



International Journal of Research Publication and Reviews

Journal homepage: www.ijrpr.com ISSN 2582-7421

Green Analytical chemistry

Dr.E Naga Deepthi.¹, Mohammed Aseef²

¹ M.Pharm(Ph.D), Associate Proffesor, Department of Pharmaceutical Analysis, Dr.K.V.SubbaReddy Institute of Pharmacy, Dupadu,Kurnool-518218,Andhra Pradesh, India. E-mail ID: deepthi.bpl@gmail.com

² Student, Dr.K.V.Subba Reddy Institute of Pharmacy, Dupadu, Kurnool-518218, Andhra pradesh, India. E-Mail ID: aseefmohammed815@gmail.com.

Abstract:

Green Analytical Chemistry (GAC) is a new field that combines safety, sustainability, and analytical accuracy. It tries to lessen the effects of analytical processes on health and the environment while still being accurate and dependable. GAC uses the twelve principles of green chemistry to make analytical methods that focus on reducing waste, using safer chemicals, being energy-efficient, and monitoring in real time. This review gives a full picture of GAC's goals, principles, technological advances, and future trends. It also shows how important it is for sustainable development and modern chemical analysis. The shift toward greener analytical methods not only helps the environment, but it also makes sure that businesses are following the rules and are cost-effective.

Keywords: Sustainability, analytical method development, miniaturization, eco-efficiency, green solvents, environmental monitoring

1. Introduction

Analytical chemistry has long been a crucial part of science. It helps us understand the makeup of materials and ensures quality control in pharmaceuticals, food, and environmental monitoring. However, traditional analytical methods often depend on toxic solvents, create large amounts of waste, and use up a lot of energy. In the late 20th century, as environmental awareness grew, scientists noticed the contradiction of using polluting methods to study pollution. This realization led to the rise of Green Analytical Chemistry (GAC). GAC merges green chemistry principles with analytical sciences to promote eco- friendly and sustainable practices.

The goal of GAC is to create analytical methods that are efficient, reliable, and gentle on the environment. By using alternative solvents, renewable reagents, and new technologies, GAC offers a way to achieve accuracy while being ecologically responsible. The field has gained worldwide attention as laboratories and industries work to meet environmental policies like the European Green Deal and the United Nations Sustainable Development Goals (SDGs).

2. Scope of Green Analytical Chemistry

The scope of Green Analytical Chemistry includes using sustainable practices throughout all stages of analytical processes. This covers sample collection, preparation, analysis, and waste management. It combines several disciplines, including chemistry, materials science, environmental engineering, and data analytics, to achieve sustainability.

Key areas include:

- Developing greener solvents and reagents, such as water, ionic liquids, or deep eutectic solvents as eco-friendly options.
- Miniaturizing analytical systems, like microfluidics and lab-on-a-chip devices that greatly lower reagent volumes.
- Implementing real-time monitoring techniques with spectroscopic or electrochemical tools for in-situ measurements.
- Conducting Life Cycle Assessment (LCA) to assess the overall environmental impact of analytical methods.

In practice, GAC principles are used in pharmaceutical analysis for drug purity testing, in food industries for contaminant detection, and in environmental laboratories for real-time pollutant tracking. The field is evolving rapidly due to new technologies and a growing focus on regulatory compliance.

3. Principles of Green Analytical Chemistry

The twelve principles of green chemistry, introduced by Anastas and Warner, were adapted for analytical chemistry to help design eco-friendly analytical processes. These principles focus on reducing waste, minimizing harmful materials, conserving energy, and maximizing safety. In analytical contexts, they include:

1. **Direct Analytical Techniques:** Use direct analytical methods to avoid or minimize sample preparation steps, which often involve the most toxic reagents and generate the most waste.
2. **Minimal Sample and Reagent Size:** Aim for minimal sample size and the minimal amount of chemicals and reagents (miniaturization).
3. **In-Situ Measurements:** Perform in-situ measurements (on-site and directly in the system of interest) to reduce sample transportation, stabilization, and potential for contamination or degradation.
4. **Integration of Operations:** Integrate analytical processes and operations to save energy, reduce the number of steps, and decrease the use of reagents. Automation and flow systems are preferred.
5. **Automated and Miniaturized Methods:** Select automated and miniaturized methods (e.g., microfluidics, microextraction techniques) to save energy, reduce reagent consumption, and minimize waste.
6. **Avoid Derivatization:** Avoid or minimize derivatization or other temporary modifications, as these steps require additional reagents and generate waste.
7. **Waste Management:** Avoid generating a large volume of analytical waste and ensure proper management of all analytical waste when it is unavoidable.
8. **Multi-Analyte Methods:** Prefer multi-analyte or multi-parameter methods over methods that measure only one analyte at a time, increasing efficiency and throughput.
9. **Minimize Energy Consumption:** The use of energy should be minimized. Prefer methods that operate at ambient temperature and pressure.
10. **Renewable Reagents:** Prefer reagents obtained from renewable sources (e.g., bio-based solvents) rather than non-renewable, depleting resources.
11. **Eliminate Toxic Reagents:** Eliminate or replace toxic reagents with less hazardous or non-hazardous alternatives, such as water or green solvents (e.g., supercritical CO₂, ionic liquids).
12. **Increase Operator Safety:** The safety of the operator should be increased by minimizing exposure to hazardous substances and reducing the potential for chemical accidents.

These principles give analytical chemists a practical way to balance scientific rigor with care for the environment. They have been included in key regulatory frameworks and recognized best practices in the industry.

4. Goals of Green Analytical Chemistry

The goals of GAC go beyond just analyzing data. They also focus on environmental, economic, and social targets. Analytical methods need to be strong, reliable, and repeatable, while also being sustainable and affordable. GAC aims to achieve these objectives:

- Minimize the use of toxic substances and reagents.
- Optimize energy use with efficient instruments.
- Reduce sample sizes and waste produced.
- Use renewable and biodegradable materials.
- Follow international green standards (EPA, REACH, ICH Q14).

Furthermore, using greenness assessment tools like the Analytical Eco-Scale, GAPI (Green Analytical Procedure Index), and AGREE (Analytical GREENness metric) offers standardized ways to evaluate the sustainability of methods.

5. Green Analytical Method Development

Developing a green analytical method requires a clear approach that improves each step to reduce its environmental impact. This includes preparing samples, measuring them, and interpreting data. Techniques like solid-phase microextraction (SPME), single-drop microextraction (SDME), and supercritical fluid extraction (SFE) have changed sample handling by cutting down or removing the use of solvents.

Instruments such as capillary electrophoresis (CE) and ultra-high-performance liquid chromatography (UHPLC) help save energy and lower solvent use. Modern data analysis tools, like AGREE software, allow for thorough greenness assessment using visual scoring systems.

Steps for Developing a Green Analytical Method

The development process should consider the greenness aspects at every stage, not just the final measurement.

1. Define Analytical Goal and Initial Assessment:

- Determine the required analytical performance (sensitivity, accuracy, precision, etc.).
- Perform an initial assessment of a reference or traditional method's "greenness" to establish a benchmark for improvement.

2. Sample Preparation Greening (The most crucial step):

- Reduce Sample Size: Use minimal sample volume as per the GAC principles.
- Choose Green Extraction: Replace large-volume liquid-liquid extraction (LLE) with greener alternatives like:

i. Solid-Phase Microextraction (SPME)

ii. Supercritical Fluid Extraction (SFE)

iii. Microwave-Assisted Extraction (MAE)

iv. Miniaturized techniques (e.g., Dispersive Liquid–Liquid Microextraction).

- Eliminate Preparation: Where possible, use direct analysis techniques to bypass sample pretreatment entirely.

3. Instrumentation and Reagents Greening:

- Mobile Phase/Solvents: Select safer, less toxic, or renewable solvents (e.g., water, ethanol, isopropyl acetate) for chromatography.
- Miniaturize Separation: Opt for green versions of traditional techniques, such as Capillary Electrophoresis (CE), Micro-HPLC, Nano-HPLC, or Ultra-High Performance Liquid Chromatography (UHPLC), which use significantly less mobile phase.
- Energy Consumption: Choose instruments and procedures that minimize energy use (e.g., ambient temperature operation, energy-efficient equipment).

4. Method Optimization (Integrating Greenness):

- Use Chemometric Tools and Experimental Design (e.g., Quality by Design, Response Surface Methodology) to simultaneously optimize analytical performance *and* greenness factors.
- The optimization objective should include:

i. Maximizing resolution/performance.

ii. Minimizing analysis time.

iii. Minimizing solvent/reagent consumption.

iv. Minimizing mobile phase environmental impact.

5. Greenness Assessment and Validation:

- After the method is developed and validated for performance metrics (LOD, LOQ, precision, accuracy), its greenness must be quantitatively evaluated.
- Use dedicated Greenness Assessment Tools (Metrics) to assign a score or visual profile to the method:

i. Analytical GREENness Metric (AGREE): A versatile tool represented by a circular pictogram with 12 segments corresponding to the GAC principles.

ii. Green Analytical Procedure Index (GAPI): Uses a color-coded pictogram (green, yellow, red) to assess the eco-friendliness of each step of the entire analytical procedure (sample to result).

iii. Analytical Eco-Scale (ESA): Assigns a numerical score based on subtracting penalty points for deviations from an ideal green process.

iv. National Environmental Method Index (NEMI): A qualitative pictogram (green or blank) focused on four criteria: PBT, hazard, corrosivity, and waste volume (<50g).

6. Continuous Improvement:

- A method is never "perfectly green." The process requires a continuous cultural shift to prioritize sustainability and seek better, greener alternatives as new technologies and solvents become available.

6. Miniaturization and Microscale Techniques

Advancements in miniaturization technologies have transformed analytical chemistry with rapid, efficient, and cost-effective analysis. By incorporating microfluidic systems into integrated systems (such as LOC devices and micro total analysis systems (μ TAS)), all analytical stages including sample handling, separation and detection can be accommodated onto a single chip. Miniaturized platforms have the potential to greatly reduce the sample and reagent volume needed to complete the analysis, consistent with GAC principles.

Among the benefits of miniaturization include waste reduction, better portability, real-time sampling, and efficiency. Applications include, but are not limited to: environmental testing, pharmaceutical assessments, and clinical diagnosis.

7. Green Analytical Instrumentation

Portable & Handheld Analyzers in GAC

These compact devices bring the "lab to the sample," offering significant green advantages:

- Real-time, On-site Analysis: Eliminates the need for sample transportation, saving time, energy, and resources.
- Reduced Sample Size and Reagents: Minimizes waste and the use of hazardous chemicals.
- Energy Efficiency: Low-power, often battery-operated designs reduce energy consumption.
- Operator Safety: Enhances user safety by minimizing hazardous chemicals and complex manual procedures.

Types of Portable Analyzers

- Handheld X-ray Fluorescence (XRF): Rapid, non-destructive elemental analysis without chemical reagents.
- Portable Raman & FTIR Spectrometers: Fast, non-destructive identification of substances, often through packaging.
- Portable Gas Chromatography-Mass Spectrometry (GC-MS): Miniaturized systems for on-site analysis of volatile/semi-volatile organic compounds.
- Handheld Laser-Induced Breakdown Spectroscopy (LIBS): Uses a laser for elemental analysis, particularly good for lighter elements.

Greening Analytical Methods Spectroscopic Methods

- Miniaturization and Automation: Reduces sample and reagent consumption.
- Reduced Solvent Use: Prioritizes solids sampling or greener solvents (e.g., ionic liquids, water).
- On-site Analysis: Uses portable instruments (like XRF, Raman) to eliminate lab-related waste and transport.
- Chemometric Tools: Allows analysis of complex samples without extensive, solvent-intensive preparation.

Electrochemical Methods

- Clean Reagent (Electrons): Uses the electron as the primary reagent, which does not produce toxic waste.
- Low Energy Consumption: Reactions often occur at room temperature and pressure.
- Miniaturization and Portability: Sensors (e.g., screen-printed) are ideal for on-site analysis.
- Low Sample/Reagent Volumes: Operates with microliter/nanoliter volumes, minimizing waste.

Energy Efficient Chromatography & Mass Spectrometry

The goal is to reduce high temperatures, long run times, and solvent/material consumption.

- Chromatography:
 - Miniaturization: Using smaller columns reduces the required mobile phase flow and energy for pumping/heating.
 - Supercritical Fluid Chromatography (SFC): A green technique using CO₂ as the primary mobile phase, reducing organic solvent use.
 - Fast Chromatography (e.g., UHPLC): Reduces run time, leading to lower energy and solvent consumption per analysis.
- Mass Spectrometry (MS):
 - Atmospheric Pressure Ionization (API): Simplifies the vacuum system (the major energy consumer) and reduces required power.
 - Miniaturization: Low-flow/nano-flow sources increase efficiency, reducing the need for strong signals and reducing the vacuum pump

workload.

- Alternative Ionization Sources: Research in ambient ionization aims to analyze samples in open air, bypassing the complex vacuum system.

Applications of Green Analytical Chemistry :

GAC has applications in multiple fields such as environmental science, pharmaceuticals, food safety and clinical analysis. In the field of environmental monitoring, solvent-free and in-situ methods such as SPME and portable spectroscopic devices can assess heavy metals and organic pollutants. For pharmaceuticals, GAC can investigate safer drug development while substituting hazardous reagents with greener alternatives. For the food industries, near-infrared (NIR) and Raman spectroscopy can measure food quality, authenticity and contamination, without chemical reagents. Likewise, in clinical diagnostics, miniaturized, portable electrochemical sensors facilitate faster and less invasive analysis.

9. Challenges and Limitations

Despite its benefits, GAC is associated with several limitations which hinder its wider sporadic implementation. These limitations reflect trade-offs between methods' sensitivity and environmental impact, high initial costs of green instrumentation, and a lack of standardization in greenness assessment protocols. Also, regulatory acceptance continues to be slow, as conventional validation criteria may not thoroughly support new green methodologies. Collaboration across regulatory agencies, academic, and industry stakeholders will be needed to break down barriers.

10. Future Trends in Green Analytical Chemistry

The future of GAC is in the intersection of AI, automation, and bio-based technologies. AI-driven optimization allows chemists to make predictions about ideal analytical conditions, eliminating the need for inefficient trial and error experiments. Next-generation solvents such as bio-based and deep eutectic solvents (DESs) will provide additional benefits of increased biodegradability and lower toxicity. The embrace of machine learning by GAC will enhance efficiency in method development and data evaluation. As industries increasingly move towards digitized laboratories and more sustainable methods of production, GAC will remain an essential component of environmentally responsible science.

11. CONCLUSION

Green Analytical Chemistry has transformed the analytical sciences into an environmentally responsible space while maintaining the principles of analytical rigor. Its inclusion into laboratories of all type will ensure safer practices, save funds for laboratories, and provide a valuable framework for sustainability compliance. The continued application of green technologies, AI-based optimization, and life cycle assessments will solidify GAC as the leading example for future analytical development.

References:

1. Armenta S, Garrigues S, de la Guardia M. Green analytical chemistry. *TrAC Trends Anal Chem.* 2008;27(6):497–511.
2. Koel M. Principles of Green Analytical Chemistry. *Anal Bioanal Chem.* 2016;408:861–868.
3. Plotka-Wasyłka J. Green Analytical Procedure Index. *Talanta.* 2018;181:204–209.
4. Pena-Pereira F, Wojnowski W, Tobiszewski M. AGREE—Analytical GREEnness metric. *Anal Chem.* 2020;92(14):10076–10082.
5. Gałuszka A, Migaszwski Z, Namieśnik J. The 12 principles of green analytical chemistry. *TrAC Trends Anal Chem.* 2013;50:78–84.
6. Khairmode PR, Darade RB. Recent Advances in Green Analytical Chemistry. *Asian J Pharm Res Dev.* 2025;13(1):231–237.
7. Miladinović SM, et al. Green Analytical Chemistry: Integrating Sustainability into Chemical Analysis. 2024.
8. Yadav S, Yadav A, Mohan C, et al. Green Chemistry Approaches to Environmental Sustainability. Elsevier; 2024.
9. Bhukya VN, Beda DP. Implementation of Green Analytical Principles. *Green Anal Chem.* 2024;8:100090.
10. Wong PM, Cernak T. Artificial Intelligence and Machine Learning for Chemical Synthesis. *Chem Rev.* 2022;122(20):16421–16469.
11. Xin T, Yu L, Zhang W, Guo Y, et al. Greenness evaluation metrics for analytical methods. *J Pharm Anal.* 2025;14(11):101013.
12. Santana AAC, et al. Green Analytical Chemistry approaches in food analysis: A review. *Microchem J.* 2021;170:106670.