



Introduction to latest LC-MS/MS technology innovations

SCIEX 7500 system and ZenoTOF 7600 system

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RUO-MKT-11-11650-C and RUO MKT-11-13113-A

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NEW SOFTWARE, NEW ION OPTICS, NEW SOURCE



- SCIEX 7500 system enables new levels of quantification across a large suite of sample types and workflows
- SCIEX OS software is the modern mass spectrometry software platform that transforms your samples into meaningful analytical answers

The Turbo V ion source evolution

FOR ROBUSTNESS AND RUGGEDNESS



Turbo V ion source



IonDrive Turbo V ion source



OptiFlow ion source



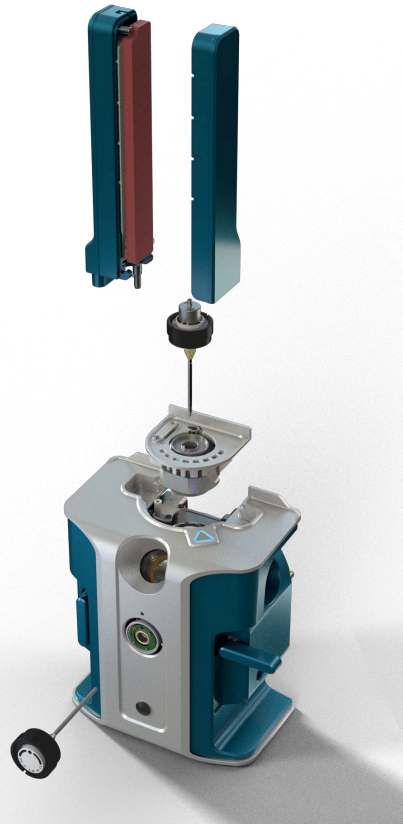
OptiFlow Pro ion source

DESIGNED FOR MODULARITY

ESI



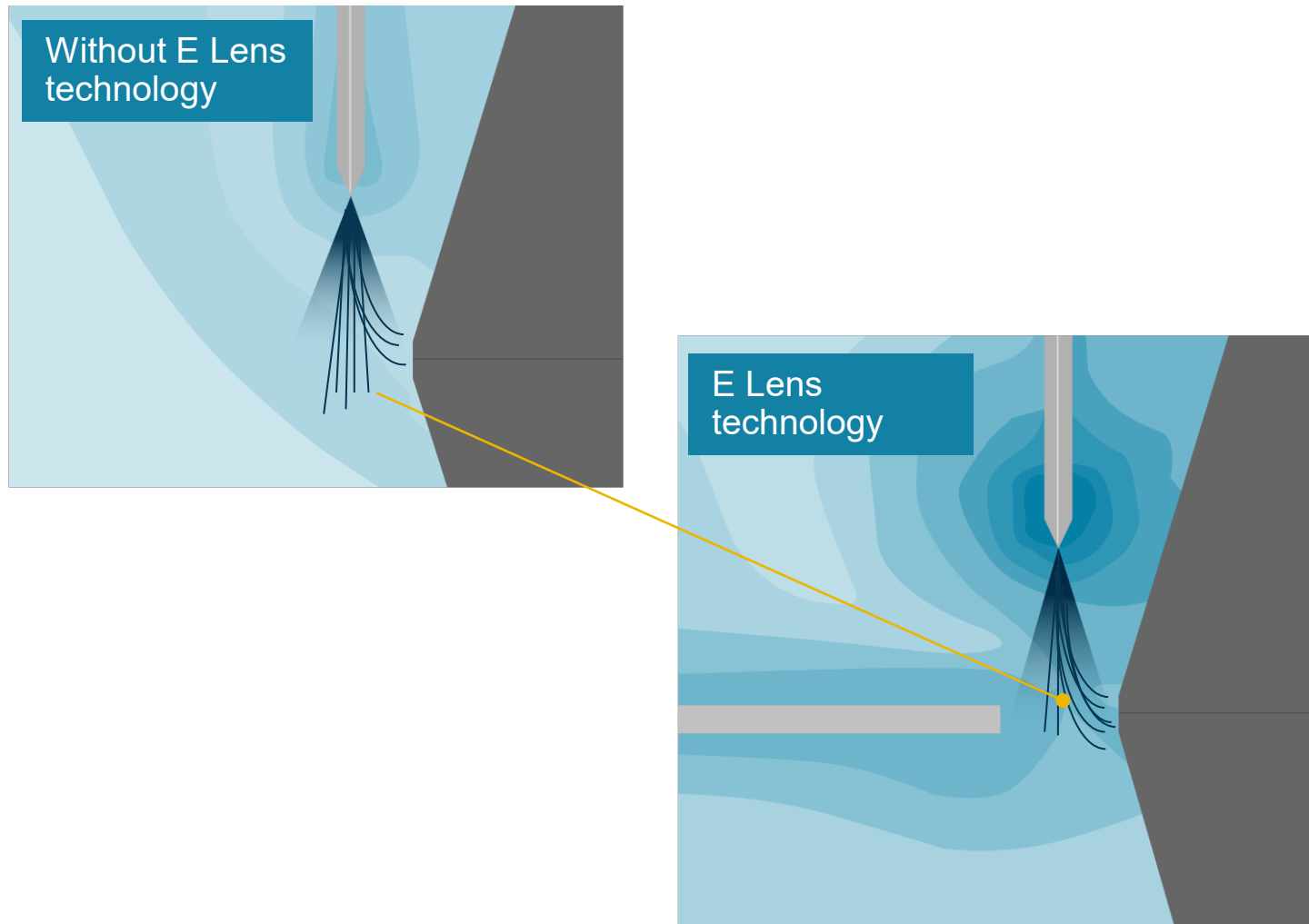
Microflow



APCI

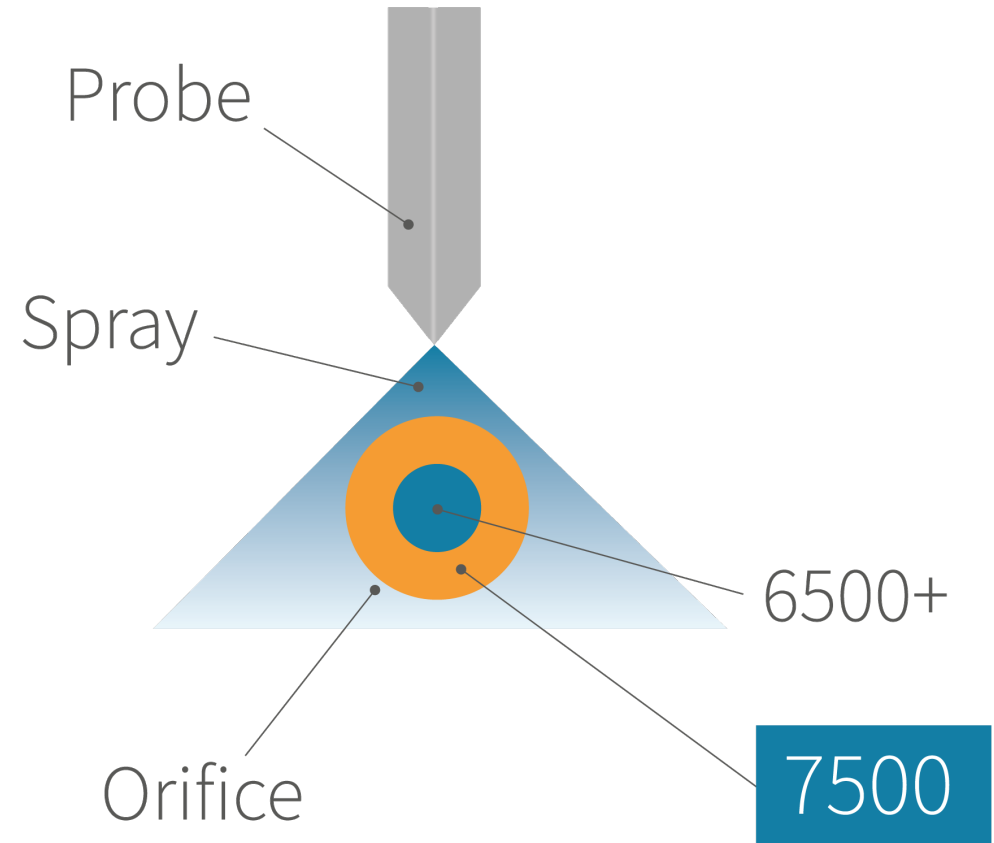


- Maximum performance from 1 $\mu\text{L}/\text{min}$ to 3 mL/min without adjustment with drop in probes and electrodes
- A versatile ion source with wide compound coverage with interchangeable ESI and APCI towers
- E Lens Technology for both micro and analytical flow for enhanced sensitivity compared to ESI alone
- Reduced user-to-user variation
- Designed with extensibility in mind



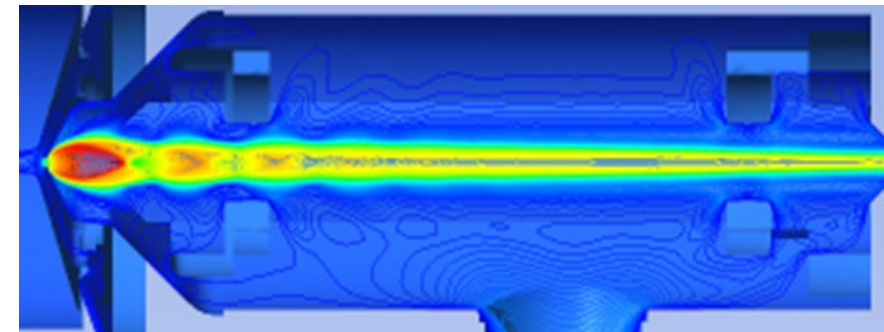
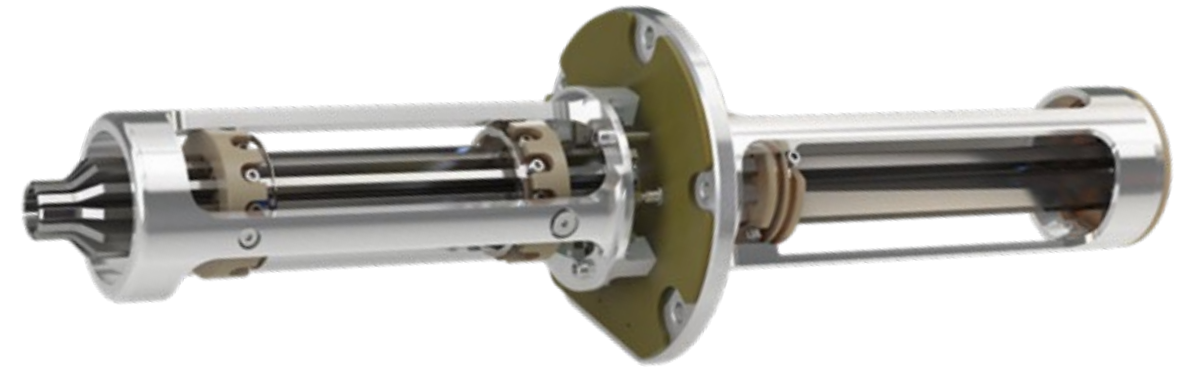
- In ESI the E Lens technology drives ions towards the orifice
- The E Lens technology creates a stronger field that the droplets must traverse leading to more efficient break-up and release of ions from the droplet
- Gains in performance are up to 2-fold with the largest gains at microflow

- Enabling greater sensitivity can be achieved through gains in the generation of ions, capturing and transmitting ions and detecting ions
- QTRAP 6500+ system with IonDrive technology delivered performance improvements in these key areas
- SCIEX 7500 system makes another leap forward in the capture and transmission of ions
- Sampling area of the SCIEX 7500 system orifice is 4.3x larger than the QTRAP 6500+ system orifice



THE D JET ION GUIDE IS A DUAL STAGE RF ION GUIDE

- The D Jet ion guide efficiently captures and transmits the ions in the high gas flow behind the orifice plate
- The tapered dodecapole geometry of the D Jet ion guide focuses the ions into the second stage QJet Ion Guide
- Gas flow for Curtain Gas™ Interface is typically 18 L/min



Roughing Pump 1
MS 120

Roughing Pump 2
MS 40

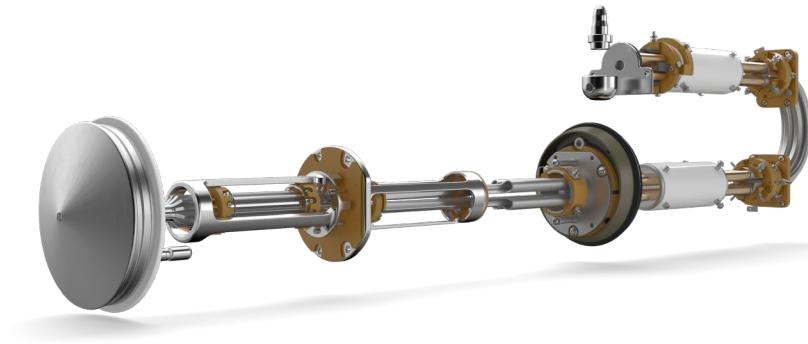
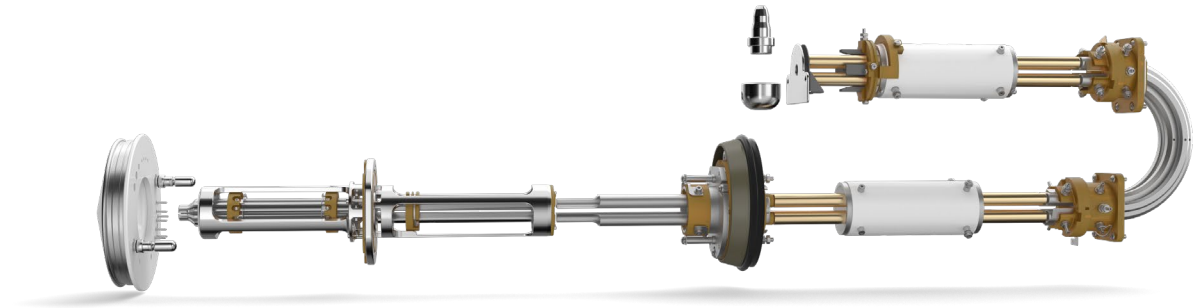
FUTURE PROOF YOUR LAB

- Allows users to future proof their lab's capabilities by providing an easy upgrade path to add linear ion trap scan features
- Uses LINAC electrodes on Q3 to enhance trap performance in LIT mode - in QQQ mode these electrodes are at same potential as the collar voltage and are therefore "invisible to the ions"



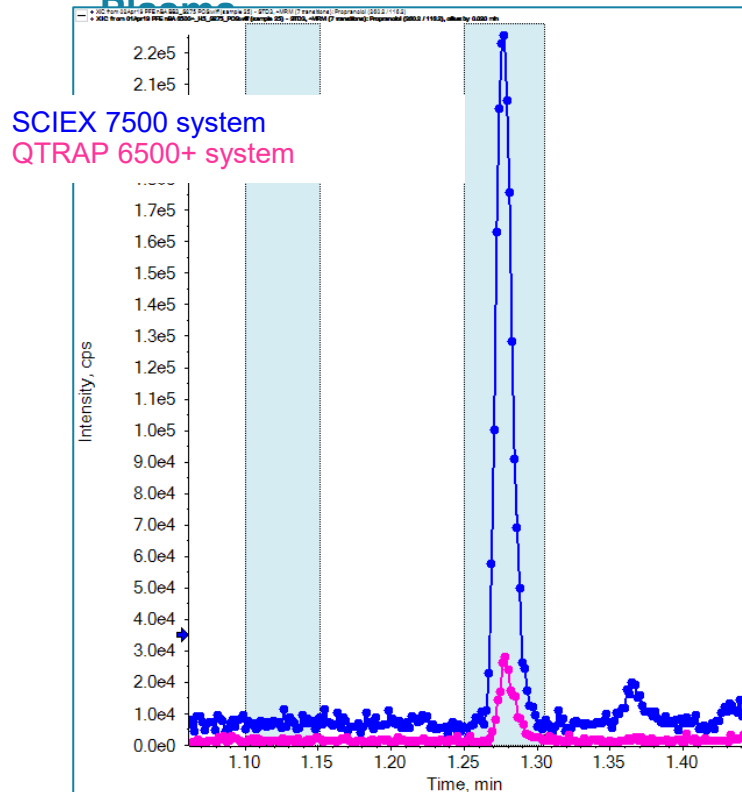
SCIEX 7500 SYSTEM FEATURES:

- Same rodsets as QTRAP 6500+ system, dual frequency RF drive with triple quadrupole mass range 5 – 2000 Da
- High Energy Detector system for fast polarity switching @ 5 msec and linear dynamic range up to 6 orders from LLOQ
- Curved LINAC collision cell for high speed analysis with no cross-talk
- Simplification: commonality with SCIEX platforms means no re-tuning of compound parameters CE and CXP and no DP tuning required
- QTRAP ready



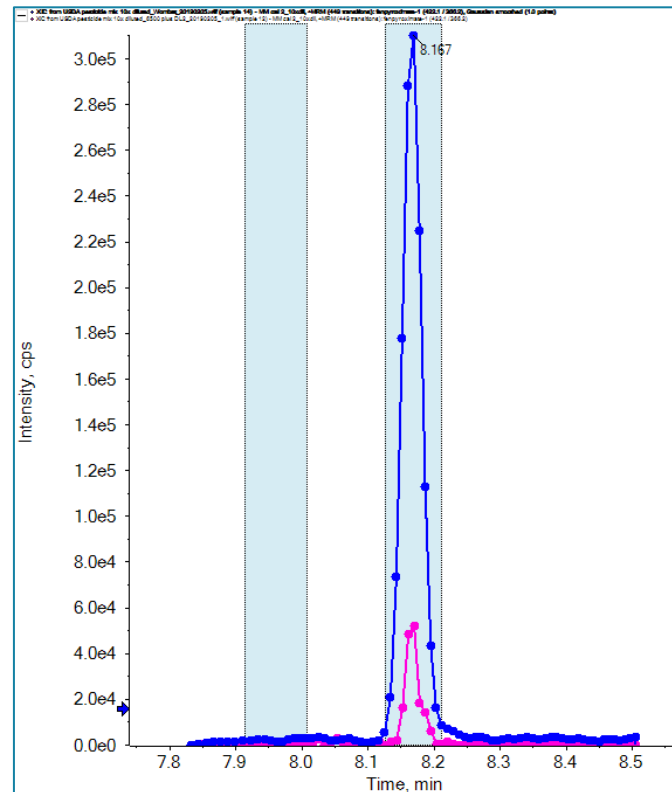
SENSITIVITY GAINS IN MATRIX SAMPLES >300 µL/MIN

Propranolol, 1 pg/mL, Rat



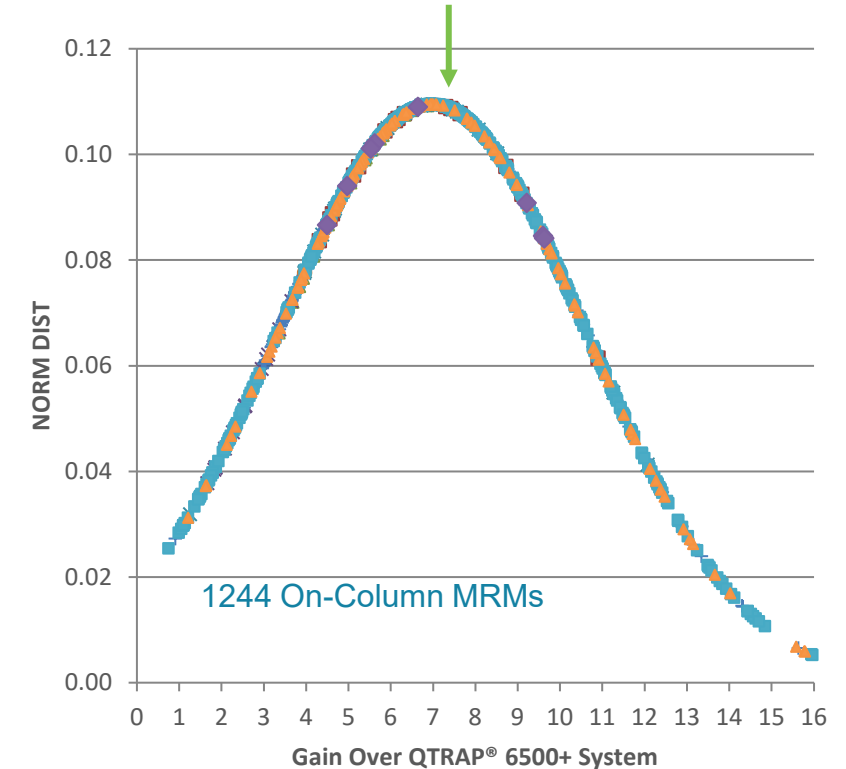
Area Gain - 9.1x
S/N Gain – 3.0x (RMS)

Fenpyroximate, Catfish



Area Gain – 8.1x
S/N Gain – 2.9x (RMS)

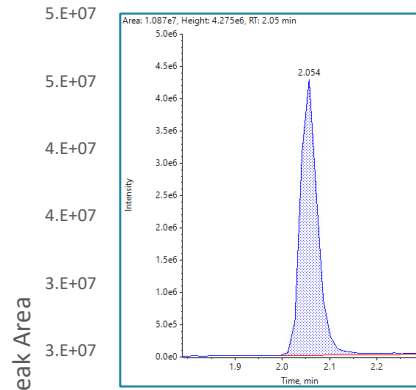
Average Peak Area Gain = 7.0x



1. Pesticides POS and NEG
2. Pharma small molecule POS and NEG
3. Vet drugs POS and NEG
4. Peptides

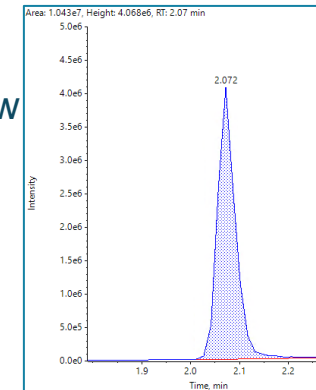


SCIEX 7500 system robustness - black tea matrix

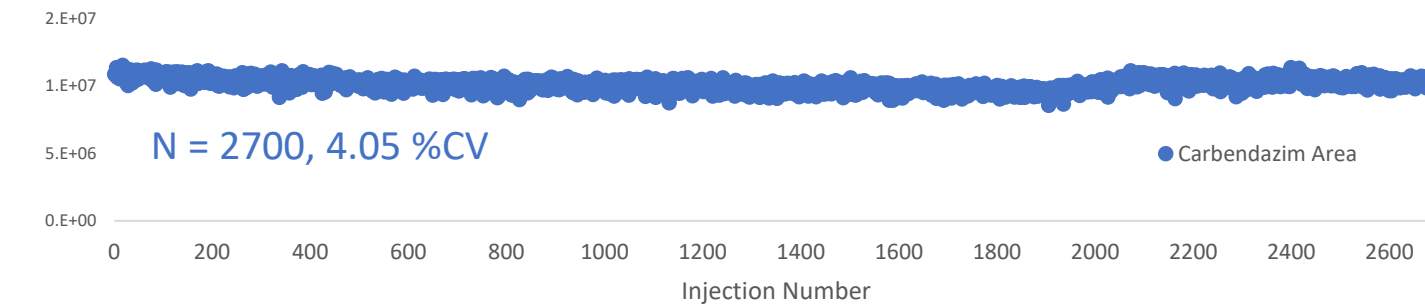


Injection 1

- Short LC-MS run was performed (4 min gradient at 400 μ L/min flow rate)
- Raw peak area plotted
- S/N remained constant across injections
- 1 ppb carbendazim



Injection 2700



QUALITATIVE FLEXIBILITY COMBINED WITH QUANTITATIVE POWER



- High sensitivity MS/MS with the ZenoTOF 7600 system
- ZenoTOF 7600 system combines the flexibility of multiple fragmentation options
- SCIEX OS software provides an intuitive workflow interface for easy acquisition and data processing

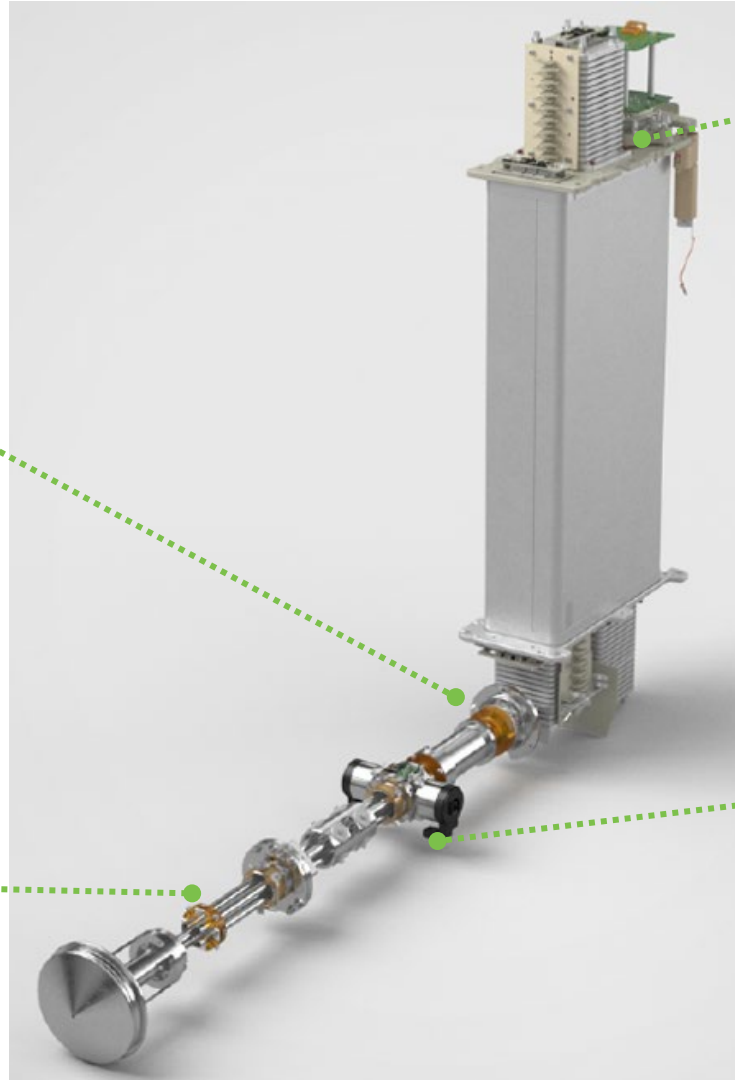
HARDWARE ADVANCEMENTS



Zeno trap
Improved MS/MS
duty cycle gain $\geq 90\%$



New Q0 design for
improved ion transmission
and maintenance



Wide dynamic range

- 5GHz, 10bit ADC with 40GHz TDC timing with 25 psec detection rate. High speed pulse counting to maintain resolution and mass accuracy $>130\text{Hz}$ and over 5 orders LDR



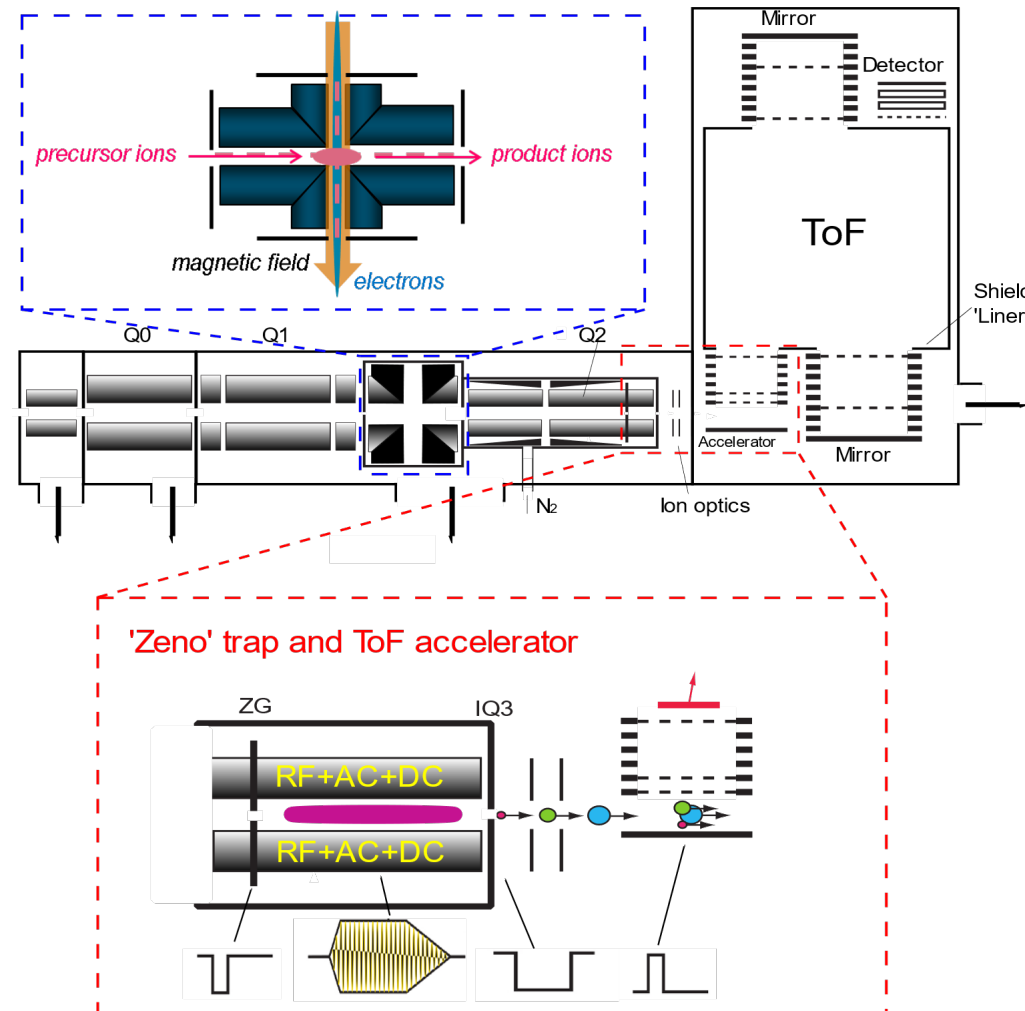
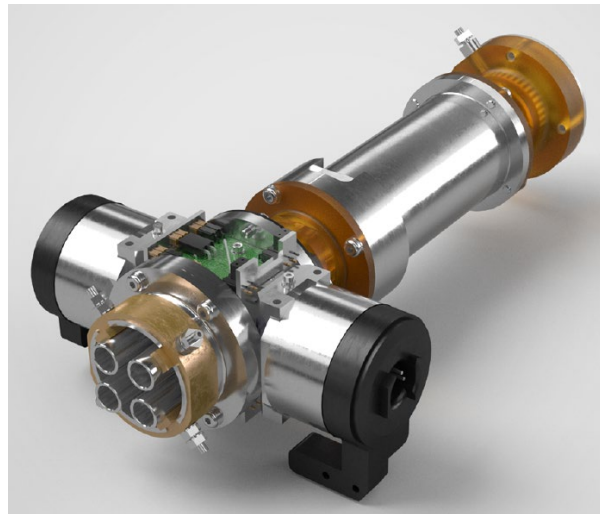
Complementary fragmentation with increased sensitivity using the EAD cell

Electron activated dissociation (EAD Cell)

ELECTRON ENERGY TUNABLE
FRAGMENTATION



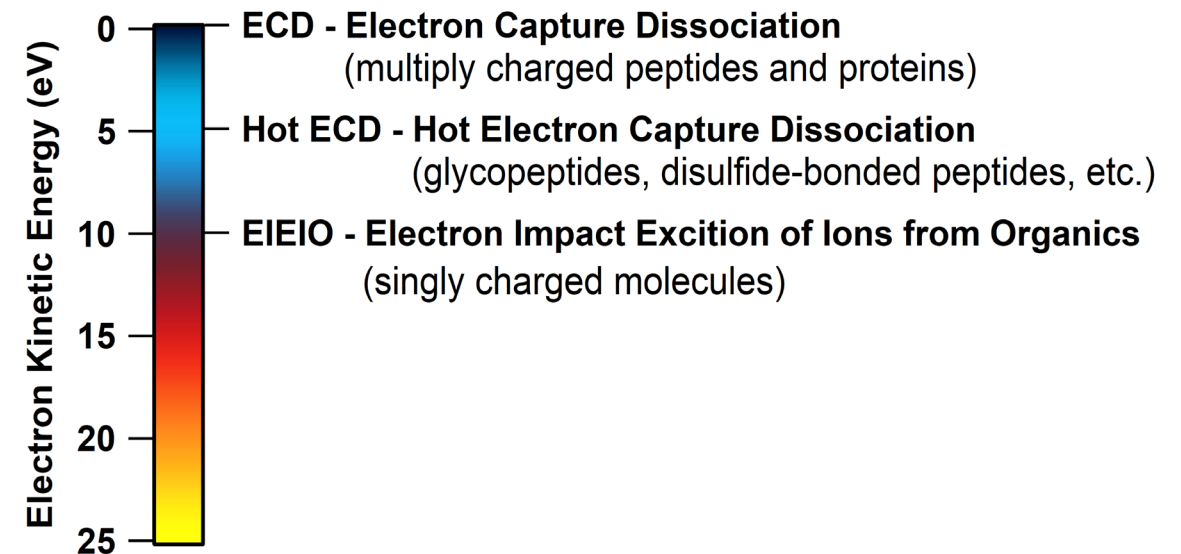
INTEGRATED MS/MS ASSEMBLY WITH ELECTRON ACTIVATED DISSOCIATION (EAD)



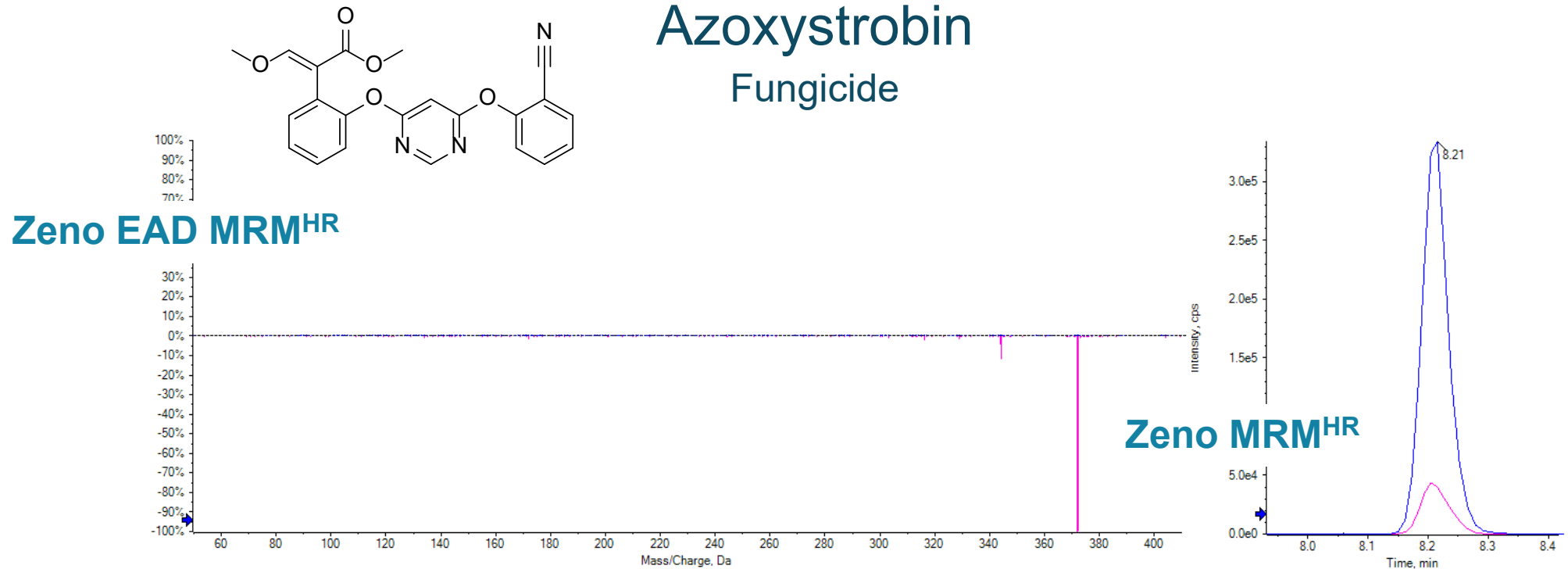
- EAD cell for electron based fragmentation
- Zeno trap for enhancement of low abundant fragment ions



- Free electrons are captured by ions and form a radical state which then fragments
 - Electrons introduced with different energies will induce fragmentation in different molecule types



IMPROVED SPECIFICITY FOR LIBRARY MATCHING AND ION RATIOS



- Zeno EAD MS/MS spectrum via MRM^{HR} contains >200 peaks
 - KE = 10 eV

- CID MS/MS spectrum typically contains only two peaks
 - CE = 35 V

Zeno trap



What is duty cycle?

... AND WHY IS DUTY CYCLE IMPORTANT?

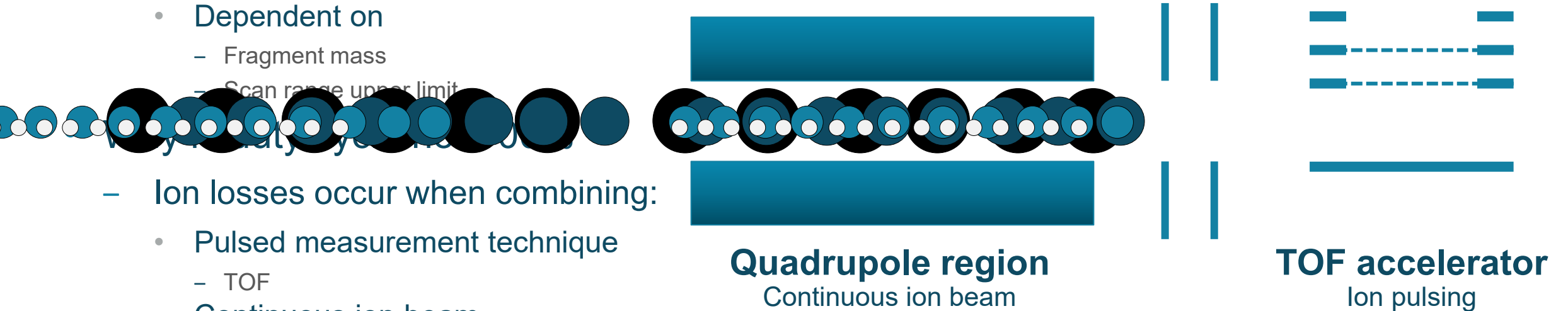
- What is duty cycle?

- % of ions injected into the TOF
- Typically, ~5-25%

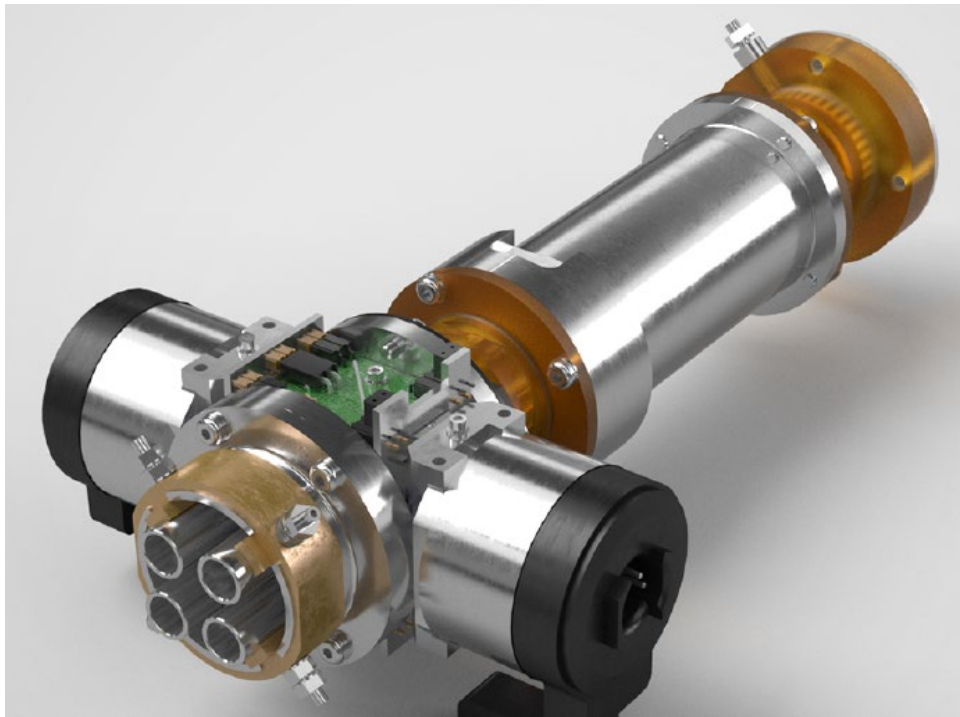
- Dependent on
 - Fragment mass
 - Scan range upper limit

- Ion losses occur when combining:

- Pulsed measurement technique
 - TOF
- Continuous ion beam
 - Quadrupole



ZENO TRAP DELIVERS FULL MASS RANGE TO ACCELERATOR ELECTRODE

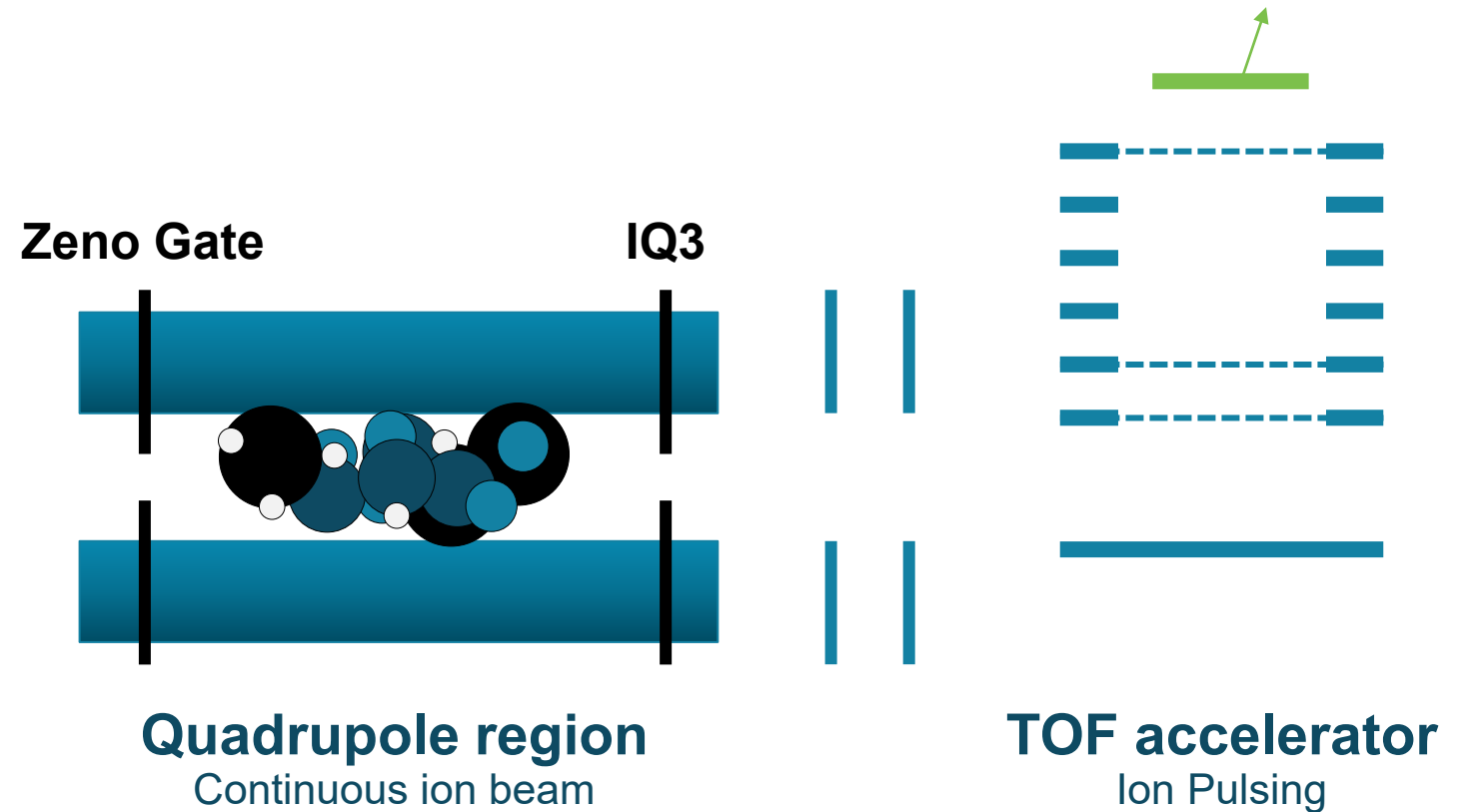


- The Zeno trap addresses the duty cycle problems with QTOFs
 - Improves MS/MS sensitivity across the entire mass range
 - $\geq 90\%$ of all ions injected into TOF region
- Improved sensitivity means
 - Lower LOQs
 - Lower sample loading
 - Improved MS/MS spectral quality
 - Improved structural elucidation / Met ID
 - Greater quality for low level metabolites
- Zeno trap capacity $\sim 7e^6$ ions

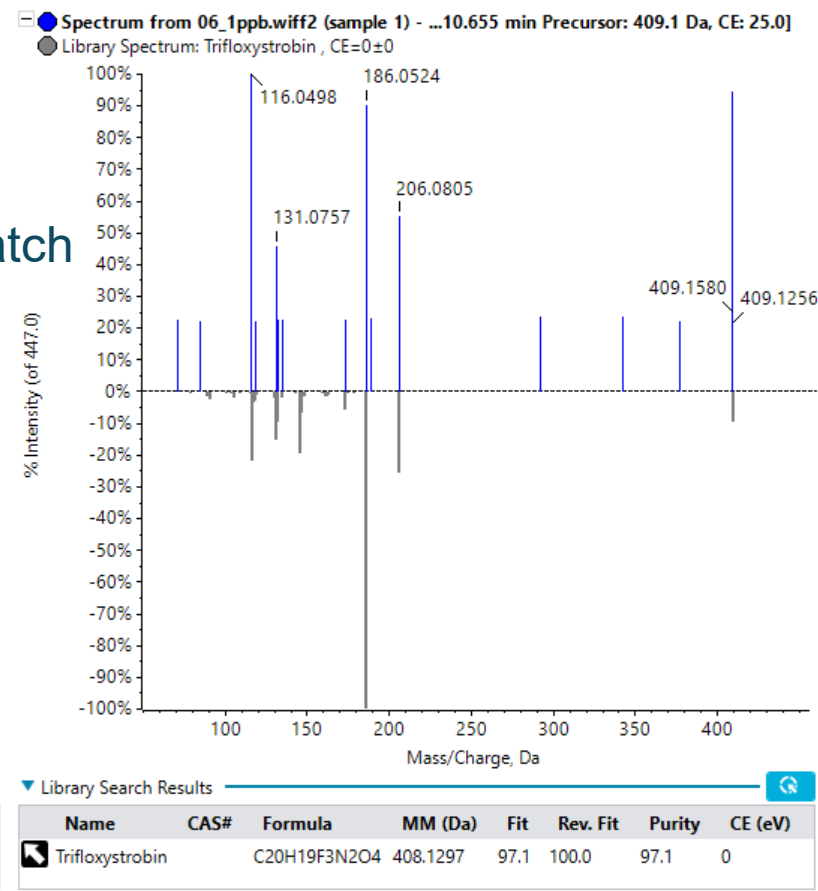
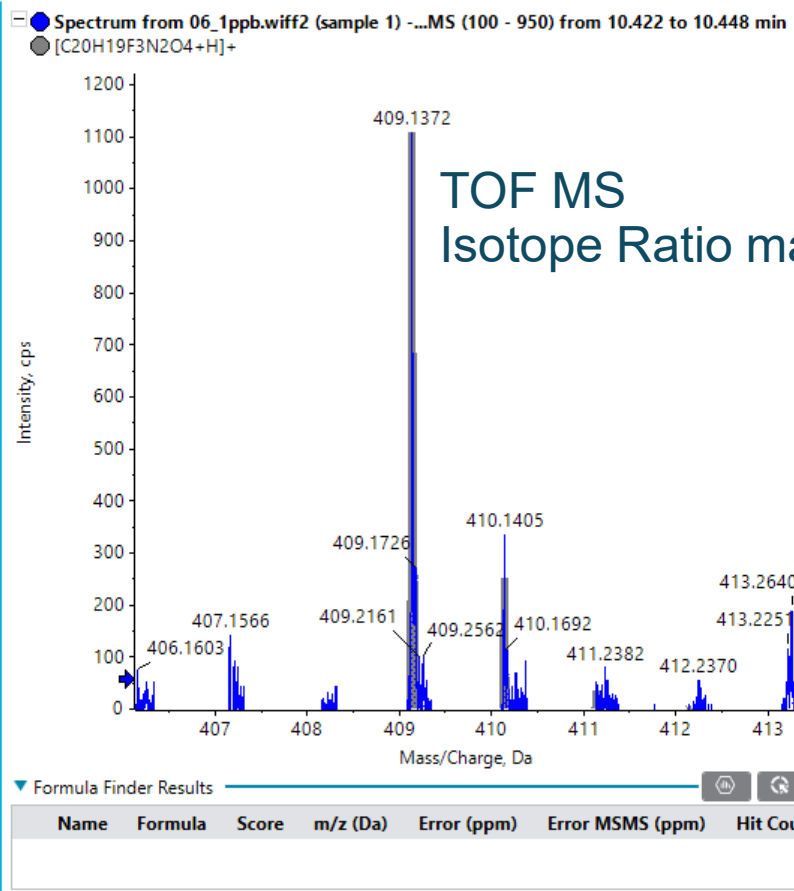
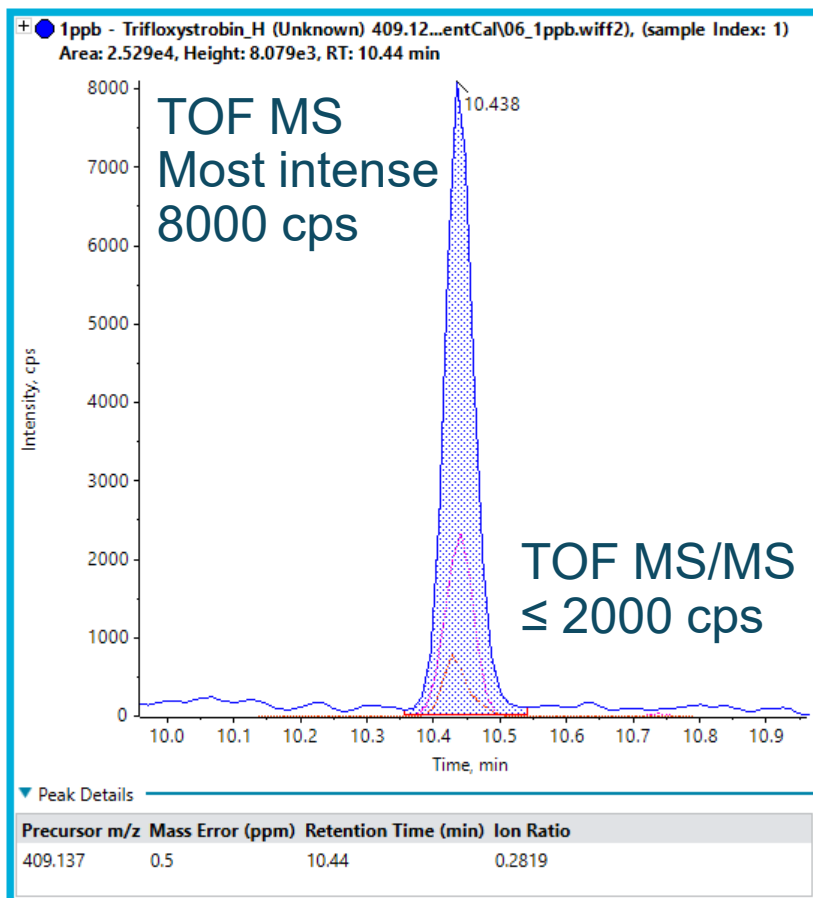
J. Am. Soc. Mass Spectrom. (2017) **28**: 2143-2150)

FOR SENSITIVITY GAINS IN MS/MS

- The Zeno trap provides control of the ion beam from the collision cell into the accelerator
- Ions are gated then released in a mass dependent manner
 - Higher m/z ions are released first then followed by lower m/z ions
 - All ions now arrive in the accelerator at the same time and location



MRM^{HR} – ZENO TRAP DEACTIVATED



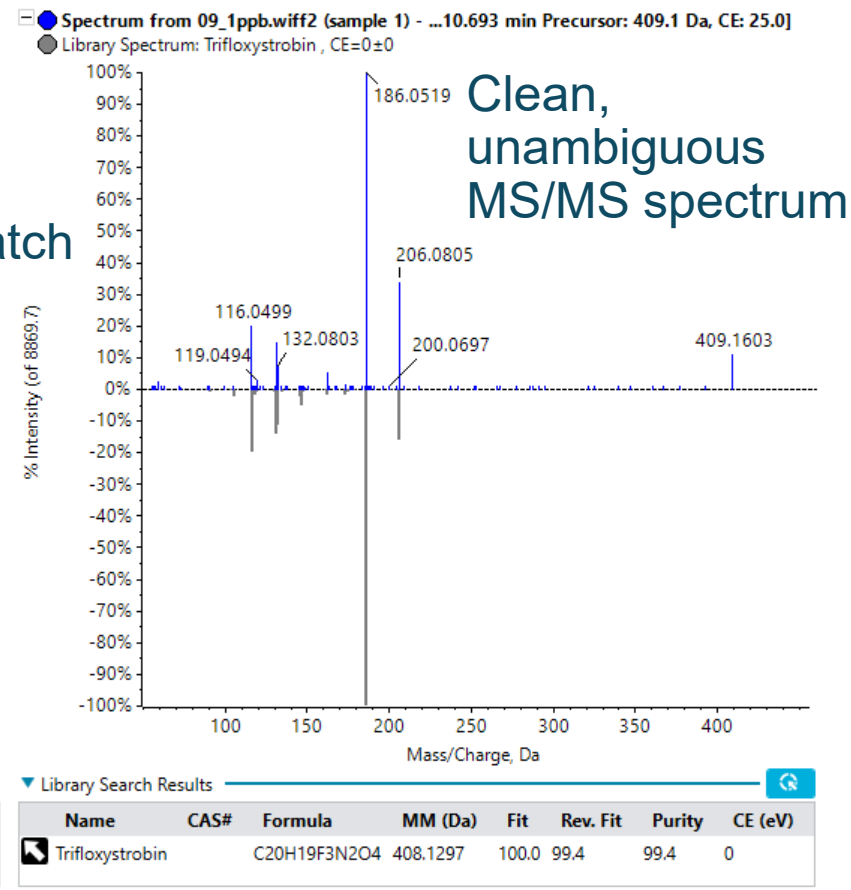
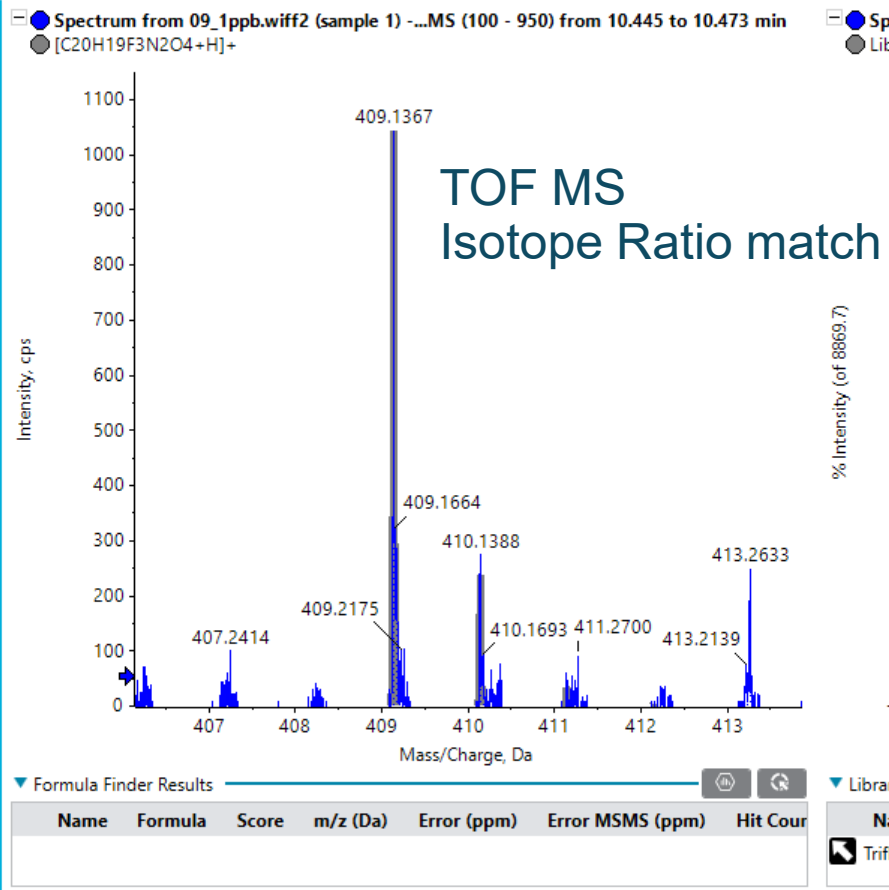
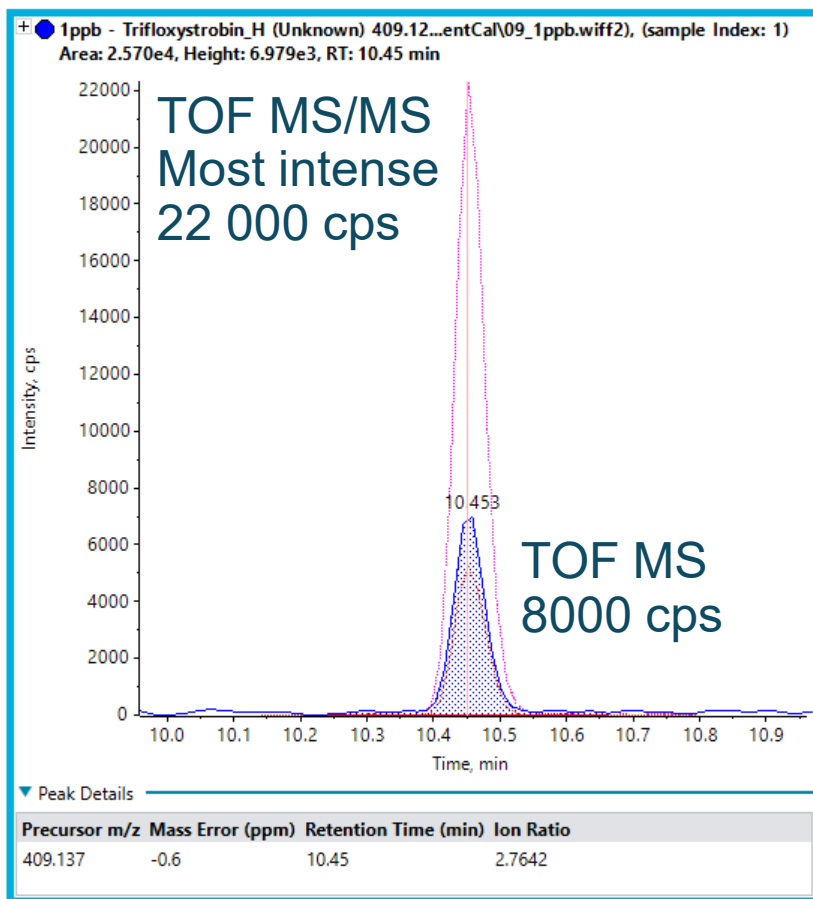
Mass error = 0.5 ppm

Library Score
Purity = 97%



The Power of Precision

MRM^{HR} – ZENO TRAP ACTIVATED



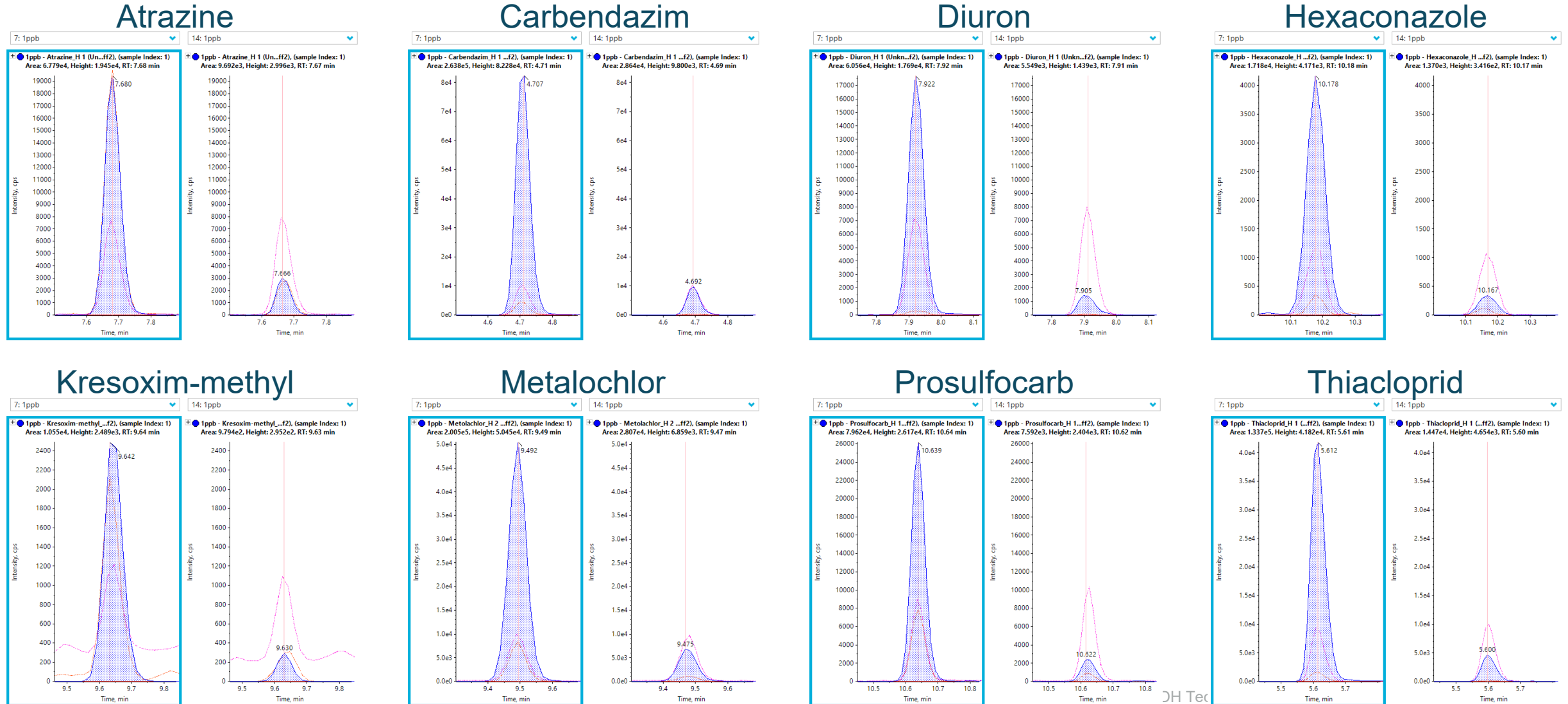
Mass error = -0.6 ppm

Library Score
 Purity = 99.4%

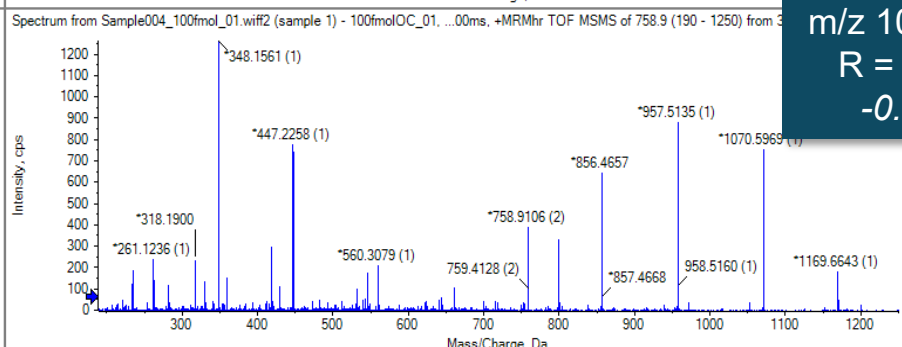
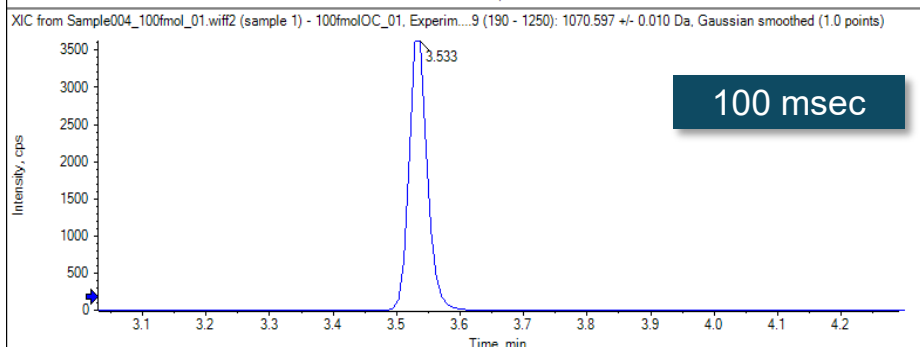
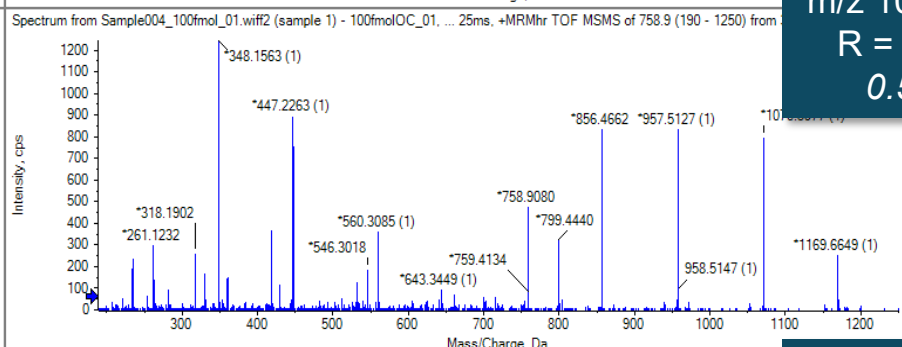
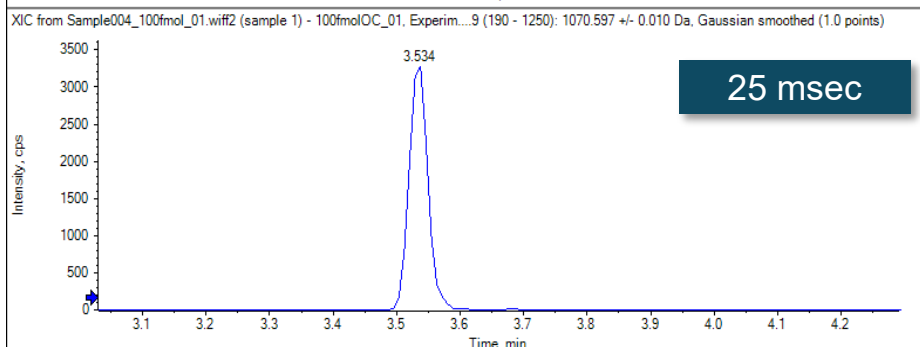
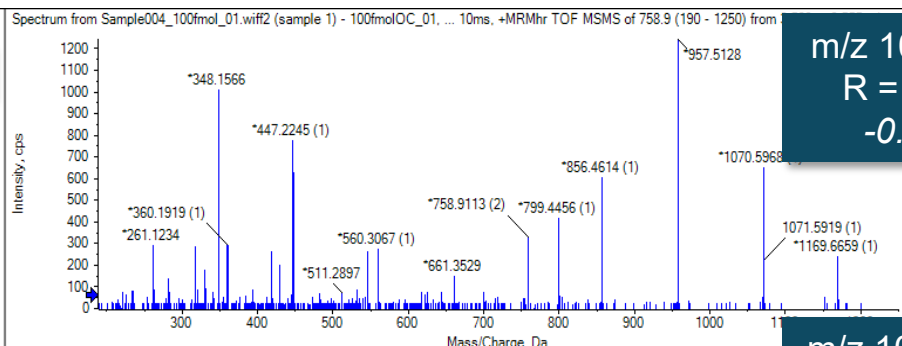
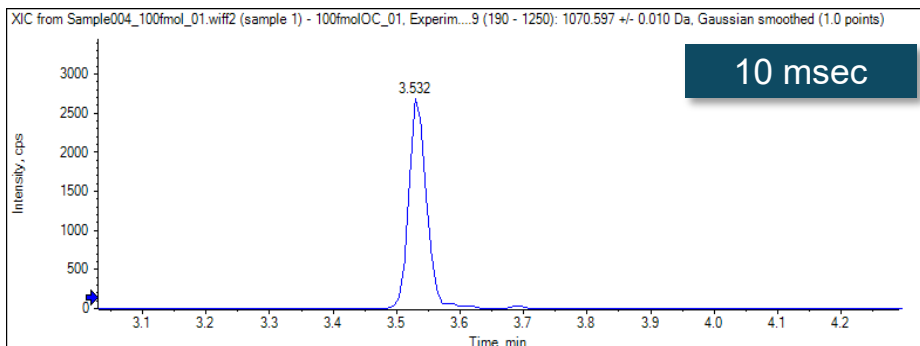


Gains depend on fragment m/z

1 PPB STANDARD: LEFT – ZENO TRAP ON, RIGHT – ZENO TRAP OFF (FRAGMENT XIC FILLED)



ACQUISITION RATES SUITABLE FOR HIGH MULTIPLEXING AND RAPID LC SEPARATION

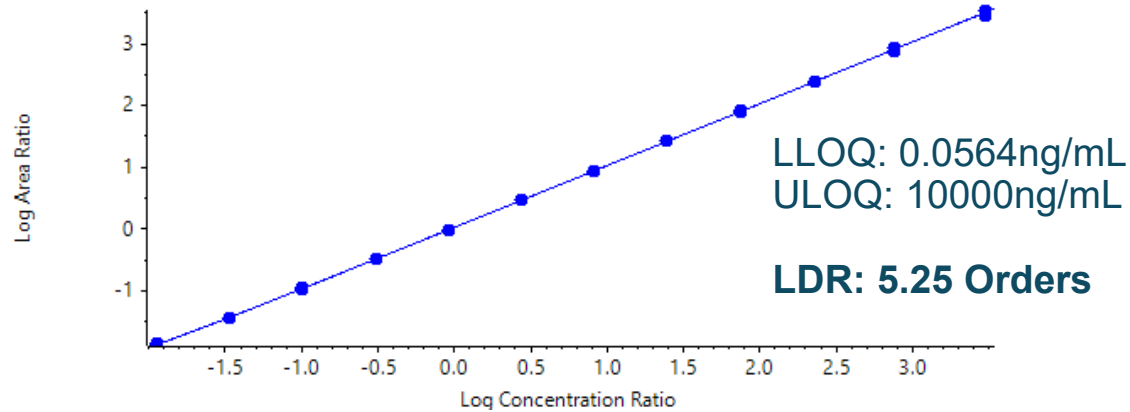


These data show how mass resolution and peak intensity is maintained when reducing the scan rate from 100 msec to 10 msec

The high-speed nature of the detector enables this very effectively

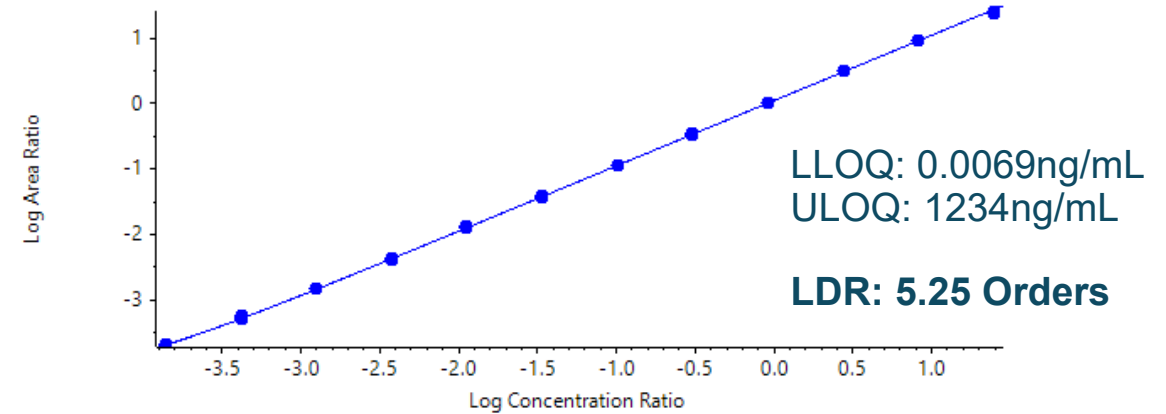
TOF MS LDR Positive Mode

Calibration for atorvastatin: $y = 1.06941x + 0.00158$ ($r = 0.99930$, $r^2 = 0.99859$) (weighting: $1/x^2$)



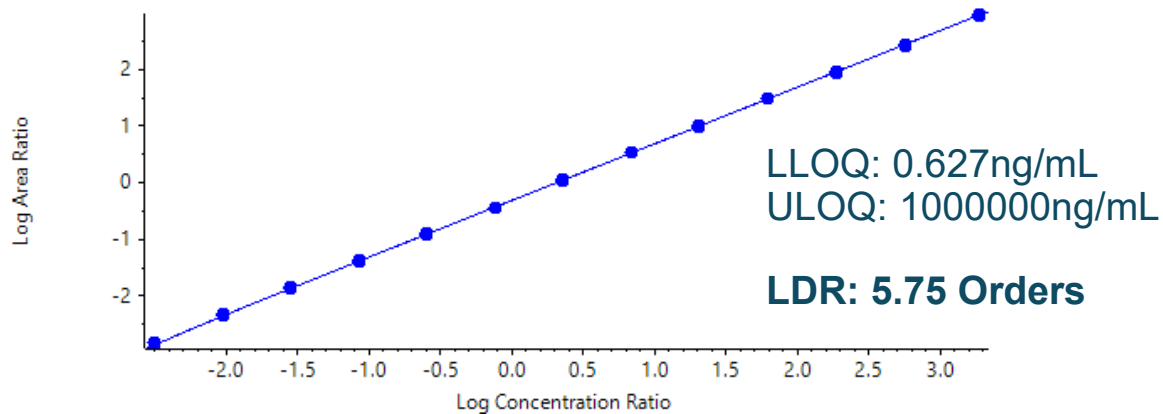
Zeno MRM^{HR} LDR Positive Mode

Calibration for Atorvastatin MSMS 004: $y = 1.11733x + \dots e-5$ ($r = 0.99904$, $r^2 = 0.99808$) (weighting: $1/x^2$)



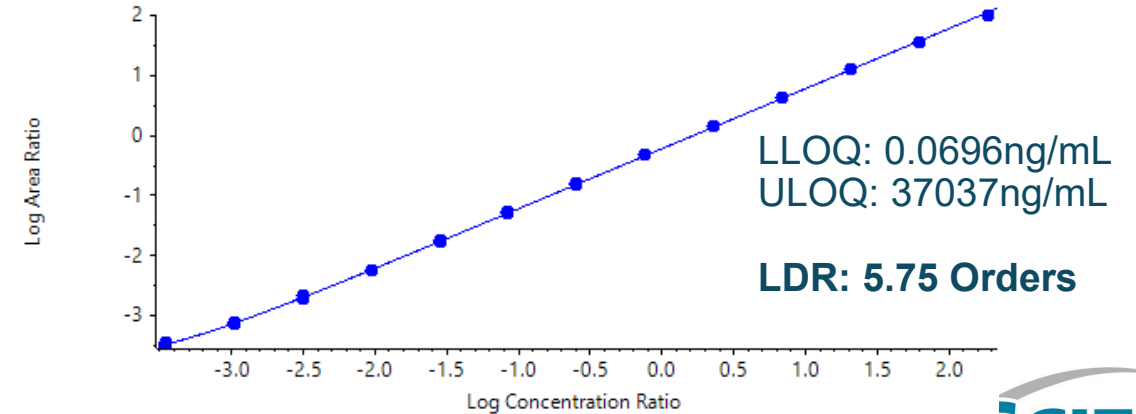
TOF MS LDR Negative Mode

Calibration for chloramphenicol: $y = 0.48623x + -1.75\dots 3e-4$ ($r = 0.99915$, $r^2 = 0.99831$) (weighting: $1/x^2$)



Zeno MRM^{HR} LDR Negative Mode

Calibration for chloramphenicol MSMS: $y = 0.60449x + \dots e-4$ ($r = 0.99775$, $r^2 = 0.99551$) (weighting: $1/x^2$)



LDR defined as $\log[\text{ULOQ} / \text{LLOQ}]$



Thank you!



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