

## A REVIEW OF MASS SPECTROMETRY: NEW TOOLS AND TECHNIQUES FOR ANALYSIS

**Amit Sukhdev Ghule, Om Ramesh Thorat, Samrat Dinesh Raut, Nikhil Rameshvar Kale**

Gajanan Maharaj Collage Of Pharmacy, Sambhajinagar.

**ghuleamit2003@gmail.com**

**Dr. Kavita Kulkarni** (phd. Mpharm), Department of Quality Assurance

Gajanan Maharaj Collage Of Pharmacy, Sambhajinagar.

### Abstract:

*Mass spectrometry is one of the best techniques for studying the structure of a molecule. It usually provides information about molecular weight of a substance, and it can present atomic mass units and up to ten thousandths of atomic mass units depending on the accuracy of the mass analyzer.*

*In addition, it provides information on the positive ions formed in the ionization process, which is linked to the chemical structure of the molecule and the nature of the bonds. This technique is widely used for analyzing compounds from natural products.*

*The development of the technique combined with the use of software and databases has been remarkable in recent years, improving the ionization processes and the ion analysis. Since natural products generally constitute a mixture of a complex quantity of components, mechanisms have been developed for coupling to chromatographic techniques of various kinds. It is also an analytical technique that identifies the chemical composition of a compound or sample based on the mass-to-charge ratio of charged particles*

*A mass spectrometer has three essential modules, an ion source-which transforms the molecules in a sample into ionized fragments, a mass analyser-which sorts the ions by their masses by applying electro- magnetic field and a detector-which measures the value of some indicator quantity and thus provides data for calculating the abundances each ion fragment present The technique has both qualitative and quantitative uses.*

**Key Words:** Ionization, Molecular Weight, Mass Analyzer, Mass-to-Charge Ratio ( $m/z$ ), Natural Products, techniques

**Introduction:** Mass spectrometry is an analytical technique whose purpose is discovering new molecules, determining quantities of known components and determining structural and chemical properties of a molecule. It involves the measurement of the mass-to-charge ratio of ions.

The definition of a mass spectrometer may seem simple: it is an instrument that can ionize a sample and measure the mass-to-charge ratio of the resulting ions. However, the versatility of this function has allowed it to become a vital tool in a wide range of fields, including biological research. This versatility arises from the fact that mass spectrometers can give qualitative and quantitative information on the elemental, isotopic, and molecular composition of organic and inorganic samples.

The technique consists of three main steps: 1) ionization, 2) mass analysis, 3) detection. During ionization, molecules are converted into ions, which can be done through various methods which are electron impact, electrospray ionization. These ions are then separated in a mass analyzer based on their mass-to-charge ratios. Finally, detector measures the abundance of these ions, generating a mass spectrum that represents the molecular composition of the sample.

**History of mass spectrometry:**

A few of the great people and major discoveries that have shaped this century-old technique.

As the applications of MS rapidly expand, so does the number of mass spectrometrists. Development of mass spectrometry was pushed first by a few dedicated promoters. in the early 20th century, the technique was used to measure masses of atoms, and one of its first contributions to science was to demonstrate the existence of isotopes; this discovery fueled the contemporaneous ongoing debates about the structure of the atom.

Many scientists in the U.K. and in Europe were working on this topic at the time, and one major question they were trying to answer was: what is the nature of cathode rays? Some researchers thought they were made of particles and some thought they were waves, but despite 50 years of research, no one yet had a definitive answer. For those who believed in the particle theory, the race was on to measure the mass of these unknown particles.

**Table 1. Other important 20th-century advances in MS instrumentation.**

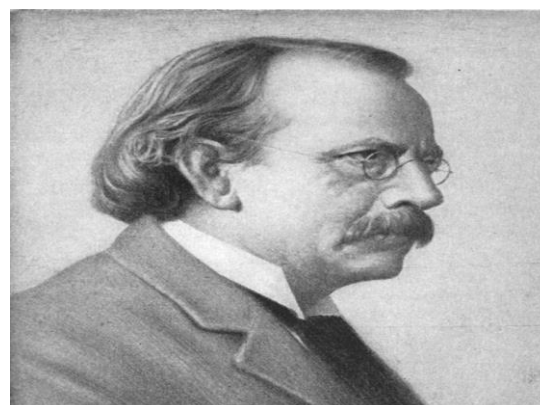
Instruments	Name of scientist	year
The first electron impact source (solids)	Dempster	1918
The first electron impact source (gases)	Bleakney	1929
TOF mass analyzer	Stephens	1946

(time of mass)		
Chemical ionization	Field	1966
Field desorption ionization	Beckey	1969
Triple-quadrupole mass analyzer	Yost and Enke	1978
FAB (fast atom bombardment)	Barber	1981

Scientist and their inventions:

- 1) Joseph John Thomson

Thomson first used his apparatus to measure  $e/m$ . (ratio of charge and mass). rather than the present MS standard of  $m/z$ ) of these fundamental particle electrons in 1897. Two years later, again with the assistance of Everett, he built an instrument that could simultaneously measure  $e/m$  and  $e$ , thus indirectly measuring the mass of the electron. For this work in “discovering” the electron, he received the 1906 Nobel Prize in Physics.



Thomson's early work on cathode rays laid the foundation of the MS field. Ultimately, Thomson with the help of his protege Francis Aston (who would go on to win his

own Nobel Prize in Chemistry in 1922), built what later would be recognized as the first mass spectrometer to measure the masses of charged atoms. This instrument used gas discharge tubes to generate ions, which were then passed through parallel electric and magnetic fields. The ions were deflected into parabolic trajectories and then detected on a photographic plate.

### Joseph John Thomson

#### 2) Alfred Nier:

Alfred Nier's journey into mass spectrometry is a great example of how adaptability and a mix of skills can lead to success. He began his career in electrical engineering, but due to a lack of relevant courses, he switched to physics for his graduate studies at the University of Minnesota. This change turned out to be a smart move; his engineering background worked well with his physics training, allowing him to thrive as an experimentalist.

Michael Grayson points out that Nier's unique combination of skills—his knowledge of electronics paired with his physics expertise—enabled him to make significant innovations in mass spectrometer design. This blend not only boosted his technical abilities but also had a lasting impact on the advancement of mass spectrometry, making Nier a key figure in the field. His story highlights the importance of drawing from multiple disciplines in scientific research and innovation.

Alfred Nier was known not just for creating some of the most advanced mass spectrometers of his time, but also for his efforts to promote mass spectrometry

beyond the realm of physicists. According to Dennis Schlutter, who worked closely with Nier at the University of Minnesota, he played a key role in making the technique more relevant and practical for other fields. Nier didn't just focus on building sophisticated instruments; he aimed to make them easier to use and applicable in various areas of research.

Nier's generosity was well-known among his peers. Mark Kurz from the Woods Hole Oceanographic Institution recalls how



selflessly Nier shared his machines and ideas with others. His willingness to help and his kind, gentlemanly nature earned him deep respect and loyalty from many in the scientific community. Nier's commitment to collaboration and his desire to advance mass spectrometry left a lasting mark on the field and on those who knew him.

Nier's impact was his work with biologists, where he prepared  $^{13}\text{C}$ -enriched carbon. He later reflected, "The material was of great

interest to biologists, who could use it for tracer studies. As a result, I gained many new friends.” His contributions didn't stop there; he also supported geochemists in determining the age of the Earth by measuring the  $^{207}\text{Pb}/^{206}\text{Pb}$  ratio in the planet's crust, among many other achievements. Nier's willingness to collaborate across disciplines helped bridge the gap between physics and other scientific fields, fostering valuable connections and advancing research.

### **Alfred Nier**

#### **MS GROWS ORGANICALLY:**

By the 1940s, mass spectrometers were available for commercial use, and mass spectrometry had become a valuable technique for physicists and industrial chemists. Carsten Reinhardt explains that when industrial chemists used mass spectrometry, they focused on quantitative measurements. This meant they used it to control production processes and determine how much of each component was in a mixture. Since they were already familiar with the identities and structures of most molecules, they primarily relied on mass spectrometry to measure concentrations rather than to identify substances.

From the mass spectrum to the molecular structure was a topic of research in industrial and government laboratories from the earliest Annual Conference on Mass Spectrometry and Allied Topics in 1954,” says Grayson. “Significant work was reported, both in the literature and at these conferences.”

#### **MASS SPECTROMETRY IN THE TWENTY FIRST CENTURY:**

Mass spectrometry (MS) is a vital tool in drug discovery because it is both sensitive and fast. It's especially good at identifying proteins that are often drug targets, even when they are present in low amounts in complex mixtures. MS also makes it easier to analyse large groups of drug candidates quickly, helping researchers measure concentrations and track chemical reactions. Mass spectrometers can be automated, which boosts their efficiency. In addition to drug discovery, MS can identify and track biomarkers in bodily fluids, helping to evaluate how effective and safe drugs are. It can also quickly assess how drugs bind to their targets, including the strength and specific sites of these interactions.

As MS becomes more widely used, some users may find it confusing, as they might not understand how it works. This review aims to explain current MS technology, helping those involved in drug discovery use it effectively and avoid misinterpreting results. For more specific examples of how MS is applied in drug discovery, readers are encouraged to look at other recent reviews.

It is also worth noting that the term "mass spectrometry" can be misleading. MS measures the mass-to-charge ( $m/z$ ) ratio of ions, not just mass alone. A mass spectrum shows how much of each ion is present against the  $m/z$  ratio, although the x-axis is often labeled "mass." For example, when benzene (with a molecular mass of 78 g/mol) is ionized, it produces an ion with an  $m/z$  of 78. The accuracy of this measurement can vary depending on the type of mass analyzer, but usually only the whole number is reported.

Basically, any information gathered from a mass spectrometer comes from the analysis

of gas-phase ions. There are three main components of a mass spectrometer which is ionization source, a mass analyser and a detector. The following discussion will provide an overview of the different ionization sources and mass analysers that are most often used in the analysis of biologically relevant samples. Although the raw data that mass spectrometers provide are  $m/z$  ratios and abundances of the components in a sample, the information that can be gathered from MS is not just limited to molecular weight and sample amount. Mass analysers can also be used in certain ways to gather structural information, such as the connectivity of atoms in a molecule. Structural analysis combined with its inherent sensitivity and speed is where MS realizes much of its power.

### **Mass Spectrometry in the Postgenomic Era:**

In Cellular systems are incredibly complex, made up of many different structures that interact in various ways. To understand these systems better, researchers need effective tools to identify all the different molecules involved and how they are organized. While we have made good progress in quickly sequencing entire genomes and analyzing mRNA in cells, finding effective methods to study other important molecules like proteins and lipids—and how they work together—has been more challenging.

Electrospray ionization (ESI) is a technique used to create charged molecules from a solution by applying a strong electric field at the tip of a tiny tube. This method is particularly useful for biological applications for several reasons:

1. Gentle Process: ESI can ionize delicate molecules without breaking them apart, which helps preserve their structure and even some weak interactions.
2. Easy Integration with Liquid Chromatography: ESI can be easily connected to liquid chromatography systems. This allows researchers to directly spray the separated components from a mixture into the mass spectrometer for analysis.
3. Production of Multiple Charges: ESI often generates ions with multiple charges. This is beneficial because it enables the measurement of larger biomolecules using mass spectrometers that might otherwise have limitations on the size of ions they can analyze.

Additionally, having multiple charges on a molecule can make it easier to break apart (fragment) for further structural analysis and identification.

Recent advancements in mass spectrometry (MS) have made it a preferred method for various biological applications, especially in protein analysis. There are two main approaches for analyzing proteins using MS: the bottom-up and the top-down methods.

1. Bottom-Up Approach: In this method, proteins are first broken down into smaller pieces called tryptic peptides using an enzyme like trypsin. These smaller peptides are then analyzed by MS and a more detailed technique known as MS/MS. The benefits of this approach include:

- Uniform Size: The smaller peptides are easier to work with and can be more uniformly handled than whole proteins.

- **Accurate Mass Measurement:** It's simpler to accurately measure the mass of these small peptides.
- **Easier Fragmentation:** Smaller peptides are more likely to break apart in a controlled way for further analysis.

Because of these advantages, the bottom-up approach is commonly used in most proteomic studies, even though it may not capture every detail of the original protein.

2. **Top-Down Approach:** This method involves analyzing the intact proteins directly without breaking them down first. It has the potential to provide a complete view of the protein and its modifications. However, this approach has its challenges:

- **Handling Whole Proteins:** Whole proteins can be more difficult to manage compared to smaller peptides.
- **Heterogeneity:** Proteins often vary in structure, which complicates the analysis.
- **Complexity:** Analyzing intact proteins can be technically challenging and is usually suited for studies focused on single proteins rather than large-scale analyses.

There is an emerging "middle-down" approach that analyses larger proteolytic peptides, which is proving useful for studying specific modifications, such as those found on histone tails. This method aims to combine some of the advantages of both the bottom-up and top-down approaches.

Identifying proteins in gel bands by MS also has been used to great effect for the definition of components of protein complexes, especially for stoichiometric complexes that have been relatively cleanly isolated. An alternative strategy for such

cases involves shotgun MS sequencing of the peptide mixture, generated by digesting the entire complex without prior separation of the proteins, and mapping these peptides onto the various proteins that are present using computer algorithms.

MS is now a preferred method for elucidating protein posttranslational modifications. Mass changes, characteristic of modifications, can be rapidly pinpointed to specific amino acids in the sequence using relatively low amounts of protein. Such applications have been greatly facilitated by improved MS instrumentation and fragmentation technology [using, e.g., so-called higher-energy collisional dissociation or electron transfer dissociation in addition to the more standard lower-energy collisional dissociation] as well as by the development of methods for enriching specifically modified proteolytic peptides, via either affinity isolation or chemical derivatization. For example, immobilized metal ion affinity chromatography (IMAC) has been used to enrich and facilitate identification of literally thousands of phosphorylation sites within whole proteomes as a function of cell state.

This difficulty arises because these studies are usually performed on proteolytic fragments, using the bottom-up approach, and for a number of reasons (IMAC isolation bias, peptides too small, too large, too hydrophilic), it can be very difficult to detect all of the peptide fragment ions (or even assign the peptides unambiguously to a particular splice variant). The resulting gaps in the coverage of the protein are further exacerbated when the amount of the available protein is limited or the

stoichiometry of the phosphorylated forms of the protein is low.

In such cases, it may be necessary to resort to the classical approach of subjecting the protein of interest to multiple digestions with proteases having different specificities, provided of course that sufficient protein is available. An alternative line of attack that is just beginning to gain traction relies on the top-down or middle-down approaches. Instead of analyzing small proteolytic fragments, one obtains MS and MS/MS of the whole protein or large peptide components of it. Under ideal conditions, the problem then reduces to assembling pairwise puzzle pieces (protein ion fragments) that each add up to the mass of the protein (or that of a relatively large component of the protein) rather than trying to assemble a great many small puzzle pieces, many of which may be missing. Although considerable progress has been made in this endeavor, the top and middledown approaches are still limited by difficulties relating to the isolation and handling of small amounts of protein as well as to introducing them into the mass spectrometer and fragmenting them efficiently.

### **Mass spectrometry and natural products:**

The high development achieved in the last hundred years by mass spectrometry with a great variability of techniques and instruments, makes possible that basically all molecules that are part of natural products may be analyzed, both qualitatively and quantitatively. The extracts coming from biological matrices with natural products are generally a mixture of various compounds, and thus it

is very common the use of GC or LC coupled systems.

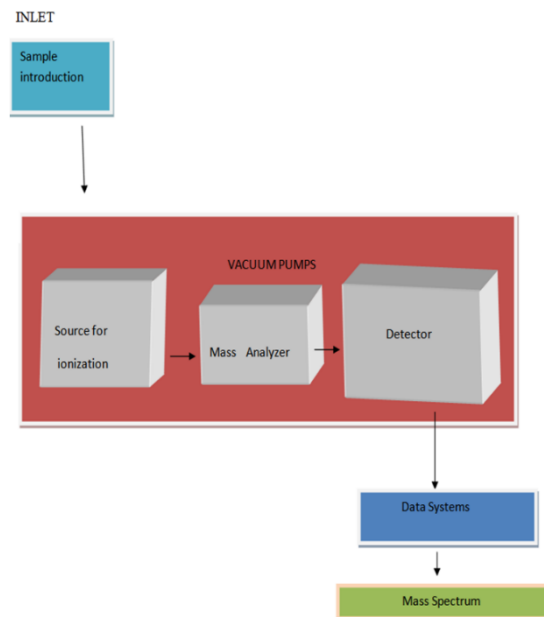
New techniques such as Electrospray Ionization, have increased the number of possible biomolecules to be analyzed, including those of high molecular weights. Similarly, the use of powerful mass analyzers makes it possible to analyze molecular ions or fractionated ions with an extremely efficient resolution.

- Essential oils
  - Fatty acids
  - Aromas and flavors
  - Phenols and polyphenols
  - Alkaloids
1. Essential oils: Essential oils are made up of volatile compounds, which makes studying their chemical composition relatively straightforward. Researchers often find that over 90% of the components in these natural products can be identified. By analyzing mass spectra and retention indices, scientists can gather information that can be cross-referenced with specific databases dedicated to these compounds.
  2. Fatty acids: Many plant species contain both saturated and unsaturated fatty acids. To study these compounds, researchers typically use gas chromatography coupled with mass spectrometry (GC/MS). Since high molecular weight compounds are non-volatile and can't be analyzed directly, scientists use chemical derivatization techniques to convert them into more easily analyzed forms, like methyl or ethyl esters. This method is commonly applied to various plants of nutritional and pharmaceutical interest, such as *Plukenetia volubilis*, *Borage officinalis*, and fish oil.
  3. Aromas and flavors: The aromas and flavors in fruits and vegetables are essential for identifying their unique taste

characteristics. Because these compounds are volatile, gas chromatography coupled with mass spectrometry (GC/MS) is an ideal method for studying their chemistry. Many of these volatile molecules cannot be captured in vapor form, so they are typically extracted using non-polar solvents before being analyzed in the chromatographic system. Alternatively, they can be directly injected using headspace introduction systems, which allow for the analysis of the volatile compounds present in the air above the sample.

4. Phenols and polyphenols: Only a few low molecular weight phenolic compounds can be directly analyzed using a GC/MS system. Most phenolic and polyphenolic compounds are non-volatile, so their structures need to be determined through different analytical methods. One common approach is chemical derivatization, which involves a process called silylation. This method modifies the compounds to make them volatile, allowing for their analysis using gas chromatography and mass spectrometry.
5. Alkaloids: The alkaloids are active ingredients whose structural feature is to have nitrogen in their structure, many of these molecules have a significant biological activity. Some alkaloids may be directly analyzed in GC/MS equipment, such as nicotine and other present in tobacco, caffeine and xanthine alkaloids and others whose volatility enables its separation in gas chromatography such as tropane alkaloids.

### components of a Mass Spectrometer



Mass spectrometer consists of three parts including

- 1) Inlet
- 2) mass analyzer
- 3) detector

### Schematic representations of parts of a mass spectrometer

#### 1. Inlet:

The inlet system is where samples are introduced before they are analyzed in the mass spectrometer. The choice of inlet depends on the type of sample being analyzed. In this system, sample molecules are ionized, and the ionization process is most effective when the sample is in a gaseous state. Therefore, the inlet is designed to convert various samples into gas by heating them, and then it exposes these samples to high vacuum pressures.

The inlet operates quickly and can detect even minor changes in the composition of the gas sample molecules. Samples that are thermally stable and in the gas phase are introduced into the source region through a needle valve. Mass spectrometers are often connected to chromatographic instruments,

which help purify the samples before they reach the mass spectrometer's source. This connection also facilitates the ionization process of the samples. There are several types of inlets available for introducing samples, each suited for different kinds of analyses.

**1.1.1 Direct Vapor Inlet:** the sample is in the gas state and is directly introduced into the source with the help of a needle valve and the air that is present in the sample is ejected out. This is the simplest sample introduction method.

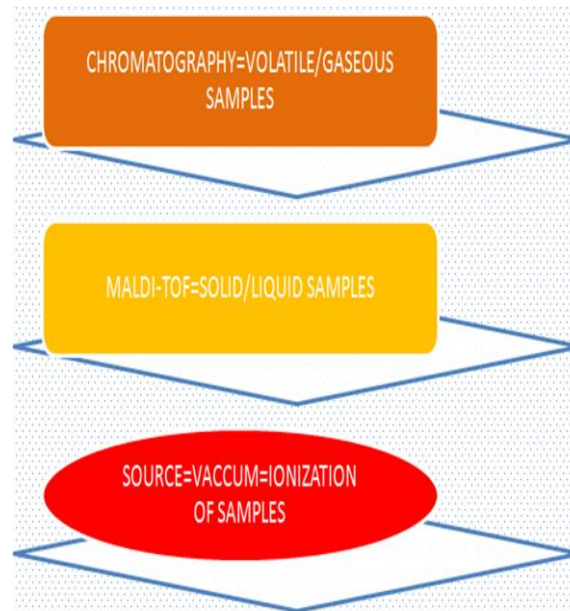
**1.1.2 Direct Insertion Probe:** Here the liquid or solid type of samples with low vapor pressure are introduced. The sample is loaded into a capillary tube which reaches the region with a high temperature that causes an increase in its vapor pressure.

**1.1.3 Gas Chromatography:** This technique is commonly used to separate complex mixtures before they are introduced into the mass spectrometer. Gas columns play a key role in controlling the flow of gas, which helps separate the different components of the sample. Once separated, these components can then be introduced into the mass spectrometer's source for analysis.

**1.1.4 Liquid Chromatography:** Thermally unstable compounds and those that don't separate well using gas chromatography are introduced into the mass spectrometer using liquid chromatography instead. This method is more suitable for handling such compounds, allowing for effective analysis without compromising their stability.

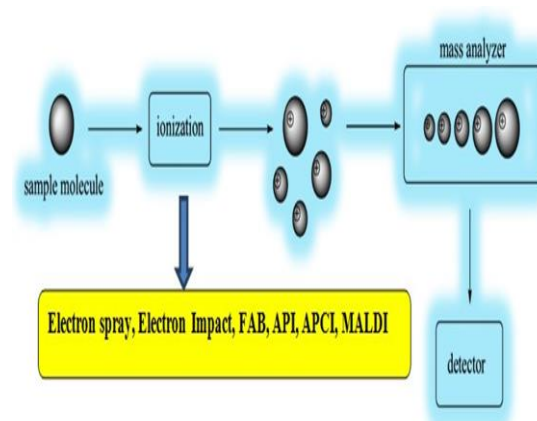
**1.1.5 Direct Ionization of Sample:** For samples that decompose quickly or have low vapor pressure, they can be directly ionized from

their liquid form instead of needing to be vaporized. This method is often used in combination with liquid chromatography, allowing for effective analysis of these types of samples.



**Diagrammatic representation of sample inlet for different types of samples**

**1.2 Ionization Methods:** Once samples enter through the inlet, they undergo a process called ionization before reaching the mass analyzer. The choice of ionization method depends on the type of sample being analyzed. During ionization, samples are broken down into either cations, anions, or adducts.



Different ionization techniques are used for various types of samples, and the energy applied during ionization plays a significant role in how the ions fragment. This

fragmentation can be observed in the resulting mass spectrum. Ionization can be classified as either soft or hard, depending on the amount of energy used. Soft ionization requires less energy and is often an incomplete process, while hard ionization uses more energy and typically results in more extensive fragmentation. The various ionization techniques will be discussed in the following sections.

### **Diagrammatic representation of Ionization methods of mass spectrometers**

#### **1.2.1. Electrons Spray Ionization Method:**

This is a commonly used hard ionization method for liquid samples. In this process, liquid samples are subjected to high voltage, atmospheric pressure, and a continuous flow of gas, which breaks them down into tiny droplets. These droplets are then heated under vacuum pressure, and the gas flow helps eject ions from the droplets. The ejected ions are accelerated toward the mass analyzer.

This basic principle has been refined to enhance sensitivity and is now applied in various techniques, such as nanospray, picospray, static nanospray, and desorption electrospray ionization.

#### **1.2.2. Electron Impact Ionization Method:**

This ionization method is used for gaseous samples and easily interfaces with gas chromatography. Once the gas sample is introduced into the inlet, it is bombarded with an electron beam generated from a filament. This process produces ions that are electrically neutral (known as radical ions), which then move on to the mass analyzer. This is a hard ionization technique, meaning that the ions are completely fragmented during the process.

**1.2.3. Fast Atom Bombardment (FAB):** This method is used for low-volatility and large compounds, such as carbohydrates, proteins, and peptides, whether they are in solid or liquid form. The samples are mixed with non-volatile matrix compounds like glycerol, ether, or nitrobenzyl alcohol, and then bombarded with

atoms like argon or xenon. This process ejects charged ions from the samples, which are then captured by the mass analyzer as adducts.

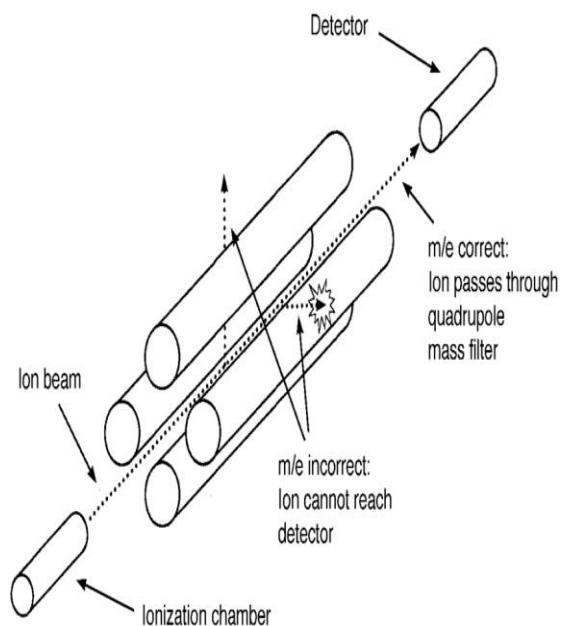
#### **1.2.4. Atmospheric Pressure Ionization (API):**

This method is useful for ionizing a wide variety of liquid compounds with different flow rates. In this process, ionization or fragmentation occurs partially outside the vacuum region, so the samples need to be analyzed further with a mass spectrometer. This ionization technique produces highly charged ions, making it ideal for coupling with liquid chromatography or mass spectrometry for detailed analysis.

## **2. Mass Analyzer:**

The mass analyzer is also described as an ion collector and is an important component of the mass spectrometer. The mass analyzer collects the ions formed in the ion source and separates them based on their charge to mass ratio and sends them to a detector where these separated ions are converted into a digital output

**2.1. Quadrupole Mass Analyzer:** The quadrupole mass analyzer is a popular and cost-effective tool used to analyze ions based on their mass-to-charge ratios. It works by creating oscillating electric fields that help separate ions, making it particularly effective for ions produced through electrospray ionization, which typically have mass-to-charge ratios under 300. One of the great things about the quadrupole mass analyzer is that it can be easily connected to gas and liquid chromatography systems, allowing for efficient analysis of complex mixtures.



It contains four parallel metal rods with specific diameter-to-spacing and is connected electrically with a radio frequency voltage. The ions are made to pass through these rods and the ion which has a specific mass to charge ratio at a specified voltage reaches the detector. The ions with varied mass to charge ratios collide with the walls of the rods and are removed by the vacuum system. Thus, ions with a particular mass to charge ratio and at a specific voltage can be isolated easily by this type of mass analyzer.

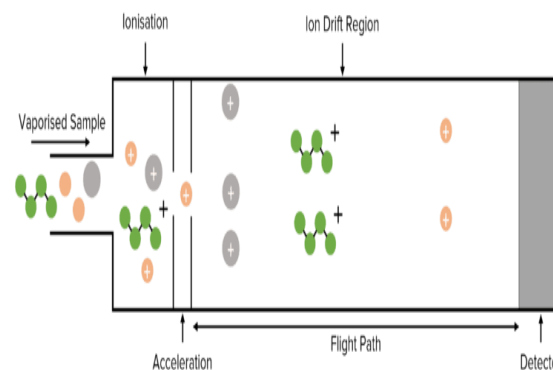
### Working apparatus of a quadrupole mass analyzer

**2.2. Time-Of-Flight (TOF) Mass Analyzer:** In a time-of-flight (TOF) mass analyzer, ions are generated by hitting the sample with short bursts of electrons or photons from a laser. Once created, these ions are separated based on how long it takes them to travel to the detector. Unlike some other types of mass analyzers, TOF doesn't use a magnetic field. Instead, ions with higher mass-to-charge ( $m/z$ ) ratios take longer to reach the detector because they move more slowly.

This technique is straightforward and works well with ions formed on surfaces, such as in laser desorption and plasma desorption mass spectrometry. The pulsed nature of TOF allows

for precise timing of ionization and helps define the area where ions are generated.

To improve resolution, TOF analyzers often use devices called reflectrons, which act like mirrors for ions. High-energy ions bounce off the reflectron and take longer to reach the detector, while lower-energy ions travel more quickly. For accurate detection, an electron multiplier detector is used, providing high resolution and sensitivity in measuring the ions.



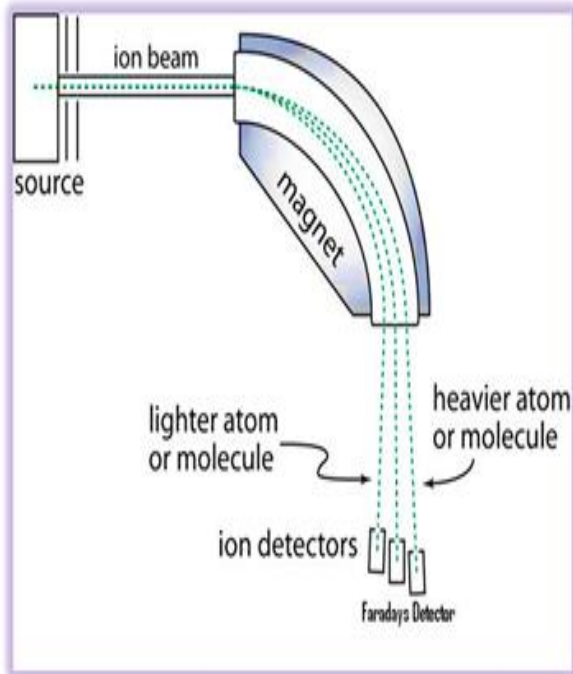
### Apparatus of a Time-Of-Flight (TOF) mass analyzer

**2.3. Magnetic Sector Mass Analyzers:** A magnetic sector mass analyzer uses both a static electric field and a magnetic field to influence the movement of charged ions. In this setup, ions with higher velocities, more charge, and lower mass are directed towards the detector. Essentially, the electric and magnetic fields work together to bend the path of the ions, allowing for their separation based on these characteristics. As a result, this type of analyzer can effectively identify and measure different ions based on their mass-to-charge ratios.

A magnetic sector mass analyzer features a curved, horseshoe-shaped tube where sample ions in vapor form are passed through a magnetic field. When the sample is bombarded with a positively charged electric field, it knocks ions out of the source, allowing them to move through the analyzer.

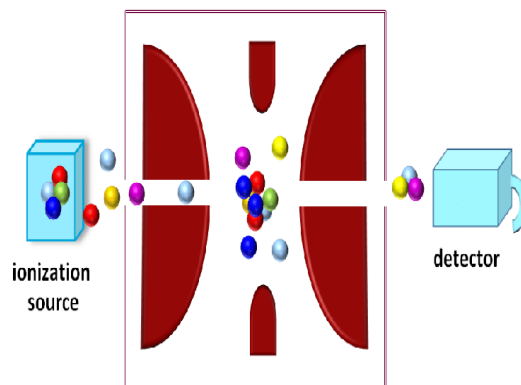
As these charged ions enter the magnetic field, they are deflected into circular paths. The ions

that are fast, light, and highly charged are directed towards the detector. Before reaching the detector, these ions follow a straight path, thanks to accelerating plates that keep their speed constant.



### Working model of a magnetic sector mass analyzer

**2.4 Ion Trap Mass Analyzers:** An ion trap mass analyzer is a device that captures ions using changing electric fields. It's also known as a radiofrequency (RF) trap or Paul trap, named after the scientist who created it. These analyzers use both electric and magnetic fields to hold ions inside. Unlike other mass analyzers, which often use a steady direct current, ion traps rely on oscillating radio frequency electric fields and a special arrangement of rods. In a 3D ion trap, there are parallel rods with electrodes, and a ring electrode is positioned between two end cap electrodes. This setup allows ions to move in a circular path due to the electric fields applied. In contrast, a linear ion trap doesn't have the circular or ring electrodes, resulting in a different way of trapping ions.



### The working apparatus of an ion trap mass analyzer

#### 3. Detector:

In a mass analyzer, the detector picks up signals from ions that come in. It does this either by creating secondary electrons that get amplified or by generating a current from the moving charged ions (known as ionization products). One of the oldest types of detectors is the photographic plate. At the end of the mass analyzer, when ions hit the photographic plate, they create spots based on their mass-to-charge ratio. The darker the spot, the more intense that particular ion is. So, by looking at these spots, scientists can determine which ions are present and in what amounts.

The choice of a detector is based on the required sensitivity, speed, and other application-specific requirements like heat stability, and available space, among others.

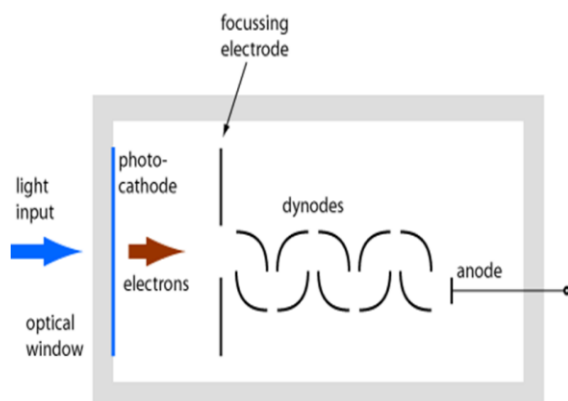
#### 3.1. Electron Multiplier Detectors:

The electron multiplier detectors are used for ions with less than 10-15 amperes. These detectors analyze or detect secondary electrons produced after the primary ion particles strike the detector surface. The signal generated is solely dependent on the intensity of the strike. The electron multiplier detectors may use a discrete dynode electron multiplier or a continuous dynode electron multiplier

### 3.2. Photomultiplier Tube (PMT):

Photomultiplier tubes (PMTs) are very sensitive detectors that can pick up light across the visible spectrum, as well as ultraviolet and near-infrared light. They work by significantly amplifying the electrical current generated by incoming light, boosting it by up to a million times.

Inside a PMT, there are several electrodes called dynodes, arranged in a vacuum. When light hits the detector, it causes electrons to be released. These electrons then strike the first dynode, which produces even more electrons through a process called secondary emission. This multiplication process continues from one dynode to the next, creating a large current by the time it reaches the final dynode. This allows PMTs to detect even very faint light signals effectively.



### Working apparatus of a photographic plate detector

#### 4. Data System:

After the ions are created through ionization, they are sent to a detector that is connected to a computer. The computer processes the information from the detector and presents it as a mass spectrum. This spectrum shows the different ions that were detected, along with their quantities,

allowing scientists to analyze the composition of the sample.

#### 5. Mass spectrum:

The mass spectrum is a visual representation that shows how ions are distributed by mass in a sample. It's the most common way to display data from a mass spectrometer. The mass spectrometer includes a histogram that provides a graphical approximation of this data, either in numerical values or categories.

Different samples create different patterns in their mass spectra, primarily due to the various ionization methods used to break down the samples. For example, straight-chain alkanes and alkyl groups will show different peaks in their mass spectra.

In the graph of the mass spectrum, the x-axis represents the mass-to-charge ratio ( $m/z$ ), while the y-axis shows the intensity of the detected signal. To identify unknown samples, this graph is compared to a standard curve made from samples with known patterns. This comparison helps determine the identity of the unknown sample detected by the spectrometer.

### Application of mass spectroscopy

- Proteomics - it is the large scale study of proteins.
- Genomics - studying all DNA of an organism.
- Monitoring volatile anesthetics in patients breathe during surgery by using end-tidal gas analyzer.
- Determination of pesticide residue in food.
- Identification of environmental pollutant.

### Conclusion:

Mass spectrometry (MS) is a versatile analytical technique used to measure

known chemical compounds and identify unknown substances in both biological and non-biological samples. Modern mass spectrometers are now portable and compact, allowing for their application in various fields, including biomedicine and biotechnology. They are instrumental in identifying proteins, estimating molecular size and weight, studying protein-protein interactions, and analyzing amino acid sequences. At the genetic level, MS verifies nucleic acids and DNA sequences, helping to understand genetic variations and changes influenced by environmental factors.

In agriculture, MS can detect pesticide residues in food, which is essential for preventing excessive pesticide use in developing countries like India. Additionally, MS monitors air quality by checking particulate content, alerting local governments to pollution levels that affect public health. The technique is also valuable in forensic science for measuring toxic substances and drugs in bodily fluids, as well as assessing gases in human breath during medical emergencies and surgeries. Geologists utilize MS to date fossils and rocks, showcasing its wide-ranging applications in research and healthcare.

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