

Green Synthesis, Characterization, Antimicrobial Activity and Applications of Cu, and CuO, Nanoparticles

D. T. Sakhare

*U.G., P.G. & Research Centre, Department of Chemistry, Shivaji, Art's, Comm. & Science
College Kannad. Dist. Aurangabad.431103, Maharashtra, India.*

E-mail- sakharedhondiram@yahoo.com

Abstract

Green method for the synthesis of metallic and metallic oxide nanoparticles has been known to play a significant role in the advancement of nanomaterial applications in various sectors since last few decades. Green synthesis of copper and copper oxide nanoparticles (Cu and CuO, NPs) was more economical and an alternative method to many existing physical and chemical methods. Green fabricated Cu and CuO NPs finds applications in catalysis, photocatalysis, electrocatalysis, organic dye degradation, biomedicine, pharmaceuticals, cosmetics and energy. The morphology and topography of biogenic NPs widely depend on the bioactive components of the plant extracts and preparative conditions. The present review aims to bring awareness about various plants applied in the biogenic synthesis of Cu and its oxide NPs for multifunctional applications in antimicrobial activity evaluation and catalysis. The characterization of biogenic Cu NPs by number of analytical tools for their compositional, morphological and topographical features has also been discussed.

Keywords:

Green Synthesis, Copper Nanoparticles, Antimicrobial Activity, Morphology, Catalysis, Application.

1. Introduction

Copper (Cu) is one of the indispensable microelements obligatory for the growth and development of plant. It can be present as Cu^{2+} and Cu^+ under natural conditions. Optimum concentration is regularly involved in the plants, ranging from 10^{-14} to 10^{-16} M. In addition to

many of its important functions such as cell wall metabolism and protein regulation, it also acts as secondary signaling molecule in plant cells. It takes part in the mitochondrial respiration, photosynthetic electron transport, iron mobilization, hormone signaling, oxidative stress response, and also acts as cofactor for many enzymes [1, 2]. However, a higher dose of Cu leads to oxidative stress generation, growth inhibition, cellular malfunctioning, and photosynthesis retardation [2–4].

Though, nanotechnology is considered to be the next industrial revolution, the search is still ongoing for better nontoxic, hazard free, and eco-friendly approach for synthesis of nanomaterials [5]. In comparison to chemically synthesized nanoparticles, green synthesized nanoparticles are more effective and eco-friendly [6]. Until now various biological organisms are reported to have their potentiality for the production of metallic nanoparticles. However, the rate of metal nanoparticle synthesis with the help of plant extract is stable [7], much faster [8,9], and extremely mono-dispersive [10] in respect to other biological methods. Moreover, in the case of copper, different organisms such as microbes [11,12], algae [13], fungi [14,15], and angiosperm plant extracts [16,17], are utilized for the nanoparticles production.

A metallic nanoparticle are multifunctional in nature and hence finds huge number of applications in various sectors for environmental, biomedical and antimicrobial, solar power generation and catalytic causes.[18]. Application of plant extracts to synthesize copper and its oxide nanoparticles is a green chemistry methodology which establishes strong relationship between natural plant material and nanosynthesis. [19].

It has been reported in the past study that copper, gold and silver nanoparticles exhibited excellent antimicrobial activity against various disease causing pathogens. In recent years, copper nano particles (Cu NPs) have gained significance due to their multifunctional uses in industries and medicine. However, other nanoparticles, such as platinum, gold, iron oxide, silicon oxides and nickel have not shown bactericidal effects in studies with *Escherichia coli*. [20] The antibacterial study on *E. coli* and *Bacillus subtilis* using Cu and Ag NPs, revealed the fact that Cu exhibited superiority over Ag.[21] Cu NPs have wide applications as heat transfer systems[22] antimicrobial materials[23], sensors[24], and catalysts.[25]. In addition, copper and its compound have been applied as antifungal, antiviral, and molluscicidal agents. The synthesis of Cu NPs by using extracts of various plants found all over the globe have been reported by many researchers in the past.[26,27].

It is essential to develop clean, reliable, biocompatible, cheap, and nontoxic green method of nanoparticle's synthesis. Many plants parts or whole plants have been used for the green synthesis of Cu NPs [28] due to the presence of large number of bioactive compounds in plants. The extracts of plants *Nerium oleander* [29], *Punica granatum*[30], *Aegle marmelos*[31] & *Ocimum sanctum*,[32] *Zingiber officinale*[33] have been efficiently applied for this purpose.

The present work concentrates on biogenic synthetic processes for Cu NPs using extracts of diverse range of plant species including medicinal plants found across the globe and their applications in electronic, magnetic, optoelectronic, biomedical, pharmaceutical, cosmetic, energy and catalysis.

2. Nanoparticle Synthesis Methods

Bottom-up and top-down are the two approaches recommended for the biosynthesis of nanoparticles [34]. In case of a bottom-up approach, the most important reaction occurred is oxidation/reduction. The synthesis of nanoparticles is currently an important area of research, which seeks an eco-friendly approach and green materials for current scenario [35]. The major steps involved in the preparation of nanoparticles that have to be evaluated from the point of green chemistry are (i) the solvent medium used for the synthesis, (ii) environmentally benign reducing agent, and (iii) the nontoxic material for the stabilization of the nanoparticles. The majority of the chemical and physical methods mentioned so far largely depend on organic solvents. This is principally due to the hydrophobicity of the capping agents used [36]. Synthesis with bio-organisms is compatible with the principles of green chemistry: (i) ecofriendly approach, (ii) the reducing agent used, and (iii) the capping agent in the reaction. The synthesis of inorganic metal oxide nanoparticles using biological elements has received immense attention due to their unusual properties (optical, electronic, chemical, etc.) [37].

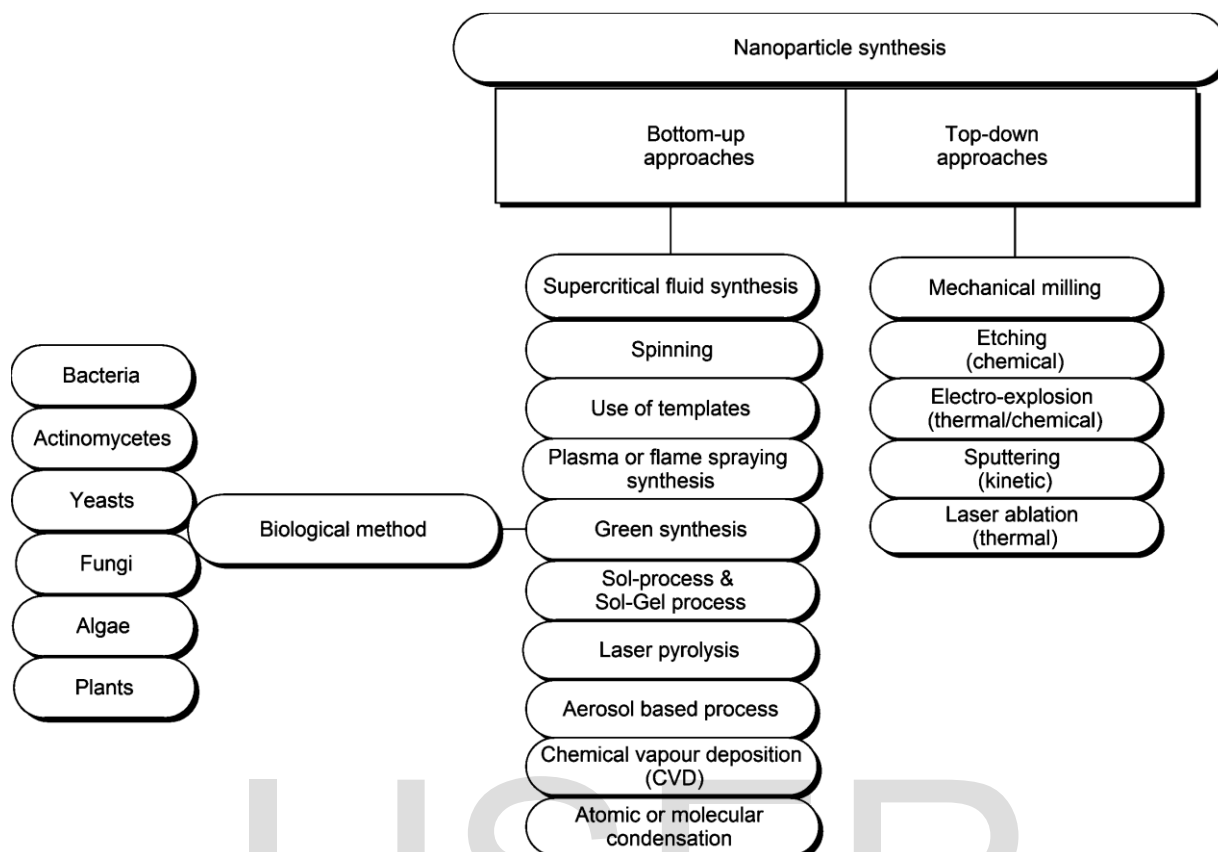


Fig. 1 Some important manufacturing methods used in nanoparticle synthesis.

3. Green Synthesis of Cu and CuO NPs

Plants consist of large number of biologically active compounds and hence, most of the plants have proven record for their anthelmintic, antitumor, antimutagenic, antibacterial and fungicidal properties. The synthesis of metallic NPs involves simple mixing of metal solution with extract of plant. Nanoparticles are produced in the medium due to reduction of metal ions. The scheme of synthesis of metallic NPs is as shown in Fig. 2.

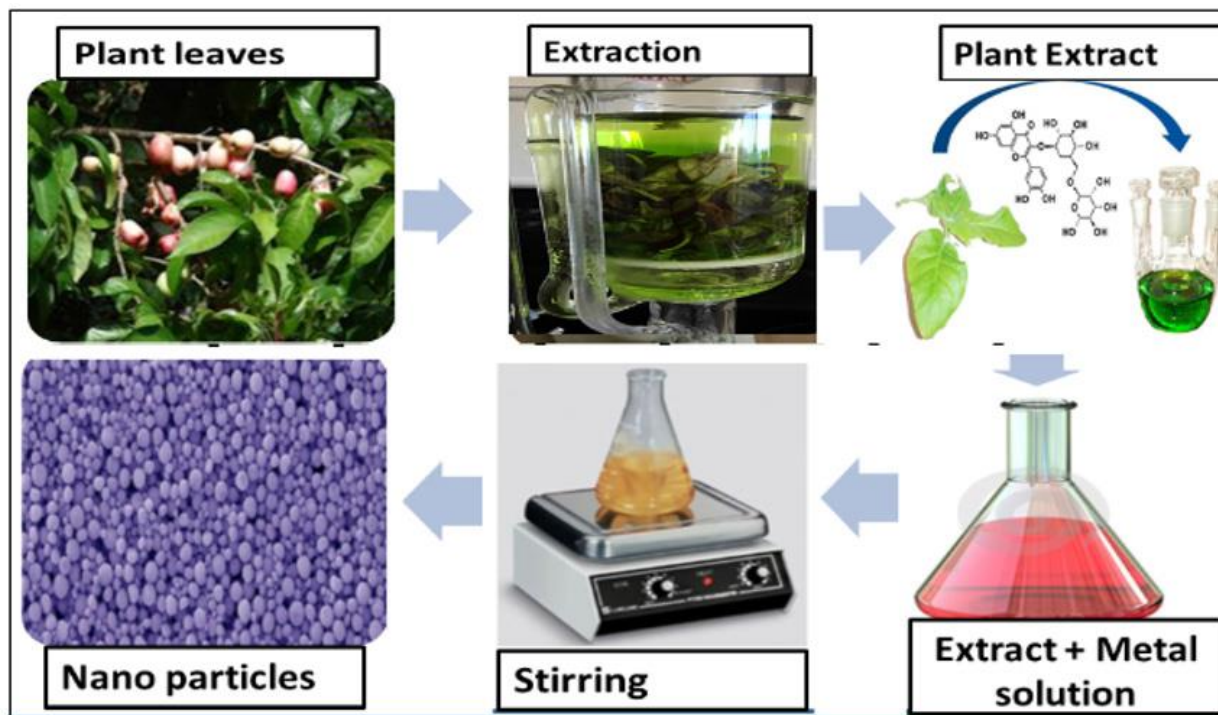








Fig. 2: A schematic diagram of green synthesis of metal nanoparticles from plant extracts










Many earlier investigations revealed that Cu NPs can be synthesised by the application of most common precursor copper salts namely, cupric acetate (monohydrate) $((CH_3COO)_2Cu \cdot H_2O)$ [38], Copper chloride di-hydrate $(CuCl_2 \cdot 2H_2O)$ [39] and Copper sulfate pentahydrate $(CuSO_4 \cdot 5H_2O)$ [40]. Various factors such as concentration, pH, temperature, influence the nature and properties of synthetic Cu NPs as well as CuO NPs.

The reduction of copper ions to get stable copper nanoparticles can be attributed to the presence of biologically active compounds present in the leaf broth of *Azadirachta indica*. [41]. It was found in this study that the rate of production varied linearly with percentage of leaf broth. The other optimum conditions for the synthesis are; $[CuCl_2] = 7.5 \times 10^{-3} \text{ M}$, $pH = 6.6$ and temperature = 85°C .





Formation of green synthesized copper nano particles capped with *T. cordifolia* (Cu NPs@Tc) was also reported.[42]. Synthesis of Cu nanoparticles has been successful with extracts of various parts of plant species that include, *Citrus medica* Linn. (Idilimbu) juice,[43] *Ziziphus spina-christi* (L.) Willd,[44] *Asparagus adscendens* Roxb. Root and Leaf,[45] *Eclipta prostrata* leaves,[46] *Ginkgo biloba* Linn,[47] *Plantago asiatica* leaf, [48] *Thymus vulgaris* L,[49] black tea leaves[50], *Terminalia catappa* leaf[51] presented in Table 1.

Table 1: Various plants extracts used in the synthesis of Cu and CuO NPs and their applications

Sr, No	Plant	Image	Precursor	Size, morphology, surface plasmon vibration (SPV)	Applications	Ref
1	Syzygium aromaticum bud		Cupriacetate (monohydrate) $((CH_3COO)_2Cu \cdot H_2O)$	-12 nm, spherical SPV@ <580 nm	Antimicrobial properties	38
2	Stachys lavandulifolia		Copper chloride di-hydrate $(CuCl_2 \cdot 2H_2O)$	80 ± 8 nm, near spherical, SPV@ ~ 590 nm	antibacterial activity	39
3	black bean		Coppersulfate pentahydrate $(CuSO_4 \cdot 5H_2O)$	~26.6nm, spherical, hexagonal and uneven shapes,	anticancer activity	40
4	Azadirachta indica leaves		Cupric chloride di-hydrate $(CuCl_2 \cdot 2H_2O)$	48nm, cubical, SPV@ ~ 506 nm	-	41
5	Tinospora cordifolia		Copper chloride $(CuCl_2(II))$	50–130nm, spherical, SPV@ ~250 nm	Catalytic Degradation	42
6	Citrus medica Linn. (Idilimbu) juice		Copper sulphate $(CuSO_4)$	33nm, SPV@ ~ 631 nm	antimicrobial activity	43
7	Ziziphus spina-christi (L.) Willd		Copper sulphate $(CuSO_4)$	8–15nm, spherical, SPV@ ~ 551 nm	triphenylmethane dye and antibacterial assay	44

8	Asparagus adscendens Roxb. Root and Leaf		Copper sulphate (CuSO ₄)	10–15 nm, Spherical, SPV@ ~ 500 to 700 nm	Antimicrobi al Activities	45
9	Eclipta prostrata leaves		Copper acetate (Cu(OAc) ₂)	31±1.2nm, spherical hexagonal and cubical SPV@ ~ 565 nm	antioxidant and cytotoxic activities	46
10	Ginkgo biloba Linn		Copper chloride (CuCl ₂)	15-20nm, spherical, SPV@560 to 580 nm	catalytic activity	47
11	Plantago asiatica leaf		Cupric chloride di-hydrate, (CuCl ₂ · 2H ₂ O)	7–35nm, spherical, SP V@ 565 nm	catalytic activity	48
12	Thymus vulgaris L.		Copper sulphate (CuSO ₄)	various sizes, sheeted,SPV @ ~ 520 nm	catalytic activity (MB)	49
13	Black tea leaves		Copper sulphate (CuSO ₄)	26–40nm, Spherical,	antibacteria l, antifungal aflatoxin B ₁	50
14	Terminalia catappa leaf		Copper sulphate pentahydrate (CuSO ₄ 5H ₂ O)	21–30 nm, Spherical,	Antibacteri al test	51
15	Rheum palmatum L.		Copper chloride (CuCl ₂)	10–20 nm, Spherical, SPV@ ~ 250–300 nm	catalytic activity	52
15	Aloe vera extract		Copper sulphate CuSO ₄	15 and 30 nm, dispersed, versatile and spherical, SPV@ ~ 265 and 285 nm	-	53

16	Oak Fruit Hull (Jaft)		Copper acetate ($\text{Cu}(\text{CH}_3\text{COO})_2$)	34nm, quasi-spherical, SPV@ ~ 590 nm	Photocatalytic Degradation (Violet 3)	54
17	Ixoro coccinea leaf		Copper sulphate CuSO_4	80–110 nm, Spherical, SPV@ ~ 191 nm	-	55
18	Syzygium alternifolium (Wt.) Walp.		Copper sulphate pentahydrate ($\text{CuSO}_4 \cdot 5\text{H}_2\text{O}$)	17.5 nm, spherical, SPV@ ~ 285 nm	Antiviral Activity	56
19	Ferulago angulata (schlecht) boiss		Copper acetate ($\text{Cu}(\text{CH}_3\text{COO})_2$)	~44nmSPV @ ~ 554 nm	Photocatalytic degradation of Rhodamine B	57
20	Rosa canina fruit		Cupric acetate, $\text{Cu}(\text{OAc})_2$	Spherical 15-25SPV@ ~ 262 nm	C-N Ullmann coupling reactions	58
21	Azadirachta indica		Copper nitrate Trihydrate ($\text{Cu}(\text{NO}_3)_2 \cdot 3\text{H}_2\text{O}$)	28-35 nm, spherical SPV@ ~ 262nm	Antibacterial activity (E. coli)	59
22	Olea europaealeaf		Copper sulphate CuSO_4	20–50 nm, Spherical, SPV@ ~ 289 nm	toxicity activities	60
23	Malus Domestica leaf		Copper sulphate CuSO_4	18 - 20 nm, spherical and crystalline, SPV	antibacterial, antioxidant, DNA	61
24	Bauhinia tomentosa leaf		Copper sulphate CuSO_4	22-40nm, Clustered & spherical, SPV@ ~ 384 nm	antibacterial 1	62
25	Moringa oleifera Leaves		Copper sulphate pentahydrate ($\text{CuSO}_4 \cdot 5\text{H}_2\text{O}$)	6 and 61 nm	Nitrates Removal	63

26	Abutilon indicum leaf		Copper(II) nitratetri hydrate ($\text{Cu}(\text{NO}_3)_2 \cdot 3\text{H}_2\text{O}$)	nm range agglomerated hexagonal wurtzite, SPV@ ~ 725nm	Antimicrobial, antioxidant and photocatalytic dye degradation activities	64
27	Eclipta prostrata leaves		Cupric acetate, $\text{Cu}(\text{OAc})_2$	$31 \pm 1.2\text{nm}$, face-centered cubic structure, SPV@ ~565 nm	antioxidant and cytotoxic activities	65
28	Calotropis procera		Copper sulphate pentahydrate ($\text{CuSO}_4 \cdot 5\text{H}_2\text{O}$)	15–20 nm, quasi-spherical, SPV@ ~ 565 nm	adsorptive of Cr(VI)	66
29	Euphorbia Chamaesyce leaf		Copper chloride (CuCl_2)	~ 36–40 face-centered cubic nm, Spherical (fcc), SPV@ ~ 325 nm	catalytic activity (4nitrophenol)	67

The organic hydrocarbon part of citron²⁴ facilitates the protection of copper. The antioxidant nature and acidic property of citron also prevents oxidation of copper as the protons present in the medium influence electro-deposition of copper at low pH range. Cu colloid formation of non-oxidized Cu NPs was observed with uv-absorption peak at 565 nm during the green synthesis of Cu NPs by Eclipta prostrata leaves extract.[46]

The reduction of Cu^{2+} ions to Cu NPs by the phenolics and other chemicals of Thymus vulgaris L. leaf extract was reported.[49] These biomolecules cogently reduce copper salts but also avoid agglomeration. The hydroxyl and ketonic groups of phenolic compounds bind to metals and show chelate effect. Flavonoids can directly scavenge molecular species of active oxygen. Antioxidant action of flavonoids resides mainly in their ability to donate electrons or hydrogen atoms.

The reduction of copper ions to give Cu NPs was attributed to the phenolic compounds present in the Rheum palmatum L. root extract.[52] The extract constituents believed to function both as reducing and capping agents in the stabilization of prepared Cu NPs.

The bioactive molecules present in the Carica papaya leaves extract⁶¹ also found to reduce precursor copper sulphate to form copper oxide nanoparticles.

In the similar way CuO NPs were also synthesised by using plant extracts of Aloe vera[53], Oak fruit hull (Jaft),[54] Ixoro coccinea leaf,[55] Syzygium alternifolium (Wt.) Walp,[56] Ferulago angulata (schlecht) boiss,[57] Rosa canina fruit,[58] Azadirachta indica,[59] Olea europaea leaf extract,[60] Malus Domestica leaf extract,[61] Bauhinia tomentosa leaves extract,[62] Moringa oleifera leaves Extract,[63] Abutilon indicum leaf extract,[64] Eclipta prostrata leaves extract,[65] Calotropis procera[66], Euphorbia Chamaesyce leaf extract[67] as above Table 1.

4. Characterization of Cu and CuO NPs

The Characterization of biogenically synthesized Cu and CuO nanoparticles has been carried out by using analytical tools namely, uv-visible spectroscopy (uv-vis), x-ray diffraction (XRD), energy dispersive x-ray spectroscopy (EDS), dynamic light scattering (DLS), scanning electron microscopy (SEM), tunneling electron microscopy (TEM), Fourier transform infra-red spectroscopy (FTIR), particle analyzer, Surface Plasmon resonance etc. The uv– vis absorption spectroscopy was applied to detect color change in Cu nanoparticle synthesised by using Ziziphus spina-christi leaves [44] which is possibly due to the surface plasmon vibrations. The surface plasmon vibration bands for Cu-NPs synthesised by many plant extracts was found to be between 191 nm and 721 nm as given in Table 1.

XRD patterns (Fig. 3) obtained for the Cu NPs synthesized using citron juice and Aloe vera extract showed intense peak confirming crystalline copper [43] and crystalline CuO NPs[33,50] (Fig.4) respectively.

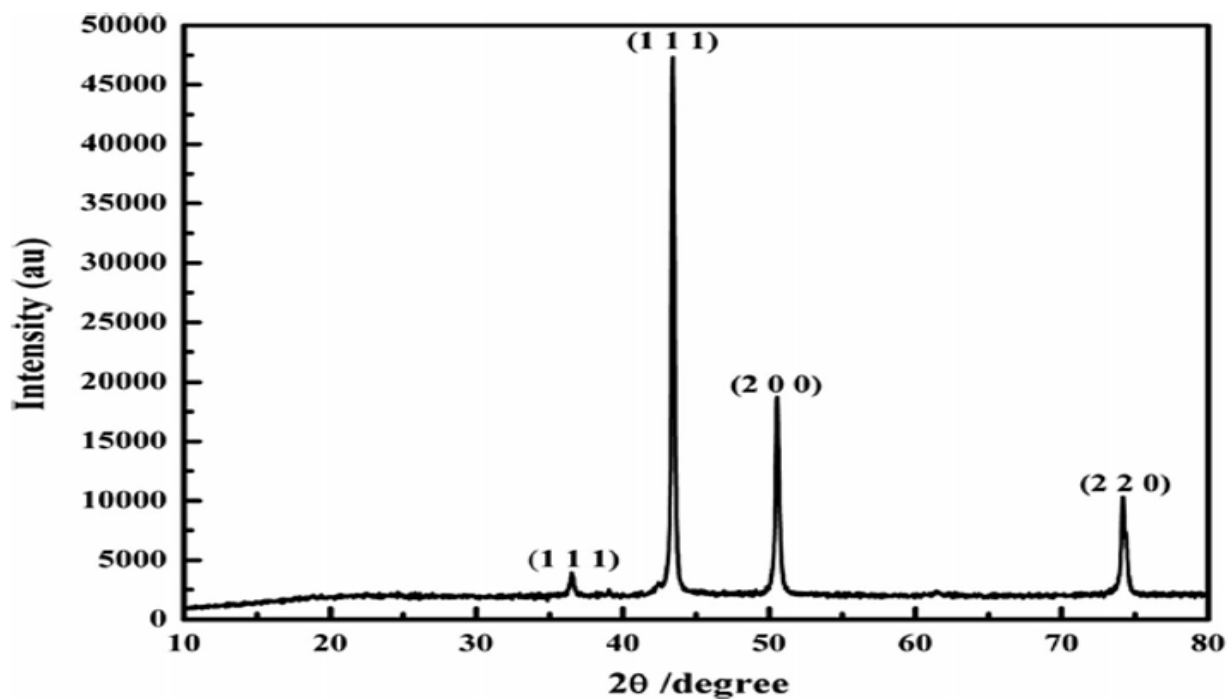


Fig. 3: XRD pattern of Cu NPs showing the FCC structure of crystallite

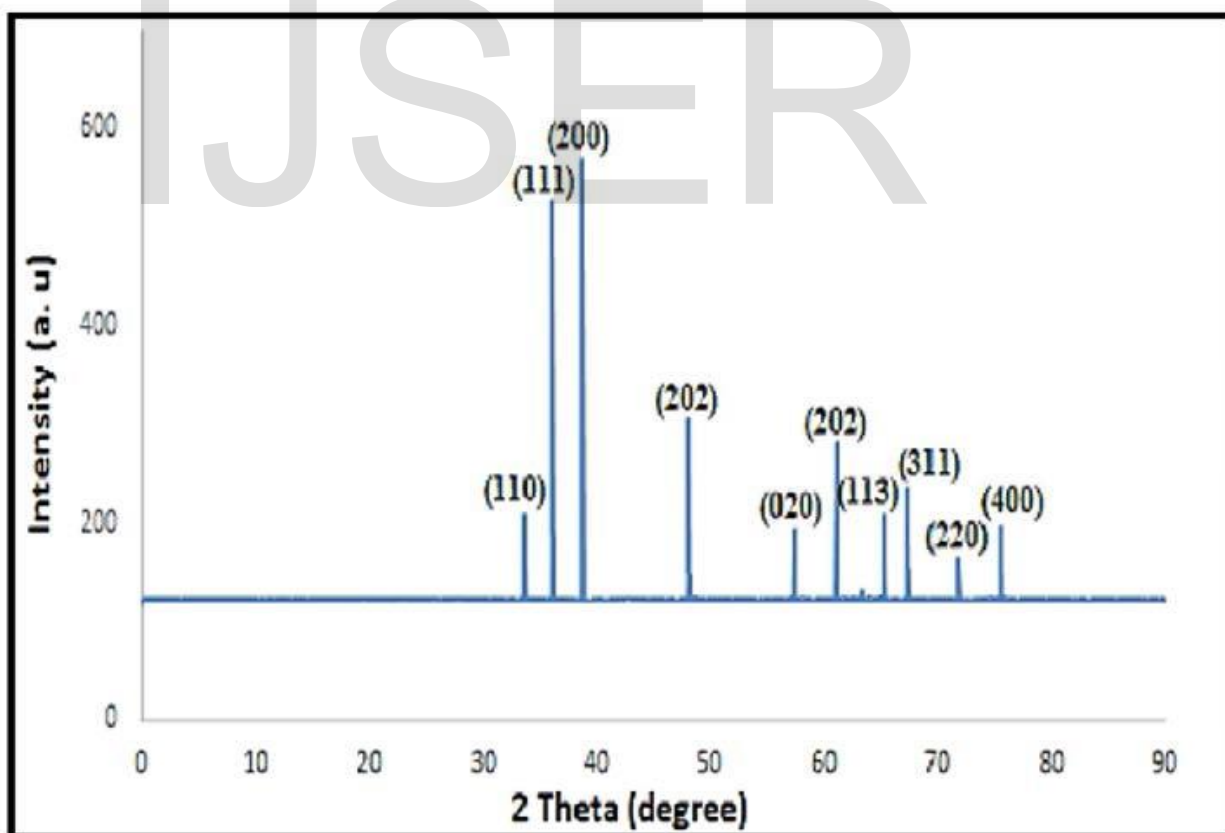


Fig. 4: XRD pattern of green synthesised CuO NPs

The analysis of FTIR spectra provides information about functional groups of biomolecules present in plant extracts. IR peaks were observed at 3,333 cm^{-1} for the hydroxyl group (H-bonded OH stretch); 2,917 cm^{-1} for methylene C-H asym. / sym. stretch; 1,615 cm^{-1} for aromatic ring stretch.[46] The peaks at about 3400, 1650, 1595, 1400 and 1100 cm^{-1} corresponds to -OH, C=O, C=C, C-OH and C-H vibrations.[47] The common IR bands[51] for cellulose were usually found at 3304 cm^{-1} , 2891 cm^{-1} , 1664 cm^{-1} , 1011 cm^{-1} corresponds to vibrations of -OH, CH₂, H₂O, and C-OH groups. Peaks at 610, 499 and 415 cm^{-1} confirmed Cu-O bond vibrations that support the presence of monoclinic phases of CuO synthesised by Aloe barbadensis Miller extract.[53]. The IR band recorded at about 800 cm^{-1} corresponds to C-H out of plane bending vibrations due to adsorbed phenolic compound on to the CuO NPs.[54] Electron microscopy is used for morphological characterization and internal composition of biogenic copper and silver nanoparticles. Homogeneous and spherical morphology of biogenic Cu NPs were revealed by FESEM image [44] as shown in Fig.5.

The average particle size varied from 20 nm to 500 nm. TEM micrographs also revealed spherical nature for NPs with least tendency towards agglomeration. DLS studies reveal size distribution of Cu NPs. The average particle size of NPs synthesised by using Azadirachta indica leaves was found to be around 50 nm.[41]

The spherical morphology and narrow diameter distributions [44,47] of Cu NPs were also confirmed by TEM image (Fig. 6). Biomolecules present in Aloe vera extract [53] believed to act as stabilizing and capping agent for copper caused a raise in the size of NP up to 30 nm but without change in shape.

FESEM images of CuO NPs too confirmed their spherical nature (20 nm to 300 nm). CuO NPs synthesised by Oak fruit extract exhibited average diameter of 34 nm.[54]. CuO NPs were found to exhibit agglomeration tendency thus enhancing average particle size to as high as 300 nm.[55]. HRTEM studies involving in-depth analysis of CuO NPs synthesised by using fruit extract of Syzygium alternifolium[56] recorded particle size of 2 nm. HRTEM image of CuO NPs[65] is shown in Fig.7 given below,

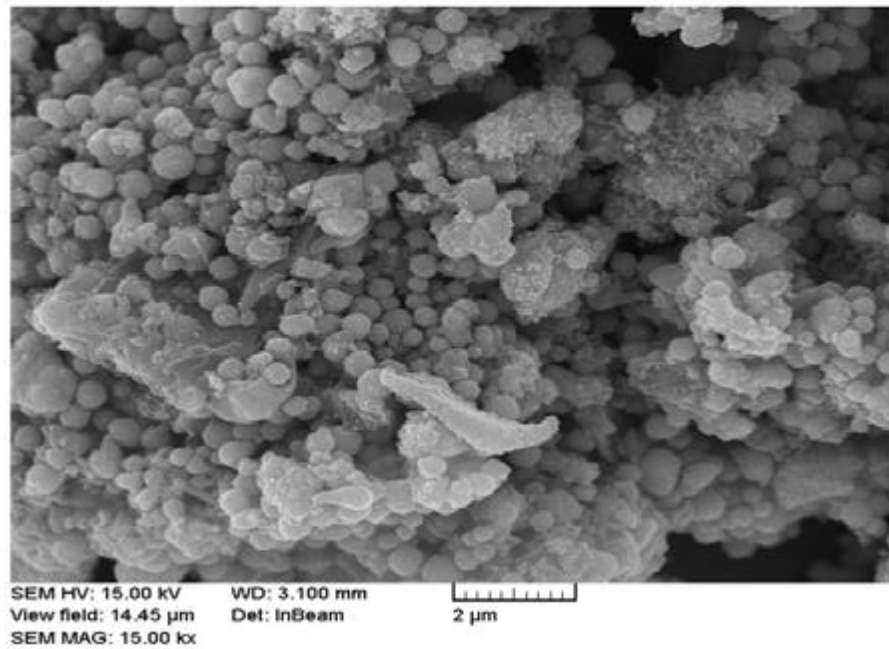


Fig. 5: FESEM image of biosynthesized Cu-NPs

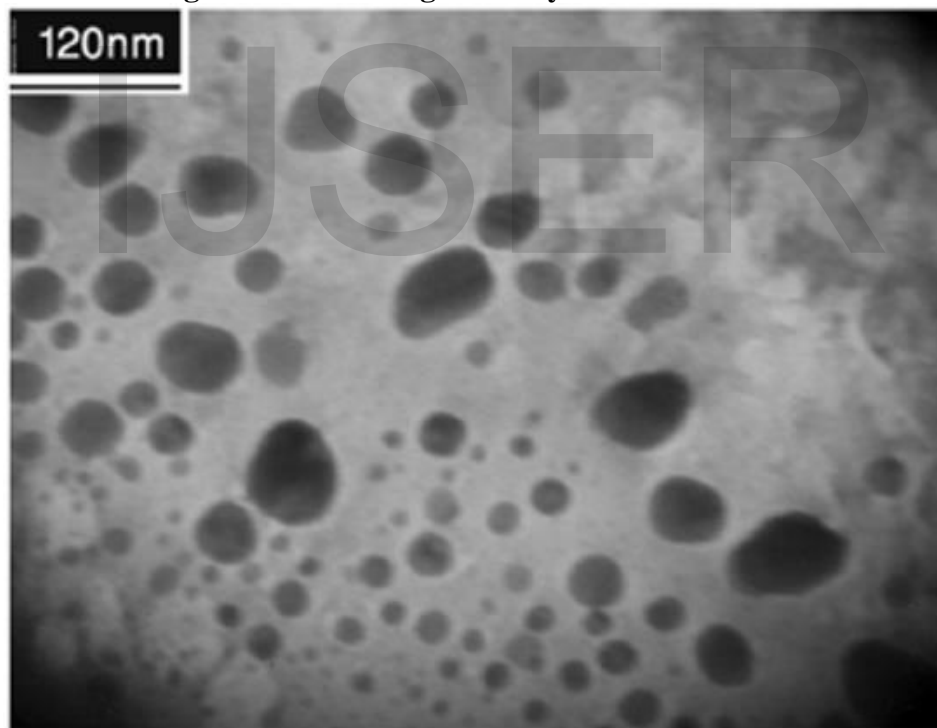


Fig. 6: TEM image of biosynthesized Cu-NPs

CuO NPs synthesised by *ferulago angulata* (schlecht) boiss extract[57] revealed shell like sheet structure. CuO NPs with relatively good monodispersed and virtually spherical structures were obtained with size range of 15 to 25 nm without agglomeration.[58] TEM analysis of CuO

NPs synthesized using *B. tomentosa* leaf extract[62] also showed spherical morphology (size of 22 to 40 nm). EDS will reveal the elemental composition of the particles. EDS spectrum of green synthesised CuO NPs is as shown in Fig. 8.

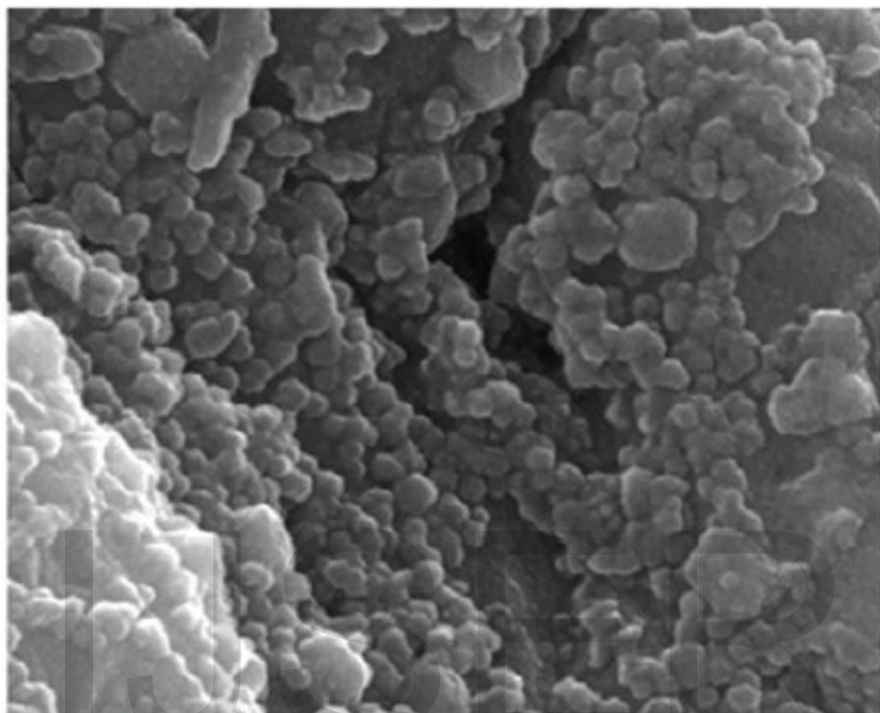


Fig. 7: HRTEM image of biosynthesized CuO-NPs

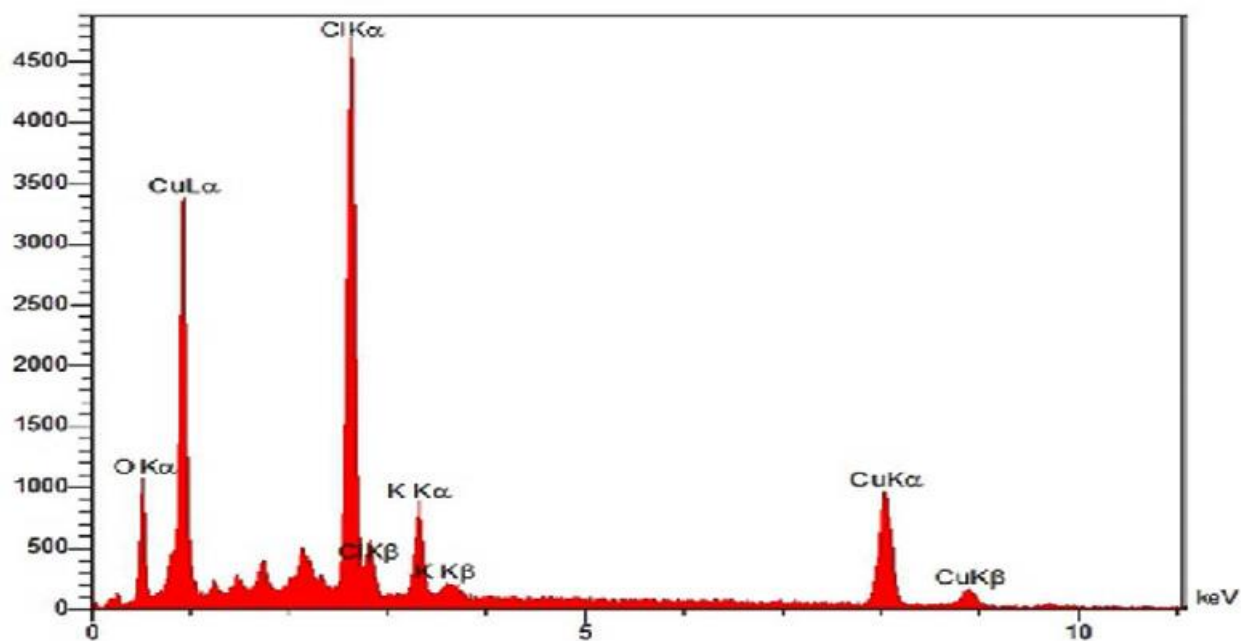


Fig. 8: XRD pattern of green synthesised CuO NPs

5. Biological Behavior: Antimicrobial Activity

Since a decade, the metal and metal oxide nanoparticles such as silver, zinc, gold, or titanium dioxide have been used as antimicrobial agents. Currently, the behavior of nanometals at nanometric sizes against pathogenic organisms is still being studied [68]. The high antimicrobial activity of the Cu NPs has been shown in multiple studies focusing on the optimal size range of about 1 to 10 nm [69, 70].

The Cu NPs have been promising for medicine and dentistry fields due to their properties, specially their interaction with pathogens, their large active surface area, and their high chemical and biological reactivity [71].

Copper is an essential element for some metabolic processes but at low concentrations because large doses could be serious consequences in the metabolic performance. It can act as electron donor or electron acceptor in some enzymes due to its redox properties increasing its toxicity for bacterium, which is induced by Cu^{+1} and Cu^{+2} ions [72–74]. Some bacteria such as *Clostridium difficile* developed important mechanisms to protect themselves from the toxic effects of copper ions while they are in contact with the surface of the Cu NPs. A possible mechanism of bacterial resistance to copper is the formation of endospores, which allows its rapid diffusion. However, the antibacterial response of the copper is being studied, mainly in human pathogens [75].

The most analyzed bacteria species in contact with Cu NPs are *C. difficile*, *E. coli*, and *P. aeruginosa*, which were significantly inhibited with a particle size between 22 and 90 nm, finding a decrease in the viability cells of these microorganisms [76–78]. Gram-positive bacteria (e.g., *S. aureus*) are more sensitive to Cu NPs and Gram-negative bacteria (e.g., *E. coli*) are associated with ROS (Reactive Oxygen Species) expression by different sizes of Cu NPs. These features could change through free surface energy of the particles directly associated with size and morphology and the pH inner of cells too [79]. The antimicrobial effect is directly related to

the nanoparticle size and minimum inhibitory concentration (MIC) (Table 2) and the oxidation degree of the surface [80].

The cell membranes of the microorganisms interact with the medium, so metal NPs especially Cu NPs will have some interactions to release metal ions that interfere with the processes of the DNA replication, cellmembranes formation, cell division, and so forth, of certain microorganisms such as bacteria, which results in an antimicrobial effect [81, 82]. The action mechanism of the copper NPs occurs through the interaction of enzymes and -SHgroups causing damage in the DNA and therefore oxidative stress generation [83–85].

Table 2: Minimum inhibitory concentrations (MIC) related to biological effect.

Sr.No.	Metal NPs	MIC	Size (nm)	Microorganism	Reference
1	Cu	140mg/mL 140–240mg/mL	6–16	<i>S. aureus</i> <i>E. coli</i>	[86]
2	Cu	3.2 ± 0.41mg/mL 1.6 ± 0.22mg/mL 3.6 ± 0.43	5–12	<i>S. aureus</i> <i>E. coli</i> <i>Salmonella typhi</i>	[87]
3	Cu	1.875–3.75mg/mL 3.75mg/mL	50	<i>S. aureus</i> <i>C. albicans</i>	[88]
4	Cu	140 µg/mL		<i>S. aureus</i>	[89,90]

5. Applications of Cu and CuO NPs

Cu and CuO nanoparticles are multifunctional in nature and hence finds significant role in applications that include antimicrobial activities, catalytic degradation, anticancer activity, photocatalytic degradation, antiviral activity, Biofilm formation, nitrates removal, upshot against human pathogens, photoluminescent activities, organic dye degradation catalysis, etc.

Cu NPs demonstrated good antimicrobial influence on *Bacillus* spp. and prominent fungicidal influence on *Penicillium* spp. microorganisms.[38] Cu NPs exhibited greater inhibition on *Escherichia coli* in comparison with *Klebsiella pneumoniae*, *Pseudomonas aeruginosa*, *Propionibacterium acnes* and *Salmonella typhi*. *Fusarium culmorum* was found to establish more sensitive plant pathogenic fungi, [45]. Cu NPs were also used for antioxidant and cytotoxic activities.[46]

Cu NPs were found to be excellent antibacterial agent against the wide range of bacterial species nanoparticles are multifunctional in nature and hence finds significant role in applications

which could be attributed to interactions of NPs with -SH groups leading to protein denaturation.[91] Cu NPs were believed to exert effect on cell membrane due to their strong affinity towards amines and carboxyl groups present on the cell surface of organisms like *B. subtilis*. [92,93] Cu NPs bind with DNA molecules and disturb the helical structure by cross-linking within and between the nucleic acid strands.[94].

CuO NPs adsorbed on cell surface of *E.coli* interact with the cell wall and later cause damage to the cell membrane, increasing its permeability and leading to a decrease in the viability of bacteria in copper oxide solution. The exact mechanism concerning antimicrobial effect of copper nanoparticles is not understood fully and needs to be elucidated.

Cu NPs green engineered by *Ginkgo biloba* L. leaf extract [47] found catalytic application for Huisgen [3 + 2] reaction. Cu- NPs synthesized with *Z. spina-christi* fruits extract proved to be excellent nanoadsorbent for Crystal Violet removal from aqueous solution.[44]

Catalytic degradation of basic violet 3 dye in water was successful with CuO NPs obtained by using extract of Oak fruit hull.[54] The extract of *Rheum palmatum* L was also used to synthesize CuO nanoparticles which proved to be efficient catalyst for degradation of 4-nitrophenol, methylene blue rhodamine B and Congo red.[95] Effective catalytic application The role of CuO as catalyst can be attributed to high surface to volume ratio associate with large number of active sites. On the same line these particles were found to be photocatalyst,[96,97] for Congo red degradation and nanocatalyst for arylation reaction.[98,99]

CuO NPs synthesised by aqueous extracts of *Anthemis nobilis* flowers[100], *Thymus vulgaris* L. leaves[101] and *Euphorbia esula* L[102] were reported to be catalytically active for the synthesis of propargylamines via aldehyde–amine–alkyne (A3) coupling reaction, and Ullmann-coupling reaction respectively. CuO particles synthesized by *Saraca indica* Leaves exhibited photoluminescence properties.[103]. Aloe vera leaf extract mediated biogenic CuO NPs exhibited the capability to serve as antimicrobial agents against fish bacterial pathogens.[104]

Cu nanoparticles and nanobiocomposites synthesised using plants animal sources found to have potential electronic device applications.[105] Cu NPs synthesized by using peel extract of *Punica granatum*[106] have demonstrated significant antibacterial inhibition against pathogens. In general, large number of plant extracts has been applied towards the green synthesis of Cu and CuO nanoparticles for applications [107-113] such as catalytic, antimicrobial activity (urinary tract), photocatalytic, antioxidant and organic dye degradation.

6. Conclusions

Green synthesis of copper and copper oxide nanoparticles has gained great significance in the recent past due to its simplicity, cost effectiveness and environment friendly nature. Large numbers of plant extracts have been successfully applied for the biogenic synthesis of Cu and CuO NPs. The bioactive compounds in the plant extracts were found to play dual role of reduction of the copper ions and also stabilization of copper NPs. UV-Visible spectroscopy, XRD, EDS, DLS, SEM, TEM, FTIR, HRTEM, Particle analyzer and Surface Plasmon Resonance are the most applied analytical tools for the characterization of copper and its oxide nanoparticles. Cu and CuO NPs were found to exhibit spherical morphology with size range of 2 - 500 nm depending on concentration of extracts as well as on preparative conditions. Cu and CuO nanoparticles proved to be multifunctional in nature with significant applications with great future implications in the fields of catalysis, photo catalysis, organic dye degradation, cosmetics, biomedicine and pharmaceuticals.

7. Acknowledgements

Authors are great full to the management of Shivaji Arts, Comm. & Science College Kannad. Dist. Aurangabad. (M.S.) India for providing support towards this work.

References

- [1]. Passam, H.C.; Karapanos, I.C.; Bebeli, P.J.; Savvas, D. A Review of Recent Research on Tomato Nutrition, Breeding and Post-Harvest Technology with Reference to Fruit Quality. *Eur. J. Plant Sci. Biotechnol.*, 1, 1–21 (2007).
- [2]. Shobha, G.; Moses, V.; Ananda, S. Biological Synthesis of Copper Nanoparticles and its impact—A Review. *Int. J. Pharm. Sci. Invent.*, 3, 28–38, (2014) .
- [3]. Yruela, I. Copper in plants. *Braz. J. Plant Physiol.*, 17, 145–156 (2005).
- [4]. Manceau, K.L.; Nagy, M.A.; Marcus, M.; Lanson, N.; Geo_roy, T.J.; Kirpichtchikova, T. Formation of metallic copper nanoparticles at the soil-root interface. *Environ. Sci. Technol.*, 1, 1766–1772 (2008).
- [5]. Filipe, V.; Hawe, A.; Jiskoot, W. Critical evaluation of Nanoparticle Tracking Analysis (NTA) by NanoSight for the measurement of nanoparticles and protein aggregates. *Pharmaceut. Res.*, 27, 796–810 (2010).

- [6]. Yedurkar, S.M.; Maurya, C.B.; Mahanwar, P.A. A Biological Approach for the Synthesis of Copper Oxide Nanoparticles by *Ixora coccinea* Leaf Extract. *J. Mater. Environ. Sci.*, 8, 1173–1178 (2017) .
- [7]. Saif, S.; Tahir, A.; Chen, Y. Green Synthesis of Iron Nanoparticles and Their Environmental Applications and Implications. *Nanomaterials*, 6, 209 (2016) .
- [8]. Makarov, V.V.; Love, A.J.; Sinitsyna, O.V.; Makarova, S.S.; Yaminsky, I.V.; Tiliansky, M.E.; Kalinina, N.O. “Green” nanotechnologies: Synthesis of metal nanoparticles using plants. *Acta Nat.* 6, 35–44 (2014).
- [9]. Ahmed, S.; Ahmad, M.; Swami, B.L.; Ikram, S. A review on plants extract mediated synthesis of silver nanoparticles for antimicrobial applications: A green expertise. *J. Adv. Res.*, 7, 17–28 (2016).
- [10]. Küüнал, S.; Rauwel, P.; Rauwel, E. Plant extract mediated synthesis of nanoparticles. In *Emerging Applications of Nanoparticles and Architecture Nanostructures*, 1st ed.; Barhoum, A., Makhoul, A.S.H., Eds.; Elsevier: Cambridge, MA, USA, 2018; pp. 411–446.
- [11]. Singh, V.; Patil, R.; Ananda, A.; Milani, P.; Gade, W. Biological Synthesis of Copper Oxide Nano Particles Using *Escherichia coli*. *Curr. Nanosci.*, 6, 365–369 (2010).
- [12]. Usha, R.; Prabu, E.; Palaniswamy, M.; Venil, C.K.; Rajendran, R. Synthesis of metal oxide nanoparticles by *Streptomyces* sp. for development of antimicrobial textiles. *Glob. J. Biotechnol. Biochem.*, 5, 153–160 (2010) .
- [13]. Abboud, Y.; Sa_aj, T.; Chagraoui, A.; Bouari, E.; Tanane, K.B.; Ihssane, O. Biosynthesis, characterization and antimicrobial activity of copperoxide nanoparticles (CONPs) produced using brown alga extract (*Bifurcaria bifurcata*). *Appl. Nanosci.*, 4, 571–576 (2013).
- [14]. Honary, S.; Barabadi, H.; Fathabad, E.G.; Naghibi, F. Green synthesis of copper oxide nanoparticles using *Penicillium aurantiogriseum*, *Penicillium citrinum* and *Penicillium waksmanii*. *Digest J. Nanomater. Biostruct.*, 7, 999–1005(2012).
- [15]. Salvadori, M.R.; Lepre, L.F.; Ando, R.A.; do Nascimento, C.A.O.; Corrêa, B. Biosynthesis and Uptake of Copper Nanoparticles by Dead Biomass of *Hypocrea lixii* isolated from the Metal Mine in the Brazilian Amazon Region. *PLoS ONE*, 8, e80519 (2013).
- [16]. Harne, S.; Sharma, A.; Dhaygude, M.; Joglekar, S.; Kodam, K.; Hudlikar, M. Novel route for rapid biosynthesis of copper nanoparticles using aqueous extract of *Calotropis procera* L. latex and their cytotoxicity on tumor cells. *Colloids Surf. B*, 15, 284–288 (2012).

- [17]. Lee, H.-J.; Song, J.Y.; Kim, B.S. Biological synthesis of copper nanoparticles using *Magnolia kobus* leaf extract and their antibacterial activity. *J. Chem. Technol. Biotechnol.*, 8, 1971–197 (2013).
- [18]. Akhtar, M. S., Panwar, J., Pilani, S. & Yun, Y. Biogenic synthesis of metallic nanoparticles by plant extracts. *ACS Sustain. Chem. Eng.* 591–60 (2013).
- [19]. Mubarakali, D., Thajuddin, N., Jeganathan, K. & Gunasekaran, M. Plant extract mediated synthesis of silver and gold nanoparticles and its antibacterial activity against clinically isolated pathogens. *Colloids Surfaces B Biointerfaces* 85, 360–365(2011).
- [20]. Williams, D. N., Ehrman, S. H. & Holoman, T. R. P. Evaluation of the microbial growth response to inorganic nanoparticles. *J. Nanobiotechnology* 4, 1–8 (2006).
- [21]. Yoon, K., Byeon, J. H., Park, J. & Hwang, J. Susceptibility constants of *Escherichia coli* and *Bacillus subtilis* to silver and copper nanoparticles. *Sci. of the Total Environ.* 373, 572–575 (2007).
- [22]. Eastman, J. A., Choi, S. U. S., Li, S., Yu, W. & Thompson, L. J. Anomalous increased effective thermal conductivities of ethylene glycol- based nanofluids containing copper nanoparticles. *Appl. Phys. Lett.* 718, 4–7 (2012).
- [23]. Guduru, R. K., Murty, K. L., Youssef, K. M., Scattergood, R. O. & Koch, C. C. Mechanical behavior of nanocrystalline copper. *Mater. Sci. Eng.* 463, 14–21 (2007).
- [24]. Xu, Q., Zhao, Y., Xu, J. Z. & Zhu, J. Preparation of functionalized copper nanoparticles and fabrication of a glucose sensor. *Sensors Actuators B* 114, 379–386 (2006).
- [25]. Rodriguez, J. A., Liu, P., Hrbek, J., Evans, J. & Pørez, M. Water Gas Shift Reaction on Cu and Au Nanoparticles Supported on CeO₂(111) and ZnO(0001): Intrinsic Activity and Importance of Support Interactions. *Heterog. Catal.* 2, 1329–1332 (2007).
- [26]. Kumar, A., Chisti, Y. & Chand, U. Synthesis of metallic nanoparticles using plant extracts. *Biotechnol. Adv.* 31, 346–356 (2013).
- [27]. Borkow, G. & Gabbay, J. Copper , An Ancient Remedy Returning to Fight Microbial , Fungal and Viral Infections. *Curr. Chem. Biol.* 3, 272–278 (2009).
- [28]. Mondal, A. K., Parui, S. M., Samanta, S. & Mallick, S. Synthesis of Ecofriendly Silver Nanoparticle from Plant Latex used as an Important Taxonomic Tool for Phylogenetic Inter-relationship. *Adv. Biores.* 2, 122–133 (2011).

- [29]. Gopinath, M., Subbaiya, R., Selvam, M. M. & Suresh, D. Synthesis of Copper Nanoparticles from Nerium oleander Leaf aqueous extract and its Antibacterial Activity. Orig. Res. Artic. 3, 814–818 (2014).
- [30]. Kaur, P., Thakur, R. & Chaudhury, A. Biogenesis of copper nanoparticles using peel extract of Punica granatum and their antimicrobial activity against opportunistic pathogens. Green Chem. Lett. Rev. 9, 33–38 (2016).
- [31]. Science, N., Technology, N. & Paper, F. Synthesis of copper nanoparticles with aegle marmelos leaf extract. NNano Sci. Nano Technol. An Indian J. 8, 401–404 (2014).
- [32]. Kulkarni, V., Rajguru, H., Rajgurunagar, M. & Kulkarni, P. Green Synthesis of Copper Nanoparticles Using Ocimum. Int. J. Chem. Stud. 1, 1–4 (2014).
- [33]. Subhankari, I. Antimicrobial Activity of Copper Nanoparticles Synthesised by Ginger (Zingiber officinale) Extract. World J. Nano Sci. Technol. 2, 10–13 (2013).
- [34] V. .irumalai Arasu, D. Prabhu, and M. Soniya, “Stable silver nanoparticle synthesizing methods and its applications,” *Journal of Biosciences Research*, vol. 1, pp. 259–270, (2010).
- [35] G. Rajakumar, T. Gomathi, M. .iruvengadam, V. D. Rajeswari, V. N. Kalpana, and I. M. Chung, “Evaluation of anti-cholinesterase, antibacterial and cytotoxic activities of green synthesized silver nanoparticles using from *Millettia pinnata* flower extract,” *Microbial Pathogenesis*, vol. 103, pp. 123–128, (2017).
- [36] P. Raveendran, J. Fu, and S. L. Wallen, “Completely “green” synthesis and stabilization of metal nanoparticles,” *Journal of the American Chemical Society*, vol. 125, no. 46, pp. 13940–13941, (2003).
- [37] P. Dhandapani, S. Maruthamuthu, and G. Rajagopal, “Biomediated synthesis of TiO₂ nanoparticles and its photocatalytic effect on aquatic biofilm,” *Journal of Photochemistry and Photobiology B: Biology*, vol. 110, pp. 43–49, (2012).
- [38]. Rajesh, K. M., Ajitha, B., Kumar, Y. A., Suneetha, Y. & Reddy, P. S. Assisted green synthesis of copper nanoparticles using Syzygium aromaticum bud extract : Physical , optical and antimicrobial properties. Opt. - Int. J. Light Electron Opt. 154, 593–600 (2018).
- [39]. Khatami, M., Heli, H., Jahani, P. M., Azizi, H. & Nobre, A. L. Copper / copper oxide nanoparticles synthesis using Stachys lavandulifolia and its antibacterial activity. IET Nanobiotechnology Res. 11, 709–713 (2017).

- [40]. Nagajyothi, P. C., Muthuraman, P., Sreekanth, T. V. M., Hwan, D. & Shim, J. Green synthesis : In-vitro anticancer activity of copper oxide nanoparticles against human cervical carcinoma cells. Arab. J. Chem. 10, 215–225 (2017).
- [41]. Nagar, N. & Devra, V. Green synthesis and characterization of copper nanoparticles using *Azadirachta indica* leaves. Mater. Chem. Phys. 213, 44–51 (2018).
- [42]. Sharma, P. et al. Green Synthesis of Colloidal Copper Nanoparticles Capped with *Tinospora cordifolia* and Its Application in Catalytic Degradation in Textile Dye : An Ecologically Sound Approach. J. Inorg. Organomet. Polym. Mater. 0, 0 (2018).
- [43]. Shende, S., Ingle, A. P., Gade, A. & Rai, M. Green synthesis of copper nanoparticles by *Citrus medica* Linn . (*Idilimbu*) juice and its antimicrobial activity. (2015). doi:10.1007/s11274-015-1840-3
- [44]. Khani, R., Roostaei, B., Bagherzade, G. & Moudi, M. Green synthesis of copper nanoparticles by fruit extract of *Ziziphus spina-christi* (L .) Willd .: Application for adsorption of triphenylmethane dye and antibacterial assay. J. Mol. Liq. 255, 541–549 (2018).
- [45]. Thakur, S., Sharma, S., Thakur, S. & Rai, R. Green Synthesis of Copper Nano-Particles Using *Asparagus adscendens* Roxb . Root and Leaf Extract and Their Antimicrobial Activities. Int. J. Curr. Microbiol. Appl. Sci. 7, 683–694 (2018).
- [46]. Chung, I. L. L. M. I. N. et al., Green synthesis of copper nanoparticles using *Eclipta prostrata* leaves extract and their antioxidant and cytotoxic activities. Exp. Ther. Med. 14, 18–24 (2017).
- [47]. Nasrollahzadeh, M. & Sajadi, S. M. Green synthesis of copper nanoparticles using *Ginkgo biloba* L . leaf extract and their catalytic activity for the Huisgen [3 + 2] cycloaddition of azides. J. Colloid Interface Sci. (2015). doi:10.1016/j.jcis.2015.07.004
- [48]. Cn, K. F., Nasrollahzadeh, M., Samaneh, S. & Sajadi, S. M. Green synthesis of copper nanoparticles using *Plantago asiatica* leaf extract and their application for the cyanation of aldehydes using. J. Colloid Interface Sci. 506, 471–477 (2017).
- [49]. Issaabadi, Z., Nasrollahzadeh, M. & Sajadi, S. M. Green synthesis of the copper nanoparticles supported on bentonite and investigation of its catalytic activity. J. Clean. Prod. 142, 3584–3591 (2017).

- [50]. Asif, M. et al., Iron , copper and silver nanoparticles : Green synthesis using green and black tea leaves extracts and evaluation of antibacterial , antifungal and aflatoxin B₁ adsorption activity. LWT - Food Sci. Technol. 90, 98–107 (2018).
- [51]. Muthulakshmi, L., Rajini, N., Nellaiah, H., Kathiresan, T. & Jawaidd, M. Preparation and properties of cellulose nanocomposite films with in situ generated copper nanoparticles using Terminalia catappa leaf extract. Int. J. Biol. Macromol. 95, 1064–1071 (2017).
- [52]. Bordbar, M. & Shari, Z. Green synthesis of copper oxide nanoparticles / clinoptilolite using Rheum palmatum L . root extract : high catalytic activity for reduction of 4-nitro phenol , rhodamine B , and methylene blue. (2016). doi:10.1007/s10971-016-4239-1
- [53]. Gunalan, S., Sivaraj, R. & Venckatesh, R. Aloe barbadensis Miller mediated green synthesis of mono-disperse copper oxide nanoparticles: Optical properties. Spectrochim. ACTA PART A Mol. Biomol. Spectrosc. 97, 1140–1144 (2012).
- [54]. Sorbiun, M., Shayegan, E. & Ali, M. Green Synthesis of Zinc Oxide and Copper Oxide Nanoparticles Using Aqueous Extract of Oak Fruit Hull (Jaft) and Comparing Their Photocatalytic Degradation of Basic Violet 3. Int. J. Environ. Res. 4, (2018).
- [55]. Vishveshvar, K., Krishnan, M. V. A., Haribabu, K. & Vishnuprasad, S. Green Synthesis of Copper Oxide Nanoparticles Using Ixiro coccinea Plant Leaves and its Characterization. Bionanoscience (2018).
- [56]. Yugandhar, P., Vasavi, T., Jayavardhana, Y., Palempalli, R. & Maheswari, U. Cost Effective , Green Synthesis of Copper Oxide Nanoparticles Using Fruit Extract of Syzygium alternifolium (Wt .) Walp ., Characterization and Evaluation of Antiviral Activity. J. Clust. Sci. 1, (2018).
- [57]. Shayegan, E., Mina, M., Ali, S., Saeid, R. & Fardood, T. Plant-mediated synthesis of zinc oxide and copper oxide nanoparticles by using ferulago angulata (schlecht) boiss extract and comparison of their photocatalytic degradation of Rhodamine B (RhB) under visible light irradiation. J. Mater. Sci. Mater. Electron. 0, 0 (2017).
- [58]. Hemmati, S., Mehrazin, L., Hekmati, M., Izadi, M. & Veisi, H. Biosynthesis of CuO nanoparticles using Rosa canina fruit extract as a recyclable and heterogeneous nanocatalyst for C-N Ullmann coupling reactions. Mater. Chem. Phys. 214, 527–532 (2018).
- [59]. Green synthesis of CuO nanoparticles using Azadirachta indica and its antibacterial activity for medicinal applications. Mater. Res. Express (2018). doi:MRX-109160.R1

- [60]. Sulaiman, G. M., Tawfeeq, A. T. & Jaaffer, M. D. Biogenic synthesis of copper oxide nanoparticles using *Olea europaea* leaf extract and evaluation of their toxicity activities : An in vivo and in vitro study. *Formul. Eng. Biomater.* (2017). doi:10.1002/btpr.
- [61]. M. S. Jadhav, S. Kulkarni, P. Raikar, D. A. Barretto, S. K. V. and U. S. R. Green Biosynthesis of CuO & Ag-CuO nanoparticles from *Malus Domestica* leaf extract and evaluation of antibacterial, antioxidant, DNA cleavage activities. *New J. Chem.* (2017). doi:10.1039/C7NJ02977B
- [62]. Sharmila, G. et al. Biogenic synthesis of CuO nanoparticles using *Bauhinia tomentosa* leaves extract : Characterization and its antibacterial application. *J. Mol. Struct.* 1165, 288–292 (2018).
- [63]. Galan, C. R., Silva, M. F. & Mantovani, D. Green Synthesis of Copper Oxide Nanoparticles Impregnated on Activated Carbon Using *Moringa oleifera* Leaves Extract for the Removal of Nitrates from Water. *Can. J. Chem. Eng.* 9999, 1–9 (2018).
- [64]. Ijaz, F., Shahid, S., Khan, S. A. & Ahmad, W. Green synthesis of copper oxide nanoparticles using *Abutilon indicum* leaf extract : Antimicrobial , antioxidant and photocatalytic dye degradation activities. *Trop. J. Pharm. Res.* April 16, 743–753 (2017).
- [65]. Chung, I. L. L. M. I. N. et al. Green synthesis of copper nanoparticles using *Eclipta prostrata* leaves extract and their antioxidant and cytotoxic activities. *18 Exp. Ther. Med.* 14 14, 18–24 (2017).
- [66]. Dubey, S. & Chandra, Y. *Calotropis procera* mediated one pot green synthesis of Cupric oxide nanoparticles (CuO - NPs) for adsorptive removal of Cr (VI) from aqueous solutions. *Appl. organometallic Chem.* 1–15 (2017). doi:10.1002/aoc.3849
- [67]. Maham, M., Sajadi, S. M., Kharimkhani, M. M. & Nasrollahzadeh, M. Biosynthesis of the CuO nanoparticles using *Euphorbia Chamaesyce* leaf extract and investigation of their catalytic activity for the reduction of 4-nitrophenol. *IET Nanobiotechnology* 11, 766–772 (2017).
- [68] A. Thit, H. Selck, and H. F. Bjerregaard, “Toxicity of CuO nanoparticles and Cu ions to tight epithelial cells from *xenopus laevis* (A6): effects on proliferation, cell cycle progression and cell death,” *Toxicology in Vitro*, vol. 27,no. 5, pp. 1596–1601, (2013).
- [69] J. Verran, G. Sandoval, N. S. Allen, M. Edge, and J. Stratton, “Variables affecting the antibacterial properties of nano and pigmentary titania particles in suspension,” *Dyes and Pigments*, vol. 73, no. 3, pp. 298–304, (2007).

- [70] N. Cioffi, L. Torsi, N. Ditaranto et al., “Copper nanoparticle/ polymer composites with antifungal and bacteriostatic properties,” *Chemistry of Materials*, vol. 17, no. 21, pp. 5255–5262, (2005).
- [71] R. P. Allaker and G. Ren, “Potential impact of nanotechnology on the control of infectious diseases,” *Transactions of the Royal Society of Tropical Medicine and Hygiene*, vol. 102, no. 1, pp. 1–2, (2008).
- [72] K.D. Karlin, “Metalloenzymes, structural motifs, and inorganic models,” *Science*, vol. 261, no. 5122, pp. 701–708, (1993).
- [73] G. Grass, C. Rensing, and M. Solioz, “Metallic copper as an antimicrobial surface,” *Applied and Environmental Microbiology*, vol. 77, no. 5, pp. 1541–1547, (2011).
- [74] L. Macomber and J. A. Imlay, “The iron-sulfur clusters of dehydratases are primary intracellular targets of copper toxicity,” *Proceedings of the National Academy of Sciences of the United States of America*, vol. 106, no. 20, pp. 8344–8349, (2009).
- [75] L. J. Wheeldon, T. Worthington, P. A. Lambert, A. C. Hilton, C. J. Lowden, and T. S. J. Elliott, “Antimicrobial efficacy of copper surfaces against spores and vegetative cells of *Clostridium difficile*: the germination theory,” *Journal of Antimicrobial Chemotherapy*, vol. 62, no. 3, pp. 522–525, (2008).
- [75] J. P. Ruparelia, A. K. Chatterjee, S. P. Duttagupta, and S. Mukherji, “Strain specificity in antimicrobial activity of silver and copper nanoparticles,” *Acta Biomaterialia*, vol. 4, no. 3, pp. 707–716, (2008).
- [77] K. Y. Yoon, J. Hoon Byeon, J.-H. Park, and J. Hwang, “Susceptibility constants of *Escherichia coli* and *Bacillus subtilis* to silver and copper nanoparticles,” *Science of the Total Environment*, vol. 373, no. 2-3, pp. 572–575, (2007).
- [78] S. Mehtar, I. Wiid, and S. D. Todorov, “The antimicrobial activity of copper and copper alloys against nosocomial pathogens and *Mycobacterium tuberculosis* isolated from healthcare facilities in the Western Cape: an in-vitro study,” *The Journal of Hospital Infection*, vol. 68, no. 1, pp. 45–51, (2008).
- [79] H. L. Karlsson, M. S. Toprak, and B. Fadeel, *Handbook on the Toxicology of Metals*, Elsevier, (2015).

- [80] G. Steindl, S. Heuberger, and B. Springer, *Antimicrobial Effect of Copper on Multidrug-Resistant Bacteria*, vol. 99, Austrian Agency for Health and Food Safety (AGES), Graz, Austria, (2012).
- [81] J. R. Morones, J. L. Elechiguerra, A. Camacho et al., "The bactericidal effect of silver nanoparticles," *Nanotechnology*, vol. 16, no. 10, pp. 2346–2353, (2005).
- [82] M. Souli, I. Galani, D. Plachouras et al., "Antimicrobial activity of copper surfaces against carbapenemase-producing contemporary Gram-negative clinical isolates," *Journal of Antimicrobial Chemotherapy*, vol. 68, no. 4, Article ID dks473, pp. 852–857, (2013).
- [83] N. M. Zain, A. G. F. Stapley, and G. Shama, "Green synthesis of silver and copper nanoparticles using ascorbic acid and chitosan for antimicrobial applications," *Carbohydrate Polymers*, vol. 112, pp. 195–202, (2014).
- [84] C. E. Santo, D. Quaranta, and G. Grass, "Antimicrobial metallic copper surfaces kill *Staphylococcus haemolyticus* via membrane damage," *MicrobiologyOpen*, vol. 1, no. 1, pp. 46–52, (2012).
- [85] G. Ren, D. Hu, E. W. C. Cheng, M. A. Vargas-Reus, P. Reip, and R. P. Allaker, "Characterisation of copper oxide nanoparticles for antimicrobial applications," *International Journal of Antimicrobial Agents*, vol. 33, no. 6, pp. 587–590, (2009).
- [86] J. P. Ruparelia, A. K. Chatterjee, S. P. Duttgupta, and S. Mukherji, "Strain specificity in antimicrobial activity of silver and copper nanoparticles," *Acta Biomaterialia*, vol. 4, no. 3, pp. 707–716, (2008).
- [87] M. Valodkar, P. S. Rathore, R. N. Jadeja, M. Thounaojam, R. V. Devkar, and S. Thakore, "Cytotoxicity evaluation and antimicrobial studies of starch capped water soluble copper nanoparticles," *Journal of Hazardous Materials*, vol. 201–202, pp. 244–249, (2012).
- [88] T. Kruk, K. Szczepanowicz, J. Stefańska, R. P. Socha, and P. Warszyński, "Synthesis and antimicrobial activity of monodisperse copper nanoparticles," *Colloids and Surfaces B: Biointerfaces*, vol. 128, pp. 17–22, (2015).
- [89] T. D. Pham and B.-K. Lee, "Disinfection of *Staphylococcus aureus* in indoor aerosols using Cu–TiO₂ deposited on glass fiber under visible light irradiation," *Journal of Photochemistry and Photobiology A: Chemistry*, vol. 307–308, pp. 16–22, (2015).

- [90] D. Chudobova, S. Dostalova, B. Ruttkay-Nedecky et al., "The effect of metal ions on *Staphylococcus aureus* revealed by biochemical and mass spectrometric analyses," *Microbiological Research*, vol. 170, pp. 147–156, (2015).
- [91]. Schrand AM, Rahman MF, Hussain SM, Schlager JJ, Smith DA, Syed AF. Metal-based nanoparticles and their toxicity assessment. *WIREs Nanomed Nanobiotechnol* 2:554–568 (2010).
- [92]. Beveridge TJ, Murray. RGE Sites of metal deposition in the cell wall of bacillus subtilis. *J Bacteriol* 141:876–877 (1980).
- [93]. Ren G, Hu D, Cheng EWC, Vargas-ReusMA, Reip P, Allaker RP. Characterization of copper oxide nanoparticles for antimicrobial applications. *Int J Antimicrob Agents* 33:587– 590 (2009).
- [94]. Avinash P. Ingle, Nelson Duran, Mahendra Rai. Bioactivity, mechanism of action, and cytotoxicity of copper-based nanoparticles: A review, *Appl Microbiol Biotechnol*, (2013).
- [95]. Herrera, A. & Reyes, A. Biosynthesis of Copper Oxide nanoparticles from *Drypetes sepiaria* Leaf extract and their catalytic activity to dye degradation Biosynthesis of Copper Oxide nanoparticles from *Drypetes sepiaria* Leaf extract and their catalytic activity to dye degradation. *Mater. Sci. Eng.* 263, (2017).
- [96]. Aminuzzaman, M., Kei, L. M. & Liang, W. H. Green Synthesis of Copper Oxide (CuO) Nanoparticles using Banana Peel Extract and Their Photocatalytic Activities. *Am. Inst. Phys.* 1828, (2017).
- [97]. Sankar, R., Manikandan, P., Malarvizhi, V. & Fathima, T. Green synthesis of colloidal copper oxide nanoparticles using *Carica papaya* and its application in photocatalytic dye degradation. *Spectrochim. Acta Part A Mol. Biomol. Spectrosc.* 121, 746–750 (2014).
- [98]. Veisi, H., Hemmati, S. & Javaheri, H. N -Arylation of indole and aniline by a green synthesized CuO nanoparticles mediated by *Thymbra spicata* leaves extract as a recyclable and heterogeneous nanocatalyst. *Tetrahedron Lett.* 58, 3155–3159 (2017).
- [99]. Manjari, G., Saran, S., Arun, T., Bhaskara, A. V. & Devipriya, S. P. Catalytic and recyclability properties of phytogenic copper oxide nanoparticles derived from *Aglaia elaeagnoidea* flower extract. *J. Saudi Chem. Soc.* 21, 610–618 (2017).

- [100]. Nasrollahzadeh, M., Sajadi, S. M. & Rostami-vartooni, A. Green synthesis of CuO nanoparticles by aqueous extract of *Anthemis nobilis* flowers and their catalytic activity for the A 3 coupling reaction. *J. Colloid Interface Sci.* 459, 183–188 (2015).
- [101]. Nasrollahzadeh, M., Sajadi, S. M., Rostami-vartooni, A., Mamand, S. & Rnh, A. Green synthesis of CuO nanoparticles using aqueous extract of *Thymus vulgaris* L . leaves and their catalytic performance for N -arylation of indoles and amines. *J. Colloid Interface Sci.* 466, 113–119 (2016).
- [102]. Nasrollahzadeh, M., Sajadi, S. M. & Khalaj, M. Green synthesis of copper nanoparticles using aqueous extract of the leaves of *Euphorbia esula* L and their catalytic activity for ligand-free Ullmann-. *RSC Adv.* 4, 47313– 47318 (2014).
- [103]. Prasad, K. S., Patra, A., Shruthi, G. & Chandan, S. Aqueous Extract of *Saraca indica* Leaves in the Synthesis of Copper Oxide Nanoparticles : Finding a Way towards Going Green. *J. Nanotechnol.* 2017, (2017).
- [104]. Kumar, P. P. N. V., Shameem, U. & Kollu, P. Green Synthesis of Copper Oxide Nanoparticles Using Aloe vera Leaf Extract and Its Antibacterial Activity Against Fish Bacterial Pathogens. *BioNanoSci.* 5, 135–139 (2015).
- [105]. Cheirmadurai, K., Biswas, S., Murali, R. & Thanikaivelan, P. Green synthesis of copper nanoparticles and conducting nanobiocomposites using plant and animal sources. *Electron. Suppl. Mater. RSC Adv.* 1–11 (2016).
- [106]. Kaur, P., Thakur, R. & Chaudhury, A. Green Chemistry Letters and Reviews Biogenesis of copper nanoparticles using peel extract of *Punica granatum* and their antimicrobial activity against opportunistic pathogens. *Green Chem. Lett. Rev.* 9, 33–38 (2016).
- [107]. Shaikh, R. R. & Lettre, D. P. Biosynthesis of Copper Nanoparticles using *Vitis vinifera* Leaf Extract and Its Antimicrobial Activity. *Sch. Res. Libr.* 8, 265–272 (2016).
- [108]. Delma, M. B. T., Vijila, M. B. & Rajan, M. J. Green Synthesis of Copper and Lead Nanoparticles using *Zingiber Officinale* stemextract. *Int. J. Sci. Res. Publ.* 6, 134–137 (2016).
- [109]. Sivaraj, R., Rahman, P. K. S. M., Rajiv, P., Abdul, H. & Venckatesh, R. Biogenic copper oxide nanoparticles synthesis using *Tabernaemontana divaricate* leaf extract and its antibacterial activity against urinary tract pathogen. *Spectrochim. ACTA PART A Mol. Biomol. Spectrosc.* 133, 178–181 (2014).

- [110]. Nasrollahzadeh, M., Atarod, M. & Sajadi, S. M. Green synthesis of the $\text{Cu} / \text{Fe}_3\text{O}_4$ nanoparticles using *Morinda morindoides* leaf aqueous extract : A highly efficient magnetically separable catalyst for the reduction of organic dyes in aqueous medium at room temperature. *Appl. Surf. Sci.* 364, 636–644 (2016).
- [111]. Nethravathi, P. C., Kumar, M. A. P., Suresh, D. & Lingaraju, K. *Tinospora cordifolia* mediated facile green synthesis of cupric oxide nanoparticles and their photocatalytic, antioxidant and antibacterial properties. *Mater. Sci. Semicond. Process.* 33, 81–88 (2015).
- [112]. Nazar, N. et al. Cu nanoparticles synthesis using biological molecule of *P. granatum* seeds extract as reducing and capping agent : Growth mechanism and photo-catalytic activity. *Int. J. Biol. Macromol.* 106, 1203–1210 (2018).
- [113]. Nasrollahzadeh, M., Sajjadi, M. & Sajadi, S. M. Biosynthesis of copper nanoparticles supported on manganese dioxide nanoparticles using *Centella asiatica* L. leaf extract for the efficient catalytic reduction of organic dyes and nitroarenes. *h*, 109–117 (2018).

IJSER