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Synthesis of Nanomaterials Using Different Methods

Broadly, nanomaterials can be created via **top-down** or **bottom-up** approaches. Top-down methods reduce bulk materials to the nanoscale (e.g., mechanical milling), whereas bottom-up methods assemble nanoparticles from atomic or molecular precursors. These methods can be **physical**, **chemical**, or **biological/green**.

Physical Methods

1. Evaporation-Condensation (Inert Gas Condensation)

- Process: A metal or other material is heated in a low-pressure chamber until it evaporates; the vapor then cools and condenses into nanoparticles.
- **Example**: Production of metal oxide nanoparticles (e.g., ZnO, TiO₂) in a high-temperature furnace, collected on a cold surface.
- Advantages: Relatively pure particles, controlled size by tuning pressure/temperature.
- **Disadvantages**: High energy consumption, specialized equipment.

2. Laser Ablation

- Process: A pulsed laser beam strikes a solid target immersed in a liquid or gas, ejecting nanoscale material.
- **Applications**: Generating stable colloidal solutions of gold or silver nanoparticles (AuNPs, AgNPs) without chemical reducing agents.
- **Challenges**: Equipment cost, difficulty scaling up.

3. Sputtering and Mechanical Milling

- Sputtering: High-energy ions knock atoms off a solid target, depositing them as a nanomaterial on a substrate.
- **High-Energy Milling**: Ball milling bulk materials into nanosized powders. Top-down approach with less control over uniformity and morphology.

Chemical Methods

1. Sol-Gel Process

- **Key Idea**: Hydrolysis and condensation of precursors (e.g., metal alkoxides) form colloidal sol, which transitions to a gel and then is calcined to form nanoparticles.
- · Uses: Synthesis of silica, titanium oxide, and various ceramic nanomaterials with controllable porosity.

2. Chemical Reduction and Precipitation

- **Example**: Noble metal nanoparticles (Au, Ag) produced by reducing metal salts (e.g., HAuCl₄) with agents like sodium borohydride (NaBH₄), citrate, or hydrazine.
- Surfactants or stabilizers (e.g., PVP, CTAB) often used to control particle growth and prevent aggregation.
- o **Tunability**: Reaction conditions (pH, temperature, reagent ratios) adjust size and shape.

3. Hydrothermal and Solvothermal Techniques

- Process: Precursors sealed under high pressure and temperature, promoting crystal growth in water or organic solvents.
- o **Advantages**: Produces crystalline nanoparticles of metal oxides, sulfides, or composites with fewer defects.
- **Challenges**: Pressure vessel needed, optimization can be intricate.

4. Emulsion and Microemulsion Methods

- **Concept**: Use water-oil systems stabilized by surfactants. Nanoparticles precipitate in microreactors (micelles).
- Applications: Polymeric nanoparticles, controlled morphologies, and well-defined core-shell structures.

Biological (Green) Methods

1. Microbial Synthesis

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- o Bacteria, fungi, or algae secrete enzymes that reduce metal ions to nanoparticles.
- Examples: Fusarium oxysporum or Pseudomonas stutzeri producing AgNPs, AuNPs.
- Benefits: Mild conditions, often water-based, minimal toxic byproducts.

2. Phytosynthesis (Plant Extracts)

- Plant extracts (rich in polyphenols, flavonoids) act as reducing/stabilizing agents, forming metal or metal oxide nanoparticles.
- o Examples: Azadirachta indica (neem) for AgNPs, Ocimum sanctum (tulsi) for AuNPs.
- o Environmentally friendly, cost-effective, fosters synergy with Ayurvedic pharmacopeia.

Molecular Basis of Biosynthesis of Nanomaterials

The "green" or **biogenic** nanoparticle synthesis hinges on the ability of biological systems (plants, microbes) to reduce metal ions and stabilize nascent nanostructures.

1. Enzymatic Reduction

- NADH/NADPH-dependent reductases in microbes or plant cells donate electrons to metal ions (e.g., Ag⁺, Au³⁺) → elemental metal (Ag⁰, Au⁰).
- o Cytochrome or other metalloproteins may also participate, forming stable nucleation centers.

2. Phytochemicals as Reductants

- Polyphenols, flavonoids, terpenoids provide the electron-donating capacity.
- Functional Groups (-OH, -CHO) can chelate metal ions, controlling nucleation/growth.
- o Secondary metabolites (e.g., ascorbic acid, anthocyanins) further enhance or modulate particle shape/size.

3. Capping and Stabilization

- Biological macromolecules (proteins, polysaccharides) or small molecules anchor onto nanoparticle surfaces, preventing agglomeration.
- This bio-capping can impart biocompatibility, potential specificity for biological targets, and extended colloidal stability.

4. Genetic and Environmental Influences

- Microbes can upregulate reductase enzymes under metal stress (self-protection) → nanoparticle formation.
- Reaction environment (pH, ionic strength) dictates whether smaller or larger aggregates form.

Assessment of Plant, Animal, and Mineral-Based Drugs for Nanomaterials

Plant-Based (Phytogenic) Nanomaterials

1. Phytopharmaceuticals

- $\circ\,$ Traditional herbal extracts or decoctions can be directly leveraged to synthesize nanoparticles.
- Example: **Green synthesis** of gold nanoparticles using **Terminalia arjuna** bark extract, combining cardiotonic properties with plasmonic potential.

2. Advantages

- $\circ\,$ Natural capping ligands typically enhance biocompatibility and reduce toxicity.
- Possible synergy with pharmacologically active phytoconstituents, opening new avenues in drug delivery or combined therapies (e.g., herbal-metal nanoformulations).

3. Challenges

- o Standardization: Variable chemical profiles in plants due to geographic, seasonal factors.
- **Scalability**: Large-scale extraction, reproducible reaction conditions.

Animal-Based (Zoogenic) Nanomaterials

1. Chitosan and Other Biopolymers

- o Derived from crustacean shells or insect cuticles.
- $\circ~$ Biodegradable, mucoadhesive, can form nanoparticles for gene/drug delivery.

2. Proteins (e.g., Silk Fibroin, Collagen)

• Self-assembly of silk fibroin into nanoparticles, used for controlled release.

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o Collagen-based nanoparticles for regenerative medicine.

3. Challenges

• Ethical sourcing, risk of immunogenicity, ensuring consistent quality.

Mineral-Based (Geogenic) Nanomaterials

1. Classical Ayurvedic Bhasmas

- Repeated calcination of metals/minerals (e.g., Suvarna bhasma from gold, Lauha bhasma from iron).
- o Modern characterization reveals nano-/submicron-sized metal oxides or carbonates.

2. Biomedical Potential

- o Proposed enhanced bioavailability, possibly lower toxicity if properly detoxified (shodhana).
- Integration with advanced toxicity and efficacy assays crucial to meet modern regulatory standards.

3. Key Issues

- Standardization: Variation in raw materials, incomplete knowledge of reaction pathways.
- Need for mechanistic insights: identifying roles of trace herbal components, controlling final nanoparticle size and morphology.

Concluding Remarks

The **synthesis of nanomaterials** comprises a rich spectrum of techniques—ranging from physical/chemical methods (sol-gel, hydrothermal, laser ablation) to **biological/green** approaches harnessing microbial enzymes or phytochemicals. The **molecular basis** of biosynthesis involves redox-active biomolecules (enzymes, secondary metabolites) directing nucleation, growth, and capping of nanoparticles.

Simultaneously, plant-, animal-, and mineral-based pharmaceuticals—especially from the Ayurvedic tradition—present unique opportunities to create hybrid nanoformulations that merge ancient medicinal wisdom with modern nanotechnology. Rigorous characterization (particle size, morphology, surface chemistry), biocompatibility, and efficacy studies are essential to unlock safe, reproducible, and scalable nanomedicines. This synergy underscores the promise of sustainable and culturally rooted approaches in advancing next-generation healthcare solutions.

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