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### **Evaluation of** *Tussilago farfara* **L. Smoke by GC/MS: A Phytochemical Approach to a Traditional Medicine**

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Escherichia coli was not inhibited.

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#### **ARTICLE INFO** *Original paper Article history:* Received: 7 Jan 2023 Revised: 15 Feb 2023 Accepted: 24 Mar 2023 *Keywords: Tussilago farfara* Smoke GC/MS Phenol Trapping Bioautography **ABSTRACT** The smoke produced from natural substances such as medicinal plants is used in various cultures for different purposes. The use of medicinal fumes has been reported in nearly 50 countries. Among medicinal plants, *Tussilago farfara* L. known as coltsfoot has been introduced in Canon the famous book of Avicenna a Persian polymath, for chronic dry cough and various pulmonary diseases and shortness of breath. *T. farfara* is distributed in wet mountainous regions of Iran. For this study, the leaves and flowers of *T. farfara* were collected from Chalous Road in Iran. The smoke from the burning of *T. farfara* organs was prepared by homemade glassware trapping the smoke in methanol and then methanol was evaporated. In general, five grams of materials were burned and the smoke was dissolved and trapped in 100 ml of methanol. The trapped and dried materials from the smoke of extracts were filtered and injected into the GC/MS for analysis and identification of its constituents. 51 compounds representing 91.1 and 92.3 percent of smoke extracts of *T. farfara* were identified in leaf and flower. Also, 57 compounds were detected in the sample of EL and EF with 96.8 % and 97.7 %. The percentage of phenolic compounds that were identified in all extracts of smoke were SL and SF with 52.1 and 46.5, respectively. Phenol, Hydroquinone, P-Cresol and O-Cresol were the major compounds in the smoke extracts. Smoke leaves and flowers of *T. farfara* were selected to test the antimicrobial to continue. This study examined the bactericidal effect of smoke flowers. Fractions of effective constituents with the help of hexane-ethyl acetate with the method of thin layer chromatography (TLC) were isolated. The results of this experiment showed that a fraction (8:2) of hexane-ethyl acetate inhibited the bacteria Staphylococcus aureus. But

#### **1. Introduction**

The demand for medicinal uses of plant-derived smoke outnumbers all other applications [\(Pennacchio](#page--1-0)  *et al*[., 2010\)](#page--1-0). The smoke produced from natural substances is used in various cultures and regions for different purposes including incense, medicine and food maintenance [\(Fabricant and Farnsworth, 2001\)](#page--1-1). Therefore, different matters are recognized to be used to produce smoke for several utilizations [\(Pennacchio](#page--1-0)  *et al*[., 2010\)](#page--1-0). Human beings have used the smoke of medicinal plants to lead a healthy life long ago and in some cultures of the world, the smoke has been used in religious festivals and ceremonies [\(Mohagheghzadeh](#page--1-2)  *et al*[., 2006\)](#page--1-2). Plant-derived smoke at high temperatures

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exhibits fast pharmacological activity when inhaled [\(Nautiyal](#page--1-3) *et al*., 2007). Some medicinal plants are used in the form of smoke (Danial *et al*[., 2018\)](#page--1-4). Smoke mainly contains compositions that have antibacterial, antifungal, anti-inflammatory and antioxidant properties and some of them are used for the treatment of neuralgia, rheumatism, capillary bleeding, and skin disorders [\(Shafiee and Moravej-Salehi, 2015\)](#page--1-5). Researchers are looking for natural medications that have fewer side effects and the Iranian traditional medicine is a valuable reference in this respect [\(Shafiee](#page--1-5)  [and Moravej-Salehi, 2015\)](#page--1-5). The antimicrobial activity of smoke is attributed to the presence of compounds like phenols, carbonyls and organic acids [\(Holley and](#page--1-6)

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[Patel, 2005\)](#page--1-6). Many studies have presented that bacteria have increased resistance. There is an increasingly growing need for the development of new strategies for conventional antibiotic therapy [\(Shalayel](#page--1-7) *et al*., 2017). The use of medicinal smoke in the therapy of many diseases such as microbial diseases and infections in Iran, Turkey and Malaysia has long been popular [\(Danial](#page--1-4) *et al*., 2018). [Roche et al. \(1997\)](#page--1-8) reported that smoke can protect seeds and seedlings against microbial corrosion. [Kulkarni](#page--1-9) et al. (2011) indicated that smoke contains aromatic hydrocarbons, chlorine compounds and aldehydes that can reduce the infestation of endophytic fungi, as these smoke compounds show similar chemical structures to those of known fungicidal compounds. Another study result showed smoke generated by burning wood and medicinal herbs eliminated some of the bacteria that are harmful to horticultural plants [\(Nautiyal](#page--1-3) *et al*., 2007). Among the many plants whose smoke is used, *Tussilago farfara* L. known as coltsfoot has been introduced in Canon the famous book of Avicenna a Persian polymath, for chronic dry cough and various pulmonary diseases and shortness of breath [\(Mahdizadeh](#page--1-10) *et al*., 2015). In fact, the leaves of *T. farfara* are smoked like tobacco, as a domestic remedy for asthma [\(Qureshi](#page--1-11) *et al*., 2007). Coltsfoot (*Tussilago farfara* L.) from the Asteraceae family is a perennial plant [\(Ferrer](#page--1-12) *et al*., 2016). The flower buds of coltsfoot have a long history in the Chinese Pharmacopoeia for the treatment of cough, phlegm and asthmatic disorders (Qu *et al*[., 2018\)](#page--1-13). *T. farfara* is distributed in wet mountainous regions of the world and can also be found in Tehran, Azerbaijan and the Northern provinces of Iran [\(Norani](#page--1-14) *et al*., 2019). Antimicrobial resistance is a worldwide health problem associated with increased disease and mortality, while the factors associated with it are well known, unfortunately, the root causes of it continue to be declined [\(Shalayel](#page--1-7) *et al*., [2017\)](#page--1-7). There are few works on the volatile combustion products of incense and their derivatives [\(Staub](#page--1-15) *et al*., [2011\)](#page--1-15). In this study, we aim for the characterization and comparative analysis of the main volatile organic compounds (VOCs) present in the smoke of *T. farfara*  organs.

#### **2. Materials and methods**

#### *2.1. Plant materials and experimental conditions*

In this study, the roots, leaves, barks and flowers of *T. farfara* were collected from Pol-e Zangholeh of Iran (51º 20ʹ 17ʺ N, 36º 48ʹ 11ʺ E). *T. farfara* samples were cleaned and then air-dried in the shade and powdered using a milling machine and kept in a cool dry place until ready to burn. Also, Green tea and black tea (*Camellia sinensis* (L.) Kuntze) were purchased from the Fuman farms in the north of Iran. Cloves (*Syzygium aromaticum* L.), and rosemary (*Rosmarinus officinalis* L.) were bought from the grocery store. The abbreviations of all samples are shown i[n Table 1.](#page--1-16)



*2.2. Smoke collection and preparation of methanolic extract*

The smoke of roots, leaves, barks and flowers of *T. farfara* was separately prepared by homemade glassware trapping the smoke in methanol [\(Fig.](#page--1-17) 1). In general, three grams of samples were burned and the smoke was dissolved and trapped in 100 ml of methanol. Extracts of root, leaf, bark and flower of *T. farfara* along with several well-known antioxidant plants including green tea, clove, black tea and rosemary were prepared by sonication of three g of dried plant material for 30 min in 50 mL of methanol. All extracts were filtered through Whatman no.1 filter paper and then concentrated at 40 °C using a rotary evaporator. The extracts were dried and stored at 4 °C until analysis.



**Figure 1. Schematic representation of the collection the smoke of roots, leaves, barks and flowers of** *T. farfara.*

### *2.3. GC–MS analysis and Identification of the smokes extract*

GC–mass spectrometry (GC–MS) analysis was carried out by a Thermoquest–Finnigan gas chromatograph equipped with a fused silica capillary HP-5 column (60 m  $\times$  0.25 mm i.d.; film thickness 0.25 μm) coupled with a trace mass spectrometer. Helium was used as the carrier gas at a flow rate of 1. mL/minute in a split ratio of 2: 00. Ionization voltage was 70 eV. Ion source and interface temperatures were 200° and 320°C, respectively. Identification was confirmed by comparison of each component's mass spectra with that of the internal mass spectra library of the main library, Wiley 7.0 and Adams and further identification was based on the comparison of peak retention indices by using a homologous series of nalkanes (C8 to C24) recorded under the same operating conditions and the published data [\(Adams, 2007\)](#page--1-18).

#### *2.4. Assessment of antioxidant activity against DPPH*

The radical scavenging activity of all extracts against DPPH (2,2-diphenyl-2 picrylhydrazyl hydrate) was determined according to the previously described method of (Bozin *et al.*, 2007), using the  $IC_{50}$  to compare the antioxidant properties. The absorbance was recorded at 517 nm with an ELISA reader (Epoch, BioTek instrument). The radical scavenging capacity (RSC) was calculated using the formula:  $In\%=[(Ab-$ As)/ Ab]  $\times$ 100, where In is DPPH inhibition. Ab is the absorbance of the blank, and As is the absorbance of the sample extract. Butylated hydroxytoluene (BHT) was used as a positive control.  $IC_{50}$  is the concentration of the sample when the inhibition percentage is 50%.

#### *2.5. Determination of total phenolic compounds*

The total phenolic content compound was determined using the Folin- Ciocalteu method [\(Slinkard and Singleton, 1977\)](#page--1-20). A calibration curve was prepared using a series of methanolic gallic acid solutions (10, 30, 100, 250, 500, 1000 μg/ml), combined with 0.1 ml Folin–Ciocalteu reagent and after 3 min, 0.3 ml sodium carbonate (7.5%). The absorbance of the mixture was measured at 765 nm using a spectrophotometer (Smart spec plus, BIORAD). The experimental extracts  $(0.01 \text{ g/ml})$  of smoke and the other antioxidant plants were each combined with the same reagents and absorption was measured after 2 h to assess phenolic compounds, with three technical replications. Gallic acid was used as the standard for a calibration curve, and the results were expressed as mg of the gallic acid equivalent dry weight of extract (mg GAE/g DW).

#### *2.6. Bioautography of the Antimicrobial Compounds*

The methanolic flowers and leaves smoke extracts were chromatographically analyzed using the thin layer chromatography (TLC) method. The samples were spotted on  $2 \times 10$  cm<sup>2</sup> silica gel plates using spotting tubes about 1-2 cm above the bottom of the plates. Then, they were placed in a chromatography tank with selected solvents of different ratios of hexane and ethyl acetate enough to wet the lower edge of the plate, before the spotting was performed. The plates were left in the solvent for some time, during that, the solvent moved across the plate from bottom to top. The plates were removed from the tank, allowed to dry, and then visualized under ultraviolet irradiations at 254 and 366 nm by spraying with Ninhydrin. The plates with more spots were used in the bioautography test. The silica gel plates were seeded with *Staphylococcus aureus* and *Escherichia coli* and incubated for 20 hours at 37°C. The clear zones due to growth inhibition of the microorganisms indicated the location of antimicrobial compounds on the TLC plates.

#### *2.7. Statistical analysis*

Parametric data were analyzed according to the analysis of variance based on a completely randomized design with three replications, using SAS Statistical Package Program version 9.0 and SPSS software version 20. The means were compared with the Least Significant Difference (LSD) test at a 5% probability level.

#### **3. Results**

#### *3.1. Plant extraction yield*

The yields of all extractions are shown in [Fig.](#page--1-21) 2. The highest extraction yields were obtained in EL (Extract of the leaf) with 11.4 % w/w. The sample of SR (Smoke extract of the root) had the lowest extraction yield with 4.1 % w/w.

#### *3.2. Chemical composition of extracts*

In total 51 compounds representing 91.1 and 92.3 percent of smoke extracts of *T. farfara* were identified in the leaf and flower [\(Table 2\)](#page--1-22). Also, 57 compounds

were detected in the sample of EL and EF with 96.8 % and 97.7 % [\(Table 3\)](#page--1-2). 2-Ethylhexyl hexanoate was the main component in the EL (20.0 %) and EF (25.3 %). Other important compounds identified in the extracts included Hydroquinone, Catechol, Neophytadiene, 9, 12, 15-Octadecatrienoic acid and *N*-Hexadecanoic acid. The results showed that most of the compounds identified in smoke are phenolic compounds. The phenolic compounds identified in SL and SF were 52.1 % and 46.5 %, respectively. Phenol, Hydroquinone, Catechol, *P*-Cresol and hydroquinone with 19 %, 10.5 %, 10.3 % and 5.7 % respectively were the major compounds in the smoke of leaf [\(Fig. 3\)](#page--1-23). As a main compound, Phenol (19.8 %), *O*-Cresol (5.3 %), Hydroquinone (4.8 %) and Phenol, 4-ethyl (4.1 %) were identified in flower smoke [\(Fig. 4\)](#page--1-24). The other important components identified in all smokes include Phenol, 4-ethyl, 3-Methyl-1,2-cyclopentanedione, n-Hexadecanoic acid, Phenol, 2,5-dimethyl, Phenol, 2 methoxy-3-(2-propenyl) and Neophytadiene.



**Figure 2. The yields of all extractions from** *T. farfara* **root, leaf, bark and flower.**

#### *3.3. Total phenolic contents (TPC)*

Results in Fig. 5-A showed the highest TPC in EL (extract of the leaf) with 341 mg GAE/g DW. The bark extract (EB) had the lowest TPC with 119 mg GAE/g DW. In comparison with green tea, cloves, black tea and rosemary; interestingly the phenolic content of this sample was higher than rosemary but lower than the green tea, cloves and black tea.

#### *3.4. Antioxidant activity (AA)*

The results of the comparison of antioxidant activity have been demonstrated in Fig. 5-B. In the DPPH assay, the highest AA was observed in EL (extract of the leaf) with  $IC_{50}$  82 μg/ml. The lowest activity was found in SR (Smoke extract of the root) with  $IC_{50}$  164.3 μg/ml. In general, in comparison with well-known antioxidant plants such as clove, black tea, green tea

and rosemary, the smoke extracts of *T. farfara* showed comparable antioxidant activity.

**Table 2. Chemical composition (%) of different organs smoke of** *Tussilago farfara* **L.**

No RT		Components	SL%	SF%	RIc
1	8.1	3-Methyl-2-cyclopenten-1-one	0.1		973
2	8.4	Phenol	19.0	19.8	992
3	9.5	3-Methyl-1,2-cyclopentanedione	3.3	3.7	1043
4	10.1	$O$ -Cresol	0.7	5.3	1068
5	10.5	P-Cresol	5.7		1093
6	10.8	Guaiacol	1.0	2.6	1096
7	10.9	2(3H)-Furanone, dihydro-4-hydroxy 1.5		3.9	$\overline{\phantom{0}}$
8	11.5	Maltol	3.8	1.0	1129
9	11.8	3-Ethylphenol		1.1	1142
10	12.0	Phenol, 2,5-dimethyl	2.7	2.4	1150
11	12.2	Phenol, 4-ethyl	1.6	4.1	1165
12	12.7	$(S)-(+)$ -2',3'-Dideoxyribonolactone	1.7		$\overline{a}$
13	12.8	Creosol		0.5	1192
14	12.8	Butanoic acid, 1-methylhexyl ester	0.9		1197
15	13.0	Catechol	10.3	1.2	1219
16	13.1	1,5-Dioxonane, 2-ethoxy-9-methyl	2.7		$\overline{\phantom{0}}$
17	13.3	Benzofuran, 2,3-dihydro	1.7	0.9	1223
18	14.3	Hydroquinone	10.5	4.8	1241
19	15.3	Phenol, 2,6-dimethoxy		2.5	1256
20	15.5	Phenol, 2-methoxy-3-(2-propenyl)	0.6	2.2	1263
21	15.9	1H-Indole, 3-methyl	1.0		1264
22	16.0	Skatole		0.7	1296
23	16.8	6-Undecylamine	0.8		1354
24	17.3	1-Dodecene	1.7		1390
25	17.4	Tetradecane	0.6	0.7	1396
26	17.8	Benzene, 1,2,3-trimethoxy-5-methyl 2.9		1.7	$\overline{\phantom{0}}$
27	18.0	M-Dioxan-4-ol, 2,6-diethyl-5-	$0.8\,$	2.9	1450
		methyl-, acetate			
28	18.7	Hexadecane	0.4	1.6	1496
29	19.1	Quinic acid	2.5		1530
30 31	19.6	Tetradecanol		0.6	1575
	19.9 32 20.6	Heptadecane Tetradecanoic acid	0.6	0.7 1.6	1596 1659
33	21.0	Octadecane	1.7	0.6	1695
34	21.5		3.8	0.6	1810
35	21.9	Neophytadiene Phthalic acid, diisobutyl ester		0.4	1774
36	22.2	Heptadecanenitrile	0.9	0.6	1800
		Hexadecanoic acid, 15-methyl-,			
37	22.4	methyl ester		1.0	1822
	38 22.7	N-Hexadecanoic acid	1.6	4.5	1860
39	22.9	Dibutyl phthalate		0.4	1869
40	23.0	Phytol	1.0		1881
41	24.0	1-Octadecanol		0.8	1989
	42 25.0	Hexadecanamide	0.6		2086
43	26.0	Heneicosane		0.7	2195
44	26.3	$(Z)$ -13-Octadecenal	0.8		2225
45	27.8	Tricosane		0.7	2300
		Phthalic acid, bis (2-ethylhexyl)			
	46 28.3	ester	0.7	7.3	2531
47	29.4	N-Hexacosane		0.9	2600
		1,3-Benzenedicarboxylic acid, bis			
48	29.8	(2-ethylhexyl) ester		0.6	2704
	49 30.4	Squalene	0.3	1.5	2780
	50 30.8	$N$ -Octacosane	0.6	0.7	2800
		Benzenepropanoic acid, 3,5-bis(1,1-			
51	38.5	dimethylethyl)-4-hydroxy-,		4.5	3800
		octadecyl ester			
		Total compounds		91.1% 92.3%	

of Tussilago farfara L.								
	No RT	Components	EL%	EF%	RIc			
	7.1	Trimethylene oxide	2.9		543			
		7.12 Propylene oxide		4.6	477			
3		7.15 3-Chloropropanoic acid		1.6				
	7.19	Heptadecafluorononanoic acid, dodecyl ester		1.7				
		7.21 4-Amino-2-fluoro-N-methylbenzamide		1.3				
6	7.25	<i>n</i> -Dodecanoyl chloride	2.3					
		7.28 Tetrahydroxypteridine	19					

**Table 3. Chemical composition (%) of different organs extracts** 





**Figure 3. Gas chromatography–mass spectrometry (GC–MS) chromatogram of leaves's smoke of** *T. farfara.*







**Figure 5. Comparison of total phenolic compounds (A) and antioxidant activity (B) in all samples with green tea, cloves, black tea and rosemary.**

#### *3.5. Correlation between AA and TPC*

The correlation between AA and TPC in smoke extracts is presented in [Table 4.](#page--1-25) The results showed the

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strongest positive correlation with total AA and TPC  $(r= 0.75, p \le 0.01)$ , [\(Fig.](#page--1-26) 6).

#### **Table 4. Simple correlation among antioxidant, total phenolic content and total flavonoid of smoke extracts.**



and ns significant at 1%, level of probability significant, respectively.



**Figure 6. Correlation among antioxidant and total phenolic content in all samples.**

#### *3.6. Antimicrobial activity of smoke extracts*

[Fig.](#page--1-27) 7 refers to the flower's smoke extract which has 1,2,3,4 and 5 fractions on the plate. Each spot was 20 mg of the sample that dissolved in 200 μl of solvent. For staining, 20 microliters are laid out for each spot. It was observed that *S. aureus* growth was inhibited around samples 2, 3 and 4 [\(Table 5,](#page--1-28) [Fig.](#page--1-29) 8-A) While the growth of *E. coli* has not been inhibited [\(Fig.](#page--1-29) 8-B).



**Figure 7. TLC separation of the flower and leaf smokes of** *T. farfara* **in the present study.**

**Table 5. Antibacterial activity against human pathogens of smoke extracts against** *Staphylococcus aureus* **and** *Escherichia coli*

	Samples Main compounds	Staphylococcus Escherichia		
		aureus	coli	
SL.	Phenol, Catechol, Hydroquinone			
<b>SF</b>	Phenol, O-Cresol, Hydroquinone			



**Figure 8. Detection of the antibacterial activity of smoke of** *T. farfara* **by direct bioautography against** *Staphylococcus aureus* **(A) and** *Escherichia coli* **(B).**

#### **4. Discussion**

The results of extract yields showed a considerable difference in all samples. The differences in the extract yields among these samples might be related to the different availability of extractable components, resulting from the varied chemical composition of plants [\(Sultana](#page--1-30) *et al*., 2009).

A comparison of volatile compounds of the flower, bark and leaves of *T. farfara* between our results in this study and previous studies (Norani *et al*[., 2019;](#page--1-14) [Judzentiene and Budiene, 2011\)](#page--1-31) revealed no similarity between the chemical compositions of the smoke and those of the essential oils. The essential oils were dominated by monoterpene hydrocarbons, oxygenated

monoterpenes, sesquiterpenes, and oxygenated sesquiterpenes, whereas the smoke extracts were marked by phenol derivatives. Catechol is a natural polyphenolic compound that occurs widely in fruits, teas, vegetables, tobacco and some of those used in traditional Chinese medicines [\(Mahdavi](#page--1-32) *et al*., 2018). Catechol has been widely studied because of its biological importance such as antivirus and antioxidant activities and effects on several enzymes (Lin *[et al](#page--1-33)*., [2009\)](#page--1-33). Generally, phenols are well-known as antimicrobial agents and some of these components have been used as disinfectants or as preservatives in cosmetic and food products (Kubo *et al*[., 1995\)](#page--1-34). In a previous study, the smoke of other matters was analyzed by GC-mass and several substances were identified such as dichloromethane, limonene, acetic acid, etc., some antibacterial agents; i.e., phenol, cresol, licochalcone A, carvacrol, etc., some anti-neoplasms such as prospidium chloride [\(Sweetman, 2009\)](#page--1-35), and monoterpenes, diterpenes, sesquiterpenes, antioxidants, etc. [\(Joharchi](#page--1-36) *et al*., 2020).

As a radical scavenging investigation (DPPH) on smoke, [Soares et al.](#page--1-37) (2016) reported that has shown activity by detaching the DPPH free radical with  $IC_{50}$ of 244 ug/mL. The antioxidant activity of smoke depends mostly on its composition, especially in phenolic compounds [\(Soares](#page--1-37) *et al*., 2016).

The antimicrobial effects of smoke may be pertinent to their phenolic and polar compounds [\(Danial](#page--1-4) *et al*., [2018;](#page--1-4) [Fouladi Fard and Farajinia, 2016\)](#page--1-38). Lignin can be converted into phenolic compounds in the form of liquid smoke [\(Chen, 2014\)](#page--1-39). Phenolic compounds have been associated with positive effects on cardiometabolic health, cognition, type II diabetes, obesity, neuroinflammation and others as well as safe and represent a new strategy to for treating skin [\(Mahdavi](#page--1-32) *et al*., 2018).

These results were in agreement wit[h Sim and Nyam](#page--1-40)  [\(2019\),](#page--1-40) [Kho et al. \(2019\)](#page--1-41) and Sim [et al. \(2019\)](#page--1-42) in which a high correlation was observed between antioxidant capacities and phenolic content of kenaf leave tea and *Hibiscus cannabinus* L. leaves. Phenolic compounds are very important natural antioxidants in various plants, which exhibit antioxidant activity through radical scavenging [\(Norani](#page--1-14) *et al*., 2019).

The result of biological activity may be due to the presence of phenols or phenolic compounds in the smoke extract, which is better known as antimicrobial agents [\(Cetin-Karaca and Newman, 2015\)](#page--1-43). In addition, different strains of tuberculosis and non-tuberculous strains such as *M. kyorinense* and *M. kansasii* were inhibited by hydroquinone (Jyoti *et al*[., 2016\)](#page--1-44). Also, some reports have demonstrated that hydroquinone possesses antioxidant properties [\(Yamaguchi](#page--1-45) *et al*., [2006\)](#page--1-45).

Beneficial bioactivities such as the antibacterial activity of phenolic compounds are related to their chemical structures, particularly the presence of an aromatic structure and hydroxyl groups able to neutralize free radicals and other reactive oxygen species (Lima *et al*[., 2019\)](#page--1-46). The published researches are not definite about the differences in the mechanisms of function of phenolic compounds on Gram-positive and Gram-negative bacteria [\(Sanhueza](#page--1-47) *et al*., 2017).

#### **5. Conclusion**

In conclusion, *T. farfara* smoke has relatively high antioxidant potency and total phenolic compounds. Also, a positive correlation between the total phenolic content and antioxidant activity of all smoke extracts was established. The phenolic compounds display a wide perspective of biological factors such as antimicrobial and anticancer activities, as well as protective effects against neurodegenerative diseases. The pleiotropy bioactivities of phenolic compounds are limited by their bioavailability. To deal with this challenge, novel delivery systems like nanoencapsulation an encouraging delivery systems for the effective forwarding and release of phenolic compounds to desired targets.

#### **Abbreviation**

AA= antioxidant activity, TPC= Total phenolic contents, DPPH= 2,2-diphenyl-2 picrylhydrazyl hydrate, TLC= thin layer chromatography

#### **Conflict of Interests**

The authors have to declare their conflict of interest.

#### **Ethics approval and consent to participate**

No human or animals were used in the present research.

#### **Consent for publications**

All authors read and approved the final manuscript for publication.

#### **Availability of data and material**

All the data are embedded in the manuscript.

#### **Authors' contributions**

The first author [M. N.]: performance of the research project and writing the article

The second author [A. C.]: cooperation in the implementation of the research project

The third author [A. A.]: statistical analysis of the data The fourth author [M. A.]: implementation of the research project

#### **Informed Consent**

The authors declare not to use any patients in this research.

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