



## Rapid synthesis of Silver Nanoparticles with *Rheum ribes* L Fruit Peels: Anticancer and Antimicrobial Effects with Biocompatible Structures

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### ABSTRACT

Silver nanoparticles (AgNPs) are substances with a wide range of uses. Utilizing extracts obtained from the peels of *Rheum ribes* L. (Rr) fruit growing in Erzurum region, silver nanoparticles were rapidly created in this study with a quick, easy, and environmentally friendly technique without harmful processes. In order to evaluate the attributes of the synthesized Rr-AgNPs, FE-SEM or TEM micrographs were utilized to characterize their morphology. A UV-visible spectrophotometer was used to assess the highest absorbance bands of RR-AgNPs. These data were used to define RR-AgNPs, which were characterized as having

exclusively negative surface charges of -25 mV, spherical shape, maximum absorbance at 428 nm wavelength, and 96 nm size distribution. The effectiveness of the produced AgNPs for use in medical applications was assessed using the MTT technique with microdilution. Minimum inhibition concentrations of Rr-AgNPs for pathogen strains ranged from 0.03 to 0.50 mg/L. Additionally, it was discovered that AgNPs effectively suppressed malignant cells, with rates of 86.27%, 74.67%, and 73.49%, in the investigation of the anticancer effects of AgNPs. Healthy cells were not subject to any inhibitory effects at the same concentrations.

Keywords: Rr-AgNPs, Fruit peels, FTIR, TGA, UV-Vis

## 1. Introduction

Among the nanosciences, green nanotechnology is one of the most interesting areas with widespread areas of use producing and controlling elements at the molecular level (SI et al. 2020). Nanoparticles (NPs) are used in both scientific study and industrial applications, and they are crucial to both (J. Singh et al. 2018). The range of use of nanoparticles is quite wide. They are products with superior physical properties such as surface area, as well as chemically important properties (Kumar et al. 2017; Baran 2019a). Metallic nanoparticles such as zinc (Zn), palladium (Pd), gold (Au), and silver (Ag) are commonly used (Ismail et al. 2017; Baran 2019; Pandiyan et al. 2019; Baran et al. 2021).

AgNPs have uses in medical applications as antimicrobial, antioxidant, anticancer and anti-inflammatory agents. In addition, AgNPs are valuable particles in many areas such as bioremediation applications, the food industry, catalysis studies, and cosmetics (Velmurugan et al. 2014; Francis et al. 2017; V. Kumar et al. 2017; 2019; Thomas et al. 2018 Mohammadi et al. 2019; Abu-Dief et al. 2020; Arroyo et al. 2020). It is possible to obtain these valuable products using various methods. Among these methods, green synthesis methods, which are among biologically-sourced synthesis methods, have various advantages over physical, chemical, and other biologically-sourced environmentally friendly methods. Some of these advantages include being economical with low cost, no toxic chemicals used in synthesis, easy process, not requiring special conditions, and high yield of product obtained as a result of the synthesis (Al-ogaidi et al. 2017; Patil et al. 2018b; Rolim et al. 2019).

AgNPs are used in medical applications as antimicrobial and anticancer agents. Their use as an antimicrobial agent can contribute to the search for a solution to antibiotic resistance, which is a very important global problem in today's world (Kumar et al. 2017; A. Singh et al. 2018; Oliveira et al. 2019; Azmi et al. 2021). A terrible disease that has been studied extensively in the search for a cure is cancer. In addition to traditional approaches, studies about different treatment processes have attracted a great deal of attention. As a result, numerous studies have been conducted about the use of Rr-AgNPs as anticancer agents (Remya et al. 2015; Chung et al. 2016; Sarkar et al. 2018; Satpathy et al. 2018; Abu-Dief et al. 2020; Zein et al. 2020).

*Rheum ribes* L. (Rr), known as "Işgın" or "Kurdish banana", is a species belonging to the Polygonaceae family. This perennial herbaceous plant grows in the Iran-Turan floristic region. It is naturally found in rocky areas in regions such as Erzurum, Hakkari, Bingöl and Elazığ in Eastern Anatolia in Turkey. The length of its shoots can reach approximately 40 cm. The plant is beneficial against ulcers, diabetes, obesity, hypertension and digestive problems. It is frequently consumed as food for these reasons. In addition, the plant forms the raw material for many medicines in the Middle East. The plant contains significant amounts of phytochemical components such as phenolic acid and flavonoids. These phytochemicals have antimicrobial, anticancer and antioxidant bioactivities ( Munzuroğlu 2000; Tosun & Kizilay 2003; Naqishbandi et al. 2009; Çınar Ayan et al. 2020).

This study used an extract from the fruit peel of Rr cultivated in Erzurum to create Rr-AgNPs by bioreduction and evaluated their antibacterial and anticancer activities in medical applications.

## 2. Material and Methods

### 2.1. Materials

Rr fruit was collected in Erzurum Çat district on May 2, 2022. The species was identified by Dr. Kenan Akbaş at Muğla Sıtkı Koçman University, Department of Biology Herbarium and was given Herbarium number K.A 1658-A.

#### 2.1.1. Preparation of extract from fruit peels and silver nitrate solution

Fruit peels were removed and rinsed with distilled water before being dried. Dried fruit peels were weighed to 20 g and placed in a 500 mL beaker. Then 250 mL of distilled water was added and the mixture was heated until boiling. After being filtered with filter paper and brought to room temperature, the extract was ready for synthesis. A solution with a concentration of 5 mM was prepared from Sigma-Aldrich silver nitrate ( $\text{AgNO}_3$ ) salt to be used to obtain Rh-AgNPs by bioreduction.

### 2.2. Methods

#### 2.2.1. Bioreduction synthesis of Rr-AgNPs with fruit peel extract

For this, 200 mL of fruit peel extract and 200 mL of 5 mM  $\text{AgNO}_3$  solution were combined 1:1 in a 1000 mL glass Erlenmeyer and left to sit at room temperature. The color shift was observed. By obtaining samples from the reaction medium at different times and performing UV-vis wavelength scans, along with color change, the presence of AgNPs at maximum absorbance was identified (Baran et al. et al. 2021; Atalar et al. 2022).

#### 2.2.2. Characteristics of the produced Rr-AgNPs

To exhibit the formation of AgNPs synthesized from *R. ribes* fruit peel extract (Rh-AgNPs), samples were taken from this medium, with reaction-related color change after mixing the extract with  $\text{AgNO}_3$  solution, and a Perkin Elmer One UV visible spectrophotometer was used (UV-vis). Maximum absorbances were measured in the 300-800 nm wavelength range. Using Fourier transformation infrared spectroscopy (FTIR-Perkin Elmer), frequency changes in the range of 4000-800  $\text{cm}^{-1}$  for both the extract and the liquid obtained at the end of the reaction were examined to evaluate bioactive functional groups in the extract responsible for bioreduction. The morphological appearance of the synthesized Rh-AgNPs was evaluated in micrographs taken using Jeol Jem FE-SEM and TEM devices. In addition, topographic structures, shape, and size distributions of Rh-AgNPs were determined by means of AFM. Also, the size distribution was determined using Marven DLS. Graphs obtained using a RadB-DMAX II computer-controlled electron dispersed X-ray (EDX) were examined, and the elemental contents of the synthesized particles were determined. Plane-reflected crystal patterns were identified for Rh-AgNPs in data acquired at  $2\theta$  using a Rigaku Miniflex 600 model X-ray diffraction (XRD). With this data, crystal nanosizes were calculated using the Debye-Scherrer equation via XRD data (Baran & Acay 2019; Umaz et al. 2019). The charge distributions of the surface structures of the synthesized Rh-AgNPs were evaluated with Marven Zeta potential. The resistance of Rh-AgNPs to temperature changes was measured between 25 and 900 °C using Shimadzu 50 thermogravimetric and differential thermal analysis (TGA-DTA).

### 2.2.3. Antimicrobial Effects of Rh-AgNPs

Minimum inhibition concentrations (MIC) of Rh-AgNPs synthesized with fruit peel extract were calculated using the microdilution method (Atalar et al. 2022) for suppression of the growth of microorganisms in pathogenic groups. Pathogenic gram-negative and positive bacteria and yeast were used in the experimental study. Each microorganism was grown in suitable media for 24-48 hours of incubation at 37 °C. Then, microorganisms were prepared according to McFarland standard 0.5 (Emmanuel et al. 2015; Patil et al. 2021; Raghavendra et al. 2022) turbidity criteria for each strain. Bacteria and fungi were transferred to 96-well microplates containing the appropriate medium. Some wells on the microplate were selected and defined for the control groups. Then, Rh-AgNPs prepared at different concentrations were transferred to the microplate wells and microdilution was applied. The microorganisms were then transferred to the wells. The prepared microplates allowed interaction between microorganisms and Rh-AgNPs for 24-48 hours in an oven at 37 °C. At the end of the period, growth control was performed and MIC values were determined. İnönü University Medical Faculty Hospital Microbiology Laboratory provided the strains of *Staphylococcus aureus* (*S. aureus*) ATCC 25923, *Escherichia coli* (*E. coli*) ATCC 25922, and *Candida albicans* (*C. albicans*) ATCC 10231. Artuklu University Microbiology Research Laboratory provided the strains of *Bacillus subtilis* (*B. subtilis*) ATCC 11774 and *Pseudomonas aeruginosa* (*P. aeruginosa*) ATCC 27853.

### 2.2.4. Anticancer effects of Rh-AgNPs

Cytotoxic effects of the synthesized Rh-AgNPs on cancer cells were evaluated by determining the % survival rate using the MTT method (Baran et al. 2021). Three different cancer cells and one healthy cell were used in experimental studies. The cells were incubated in suitable nutrient media and under optimum conditions. Cells were then suspended at different concentrations using a hemocytometer, transferred to 96-well microplates, and incubated for 24 hours. Then, solutions containing Rh-AgNPs at different concentrations were added to the wells cultured with cell lines and incubated for 48 hours. The absorbance of the cells was then read using a Multi Scan Go (Thermo) instrument at 540 nm. The viability suppressing concentrations of Rh-AgNPs in the cell lines were calculated with the following formula ( Remya et al. 2015; Mohmed et al. 2017; Sunderam et al. 2019; Chen et al. 2021).

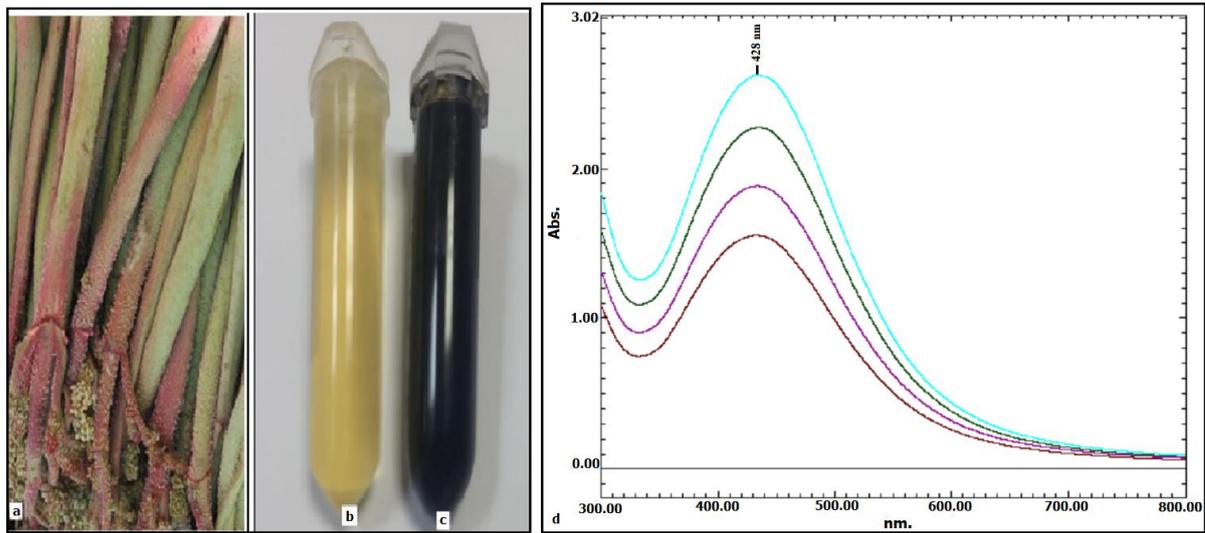
$$\text{Viability \%} = (\text{sample absorbance} / \text{control absorbance}) \times 100 \quad (2)$$

## 3. Results and Discussion

### 3.1. Maximum absorbance bands obtained via UV-vis data of Rh-AgNPs

A color change from yellow to brown was observed 35 minutes after combining Rr fruit peel extract with AgNO<sub>3</sub> solution, which is a macroscopic finding indicating that Rh-AgNPs had formed (Acay and Baran, 2019; Jebiril et al. 2020; Mamdooh & Naeem, 2021; Mani et al. 2021; Sattari et al. 2021). In addition to this finding, the maximum absorbance bands at 428 nm taken with UV-vis were characteristic, showing the vibrations (SPR) occurring on the plasma surface and the formation and presence of Rh-AgNPs (Figure 1) (Luna et al. 2015; Patra et al. 2016; Francis et al. 2017; Eren & Baran 2019; Anandalakshmi 2021).

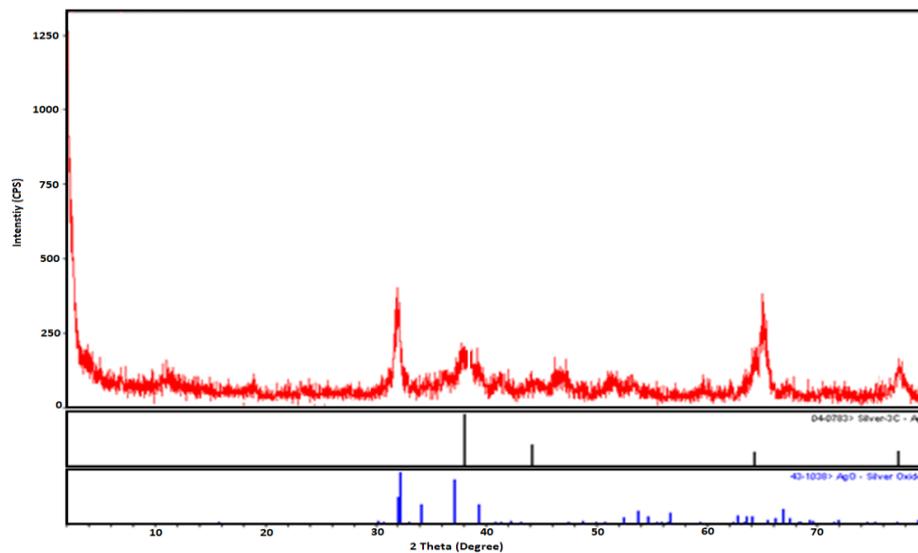
In a study in which AgNPs were synthesized using the fruit of the Rh plant and 1 mM AgNO<sub>3</sub> solution in a ratio of 10:1, respectively, the maximum wavelength absorbance of the obtained AgNPs was 425 nm (Mustafa, Mahdi Auda, Hajir, Ali Shareef & Bari 2021). In this study, the maximum absorbance of AgNPs obtained by using the fruit peels of the Rh plant and 5 mM AgNO<sub>3</sub> solution in a 1:1 ratio was 428 nm. UV-vis findings of the studies show that the maximum wavelengths were almost the same, despite the use of different parts of the plant, different extracts and metal concentrations. In another study using Rh fruit, the maximum wavelength absorbance of AgNPs obtained using the ethanol extract in an medium with 10 mM AgNO<sub>3</sub> concentration at 200 rpm at 75 °C for two days was 410 nm (Aygün et al. 2020). This study is more advantageous in terms of application conditions such as the method of obtaining the extract, synthesis temperature, shaking and metal concentration.



**Figure 1- a. Rr fruit b. Rr fruit extract, c. The appearance of the end-reaction fluid indicates the formation of Rr-AgNPs as a result of synthesis, and d. Maximum absorbance spectra dependent on SPR according to UV-vis for the presence of Rr-AgNPs**

### 3.2. XRD Analysis of Rr-AgNPs

In Figure 2, data obtained at  $2\theta$  with XRD was evaluated to calculate the crystal patterns and nanosizes of Rr-AgNPs. Widening of the Bragg angle peaks showed that the Rh-AgNPs have a central faceted cubic pattern (fcc) (A. Singh et al. 2018; Wongpreecha et al. 2018a; 2018b; Jebril et al. 2020; Anandalakshmi 2021). The FWHM values of  $111^\circ$ ,  $200^\circ$ ,  $220^\circ$ , and  $311^\circ$  Bragg angles in the data were determined as 38.72, 44.62, 64.90, and 77.48, respectively. By using Debye-Scherrer's high peak FWHM value for the crystal nano dimensions of Rr-AgNPs, the cubic pattern of Rr-AgNPs was calculated to have crystal size of 58.06 nm. In two green synthesis studies using Rh fruits, the crystal patterns of the obtained AgNPs had similar characteristics with broadening peaks occurring at  $111^\circ$ ,  $200^\circ$ ,  $220^\circ$ , and  $311^\circ$  Bragg angles (Aygün et al. 2020; Mustafa et al. 2021). In the biosynthesis study of AgNPs using *Cucurbita maxima* L. extracts, the crystal nanosizes of AgNPs were calculated as 67 and 56 nm (Ali 2020). In studies using herbal extracts for the synthesis of AgNPs, the crystal nanosizes were calculated as 24.36 nm (Khan et al. 2022), 40 nm, and 21.17 nm (Aktepe & Baran 2021).



**Figure 2- Data from X-ray diffraction showing the crystal structures of Rr-AgNPs.**

### 3.3. FTIR Spectroscopy Data

The FTIR spectra of the liquid media as a result of extract synthesis were analyzed in order to assess the groups of bioactive substances that may control bioreduction and stability. Frequency changes in the data occur at  $3337.81\text{-}3330\text{ cm}^{-1}$ . In order to assess the groups of bioactive components ensuring bioreduction, stability of  $\text{Ag}^+$  form to  $\text{Ag}^0$  form, and the stability of produced nanoparticles were investigated from  $2122.95\text{-}2122.80\text{ cm}^{-1}$ , and  $1635.48\text{-}1635.33\text{ cm}^{-1}$ . These observations demonstrated the

potential for aromatic chains with C=C stretching, flavonoid-phenolic groups with C=O stretching, and hydroxyl groups resulting from O-H stretching to affect bioreduction and durability (Figure 3) ( B. Kumar et al. 2015; Ahmad et al. 2018; A. U. Khan et al. 2018; Hemmati et al. 2019; Jogaiah et al. 2019; Jebril et al. 2020;Auda et al. 2021).

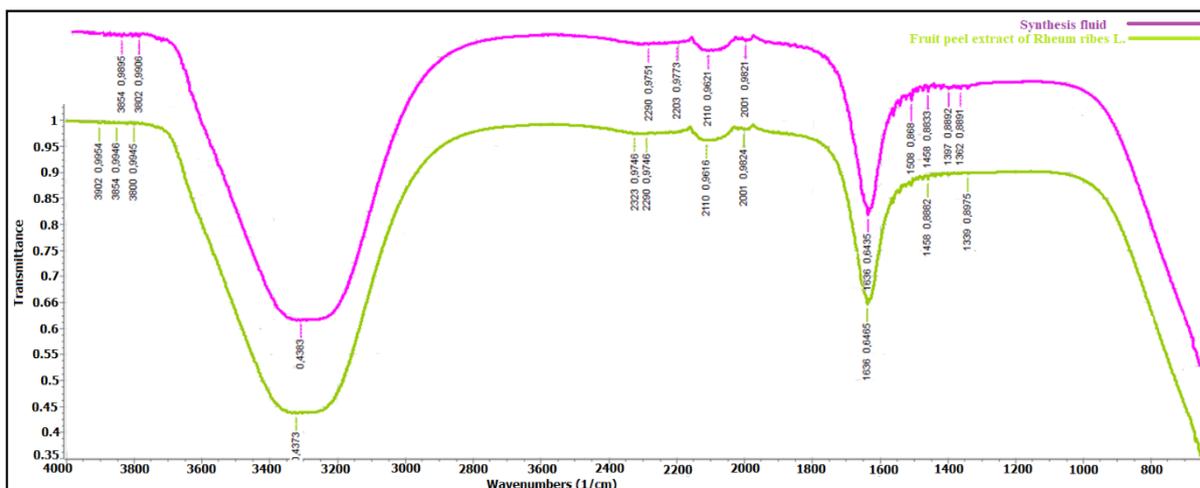


Figure 3- FTIR spectra of Rr fruit peel extract and reaction liquid obtained after synthesis

### 3.4. SEM and EDX Images of Rr-AgNPs

In SEM micrographs, the synthesized Rr-AgNPs had spherical morphology (Figure 4). According to reports, Rr-AgNPs had spherical morphology in SEM images taken during green synthesis investigations ( Satpathy et al. 2018; Jebril et al. 2020; Aktepe & Baran 2021).

The profile of the EDX graph was used to determine the elemental contents of the particles synthesized with fruit peel extract (Figure 5). Strong peaks in the areas for elemental silver indicated that the synthesized particles were Rr-AgNPs (Pallela et al. 2018; Butola et al. 2019; M. R. Khan et al. 2022; Suriyakala et al. 2022). The existence of phytochemicals that actively contribute to the stability of the medium around Rr-AgNPs was suggested by weak peaks for elements in the graph, such as carbon and oxygen (Vastrand 2016; Kumar et al. 2019; Das et al. 2021). FTIR and zeta potential results in Figures 3 and 7, respectively, further confirm this data.

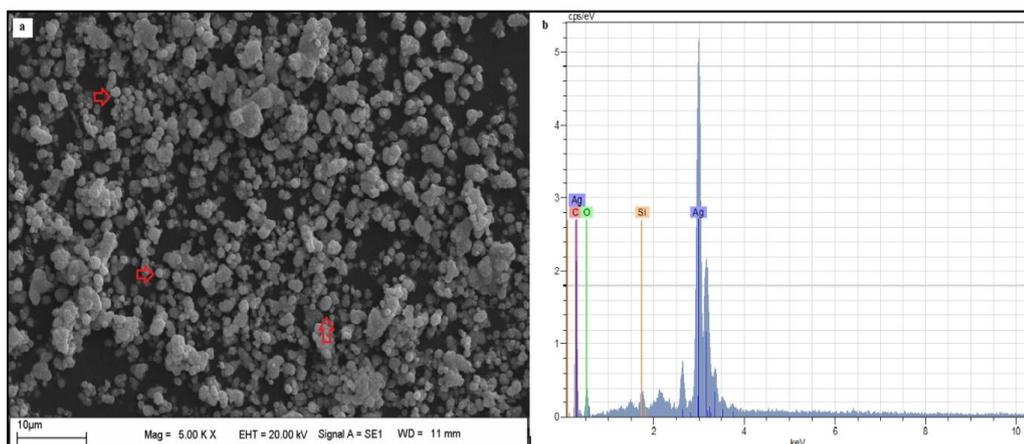
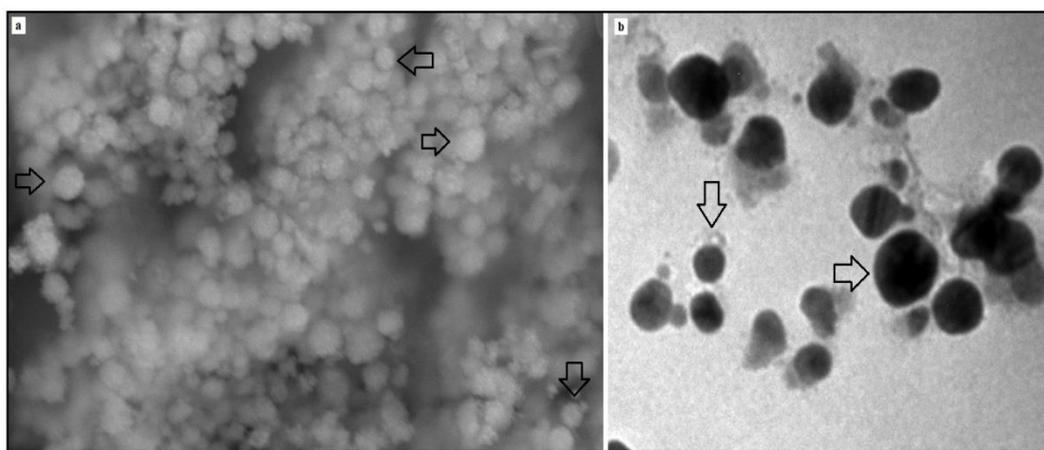


Figure 4- Rr-AgNPs after synthesis with Rr fruit peel extract; a. SEM micrograph of morphology and b. EDX profile

### 3.5. Appearance of Rr-AgNPs in FESEM and TEM Micrographs

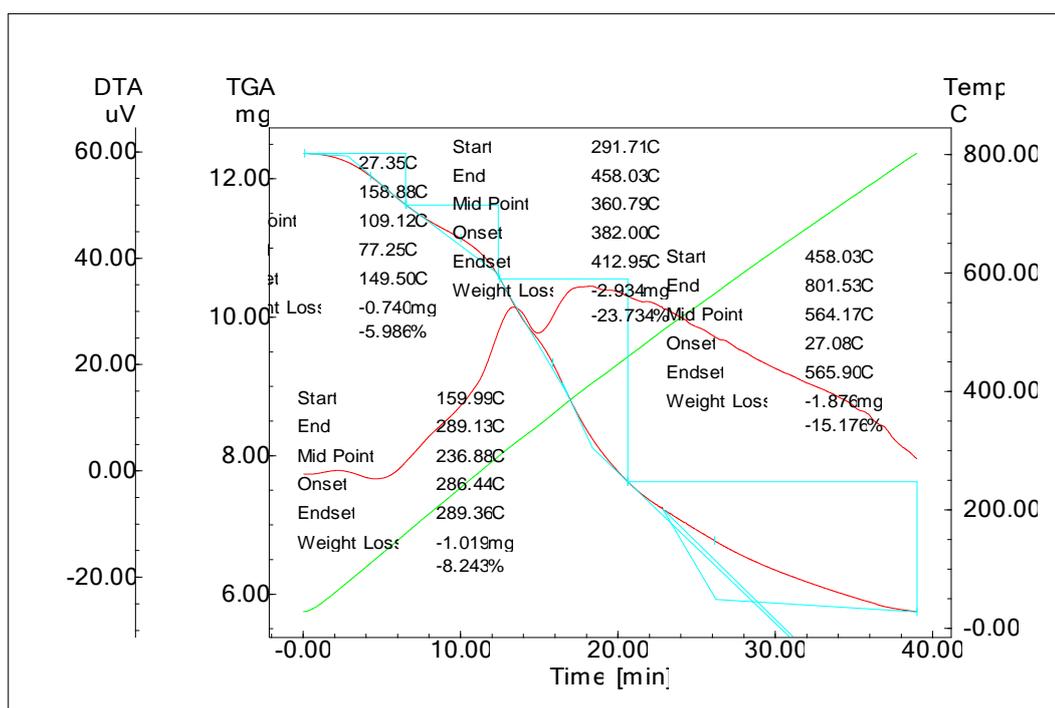
Figure 5 shows images taken with a FESEM and TEM microscope of Rr-AgNPs produced with Rr fruit peel extract. The figure shows that the synthesized Rh-AgNPs exhibited spherical morphological appearance. AgNPs had the same morphological appearance in both studies using Rh fruit (Aygün et al. 2020; Auda et al. 2021). AgNPs synthesized in other green synthesis studies using different plant sources had the same morphological appearance (Velmurugan et al. 2014; Remya et al. 2015; Mamdooh & Naeem 2021; Wang et al. 2021).



**Figure 5-** Rr-AgNPs after synthesis with Rr fruit peel extract; a. FE-SEM ve b. TEM micrograph of morphological views

### 3.6. TGA-DTA analysis of synthesized Rr-AgNPs

The thermal resistance of the synthesized Rr-AgNPs was determined by TGA-DTA analysis data at 10-800 °C. In the data, there were 5.98%, 8.24%, 23.73%, and 15.17% mass losses, respectively, occurring at 27.35-158.88 °C, 159.99-289.13 °C, 291.71-458.03 °C, and 458.03-801.53 °C (Figure 5). The loss of water absorbed was the cause of the initial mass loss (7.07%). The presence of flavonoids surrounding the surface of Rr-AgNPs caused mass losses at the other three temperatures (Baran 2019b; Rolim et al. 2019; Baran et al. 2021). The weak C, O peaks and -25 mV negative surface charge in the EDX and Zeta potential data in Figures 5 and 7, respectively, also support this finding.

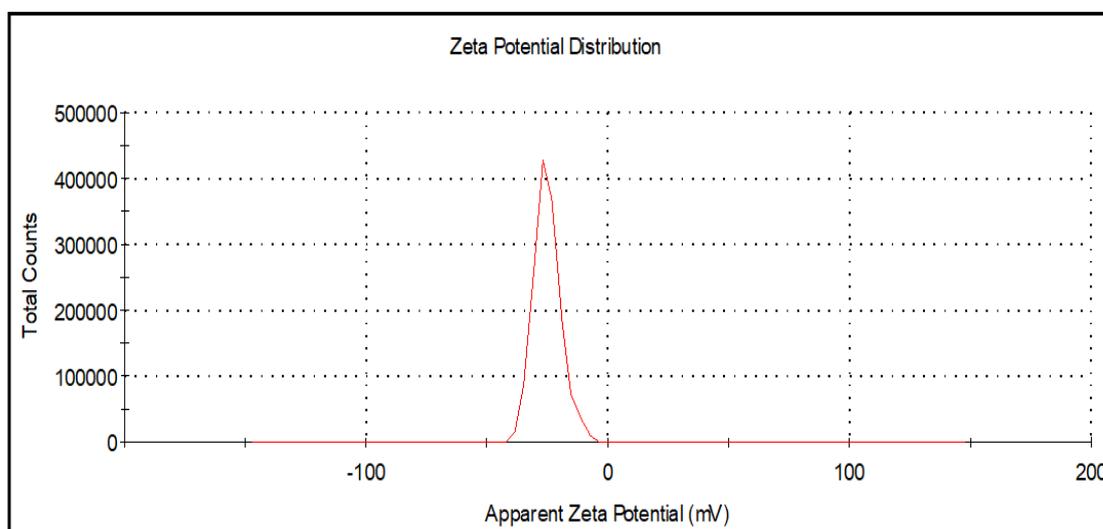


**Figure 6-** TGA-DTA data for Rr-AgNPs with mass loss points against heat treatment

### 3.7. Zeta potential analyses of synthesized Rr-AgNPs

The surface charges of Rr-AgNPs synthesized using fruit peel extract were identified as -25 mV by zeta potential analysis (Figure 7). The negative charge distribution of Rr-AgNPs is an important indicator of their stability. Green synthesized Rr-AgNPs exhibit more efficient negative charge distribution compared to those obtained by other synthesis approaches. The existence of flavonoids in the surface structure of Rr-AgNPs influences the distribution of negative charges. A negative charge arrangement stabilizes free electrons in colloidal form by oxidizing the hydroxyl groups. (Pugazhendhi et al. 2018; Wongpreecha et al. 2018b; Oliveira et al. 2019; Jebriil et al. 2020; Aktepe et al. 2022). The presence of phytochemicals around Rr-AgNPs in the findings obtained in Figure 3, Figure 5, and Figure 6 also contribute to the explanation for the surface structure. The fact that the surface

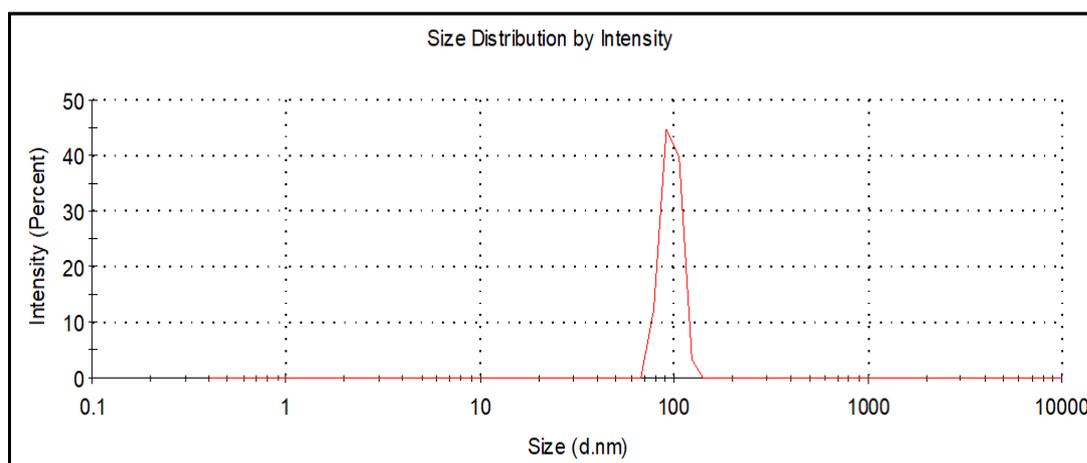
charges of the synthesized Rr-AgNPs have different charges (negative and positive) triggers the formation of negative results that affect stability, such as fluctuation and aggregation, as the nanoparticles attract with each other by electrostatic attraction (Al-ogaidi et al. 2017; Patil et al. 2018a; Satpathy et al. 2018). The -25 mV surface charge data showed that the synthesized Rr-AgNPs were stable. It was reported that Rr-AgNPs synthesized from different plant sources had surface charge distribution of 25.01 mV (Aktepe et al. 2022),  $-22 \pm 5$  mV (Ferreya Maillard et al. 2018), and -19 mV (Oliveira et al. 2019).



**Figure 7- Graphs of zeta potential charge distribution of Rr-AgNPs after synthesis**

### 3.8. Density-dependent Rr-AgNPs size distribution

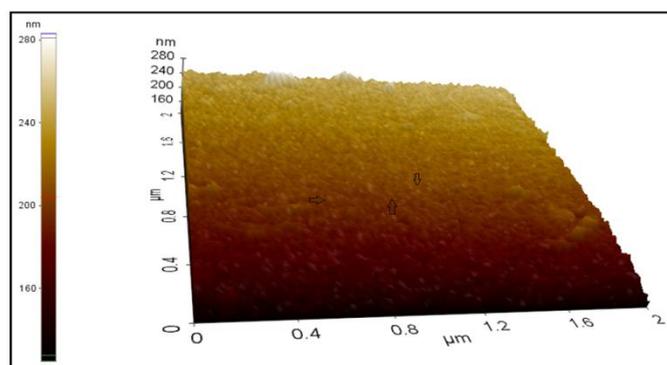
The mean size distribution of the produced Rh-AgNPs, as seen in Figure 8, is 96 nm. They had an average size distribution of 54 nm in a study about the manufacture of AgNPs utilizing extracts from *Crinum asiaticum* leaves (Shukla et al. 2022). According to Kumar et al. (2017), AgNPs made with *Prunus persica* extract had size distribution of 2-130 nm. According to reports from earlier studies about environmentally friendly synthesis, the produced AgNPs had size distribution between 59.74 nm and 268 nm (Alkhulaifi et al. 2020; Mamdooh & Naeem 2021).



**Figure 8- Density-dependent size distribution of synthesized Rr-AgNPs**

### 3.9. AFM Micrograph of the distributions of synthesized Rr-AgNPs

As shown in Figure 9, the topographical features and morphologies of the produced Rr-AgNPs were determined by analyzing AFM data. The data showed that Rr-AgNPs were less than 100 nm, exhibited spherical appearance, and were distributed in a single structure (Swamy et al. 2015; Kumar et al. 2017; Rauf et al. 2021).



**Figure 9- AFM micrograph of Rr-AgNPs synthesized using fruit peel extract**

### 3.10. Antimicrobial effects of synthesized Rr-AgNPs

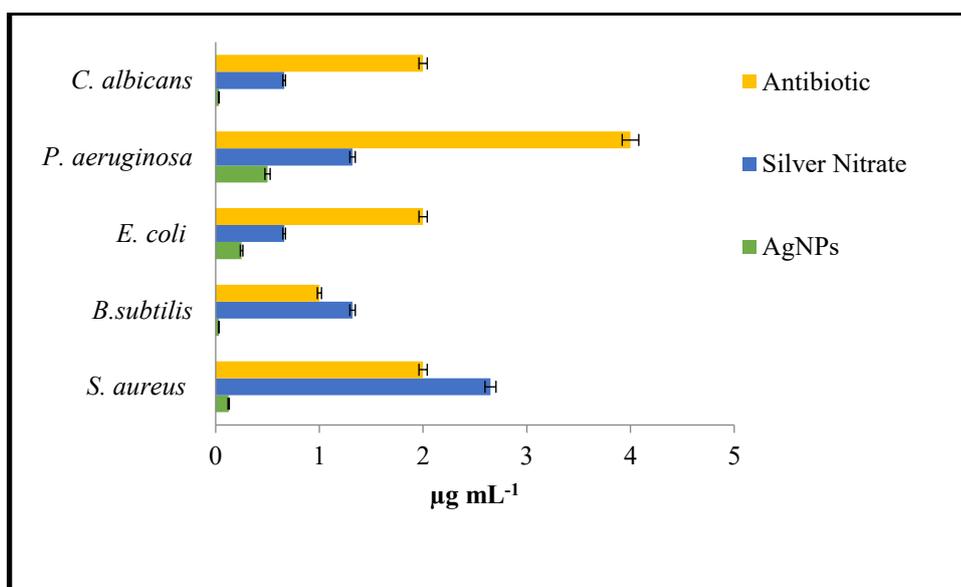
By using the microdilution method, the MIC values of the produced Rr-AgNPs for the development of pathogenic microbes were examined. In Table 1 and Figure 9, the MIC values affecting the growth of the strains are given. As seen in the data, 0.03-0.50  $\mu\text{g}/\text{L}$  concentrations had a suppressive effect on growth of microorganisms. The effective MIC values of the synthesized Rr-AgNPs were 0.03-0.13 and 0.25-0.50  $\text{mg}/\text{L}$  for gram positive and negative bacteria, respectively. These values were considerably lower than the effective concentrations of silver nitrate solution and antibiotics. The effective MIC value for the fungus *C. albicans* was found to be 0.03  $\mu\text{g}/\text{L}$ , and this value is also many times lower than the value at which silver nitrate solution and antibiotics were effective. Table 2 gives the MIC values of Rr-AgNPs obtained in some applications using green synthesis.

Rr-AgNPs ionize in liquid media and show high reactivity. They approach one other with the electrical force of attraction, interacting with organisms in the exact same environment (Narayan & Dipak 2015; Ahmed et al. 2016; Chung et al. 2016; Zhang, et al. 2010; Aina et al. 2018). They cause defects in the membrane structure of microorganisms and adversely affect the functions of DNA, RNA, and vital enzymes, which have high affinity for these species due to the increase in reactive oxygen species (ROS). The structure of the biomolecules is damaged by these species and as a result, their functions deteriorate and the microorganism dies (Emmanuel et al. 2015; Shao et al. 2018; Huq et al. 2022).

**Table 1- MIC values for Rh-AgNPs, antibiotics and silver nitrate solution with antimicrobial effect on the growth of microorganisms**

	<i>Microorganisms</i>	<i>AgNPs</i> $\mu\text{g}/\text{mL}$	<i>Silver Nitrate</i> $\mu\text{g}/\text{mL}$	<i>Antibiotic</i> $\mu\text{g}/\text{mL}$
<b>Gram Positive</b>	<i>B. subtilis</i> ATCC 11774	0.13	2.65	2.00
	<i>S. aureus</i> ATCC 29213	0.03	1.32	1.00
	<i>E. coli</i> ATCC25922	0.25	0.66	2.00
<b>Gram Negative</b>	<i>P. aeruginosa</i> ATCC27833	0.50	1.32	4.00
	<i>C. albicans</i> ATCC 10231	0.03	0.66	2.00

Commercial antibiotics used: **vancomycin** for gram (+) bacteria, **colistin** for gram (-) bacteria, and **fluconazole** for *C. albicans*.



**Figure 9-** Graph of MIC values of Rh-AgNPs, AgNO<sub>3</sub> solution, and antibiotics Commercial antibiotics used: vancomycin for gram (+) bacteria, colistin for gram (-) bacteria, and fluconazole for *C. albicans*.

**Table 2-** Antimicrobial effects of AgNPs obtained in green synthesis studies

Biological Source	Size (nm)	Gram Negative µg/mL	Gram Positive µg/mL	References
<i>Agastache foeniculum</i>	9-19	12.50	6.25	(Polivanova et al. 2021)
<i>Vitis vinifera</i>	18.53	0.31	0.07	(Acay et al. 2019)
<i>Fritillaria sp.</i>	10	1-2	1-4	(Hemmati et al. 2019)
<i>Zataria multiflora</i>	25	1.25	0.5-4	(Barabadi et al. 2021)
<i>Euphorbia hirta</i>	15.5	0.67	0.82	(V. Kumar et al. 2016)
<i>Madhuca longifolia</i>	30-50	80-90	40-60	(Patil et al. 2018b)

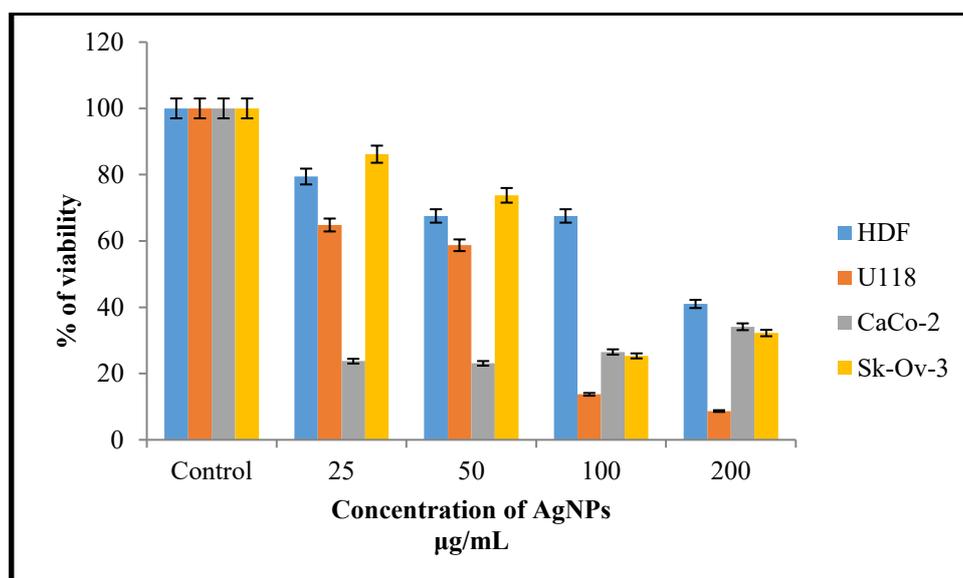
### 3.11. Anti-cancer effects of Rr-AgNPs

The MTT technique was used to investigate the viability-suppressing concentrations of Rr-AgNPs made using Rr fruit peel for healthy cell lines and the suppressive effects for cancer cell lines. Up to a concentration of 25-200 mg/L, the Rr-AgNPs were not hazardous for the normal cell line HDF. However, it had a very strong suppressive impact on lines of cancer cells, indicating a very positive outcome for use as an anticancer drug. Even at a 25 mg/L dosage, a good suppressive effect was seen. With an inhibitory rate of 86.27%, 74.67%, and 73.49% for U118, CaCo-2, and Skov-3, 100 mg/L concentration was discovered to have excellent anticancer effect (Table 3 and Figure 10). Due to the existence of extensive vascular holes in the target cancer cells, the tiny size of NPs impacts the build-up and penetration of NPs into these areas. Additionally, Rr-AgNPs have a propensity to accumulate in crucial components such as membranes of cells, proteins, and nuclei. After localization, Rr-AgNPs with strong oxidative characteristics cause the cell to die by inducing cell death mechanisms such apoptosis and an increase in ROS. As a result of these impacts, it is crucial to assess the harmful effects of Ag<sup>+</sup> ions in living organisms (Wongpreecha et al. 2018b). Some NP characteristics contribute to harmful impacts.

**Table 3-** Viability percentages after Rr-AgNP interactions with various cell lines using the MTT technique for 48 hours.

Cell Line	25 µmg/L	50 µmg/L	100 µmg/L	200 µmg/L
HDF	79.45	67.55	67.55	41.00
U118	64.82	58.71	13.73	8.68
CaCo-2	23.74	23.08	26.51	34.11
Skov-3	86.18	73.77	25.33	32.22

HDF; human skin fibroblast cells: U118; glioblastoma cells: CaCo-2; colon cancer cells: Skov-3; ovarian sarcoma cells



**Figure 10-** Viability percentages after Rr-AgNP interactions with various cell lines using the MTT technique for 48 hours. In Table 4, the cytotoxic effects on cancer cells of some nanoparticles obtained with green synthesis are compared.

**Table 4-** Amounts of Rr-AgNPs found in research employing environmentally friendly synthesis methods that contribute to the cytotoxic impact

Biological Source	Cancer Cell Line	Size (nm)	Shape	Effective Concentration µmg/L	Refereces
<i>Aloe Vera</i>	HDF	20	Spherical	100	( Zhang, et al. 2010)
<i>Aspergillus sp.</i>	CaCo-2	5-30	Spherical	3.75-5	(Mohmed et al. 2017)
<i>Pueraria tuberosa</i>	Skov-3	162.72	Spherical	29.36	(Satpathy et al. 2018)

#### 4. Conclusions

Rr-AgNPs are widely used, which makes them very valuable materials. The manufacture of these items receives a lot of attention daily for green synthesis techniques. The extract from the peel of the Rr fruit, also known as "Işgin" or "Kurdish banana," which grows in the Erzurum region, was used in this study to produce Rr-AgNPs for the first time in a quick, straightforward, ecologically benign method without harmful processes. Data from TEM, AFM, UV-vis, FE-SEM, XRD, EDX, TGA-DTA, Zetasizer, and Zeta potential analyses were used to identify the characteristics of the synthesized Rr-AgNPs. In order to evaluate the functional components of flavonoids that contribute to bioreduction during the manufacturing process, FTIR data were examined. Maximum absorbance, spherical shape, median size range of 96 nm, and charge on the surface of -25 mV at wavelength of 428 nm were all characteristics of the produced Rr-AgNPs. The efficacy of produced Rr-AgNPs in medicinal applications was studied using MTT and microdilution techniques. The minimum inhibitory concentrations of Rr-AgNPs needed to inhibit hospital pathogens were between 0.03-0.50 mg/L. These concentrations were extremely low compared to antibiotics and AgNO<sub>3</sub> solution. With inhibitory rates of 86.27%, 74.67%, and 73.49%, Rr-AgNPs were discovered to have a very good anticancer impact on U118, CaCo-2, and Skov-3 cancer cells. The healthy cell HDF was not damaged by the same concentration. In particular for biopharmacology, enhancing application processes will considerably advance the search for antibacterial and anticancer drugs.

#### Declarations

#### Conflict of interest

All authors declare there are no conflicts of interest in this research work.

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