

TURNING THOUGHT LEADERSHIP INTO LEADS: BIO-TECHNE'S SCIENCE MILESTONE

OVERVIEW //

Bio-Techne partnered with *Drug Discovery News* on a **Science Milestone** to support the launch of their new high-throughput western analysis platform. The goal was to position the company as a long-standing innovator in protein analysis while generating qualified leads through engaging, high-quality content.

CLIENT //

- Trace the evolution of western blotting and automated western techniques
- Highlight Bio-Techne's decades-long leadership in the field
- Build awareness of their new platform
- Reach and convert a targeted audience of life science professionals

milestone

Leapfrogging to quantitative, high throughput protein detection and analysis

The ability to detect and analyze proteins has long been a key quest for scientists. For decades, the pursuit of accuracy and reliability in protein detection led to a series of technological leaps. From the early days of gel electrophoresis and the invention of western blotting to the development of automated western analysis, researchers continuously push the boundaries of what's possible with protein analysis tools.

1979 A new blot in town

As scientists began to realize the potential for analyzing DNA samples, it also inspired researchers to explore its potential for other applications. In 1979, George Stark, a biochemist at Stanford University, and colleagues introduced a DNA blotting method using electrophoretically transferred (EMSA) oligonucleotide probes to detect specific DNA sequences. Stark's team further refined the blotting technique for protein analysis and published their methods in July 1979. The method allowed researchers to detect specific proteins by incorporating polyvinylidene difluoride (PVDF) membrane support papers in cell lysates. For protein transfer, they used DNA paper and to probe the target proteins, they utilized a radioactive protein. Using this method, the group successfully identified kinase enzymes in a rat brain tissue extract.

Meanwhile, two biochemists on opposite sides of the world, Harry Towbin at the French Mérieux Institute and Neil Boran at the Fred Hutchinson Cancer Center, were independently developing a similar protein blotting method. Both teams established an electrochemical blotting procedure to transfer the proteins, which involved using an electric field to drive proteins from an electrophoretic gel onto nitrocellulose membranes. Towbin utilized a primary antibody to bind the target proteins, followed by a secondary antibody of fluorescein isothiocyanate (FITC) labeled antibody to detect fluorescently. Boran's approach to protein detection was slightly different—he used a primary antibody and a secondary visualization process.

Towbin published his method in 1979, just one month after Stark's paper, while Boran's manuscript, prepared around the same time, was initially rejected. In 1980, Boran's article was accepted by the same journal, and combining the two works, he called the method western blotting.

1980s-2000s Rapid adoption, persistent challenge

Western blotting quickly captured the attention of researchers worldwide. Within the first decade of its inception, it became a key diagnostic tool for identifying disease-associated proteins. Some researchers applied western blotting analysis to identify human immunodeficiency virus antibodies (HIV antibodies), while others used it to detect *Cryptosporidium* antibodies, a diarrheal disease caused by intestinal protozoa (1).

During this time, technological advancements made western blotting more accessible. Researchers introduced more membrane materials, such as polyvinylidene fluoride and nylon, to improve blotting processes. These materials were durable, stable, and enabled protein reuse efficiently. Detection methods also evolved. Chemiluminescence, which involves the use of chemically resistant substrates that emit light when they react with enzyme-linked secondary antibodies, became the preferred method over autoradiography for many researchers.

Despite these improvements, achieving reproducibility was a significant challenge for researchers performing western blotting experiments. The multiple blotting steps—gel preparation, transfer, blotting, washing, and detection—often introduced variability and led to inconsistent results. This quest for a growing interest in building more standardized western blotting operations. Cell Bioscience, a company that introduced its *ProteinSimple* in 2011 and has become part of Bio-Techne, began exploring automated solutions.

Tom Yang, now Senior Director of Engineering at Bio-Techne, joined Cell Bioscience in 2011 after several years working on human genome sequencing. "While the western blot was the lifeblood of molecular biologists at the time," Yang said, "it was very hard for a scientist to handle a flow gel, perform blotting, and take the membrane through different incubation and washing steps." Yang and the Cell Bioscience team worked hard to simplify the process.

1970s Foundational techniques

In 1962, scientists first observed that clay particles in water moved toward the positive electrode under an electric field, a phenomenon that laid the foundation of electrophoresis. It wasn't until the 1960s, with the introduction of electrically charged substrates such as nitrocellulose and polyacrylamide, that researchers could use gel electrophoresis to separate proteins in complex mixtures (2). However, these early electrophoresis methods lacked the resolution to differentiate proteins of similar sizes.

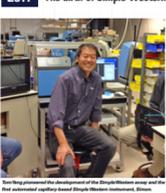
In 1970, Ulrich Laemmli, a molecular biologist studying hemoglobin (3) at the Medical Research Council Laboratory of Molecular Biology, recognized that sodium dodecyl sulfate (SDS), a protein-detergent detergent, could help separate polypeptide chains based on their molecular weight by breaking through hydrophobic interactions. By creating a discontinuous two-gel system containing SDS and polyacrylamide at different concentrations, Laemmli was able to observe distinct protein bands. This method, now known as SDS polyacrylamide gel electrophoresis (SDS-PAGE), allowed him to identify previously unknown proteins that made up the hemoglobin complex (4).

At the same time, other researchers were using gel electrophoresis to separate nucleic acids with DNA in 1970. Edward Southern, a molecular biologist at the University of Edinburgh, took the technique a step further by detecting specific DNA sequences from electrophoretically separated segments. To do this, he introduced a new step: blotting. After separating DNA fragments on an agarose gel, he transferred them onto a nitrocellulose membrane, which captured the DNA (5). He then used radioactive RNA probes that could bind to specific DNA sequences and detect them with autoradiography. Southern renamed his method after himself: Southern blotting.



Edward Southern developed Southern blotting by introducing a blotting procedure following gel electrophoresis, enabling the detection of specific DNA sequences.

2011 The birth of Simple Western



Tom Yang pioneered the development of the Simple Western system and has led automated assay development at Bio-Techne.

2014-2018 Enhancing sensitivity and multiplexing

In 2012, while Yang and his team were making progress in developing Simple Western, another scientist, now Director of R&D at Bio-Techne, was conducting groundbreaking research in Dan Fisher's laboratory at Stanford University. There, the researchers used a capillary electrophoresis-based protein separation system in collaboration with the ProteinSimple team, which led to an opportunity for him to join that product development team.

After the release of Simple Western, Demery recalled, "We got a lot of feedback from scientists who had tried the ProteinSimple but still had the high level of variability that the DNA blotting method introduced. We were working on developing a more robust, sensitive system that could handle more complex samples (6).

In 2014, ProteinSimple introduced the *Simple Western* system, a Simple Western system that could run up to 25 samples in three hours, with sensitivity 100 times greater than SDS-PAGE. This collaboration involved a partnership with advanced multiplexing capabilities, enabling assays, and the same level of automation and simplicity. In 2018, the ProteinSimple team announced the release of the new instrument, marking the *Simple Western* platform's evolution. "These advances are exciting," Demery said, "because they've got us to more places than we've ever had before. We have lots of other things we're working on, but these 20 instruments are a big step."

With experience in DNA sequencing and capillary electrophoresis, Yang was particularly interested in the western blot and the western blotting approach to protein separation. Yang recognized the need for gel.

The next goal was to control the retention of proteins during steps in the capillary tube, and the key challenge for this was introducing proteins to the capillary wall. "We applied UV light to photo-capture support proteins to the capillary wall. By developing protein surface chemistry, we were controlling that more precisely with electrophoresis and photo-capture," Yang explained. "Even though we don't talk about it, it's still a physical attachment of the protein to the wall."

To clarify the separation, they used a primary antibody and a secondary visualization process using horseradish peroxidase-conjugated antibodies and chromogenic substrates—*not* after the capillary. These researchers set a goal to have their protein detection method, which they named the *Simple Western* (7). "The original goal of the western blot publication in *Science* was to just do it in a way that a much wider audience could use," Yang said.

With Simple Western fully established, Yang's team moved ahead with automating the technique with a robot. They brought together an experienced team of scientists, engineers, and software developers. "We put together many existing technologies and invented a key technology that bridged everything, from protein immobilization to immunodetection," Yang said.

In 2017, the ProteinSimple team unveiled the first automated capillary-based Simple Western instrument, the *Simple Western* System, at the ProteinSimple Organization Research World Congress. "When we first announced and showed the instrument at that conference, there was a lot of attention. It was very exciting," Yang recalled. By automating the process, the team demonstrated significantly improved accuracy in protein analysis (8). "We're moving into the quantitative space, where our data precision, repeatability, and quality are just heads above some traditional Western blotting," Yang said.

It wasn't long before the team set its sights on a new challenge. "Our customers were wanting higher multiplexing capabilities," Demery said. "The next phase of engineering began to take shape."

Demery and a team of biochemists, engineers, software developers, and product managers worked in parallel on the instrument, software, and assays. Together, they developed dual detection capabilities—immunofluorescence and fluorescence—allowing users to detect multiple targets in their samples. The team also incorporated a large protein measurement system that could handle a wide variety of assays, including kinase assays, which require the ongoing and repeating steps of traditional western blotting but with the same quality, enabling multiplexing capabilities.

This collaboration effort resulted in a platform with advanced multiplexing capabilities, enabling assays, and the same level of automation and simplicity. In 2018, the ProteinSimple team announced the release of the new instrument, marking the *Simple Western* platform's evolution. "These advances are exciting," Demery said, "because they've got us to more places than we've ever had before. We have lots of other things we're working on, but these 20 instruments are a big step."



Jeny Demery led the scientific team involved in the development of *Simple Western* technology, which was featured in *Drug Discovery News* as a Science Milestone.



The *Simple Western* platform, powered by Simple Western Technology, is a high-throughput automated western blotting system that can handle complex samples, providing a lot of data points and reducing the time to get results.

2025 Breaking new ground

The success with Simple Western and Jeny's role led to his new appointment as President of ProteinSimple. The team has been dedicated to helping Simple Western capabilities to better support diverse research needs. For example, the *Simple Western* system was introduced in 2022, offering an accessible, budget-friendly solution for researchers seeking to automate western blotting workflows.

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