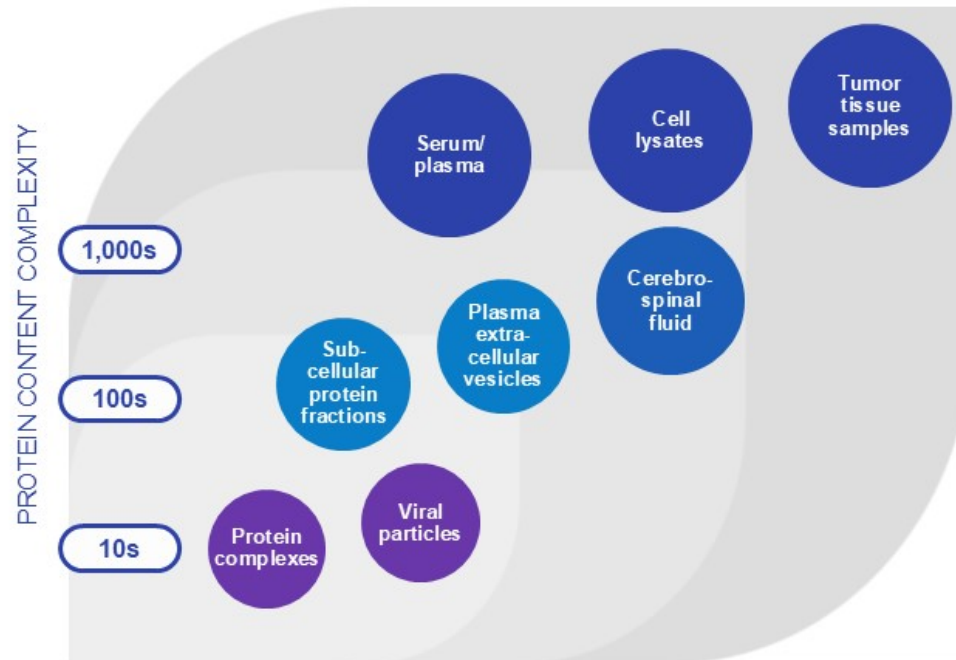
- **Affinity reagents** can be used in a bottom-up or top-down configuration
- **Example:** bottom-up recognition of pT\* on human TAU peptides, top-down detection of immobilized full-length TAU proteoforms
- **Real-time approach enables proteoform detecting** reagents to be run simultaneously
- **First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

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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



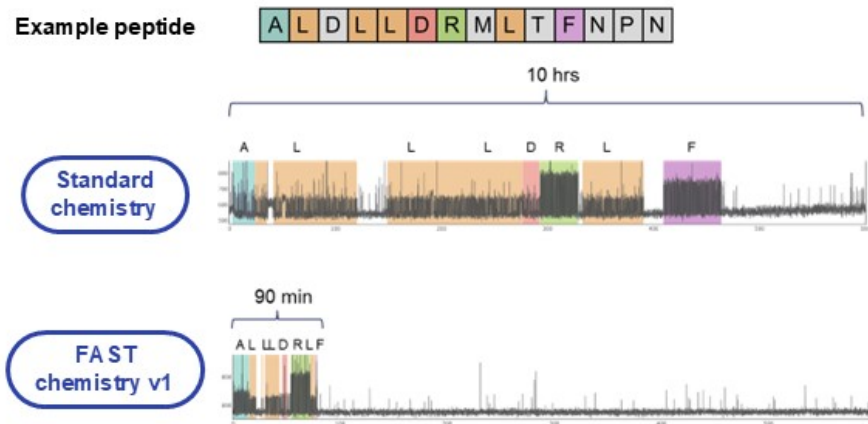
- **Biological samples like serum** contain hundreds to thousands of proteins with wide dynamic range of abundance
- **Unbiased, consistent, accessible interrogation** of these samples is a challenge in proteomics
- **Sequencing is not limited to predefined content:** enables discovery of changes in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

# Fast Sequencing for Deep Coverage and Rapid Sample-to-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

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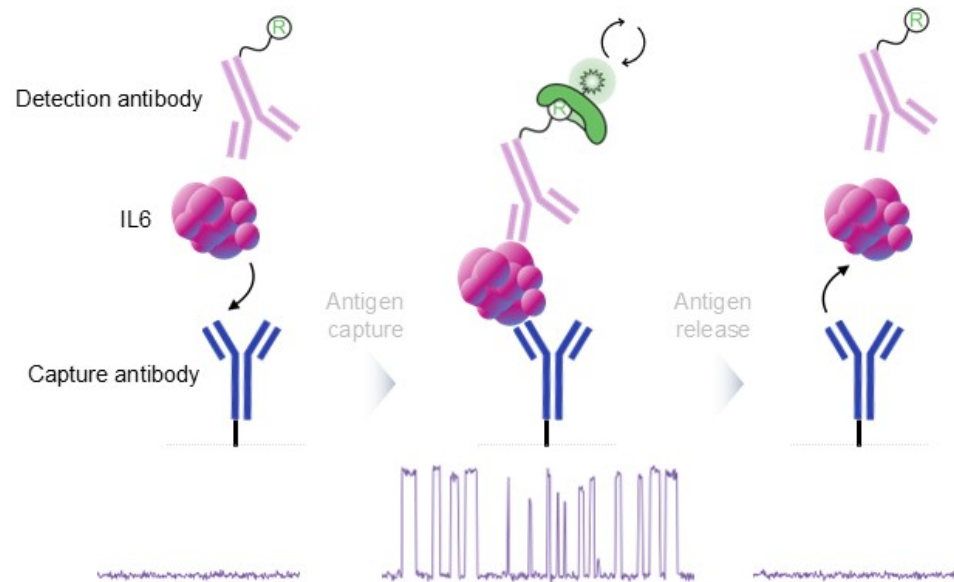
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**Beyond Sequencing: the First  
Commercially Available Platform for  
Top-Down Single-Molecule Proteomics**



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

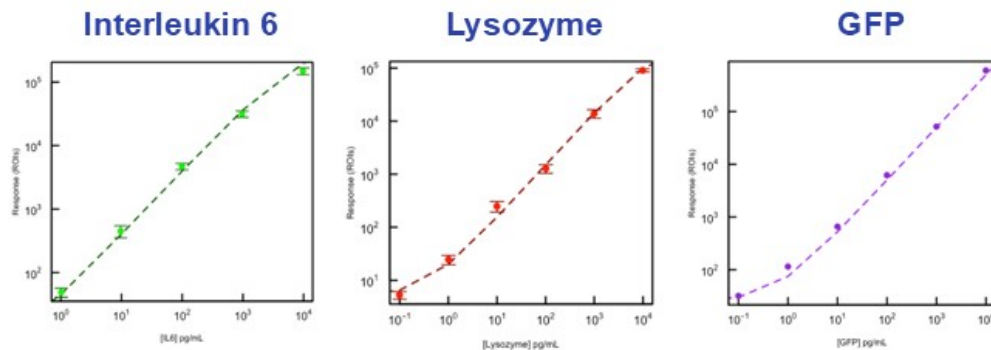
Dye-cycling enables ultrasensitive real-time detection of biomarkers



- **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

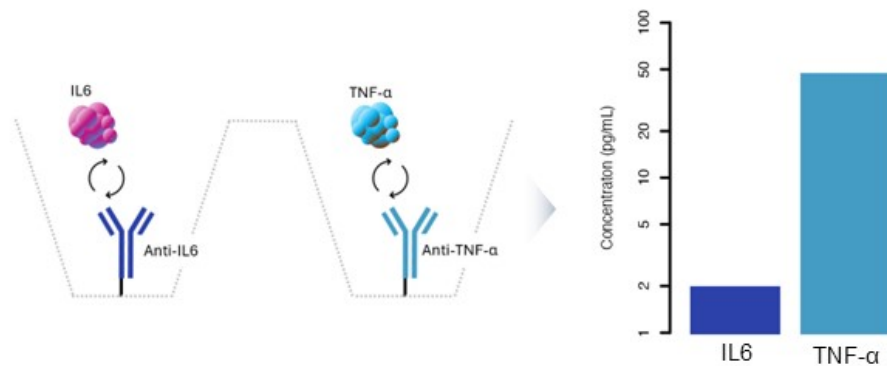
Direct detection of proteins in serum with high sensitivity



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

# Multiplexed Ultrasensitive Protein Biomarker Detection

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



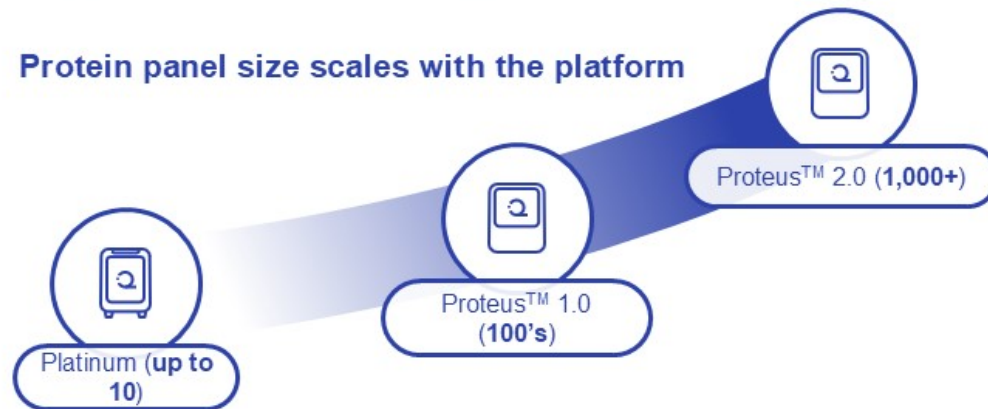
- **Affinity reagents against multiple biomarkers** can be loaded on the chip
- **Dye-cycling approach enables discrimination** of biomarkers by fluorescence and kinetic properties in multiplexed assays, along with PTMs

# A Platform for Ultrasensitive Detection of Protein Panels

Multiplexed biomarker detection directly in serum



Protein panel size scales with the platform



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- **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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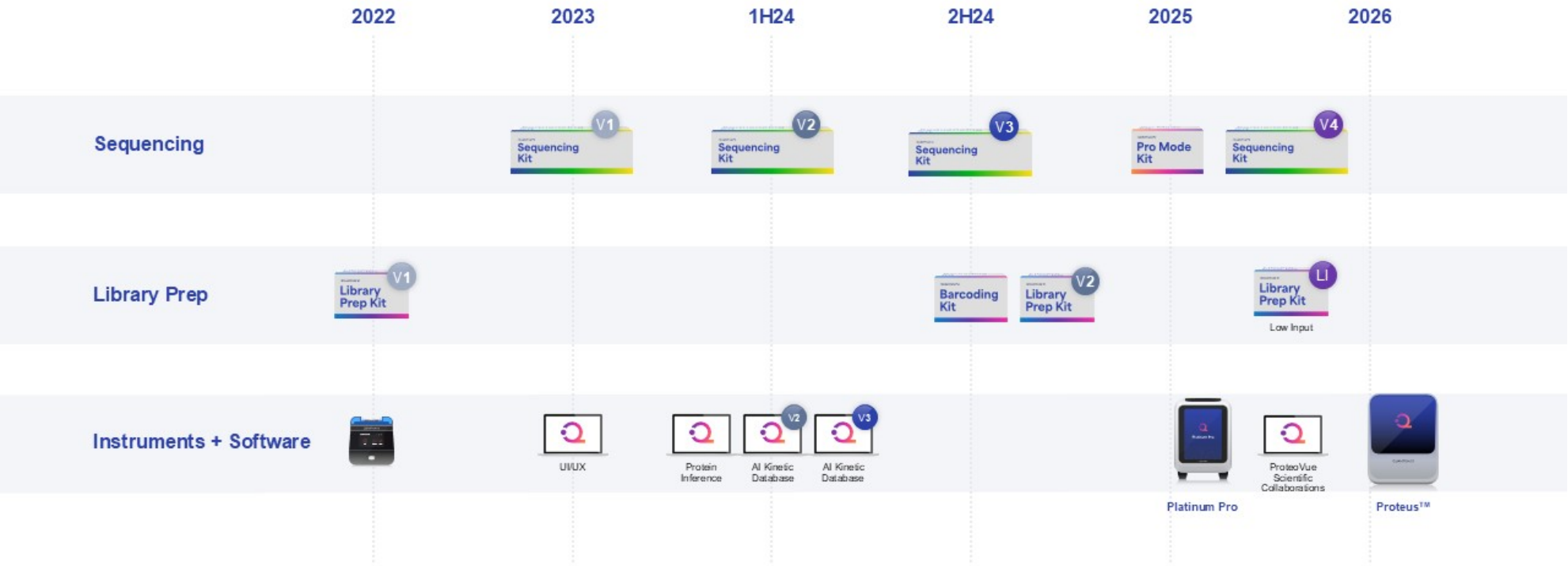
# Platform Roadmap

Nov 20, 2024

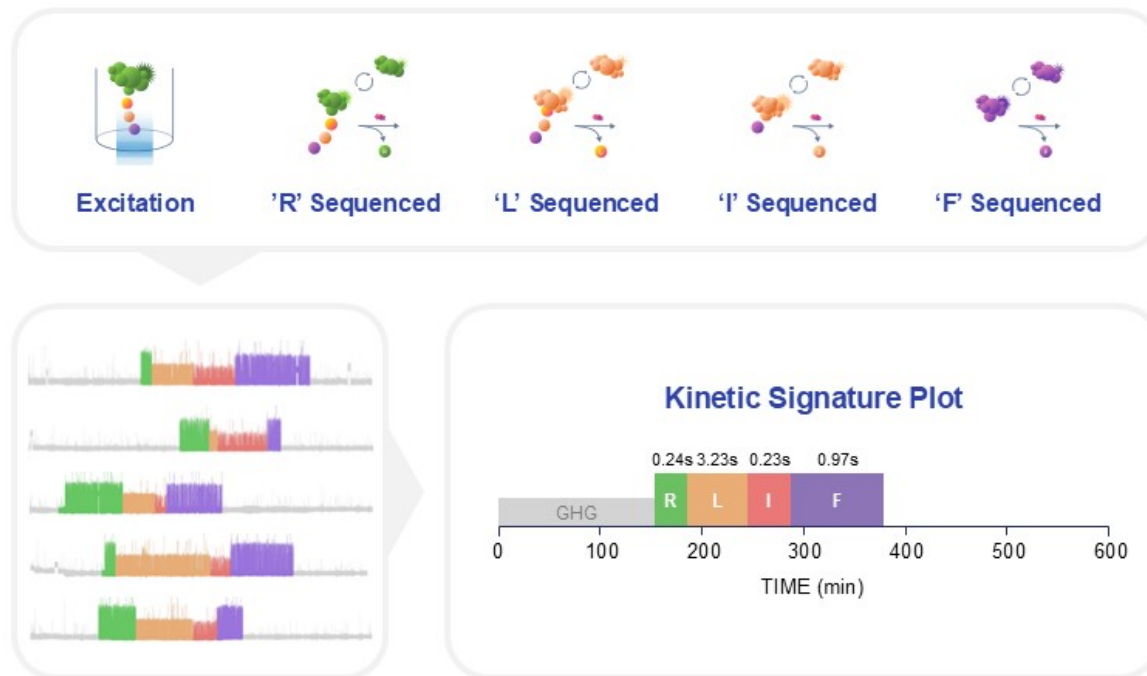
John Vieceli



# Innovation Pipeline is Robust and Accelerating



# Sequencing Analysis Software



## Sequence

Measure fluorescence from single molecule binding of N-terminal amino acid recognizers



## Pulse Caller

Assign pulses to a recognizer based on fluorescence intensity and lifetime



## Analyze

Determine amino acid sequence using kinetic signature

# Software Workflows for Next-Gen Protein Sequencing™



## Protein Inference

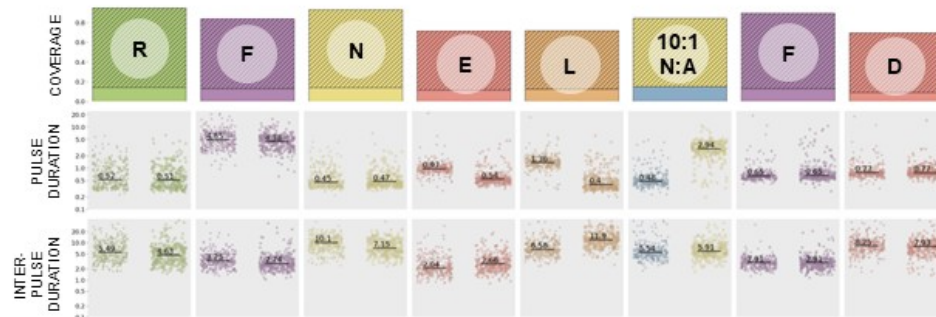
Kinetic signature enables inference of sample protein from whole human proteome panel

Rank	Inferred Protein	Score	Likelihood	Mass (kDa)	Length
1	<b>IL4</b> spIP04112IL4_HUMAN	11.035496	99.99%	17	153
2	spIP06127ICD5_HUMAN	0.593929	44.78%	55	495
3	spIQ15208ISTK38_HUMAN	0.582068	44.12%	54	465
4	spIQ96LQ0IPPR36_HUMAN	0.506878	39.76%	49	422
5	spIQ81WR1ITRI59_HUMAN	0.440162	35.6%	47	403
6	spIQ9UMR3ITBX20_HUMAN	0.428596	34.85%	49	447
7	spIQ96EU6IRRP36_HUMAN	0.403635	33.21%	30	259
8	spIQ9H2F9ICCD68_HUMAN	0.371386	31.02%	39	335
9	spIQ8IVI9INOSTN_HUMAN	0.365859	30.63%	58	506
10	spIO9BZ81IMAGB5_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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# Artificial Intelligence

# N-terminal Amino Acid Recognizer Development

 **FYW**  **LIV**  **R**

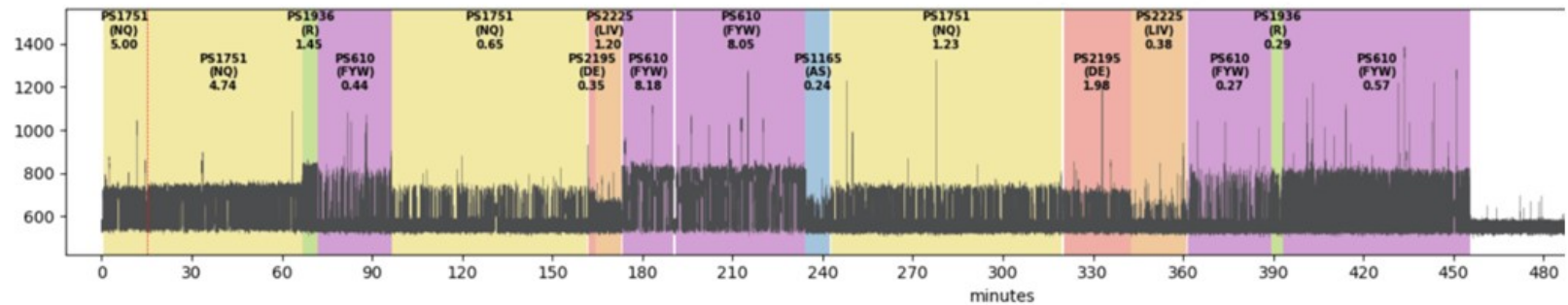
**Science Publication**  
Oct 2022

 **AS**  **NQ**

**Version 1 Kit**  
Mar 2023

 **DE**

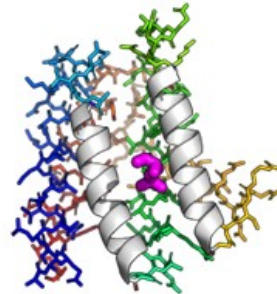
**Version 2/3 Kit**  
Jan/July 2024



# Recognizer Protein Design AI



Amino acid recognizer  
**backbone design**



Amino acid recognizer  
**sequence design**



**Orthogonal verification**  
of amino acid recognizer  
protein design



Protein design AI leverages **NVIDIA GPUs on-premises and in Amazon Cloud**

# 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 AI

Currently predicts **~4.6 million** pulse widths used in analysis



## Better Performance

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



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Platinum® 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

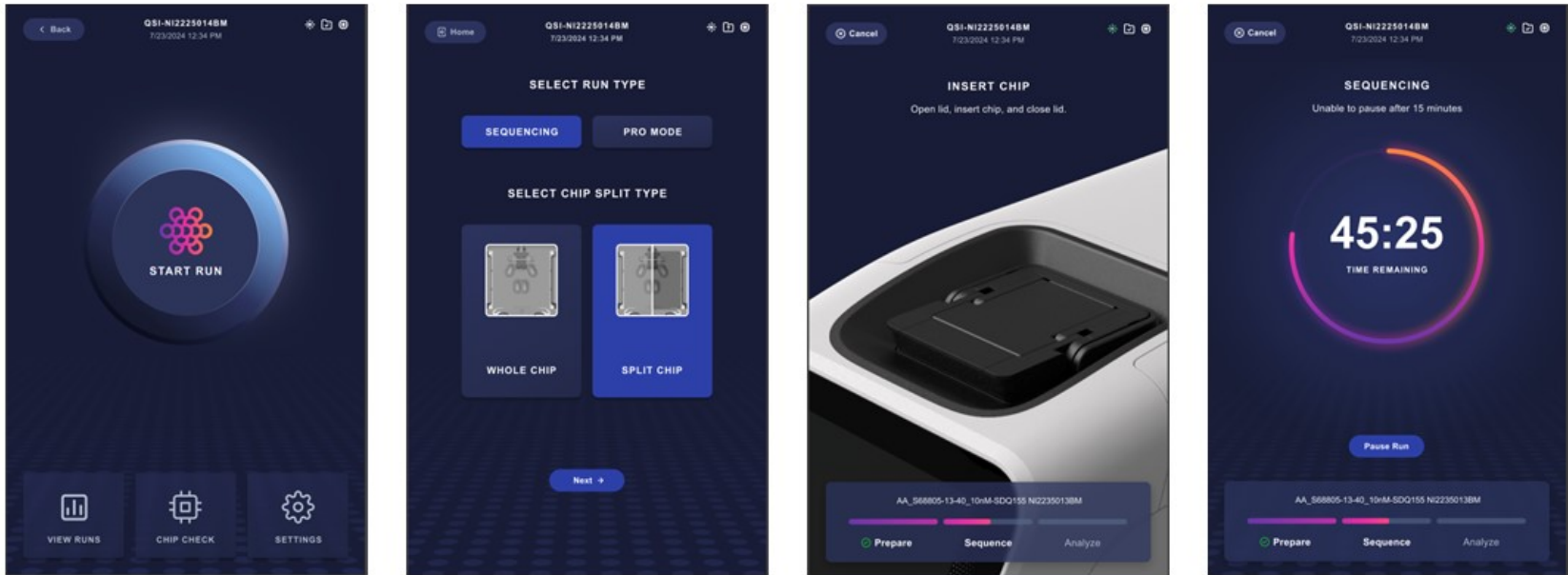


**Onboard analysis** or via the cloud



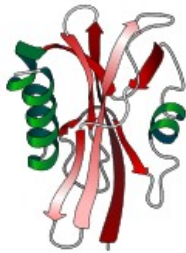
**Enhanced user interface**

# Streamlined Workflow Improves Usability





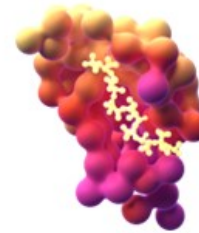
# Pro Mode Available only on Platinum Pro



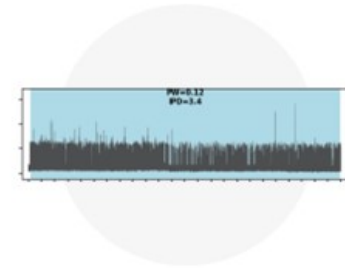
Protein of Interest



Dye-labeling Kit



Protein Binding



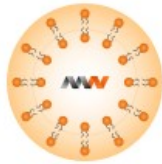
Binding Kinetics



Kit and platform enable detection of **single-molecule protein binding and kinetics**

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

IN VIVO



Equimolar 5 barcode mix encoded as mRNA and packaged into LNPs



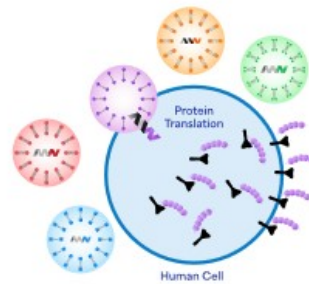
Inject into mouse model; harvest target tissues



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



IN VITRO



# 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

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Proteus™

# Proteus™ 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™ Increases Number of Samples



**Run one or two** samples simultaneously



**Reagent cartridges** with sequencing workflow automation



**Run up to 8 samples** in one sequencing run



QUANTUM.SI™

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## 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

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

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**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

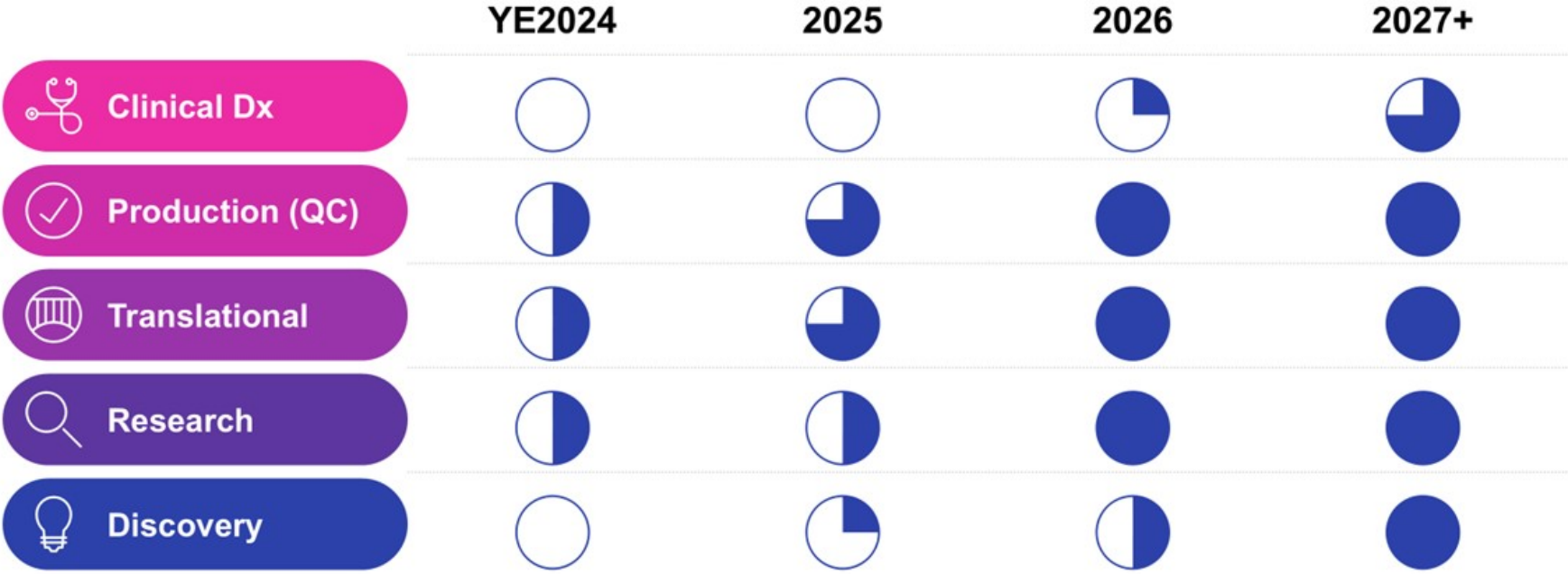


**QSI ultrarapid sequencing** can enable significant increase in sample throughput per day and unlock time-sensitive applications (e.g. clinical diagnostics) in the future



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

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



# Proteomics Lab Today

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**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



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Q&A

