

## Plant-mediated green synthesis of nanoparticles

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### Abstract

Nanoparticles are an inspiring group of nanostructured materials with broad-spectrum applications in different fields such as catalysis, antimicrobial treatment, drug delivery, nanomedicine, environmental remediation, electronics, and chemical sensors. Nevertheless, the techniques used for preparation are environmentally unfriendly. Aiming to promote the greener synthesis of nanoparticles, this chapter spotlights plant-mediated eco-friendly and sustainable development of nanoparticles. Naturally occurring plant extracts are enriched with a plethora of various biologically active biomolecules and secondary metabolites, including alkaloids, terpenoids, flavonoids, enzymes, and phenolic substances. These bioactive compounds can catalyze the reduction of metal ions into biogenic nanoparticles in an eco-sustainable single-step biosynthetic process. Additionally, the utilization of plant extracts and their derived compounds circumvents the necessity for capping and stabilizing agents and generates bioactive size and shape-dependent green nanoparticles. Herein, we have made an effort to describe the synthesis of a wide range of metal-based nanoparticles (platinum, gold, zinc oxide, silver, and titanium dioxide nanoparticles) by using plant extract as a green synthesis matrix. In addition, different parts of plants that have widely been utilized for the biosynthesis of these NPs with several sizes and shapes by biological

31 methodologies are briefly described. In conclusion, the greener synthesis approaches are safer and  
32 easier to exploit the massive preparation of nanostructured particles.

33 **Keywords;** Nanotechnology; Metal nanoparticles; Green chemistry; Plant extract; Secondary  
34 metabolites

35

## 36 **1. Introduction**

37 Nanotechnology may be described as the modification of matter by different physical or chemical  
38 tactics for the formation of substances with particular applications [1]. It can also be defined as the  
39 **microscopic-sized** particle that has at least, one dimension much lesser than one  
40 hundred nanometers in size [2]. **Nanoparticles (NPs)** possesses many interesting applications and  
41 multi-functional properties in diverse fields which includes nutrition energy and medicine [3, 4]  
42 because of their significant surface to volume ratio and abundance of surface atoms. Other  
43 considerable advantageous features of plant-based nanoparticles are shown in **Figure 1**. The  
44 biogenic synthesis of monodispersed nanoparticles with particular sizes and shapes  
45 was an undertaking in biomaterial science. Also, it has garnered prodigious interest within  
46 the industry of pharmacology for the cure of viral and bacterial infections [5]. The biological  
47 synthesis strategies are prospective alternative as compared to other classical methods of synthesis  
48 due to the usage of extra-biological compounds. The easy accessibility and rich biodiversity of  
49 natural material had been enormously studied for the biosynthesis of **nanomaterials (NMs)** [6].  
50 **Recently**, the green synthesis of nanosized tubes, particles, wires, and flowers was described.  
51 These biosynthesized NMs exhibit great prospective in many diverse fields like diagnosis,  
52 treatment, commercial product manufacturing, and improvement surgical nanodevices [7].  
53 Nanomedicines had made a great influence in the healthcare sector by  
54 curing numerous chronic ailments. Therefore, green biosynthesis of NPs is taken into  
55 consideration as the building blocks of the coming near generations by controlling various  
56 health issues [8].

57 Nowadays, nanotechnology has attracted great interest as a promptly growing scientific field.  
58 Manipulation, characterization, fabrication of NPs are the foremost aims observed in this new  
59 technology [4, 9, 10]. Plant crude extracts of plants consist of unique secondary  
60 metabolites like phenolic acid, terpenoids, flavonoids, and alkaloids, wherein these metabolites  
61 are known to be the reason **for** ions reduction during the biosynthesis of NPs from different metals

62 [11]. These plant metabolites are frequently used for redox reactions for the preparation of eco-  
63 friendly NPs. Many earlier reviews are revealing that these synthesized nanoparticles are  
64 efficiently controlling the apoptosis, genotoxicity, and oxidative stress associated changes [12].  
65 Moreover, NPs possess a wider range of applications in plant sciences and the agriculture industry.  
66 For example, with the help of bioprocessing technology, nanoparticles can convert agricultural  
67 wastes and food into energy and many other useful products.

## 68 **2. Methods for metallic nanoparticle biosynthesis**

69 Numerous procedures are utilized for the formation of different nanoparticles like chemical,  
70 biological, physical as well as enzymatic. Chemical protocols are utilized to prepare  
71 nanoparticles by way of the sol-gel, electrodeposition, and vapour deposition  
72 [13], moist chemical, and co-precipitation techniques [14], hydrolysis, catalytic route, Langmuir  
73 Blodgett approach and soft chemical technique [15]. Likewise, physical techniques that are being  
74 used include ball milling, plasma arcing, thermal evaporate, ultra-thin films, spray  
75 pyrolysis, lithographic techniques, pulsed laser desorption, layer by using layer growth, molecular  
76 beam epitaxis, sputter deposition and diffusion flame [16]. Chemical and physical techniques are  
77 using stabilizing agents, highly concentrated reductants, and high radiations that are toxic and  
78 unsafe for humans and the environment. Therefore, the biosynthesis of NPs is a single step bio-  
79 reduction procedure that uses lesser energy for the synthesis of eco-friendly nanoparticles [17].  
80 Plant extracts, enzymes, fungi, bacteria, and microalgae are the different resources that are being  
81 used for the biosynthesis of nanoparticles [18]. Table 1 depicts a recent list of biosynthesis  
82 of metallic nanoparticles from various plant sources.

## 83 **3. Green biosynthesis of metallic NPs**

84 The procedures for acquiring NPs through different naturally occurring compounds like sugars,  
85 plant extracts, vitamins, micro-organisms, and biodegradable polymers might be considered as  
86 appealing for nanotechnology. This biological synthesis has led to the fabrication of a  
87 confined variety of inorganic NPs (particularly metallic NPs, though numerous salts and  
88 metal oxides also are described). Among the material noted above, primarily plant-  
89 based substances are to be the greatest source and are appropriate for the large-scale formation of  
90 NPs [18]. For the biological synthesis of metallic NPs, different parts of plants like root, stem, leaf,  
91 seed, and latex are being utilized. Polyphenols that are present in plants for example in wine, tea,  
92 red grape pomace, and winery waste are supposed to be the active and important agent required

93 for the synthesis. Green biosynthesis of nanoparticles offers advancement as compared to other  
94 strategies as it is cost-efficient, comparatively reproducible, simple and it results in more stable  
95 products [19]. Generally, biological material offers an environmentally friendly and green  
96 chemical protocol for the production of valuable materials as the biomaterial-based routes reduce  
97 the usage of toxic chemicals [20]. Figure 2 shows a schematic illustration of the biosynthesis of  
98 metallic nanoparticles using various parts of the plant. Few different types of synthesis for metallic  
99 NPs are as follows:

### 100 3.1. Gold nanoparticles

101 Gold (Au) nanoparticles have been stated as an extensive research tool in different fields of  
102 agriculture, medicine, and health center due to their stability and biocompatibility [21]. Au  
103 is considered as one of the extraordinary metals having a melting and boiling point of 1064 °C and  
104 2808 °C, respectively. Numerous properties of Au, like its incapability to react with oxygen and  
105 water and its exceptional conductive properties, have made it very beneficial for mankind. During  
106 the 5<sup>th</sup> millennium BC, gold extraction began close to Bulgaria and is assumed that “soluble” Au  
107 come to be seen approximately the 4<sup>th</sup> or 5<sup>th</sup> centuries BC in China. The spectacular statue of  
108 Touthankamon that was built in this time, is proof. Earlier, gold was known with various terms,  
109 like drinkable and soluble Au, Graham [22] coined the term “colloid”. The colloids of Au and the  
110 stunning ruby-red color have attracted people for lots of eras [23, 24]. Gold was widely utilized  
111 for medicinal, cosmetic, and ornamental purposes [25, 26]. Drinkable Au was also used for the  
112 cure of several health problems like arthritis, heart diseases, tumors, dysentery, epilepsy, and  
113 venereal disease [27].

114 Gold nanoparticles (AuNPs) fascinated some researchers inside the area of plant-based  
115 biosynthesis due to their exceptional applications and properties in biomedical, nanodevices,  
116 nonlinear optics, catalysis, and nanodevices [28]. Gold nanoparticles offer promising scaffolds for  
117 gene and drug delivery [29]. These nanoparticles possess many useful features such as  
118 monodispersity, tunable core size, tuning, and transport of delivery processes and the large  
119 surface-volume ratio [30]. The AuNPs can be formed by green synthesis [11, 31], however, the  
120 number of gold synthesis reports is considerably lesser as compared to AgNPs. The size of AuNPs  
121 varies within a range of 20-300 nm. For example, gold nanoparticles (100 nm) have been prepared  
122 from the peel extract of banana (*Musa paradisiaca*) by using simple, eco-friendly, and non-toxic  
123 material [32]. The crushed, boiled, acetone-precipitated and dried powder of banana peel was used

124 for the reduction of chloroauric acid. In this study, the enlargement of NPs into microwire and  
125 microcubes networks to the periphery of the banana sample was seen. The contribution of amine,  
126 hydroxyl, carboxyl groups was observed during the synthesis. Using tea extracts, nanocomposites  
127 of gold nanoparticles can be prepared. The extract was made in the solution of 1-methyl-2-  
128 pyrrolidinone during the process of nanoparticles (20 nm) formation [33]. The AuNPs formation  
129 in the polyaniline matrix was confirmed through TEM. In another research by Wu and Chen [34],  
130 a facile and green route was reported, by mixing rice wine, soda, and Au (III) at pH 6.5 at a  
131 temperature of 25-55 °C without the use of any protective agent. No precipitation occurred, and  
132 the resultant solution was stable after a few months.

133 Shankar et al. [35] described the formation of AuNPs by using the leaves of *Pelargonium*  
134 *graveolens*. It was stated that present terpenoids in geranium leaves act as reducing and capping  
135 material for a quick reduction of chloroaurate ions towards the AuNPs of different sizes. Later on,  
136 these research groups determined the formation of gold nanoparticles by using different plants like  
137 *Azadirachta indica* [36] and lemongrass [37]. Prism-, trapezoid-, rod- and sphere-shaped gold  
138 nanoparticles were formed from the black tea leaf extract. The tea phenols and flavonoids are  
139 known to be the reason for efficient reduction caused by the extract [38]. Song et al. [39] stated  
140 that leaf extracts of *Diospyros kaki* and *Magnolia kobus* successfully synthesize the extracellular  
141 gold nanoparticles within a range of 5-300 nm having pentagonal, triangular, spherical and  
142 hexagonal shapes at the temperature of 95 °C within few minutes. Babu et al. [40] cited that leaves  
143 of *Mentha arvensis* (ethanolic extract) formed AuNPs of spherical and hexagonal shapes having  
144 approximately a size of  $39 \pm 15$  nm. Extract of pear fruit biosynthesized hexagonal and triangular  
145 nanoparticles at room temperature of 200-500 nm [41]. The reason for the synthesis of AuNPs is  
146 the presence of organic acids, proteins, peptides, amino acids, and saccharides in pear fruit extract.  
147 Rajan et al. [42] used *Garcinia combogia* (fruit extract) for the biosynthesis of AuNPs of  
148 anisotropic and spherical shapes. It was concluded that the shape of nanoparticles relies on the  
149 reaction temperature and quantity of extract. In 2015, Islam [43] and his research group stated that  
150 *Pistacia integerrima* (leaf galls) reduces ions into the gold nanoparticles. Carboxylic acid and  
151 hydroxyl groups of polyphenols were known to be the reason for the process of reduction. These  
152 polyphenols capped the NPs and making them more stable in varied pH solution and at high  
153 temperatures. These synthesized Au nanoparticles showed great potential in antimicrobial, enzyme

154 inhibition, muscle relaxant, and antinociceptive activities. The mechanism of antimicrobial  
155 activity of nanoparticles is shown in Figure 3.

### 156 3.2. Platinum nanoparticles

157 Noble metallic nanoparticles like silver, gold, platinum, and palladium, possess a wider range of  
158 different applications, such as material science, medicine, pharmaceuticals, and chemistry [44, 45].  
159 Amongst them, platinum has few specific properties like good resistance towards chemical attacks  
160 and corrosion, and high surface area. For the synthesis of platinum nanoparticles (PtNPs) till now  
161 numerous methods have been used such as sol-gel route, vapour deposition, and chemical  
162 precipitation. Though, these protocols have few limitations such as usage of toxic and unsafe  
163 chemical, high cost and energy requirements, multistep for preparation. To overcome these  
164 limitations, recently different plant-based synthesis of these metallic nanoparticles, especially  
165 platinum nanoparticles have gained great attention because of their simple usage, eco-friendly, and  
166 non-toxic nature.

167 Platinum (Pt) is one of the expensive, rare, and high-density metal. Platinum nanoparticles are  
168 mostly used in the form of suspension or colloid. The antioxidant ability of PtNPs is one of the  
169 reasons for their extensive research [46]. The main application areas of PtNPs are cancer therapy,  
170 polymer membranes, catalytic converters, plastics, textiles, and nanofibers. PtNPs are being  
171 extensively utilized as catalysts and in the different biomedical applications [47]. In comparison  
172 with silver, gold, platinum NPs are notably limited. The reaction between an aqueous solution of  
173 Pt and plant extracts leads to the following mechanism:



175 Specifically, the interest in PtNPs is because of their unique structural, catalytic, and optical  
176 properties making them a promising nanoparticle in catalysis and biomedical applications [48, 49].  
177 Biological approaches for the biosynthesis of PtNPs with the help of plant extracts has not been  
178 extensively used. According to literature survey, the plants that have been used for the green  
179 synthesis for PtNPs are; *Diospyros kaki* [50], *Ocimum sanctum* [51], *Anacardium occidentale* [52],  
180 *Cacumen platycladi* [53], *Bidens Tripartitus* [54], *Punica granatum* [55], *Cochlospermum*  
181 *gossypium* [56], *Azadirachta indica* and *Quercus glauca* [57]. Jae et al. [58] prepared PtNPs (2-12  
182 nm) from a solution of  $\text{H}_2\text{PtCl}_6 \cdot 6\text{H}_2\text{O}$ , where leaves extract of *Diopyros kaki* were used as the  
183 agent for reduction of ions. At a temperature of 95 °C with an extract concentration of more than  
184 10%, around 90% of platinum ions were successfully converted into PtNPs. Zheng et al. [53],

185 biologically synthesize PtNPs ( $2.4 \pm 0.8$  nm) by using an extract of *Cacumen platycladi* at 90 °C  
186 having an extract of 70% and reaction time of 25 h. During this reaction, the flavonoids and  
187 reducing sugars performed an important part in the reduction of the platinum ion as compared to  
188 proteins. Likewise, Soundarrajan et al. [51] used the leaves of *O. sanctum* that acts as the reducing  
189 agent during the biosynthesis of PtNPs of size 23 nm, whereas the solution of  $H_2PtCl_6 \cdot 6H_2O$  was  
190 used. Different compounds of plants such as gallic acid, ascorbic acid, proteins, and terpenoids  
191 acted as agents for reduction during the formation of PtNPs. Kumar et al. [59] isolated polyphenols  
192 from the extract of *Terminalia chebula* that causes the one single step synthesis of platinum  
193 nanoparticles. They demonstrated that the reduction of platinum (IV) to platinum (0) was due to  
194 the presence of polyphenols in the *T. chebula* extracts.

195 Platinum nanoparticles, which were synthesized from tea polyphenol act as both capping and  
196 reducing agents. These PtNPs were crystalline in nature, having a particle size of 30-60 nm, with  
197 a structure of face-centered cubic. Transmission electron microscopy determines that the NPs were  
198 of flower-shaped. The polyphenols present in tea include an amount of different phenolic  
199 components that form the complexes with the ions and quickly reduce them into metallic  
200 nanoparticles of various sizes and shapes [60-62]. The usage of different extracts from plant  
201 extracts for the green biosynthesis of PtNPs relies on the fact that the procedure is easier, faster,  
202 reliable, cost-effective, and eco-friendly and forms more stable nanoparticles as compared to other  
203 classical methods [63].

### 204 3.3. Silver nanoparticles

205 Usually, silver (Ag) is used as a catalyst for the oxidation of ethylene to ethylene oxide and  
206 methanol to formaldehyde [64]. Some of the features of Ag are good conductivity, catalytic,  
207 antibacterial potential, and chemical stability [65]. In the era of the Roman Empire, silver  
208 nanoparticles were utilized by the founders of glass, for the creation of the Lycurgus cup (4<sup>th</sup>  
209 century AD), now this cup is in British Museum [64, 66]. Before the 1980s, silver nanoparticles  
210 (AgNPs) were used for isolated supports that were utilized for signals in Raman spectroscopy for  
211 practical and scientific concern [65], revealing that AgNPs exhibits a unique organization of high  
212 electrical double layer capacitance [67]. Currently, the formation of AgNPs is one of the most  
213 actively rising development in the colloid chemistry and continuous increase in the scientific  
214 publications in around the last 20 years.

215 The mechanism of AgNPs synthesis is due to the occurrence of the polyphenols in the extracts of  
216 different plants, which causes the reduction of NPs. This reduction is carried out by the removal  
217 of hydrogen due to the OH groups present in polyphenols. The formation of silver nanoparticles  
218 is being performed by various biological systems [68-71]. Numerous plants and their respective  
219 parts have been utilized for the formation of silver nanoparticles. This formation involves the  
220 reaction of AgN salt with the plant. The presence of a brownish-yellow colour confirms the  
221 formation of AgNPs.

222 At present, AgNPs have gained great attention and are being considered as the most capable and  
223 useful NP for different biological applications, including biomolecular detection, therapeutics  
224 goals, food production and preservation, agricultural purposes, antibacterial agents, drug delivery  
225 [72], bio-labeling wound healings, sensing, cosmetics and water purifications [73]. These NPs can  
226 be formed through protocols [74]. Though the method of chemical reduction is a common one for  
227 the formation of silver nanoparticles, the usage of precarious and expensive chemicals has diverted  
228 the attention of researchers for the search of new and alternative methods [75]. Some potential  
229 health risks were another reason for the search for new methods and had attracted the attention of  
230 researchers worldwide [2, 76].

231 Recently, plant-based green biosynthesis of AgNPs is rising into an important subdivision of  
232 nanotechnology, as it gained importance and is developed due to its cost-effectiveness and lesser  
233 toxicity [77]. Silver has been considered and studied very extensively for plant-based synthesis  
234 and it has been known as a more rapid and easier process as compared to the monotonous and time  
235 taking microbial synthetic methods [47]. Shankar et al. [78] used leaves extract of *Pelargonium*  
236 *graveolens* for the extracellular formation of AgNPs. Through mixing AgN solution and plant  
237 extract, a quick reduction of Ag ions take place followed by the synthesis of crystalline and highly  
238 stable AgNPs (16–40 nm) in solution. Later on, in 2007, Huang [79] displayed that spherical or  
239 triangular-shaped silver nanoparticles of size 55-80 nm can be prepared from the leaf extract of  
240 *Cinnamomum camphora*. He discovered that water-soluble heterocyclic and polyol compounds in  
241 the leaves of *C. camphora* are responsible for silver ions reduction. Leela and Vivekanandan [80]  
242 compared the leaves extract of some different plants such as *Sorghum bicolor*, *Basella alba*,  
243 *Helianthus annus*, *Saccharum officinarum*, *Zea mays*, *Oryza sativa* and for the formation of  
244 AgNPs and determined that *H. annus* showed a speedy reduction of silver ions. Similarly, Ahmad  
245 et al. [81] used broth of *Ocimum sanctum* for the biosynthesis of AgNPs and observed highly stable

246 NPs with a size of  $5 \pm 1.5$  nm to  $10 \pm 2$ . In a research conducted by Jeyaraj et al. [82] the biological  
247 preparation of silver nanoparticles from the leaves extract of *Podophyllum hexandrum* was  
248 described. The complete reduction of silver ions was completed in 2.5 h at the temperature of 60  
249 °C and pH 4.5 with the formation of spherical shaped nanoparticles in the range of 12 to 40 nm.  
250 Saratale et al. [83] biosynthesized spherically and monodispersed silver nanoparticles (15 nm) by  
251 using the leaf extract of *Taraxacum officinale*. The presence of flavonoids, terpenoids, and  
252 triterpenes are known to be the active compounds present in extract for the synthesis of silver  
253 nanoparticles.

254 By using the aqueous extract of *Alternanthera dentate* a rapid and green synthesis of AgNPs  
255 having spherical shaped with a size of 50–100 nm was prepared. Within 10 min, the silver ions  
256 were reduced into silver nanoparticles by the leaf extract. These prepared AgNPs showed  
257 antibacterial potential against *Escherichia coli*, *Klebsiella pneumonia*, *Enterococcus faecal* and  
258 *Pseudomonas aeruginosa* [84]. For the formation of AgNPs, *Acorus calamus* was also used and  
259 its antibacterial, anticancer, and antioxidant effects were determined [85]. In 2014, Nakkala [86]  
260 and his research members used the extract of *Boerhaavia diffusa* plant as the reducing material for  
261 the formation of AgNPs. It was revealed from TEM and XRD that the prepared NPs possess a  
262 particle size of 25 nm, spherical shape with a face-centered cubic structure. The antibacterial  
263 potential was determined against *Flavobacterium branchiophilum*, *Aeromonas hydrophila*, and  
264 *Pseudomonas fluorescens*, the highest sensitivity was observed toward *F. Branchiophilumin* as  
265 compared to the two other bacteria. Similarly, Nabikhan et al. [87] prepared spherical AgNPs (5-  
266 20 nm) by using the callus extract of *Sesuvium portulacastrum* L. a salt marsh plant.

267 The fruit of *Tribulus terrestris* L was dried and reacted with Ag nitrate for the formation of silver  
268 nanoparticles. The newly synthesized AgNPs possess a size of 16-28 nm with spherical shape and  
269 were used to determine their antibacterial potential against few multi-drug resistant bacteria like  
270 *Streptococcus pyogens*, *Staphylococcus aureus*, *Pseudomonas aeruginosa*, *Escherichia coli* and  
271 *Bacillus subtilis* [88]. Methanolic and Ethyl acetate extracts of *Cocous nucifera* were used for the  
272 successful synthesis of Ag nanoparticles (22 nm) and exhibited good antimicrobial activity  
273 towards different bacterial strains such as *Klebsiella pneumoniae*, *Bacillus subtilis*, *Salmonella*  
274 *paratyphi* and *Pseudomonas aeruginosa* [89]. Another spherical shaped and stable AgNPs were  
275 synthesized from *Abutilon indicum* which also possesses good antimicrobial potential against *E.*  
276 *coli*, *S. typhi*, *B. subtilis*, and *S. aureus* [90]. Leaves of *Ziziphoratenuior* were also utilized for

277 the preparation of AgNPs and the nanoparticles were characterized through different techniques.  
278 TEM analysis reveals the spherical shape and size of 8 to 40 nm, and FTIR showed carbonyl,  
279 hydroxyl, amine, and some other stabilizing functional groups [91]. By using the leaves of  
280 *Acalypha indica*, a rapid and green protocol for the formation of nanoparticles was reported by  
281 Krishnaraj et al. [92], where a successful formation of nanoparticles was completed in 30 min [92].  
282 A weed *Chenopodium album* was reported in a rapid and facile biosynthesis of AgNPs. Its leaf  
283 extract was used for the synthesis and results in NPs of the size range of 10-30 nm and its shape  
284 was spherical that was inferred through TEM analysis [93]. Similarly, silver nitrate was reduced  
285 by *Azadirachta indica* (leaf extract) that results in the biosynthesis of silver nanoparticles (10-35  
286 nm) with spherical shape [94].

### 287 **3.4. Zinc oxide nanoparticles**

288 **Zinc oxide (ZnO)** is an n-type semiconducting metallic oxide. **ZnO nanoparticles (ZnONPs)** have  
289 diverted the attention of researchers in the past few years because of its wider variety of  
290 applications in the field of optics, biomedical systems, and electronics [95, 96]. Different  
291 categories of inorganic metallic oxides have been prepared in some latest findings such as CuO,  
292 ZnO, and TiO<sub>2</sub>. Of all these metallic oxides, ZnONPs have gained much interest as they are not  
293 costly to synthesize, safe, and easily prepared [97]. ZnO as a metal oxide has been registered as  
294 GRAS (generally recognized as safe) by FDA US [98]. These nanoparticles possess great  
295 semiconducting features due to the large bandgap and high exciton binding energy such as catalytic  
296 potential, wound healing, UV filtering, and optics [99, 100]. ZnONPs have been widely utilized in  
297 different cosmetics such as sunscreen lotion, depending on its UV filtering potential [101]. It also  
298 has extensive use in biomedical fields like anticancer, antifungal, antibacterial, antidiabetic, and  
299 drug delivery [102, 103]. While ZnO is being utilized in targeted drug delivery and still have the  
300 constraint of cytotoxicity, which has to be solved [104]. Nanoflower, nanoflake, nanorod,  
301 nanowire, and nanobelt are different morphologies of ZnONPs that have been reported [105, 106].  
302 The different parts of plants like leaf, fruit, stem, seed, and root, have been used for the formation  
303 of ZnONPs, as they contain a sufficient quantity of phytochemicals. The usage of natural extracts  
304 for biosynthesis is an eco-friendly and cheap process that does not require any intermediate base  
305 groups [107]. Plants produce large scale synthesis of stable nanoparticles with various sizes and  
306 shapes, it is one of the reasons that plants are the preferred source for the formation of NPs [108].  
307 The synthesis of ZnO nanoparticles is achieved by bio-reduction, which involves the reduction of

308 metallic ions into 0 valences metallic NPs. This process is assisted by different phytochemicals  
309 such as polyphenolic compounds, amino acids, vitamins, polysaccharides, terpenoids, and  
310 alkaloids secreted from several plants [107, 108].

311 Jafarirad [109] and his research team experimented and compared the results of zinc oxide  
312 nanoparticles that were synthesized through two different methods- microwave irradiation (MI)  
313 and conventional heating (CH) and results proved that MI took lesser time and faster reaction rate  
314 for the formation of nanoparticles [109]. Plants of family Lamiaceae have been widely used like  
315 *Plectranthus amboinicus* [110], *Vitex negundo* [111], and *Anisochilus carnosus* [112] showed NPs  
316 of different shapes and sizes like quasi-spherical, rod-shaped, spherical, and hexagonal. The  
317 findings demonstrate that an increase in the concentration of the plant extract causes a decrease in  
318 the size of nanoparticles [110-112]. *Azadirachta indica* (leaf extract) of family Meliaceae have  
319 been commonly used for the ZnONP biosynthesis [113, 114]. Sangeetha et al. [115] synthesized  
320 spherical and highly stable ZnONPs of size 25-40 nm from the leaf extract of *Aloe barbadensis*. It  
321 was described that the synthesized NPs were polydispersed and their particle size can be controlled  
322 by changing the concentration of extract. Similarly, spherical and highly stable NPS were  
323 synthesized using *Parthenium hysterophorus* leaf extracts. Additionally, *Plectranthus amboinicus*  
324 (leaf extract) was utilized for the formation of zinc oxide NPs by Vijayakumar et al. [116]. The  
325 synthesis of ZnONPs has also been performed by using *Sedum alfredii* [108], *Physalis alkekengi*  
326 [117], *Pongamia pinnata* [118], flowers of *Trifolium pretense* [119], *Cassia Auriculata* [120, 121].  
327 Qu et al. [117] synthesized crystalline ZnO nanoparticles (72.5 nm) from *Physalis alkekengi*, it  
328 can grow in soils with high levels of Zn and can incorporate zinc in its aerial parts.

### 329 **3.5. Titanium dioxide nanoparticles**

330 **Titanium dioxide (TiO<sub>2</sub>)** is the oxide form of Ti and occurs naturally. It can be obtained from  
331 different minerals as brookite, rutile, and anatase. The manufacturing and utilization of TiO<sub>2</sub> only  
332 in the US have stayed is 1100 thousand tons since 1997, and it has been categorized as a possible  
333 carcinogen for humans (Group 2B) by International Agency for Research on Cancer [122]. TiO<sub>2</sub>  
334 is a main and vital metallic oxide nanoparticle that is being extensively used in industrial  
335 photocatalytic processes, printing ink, paints, paper, rubber, sunscreens, cosmetics, air cleaning  
336 products, and car materials because of its biological, chemical and physical features [123]. The  
337 nanoparticles of TiO<sub>2</sub> are being utilized in a wide variety of applications such as sunscreens, drug  
338 delivery systems, food preparation, and cosmetics [124-126].

339 Due to its bright, white pigment and high refractive index titanium dioxide is an ideal material  
340 used as a whitening agent in different applications. Discovery of the super-hydrophilicity and  
341 photocatalytic properties of TiO<sub>2</sub> has also led to applications in some industries by producing self-  
342 cleaning products as well as enhancing sterilization and deodorizing processes. The biomedical  
343 applications of nanoparticles synthesized from titanium dioxide have greatly developed in recent  
344 few years. Research is being conducted on these NPs to attenuate the effects of chemotherapy by  
345 making cancer therapy more efficient and targeted [127, 128]. Additionally, food-grade titanium  
346 dioxide nanoparticles (TiO<sub>2</sub>NPs) are present in a range of food products like gum, candy, donuts,  
347 marshmallows, cookies, and some others. Toothpaste, shaving creams, deodorants, conditioners,  
348 shampoos, and sunscreens are some personal care products also containing food-grade TiO<sub>2</sub>NPs  
349 [129].

350 Sundrarajan and Gowri [130] synthesized titanium nanoparticles by using titanium isopropoxide  
351 solution and leaf extract of *Nyctanthes arbortristis* with a size range of (100-150 nm). Similarly,  
352 TiO<sub>2</sub>NPs (25–100 nm) were also biosynthesized from *Jatropha curcas* (aqueous extract) [131].  
353 An enzyme, curcain, and cyclic peptides were recognized as the possible capping and reducing  
354 compound in latex of *J. curcas*. Spherical shaped (23 ± 2 nm) nanoparticles were synthesized from  
355 the fruit peel of *Annona squamosa* at room temperature with a time of 6 h [132]. At room  
356 temperature, *Solanum trilobatum* (leaves) were used for the formation of titanium NPs that were  
357 having pediculicidal and larvicidal activities [123] (Rajakumar et al., 2013). Velayutham et al.  
358 [133] use leaves extract of *Catharanthus roseus* for the synthesis of TiO<sub>2</sub>NPs, the synthesized  
359 nanoparticles were having rough shape and size of 25-110 nm. These NPs possess great adulticidal  
360 and larvicidal potential against *Bovicola ovis* and *Hippobosca maculate*. Similarly, the synthesis  
361 of TiO<sub>2</sub>NPs from plant *Eclipta prostrata* was reported by Rajakumar et al. [134]. Santhoshkumar  
362 et al. [135] biosynthesize TiO<sub>2</sub>NPs from *Psidium guajava* (aqueous leaf extract), antibacterial, and  
363 antioxidant potential of these nanoparticles was explored. By using 20 µg/mL of nanoparticles the  
364 highest zone of inhibition was recorded against *E. coli* and *S. aureus*. In another research conducted  
365 by Priyadarshani et al. [136], TiO<sub>2</sub>NPs of *Cissus quadrangularis* having significant bactericidal  
366 potential against *Staphylococcus* and *E. coli* were also reported. Kumar et al. [137] compared the  
367 antibacterial potential of two types of biosynthesized titanium dioxide nanoparticles one from  
368 extracts of *Hibiscus rosa sinensis* and one chemically synthesized. The author concluded that

369 nanoparticles synthesized from plant extract showed higher activity as compared to chemically  
370 synthesized ones.

#### 371 **4. Different parts used for the synthesis of metallic nanoparticles**

372 Recently, green plant-based nanotechnology has diverted the attention because of its wide  
373 applications in many different fields. The metallic NPs like gold, zinc, silver, platinum, nickel,  
374 copper, titanium oxide, and magnetite were biosynthesized by using various plant parts and are  
375 being studied extensively. Stem, fruit, root, seed, peel, callus, flower, and gums are the different  
376 parts that are utilized for the synthesis of nanoparticles with various sizes and shapes by biological  
377 methodologies [138].

##### 378 **4.1. Fruit**

379 Fruit bodies of *Tribulus terrestris* and solution of silver nitrate were utilized for the eco-friendly  
380 biosynthesis of silver NPs [88]. The presence of different phytochemicals in the plant extracts  
381 causes the formation of nanoparticles in only one step of reduction. The shape of synthesized NPs  
382 was spherical and exhibit excellent antimicrobial potential against the multidrug-resistant human  
383 pathogens. Similar research was performed by Amarnath et al. [139] in which polyphenol from  
384 grapes was utilized for the formation of palladium NPs and these nanoparticles act very efficiently  
385 against different bacterial diseases. Fruit extracts of *Rumex hymenosepalus* act as reducing and  
386 stabilizing agents in the biosynthesis of silver nanoparticles.

##### 387 **4.2. Stem**

388 The methanolic extract of *Callicarpa maingayi* (stem) was used for the biosynthesis of AgNPs  
389 [140]. The prepared extract consists of a group aldehyde that mainly involves in the process of  
390 reduction of Ag into silver nanoparticles. The functional groups such as amide I and polypeptides  
391 are considered as the responsible groups for the capping of ions to metallic NPs. The molecular  
392 findings on the synthesis of Ag crystals are complicated and still not completely known. According  
393 to some latest studies, AgNPs bind with the proteinaceous outer cell of fungi and bacteria that  
394 results in the breakage of lipoproteins of the microbial cell wall. This is followed by the blockage  
395 of cell division and leads to cell death. Vanaja et al. [141] stated the photosynthesis of silver  
396 nanoparticles by using extracts of *Cissus quadrangularis*. The functional groups mainly the amine,  
397 phenolics, and carboxyl are directly involved in the reduction reaction. The synthesized NPs  
398 possess good antibacterial potential towards *Bacillus subtilis* and *Klebsiella planticola* (pathogenic  
399 bacteria).

### 400 4.3. Seeds

401 The seed extract of fenugreek contains many naturally occurring bioactive compounds like lignin,  
402 saponin, vitamins, and high content of flavonoids. In the presence of reducing agents, this extract  
403 of seed acts as a good surfactant in the process of reduction of chloroauric acid for the formation  
404 of nanoparticles. Seed extracts of different plants contain some functional groups such as COO<sup>-</sup>  
405 group, C,C, and C,N. This functional group serves as a surfactant of AuNPs and flavonoids easily  
406 stabilizes the electrostatic stabilization during the synthesis of AuNPs [74]. *Macrotyloma*  
407 *uniflorum* (aqueous extract) increases the rate of reduction of Ag ion in the biosynthesis of AgNPs.  
408 Caffeic acid present in the seed extracts is recognized to be the reason for the increase in the  
409 reduction rate. Hence, the present caffeic acid completes the reduction reaction of nanoparticle  
410 synthesis within a minute.

### 411 4.4. Flowers

412 Petals of rose were utilized for the formation of gold nanoparticles through an eco-friendly method  
413 by Noruzi et al. [142]. This extract contains an ample amount of proteins and sugars that are  
414 considered to be the main constituents during the reduction of salt tetrachloroaurate into the bulk  
415 of gold nanoparticles. Likewise, flowers of *Clitoria ternatea* and *Catharanthus roseus* are also  
416 utilized for the biosynthesis of different metallic nanoparticles of desired sizes and shapes. Vankar  
417 and Bajpai, [143] also synthesize AuNPs by using the flower extract of *Mirabilis jalapa*.

### 418 4.5. Leaves

419 Extracts of plant leaves also serve as a facilitator in the biosynthesis of different metallic  
420 nanoparticles. Leaves of several plants like *Alternanthera sessilis*, *Murraya koenigii*, and *Centella*  
421 *asiatica* have been used for the said purpose. Recently, *Piper nigrum* leaves were utilized for  
422 successful biosynthesis of silver nanoparticles through an eco-friendly protocol, as they contain  
423 important bioactive products. The silver nanoparticles have an effective role in cancer medicine  
424 for the treatment of different dreadful diseases. The presence of piper longminine and longumine  
425 in *Piper nigrum*, acts as the agent for capping in preparation of silver nanoparticles and can also  
426 improve the cytotoxic effects of the tumour cells [144]. Another green preparation of AgNPs from  
427 *Artemisia nilagirica* (leaves extract) was described by Vijayakumar et al. [145]. These metallic  
428 nanoparticles serve as a significant antimicrobial agent. Similarly, AgNPs formed from the  
429 different plant leaves can control many pathogenic problems in humans.

### 430 5. Conclusions

431 In this chapter, we have summarized information about the green and environmentally responsive  
432 production of various metal nanoparticles. The presence of biologically active molecules and  
433 secondary metabolites in plant extracts is responsible to reduce metal ions to nanostructures in a  
434 rapid single-step synthetic strategy. A large number of plant species, their parts, and derived extract  
435 has been effectively employed for the preparation of numerous kinds of nanoparticles and other  
436 nanostructured materials due to the occurrence of abundant metabolic compounds, including  
437 phenols, alkaloids, carbohydrates, terpenoids, and bio-enzymes. It is worth noting that the  
438 utilization of plant sources for nanoparticle biosynthesis obviates the requirement for capping and  
439 stabilizing agents. Taking into consideration the inescapable applications of nanotechnology and  
440 nanoscience in a range of modern everyday life, additional research is necessary to explore the  
441 unique physical and chemical attributes of newly-synthesized nanostructures. From the future  
442 development perspective, Figure 4 shows the involvement of nanotechnology to further advance  
443 the characteristics features and functionalization of nanoparticles [146,147].

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