Extraction, Isolation and Structure Elucidation of Two Phenolic Acids from Aerial Parts of Celery and Coriander

Thukaa Z. Abdul-Jalil*, Ruaa Mohammed Ibrahim and Zahraa Suhail Nassir

Department of Pharmacognosy, College of Pharmacy, University of Baghdad, Baghdad, Iraq. *Corresponding Author E-mail: thukaazuhaira@gmail.com

https://dx.doi.org/10.13005/bpj/2807

(Received: 02 December 2022; accepted: 28 August 2023)

Celery and coriander are vastly applied in modern medicine and traditionally because various medicinal and nutritional benefits depend on their medicinal characteristics. The study aimed to detect, isolate and compare extracts contents of phenolic acids (caffeic and p-coumaric acids) in ethyl acetate fraction of fresh and dry aerial parts of coriander (Coriandrum sativum L.) and celery (Apium graveolens L.) of the Apiaceae family. The extraction of these constituents was carried out by maceration method using 70% ethanol and fractionation was done by using petroleum ether, and ethyl acetate. The existence of caffeic and p-coumaric acids in aerial part extracts of two plants was identified by thin-layer chromatography (TLC) and high-performance liquid chromatography (HPLC). After qualitative determination of these active constituents, high-performance liquid chromatography analysis was also applied for quantification of caffeic and p-coumaric acid and the results showed that ethyl acetate extract of dry coriander showed higher contents of caffeic and p-coumaric acid than ethyl acetate extracts of fresh coriander and celery (dry and fresh). Depending on the results of TLC and HPLC, two bioactive compounds (caffeic acid and p-coumaric acid) were isolated and identified by Fourier transform infrared spectroscopy, proton nuclear magnetic resonance spectroscopy, and full scan product ion mode liquid chromatography mass spectrometry. This result opens the minds to the dry plant era and leads to counter fiction of what was known that fresh plant gives better results than the dry one.

Keywords: Apiaceae; Caffeic acid; Celery; Coriander; P-coumaric acid.

Several therapeutic compounds have been detected in medicinal herbs today. These herbs are recorded in pharmacopeia throughout the world and play an important role in conventional medicine¹. *Coriandrum sativum* L. (known as coriander) is a seasoning plant of the Apiaceae family, with origins in Mediterranean countries. This plant is grown commercially in Asia, Africa, and Europe². All parts of the plant are edible while the common

parts utilized in cooking are the dried seeds and the fresh leaf³. Coriander (see Figure 1) has nutritional components such as protein, water, fiber, fat, ash, sugar, essential oils, and minerals⁴. The main active ingredients of coriander are fatty oils and essential oils⁵. It also contains coumarins, phthalides, phenolic acids (chlorogenic and caffeic acids), and flavonoids (quercetin and rutin)^{6,7}. Oleic, linoleic, palmitic, ascorbic, and stearic acids are also present



in the coriander plant and are very effective in lowering the blood cholesterol level⁸. Coriander is used in Indian folk medicine for the treatment of urinary, digestive, and respiratory disease; it also has diuretic, diaphoretic, stimulant, and carminative activity³. This plant exhibits a strong antioxidant effect because it is an excellent source of phytochemicals and polyphenols². The coriander leaf contains higher antioxidant concentrations than its seeds⁵. It also possesses other pharmacological properties such as anti-mutagenic ⁹, antidiabetic ¹⁰, anti-spasmodic ¹¹, antilipidemic effects ¹², and others.

Apium graveolens L. (known as celery) is an annual plant of the Apiaceae family, native to outlying hills in Afghanistan, Punjab, and the North Western Himalayas¹³. The leaves, roots, and stems are vastly utilized in cooking such as salads, side dishes, and soups. Celery contains furanocoumarin, polyphenols, tannins, steroids, terpenoids, saponins, acids (glutamine, ascorbic, oxalic), vitamins B2, B1, A, PP, B9, B6, K, E, and mineral salts (14). It also contains essential oils as the main active constituents¹⁴. It is applied in traditional medicine to the bladder, kidney diseases, gout, gastric ulcer, gastritis, duodenal gout, obesity, diabetes, dermatitis, nephritis, rheumatism, and prostate inflammation¹⁴. The celery plant has different activities such as antidepressant, antimicrobial, antidiabetic¹³, hepatoprotective, estrogenic, antioxidant, antiestrogenic, cytotoxic^{15-17,} antiinflammatory and analgesic effects^{18,19}.

MATERIALS AND METHODS

Apparatus and Instruments

The following were used, Rotatory evaporator type Buchi attached to a vacuum pump Buchi-Germany), oven (Memmert-854, Buchi-Germany), and an electrically sensitive balance (Sartorius, Germany) with an Ultraviolet light type DESAGA HEIDELBERG of 254 nm and 366 nm waves lengths type DESAGA-Germany. Fourier transforms infrared spectra (FT-IR) spectra were scanned on Jusco/ Japan FT-IR-4200 at the University of Baghdad, College of Pharmacy, 1H NMR was carried out in al-Albayt University, Al-Mafraq, Jordan (Euro-vector EA 3000A/ Italy), LC/MS was carried out in Iraqi National Center for Drug Control and Research (LCMS-8040 series system, Shimadzu/ Japan) and HPLC analysis was carried out using SYKAM/ Germany at the directorate of the ministry of science and technology/environment and water center.

Chemicals and Reagents

Standard of p-coumaric acid and Caffeic acid were purchased from Biopurfy-China, Silica gel of GF _{254nm} with a thickness of 0.25 mm MERCK-USA while the 0.75mm from Sanpont-China, absolute both ethanol and methanol, HPLC grade acetonitrile, the ethyl acetate and petroleum ether were purchased from Schar lab S.L.-Spain and Toluene was purchased from the CDH-India while the formic acid was from GCC, U.K.



Fig. 1. Photo of Iraqi Coriandrum sativum(A) and Iraqi Apium graveolens(B)

Plant material

The aerial parts of the plants (*Apium graveolens* L. and *Coriandrum sativum*) were collected from Baghdad city, which is located in the center of Iraq, in December 2021. Some aerial parts of each plant were dried and the other aerial parts were used fresh.

Plant collection licenses

We have received ethical approval and permission to collect celery and coriander from the Pharmacognosy Department at Baghdad University, and Iraqi Medical Research Center 1120,2020.

Extraction of phenolic acids

The extraction was performed according to the process with some modifications described by Iswantini *et al.* ^{20,21}. The extraction was carried

out for dry and fresh aerial parts (leaves and stem) of *Apium graveolens* L. and *Coriandrum sativum* L., taking (100 gm) of each dry plant and fresh parts and then macerated with 70 % ethanol (1000 ml) at room temperature separately. The extracted crudes were concentrated under reduced pressure at 50 ° C. Then successively fractionated with petroleum ether (250 ml*2) and ethyl acetate (250 ml*2). The ethyl acetate fractions were concentrated separately to obtain four fractions of ethyl acetate for each part of the plant.

Identification of phenolic acids by thin-layer chromatography (TLC)

The preliminary analysis of caffeic acid $(C_9H_8O_4)$ and p-coumaric acid $(C_9H_8O_3)$ in ethyl acetate fractions from fresh and dry aerial parts of *Apium graveolens* and *Coriandrum sativum* was achieved by TLC.

Table 1. HPLC conditions

Mobile phase(isocratic)	Acetonitrile: Water (1:1)
Column	Shimadzu LC C18 (250 mm x 4.6 mm, 5 µm particle size)
Column temperature	ambient
Flow rate	1ml / min
Injection volume	20ìL
Injection concentration	1 mg/ml
Detection	UV detector at ë 265 nm
Sample	Ethyl acetate fraction
Standards	Caffeic acid and p-coumaric acid

Table 2. The percentages of ethyl acetate fractions of dry and fresh aerial parts of *Coriandrum sativum* and *Apium graveolens*

Analyzed fraction	Percentage yield (%w/w)	
Fresh Coriandrum sativum (FC)	2.6 %	
Dry Coriandrum sativum (DC)	4.7 %	
Fresh Apium graveolens (FA)	2.4 %	
Dry Apium graveolens (DA)	3.5 %	

The GF₂₅₄ plate of silica gel (readymade) was used as the stationary phase and toluene: ethyl acetate: formic acid (36:12:5) as developing solvent ²¹. The detection of developed plates is done first by visualizing under UV light at 254nm, then spraying with a chemical reagent for phenolic compounds such as ferric chloride (0.5%). The value of the retardation factor (Rf value)of the separated spots of caffeic and p-coumaric acids was calculated and matched to that of their standard.

Table 3. The Rf values of separated phenolic acids (caffeic and p-coumaric acids) and their standards in one solvent system in TLC

Rf value	Standards	Fresh Coriandrum sativum (FC)	Dry Coriandrum sativum (DC)	Fresh Apium graveolens (FA)	Dry Apium graveolens (DA)
caffeic acid	0.371	0.371	0.371	0.371	0.371
p-coumaric acid	0.513	0.513	0.513	0.513	0.513

Qualitative and quantitative estimations of phenolic acids by high-performance liquid chromatography (HPLC)

By HPLC (table 1), qualitative analysis of caffeic and p-coumaric acids in ethyl acetate fractions of celery and coriander plants is performed by matching the retention time of caffeic and p-coumaric acids in all samples with those of their standard at identical chromatographic conditions ²².

In HPLC, quantitative estimations of p-coumaric and caffeic acids in ethyl acetate fractions of celery and coriander plants are made by drawing a calibration chart for which serial dilutions of the standard caffeic and p-coumaric acids were made by methanol (10, 20,30, and 40 ppm).

Isolation and purification of phenolic acids by preparative TLC plates

The isolation of caffeic and p-coumaric acids from the ethyl acetate fraction of the dry

coriander aerial part(DC) is achieved by using preparative TLC. TLC plates (silica gel GF₂₅₄ with thick 0.75mm and 20*20) were used and activated at 120°C before the application of the sample. The ethyl acetate fraction of DC is seen by a capillary tube on the plate in the form of a row and, according to standard processes, allowed the plate to develop in toluene: ethyl acetate: formic acid (36:12:5) as a solvent system. Under UV at 254nm, the separated bands were identified and with a needle marked. The two bands were rubbed off with standards similar to those of the p-coumaric and caffeic acids standards put in a beaker and then adding a sufficient quantity of absolute methanol, the flasks were later shaken on a warm water bath, kept cool in a refrigerator for crystallization about 4-5 hrs. The crystalline constituent was then filtered through double filter paper and dried for a pure constituent 23.

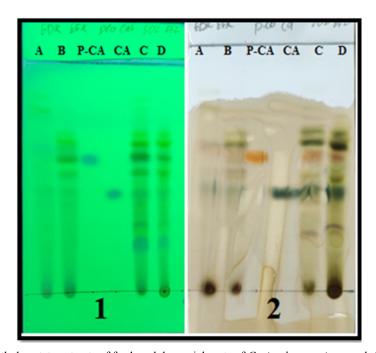


Fig. 2. TLC of ethyl acetate extracts of fresh and dry aerial parts of *Coriandrum sativum* and *Apium graveolens* using adsorbent (silica gel GF254) and a mobile phase (toluene: ethyl acetate: formic acid (36:12:5)). Detection by UV light at (1) 254 nm, (2) after spraying with ferric chloride (0.5%). (CA: caffeic acid standard, P-CA: p-coumaric acid standard, A: Dry *Apium graveolens* (DA), B: Fresh *Apium graveolens* (FA), C: Dry *Coriandrum sativum* (DC), D: Fresh *Coriandrum sativum* (FC))

Identification and characterization of the isolated constituents

Fourier transforms infrared (FT-IR) spectroscopy

The FT-IR spectroscopy was performed in Baghdad University/College of Pharmacy/Department of Pharmacognosy and Medicinal plants for each isolated constituent by using a KBr disc and then the structural assignments correlated for characteristic bands as illustrated in the results.

1H nuclear magnetic resonance spectroscopy

(NMR) analysis The spectra of the NMP were taken

The spectra of the NMR were taken by dissolving the sample in dimethyl sulfoxide

(DMSO) – d6 and later on running the NMR Spectrometer. All the chemical shifts were reported with tetramethyl silane (TMS) reference at 0 parts per million (ppm). NMR measurement was carried out at Al-Albayt University/Al-Mafraq/Jordan and the Euro-vector EA 3000A NMR spectrometer apparatus (300MHz for 1H-NMR) was used. Chemical shifts are given on a ä (ppm) scale with TMS as the internal standard.

Liquid chromatography/ mass spectroscopy LC/MS

Analytical LC/MS was done by using Shimadzu-LC / MS-8040 series system and coupled to a mass Spectrometer-Shimadzu with

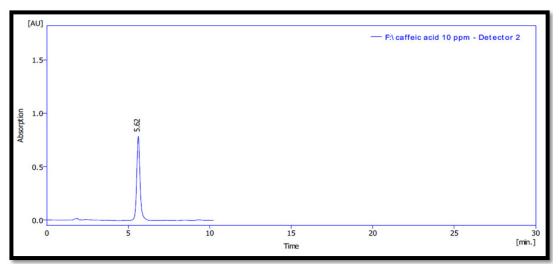


Fig. 3. HPLC analysis of caffeic acid standard

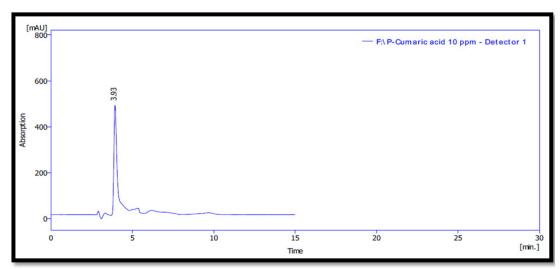


Fig. 4. HPLC analysis of p-coumaric acid standard

an electrospray interphase (ESI). The caffiec and p-coumaric acids isolated from *Coriandrum sativum* dried aerial parts were performed by C18, pore size 3.5 im, length 15 cm, id 4.6im column, the column was maintained at 42C⁰ temperature, the mobile phase was composed of 0.1% acetic acid and acetonitrile (V/V), the flow rate was 0.8 ml/min and the isocratic mode of elution was used. The volume of the injected standard and sample was 5 iL and Negative ionization mode was applied for structural elucidation ^{24,25}.

Preparation of the isolated compounds

Two milligrams of caffeic acid and 2 mg of p-coumaric acid were dissolved in 2 mL

of ethanol separately and after that filtrated using disposable filters of 0.45 im pore size before utilization of LC/MS for analysis.

MS conditions

The mass analysis was performed by a Shimadzu mass spectrometer with an electrospray interphase (ESI) and then Chem Station was used for the chromatographic data processing. MS data were acquired in both modes the negative and positive ionization at a defined condition that is: nitrogen gas flow rate 3 l/min, heat block temperature 400 °C, desolvation line (DL) temperature 250 °C, drying gas flow rate 15 l/min and mass spectra were recorded using the fill scan mode in the range of 100-800 Daltons ²⁶.

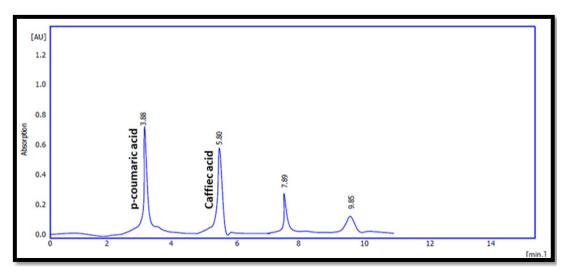


Fig. 5. HPLC analysis of ethyl acetate fractions of dry Apium graveolens

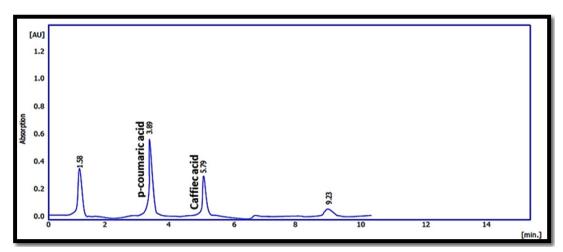


Fig. 6. HPLC analysis of ethyl acetate fractions of fresh Apium graveolens

RESULTS AND DISCUSSION

Caffeic and p-coumaric acids are curative phenolic acids that are used in nutrition and medicine. However, recent studies have reported their many pharmacological effects for the treatment of various disorders. *Apium graveolens* (Celery) and *Coriandrum sativum*(Coriander) are readily available in the supermarket, and they have many phytochemicals that are widely known for

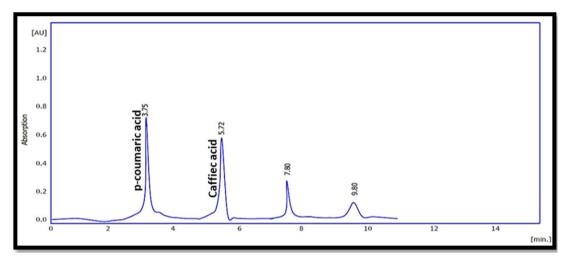


Fig. 7. HPLC analysis of ethyl acetate fractions of dry Coriandrum sativum

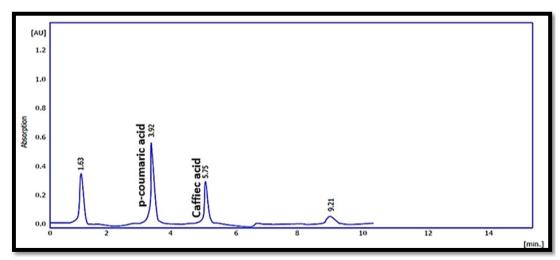


Fig. 8. HPLC analysis of ethyl acetate fractions of fresh Coriandrum sativum

Table 4. Concentration of caffeic and p-coumaric acids in fresh and dry aerial parts of coriander and celery

Ethyl acetate fraction	The concentration of caffeic acid (ppm)	Concentration of p-coumaric acid (ppm)	
Dry Coriandrum sativum (DC)	166.911	247.615	
Fresh Coriandrum sativum (FC)	166.883	211.357	
Dry Apium graveolens (DA)	120.878	181.256	
Fresh Apium graveolens (FA)	112.365	165.021	

their various applications in the medical field and other uses.

The percentage yields (%w/w) of each ethyl acetate fraction for dry and fresh aerial parts of *Apium graveolens* and *Coriandrum sativum* are shown in Table 2.

Identification of phenolic acids by TLC

The identification of caffeic and p-coumaric acids was achieved by TLC in aerial parts of the ethyl acetate extracts of *Apium graveolens* and *Coriandrum sativum* as shown in Figure (2).

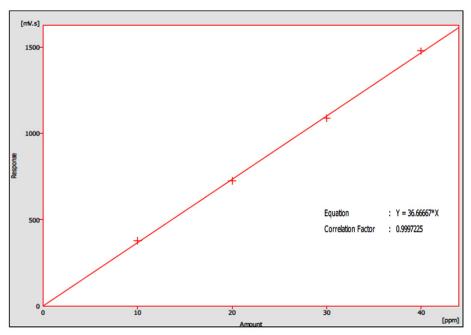


Fig. 9. HPLC calibration curve of the caffeic acids standard

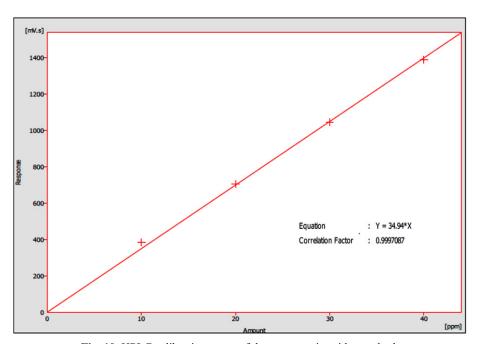


Fig. 10. HPLC calibration curve of the p-coumaric acids standard

After observation of the spots under UV at 254 nm and by spraying with ferric chloride (0.5%), caffeic and p-coumaric acids in plant samples are detected by calculating their Rf values as shown in Table (3)

Qualitative and quantitative estimations of phenolic acids by HPLC

HPLC was performed to determine the existence and quantity of caffeic and p-coumaric acids in the extracts of *Apium graveolens* and *Coriandrum sativum* (fresh and dry aerial parts).

Table 5. FTIR absorption bands (cm⁻¹) of the isolated caffeic acid

Functional group	Number of group frequency waves (in cm ⁻¹)		Main attribute
	Isolated caffeic acid	Caffeic acid standard	
Phenol O-H	3432.67	3409.5	O-H vibration of stretching
Benzene Ring	3237.9	3234.04	C-H stretching of the aromatic ring
С-Н	2791.46, 2702	2831.95	Asymmetric and symmetric stretching of CH2
C=O	1644.02	1644.98	C=O stretching vibration
C=C	1618.95, 1513.12	1618.95, 1529.27	C=C stretching of the aromatic ring
О-Н	1449.24	1449.24	O-H bending of the carboxyl group
Phenol-O-H	1367.28	1352.82	O-H bending vibration
Phenol-O-H	1216.86	1216.86	C-OH stretching vibration
C=C	974.84	974.84	C=C out-plane bending vibration
С-Н	850,816,736,649	849,815,736,647	Benzene ring C-H out-plane bending vibration

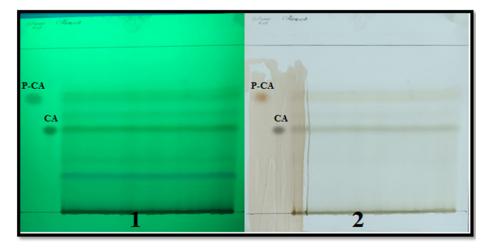


Fig. 11. Preparative TLC plates of the ethyl acetate fraction of the dry aerial part of coriander (sample = DC) developed in toluene: ethyl acetate: formic acid (36:12:5) at 254 nm (1) and after spraying with0.5% ferric chloride (2), CA: caffeic acid standard, P-CA: p-coumaric acid standard

For qualitative estimations, the result revealed that the retention time of p-coumaric and caffeic acids in different ethyl acetate extracts and the retention time of their standards were similar.

This proved the presence of p-coumaric and caffeic acids in the samples as shown in Figures (3-8).

In quantitative estimations of p-coumaric and caffeic acids in fractions analyzed from

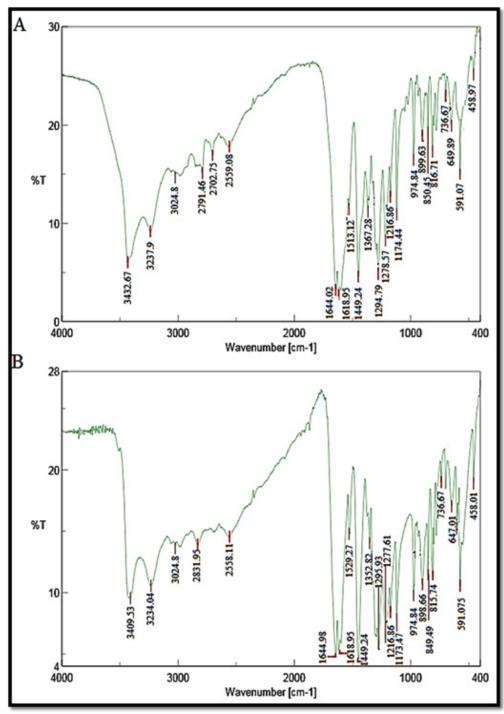


Fig. 12. FTIR spectra of A= isolated caffeic acid and B = standard caffeic acid

coriander and celery plants by HPLC, a plot of the area vs. standard concentration for both caffeic and p-coumaric acids show a linear fit. The concentration of these phenolic acids in each fraction was determined by using a straight-line equation as in Figures (9,10).

The coriander and celery plants had shown different concentrations of p-coumaric and caffeic acids, as shown in Table 4.

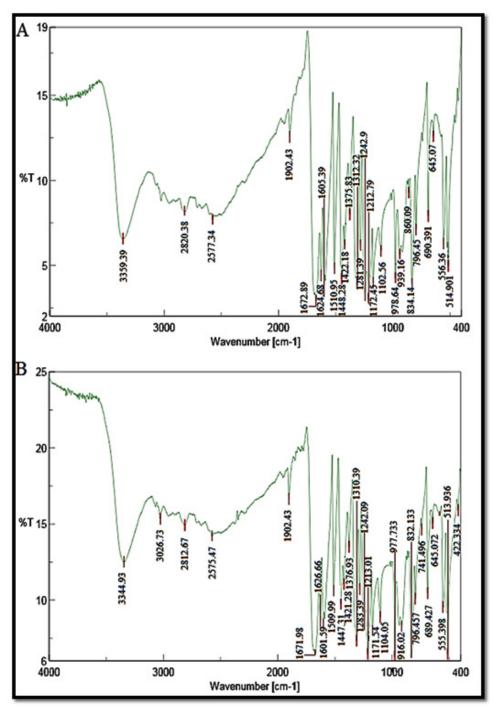


Fig. 13. FTIR spectra of A= isolated p-coumaric acid and B= p-coumaric acid standard.

The results indicated that the concentration of p-coumaric and caffeic acids is higher in the ethyl acetate fraction of the dry coriander aerial part (DC) than their concentrations in other ethyl acetate fractions. Therefore, p-coumaric and caffeic acids are isolated by preparative TLC from the ethyl acetate fraction of DC.

The result revealed that the HPLC method was efficient for the quantitative and qualitative determination of caffeic and p-coumaric acids.

Isolation of phenolic acids by preparative TLC plates

The development of the ethyl acetate fraction of the aerial part of dry coriander in

Table 6. FTIR (cm-1) absorption bands of isolated p-coumaric acid

Functional group	Number of frequency wav Isolated p-coumaric	p-coumaric acid	Main attribute
	acid	standard	
Phenol O-H	3359.39	3344.93	O-H stretching vibration
C=C-H	3024.37	3026.73	C=C-H stretching of the aromatic ring
С-Н	2820.38	2812.67	Asymmetric and symmetric stretching of CH2
C=O	1672.89	1671.98	C=O stretching vibration
C=C	1510.95	1509.99	C=C stretching of the aromatic ring
О-Н	1448.28	1447.31	O-H bending of the carboxyl group
Phenol-O-H	1375.83	1376.93	O-H bending vibration
Phenol-O-H	1212.79	1213.01	C-OH stretching vibration
C=C	978.64	977.73	C=C out-plane bending vibration
С-Н	860,834,796,690	832,796,689	Benzene ring C-H out-plane bending vibration

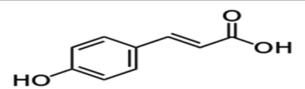


Table 7. Assignment of 1H-NMR spectral data of isolated caffeic acid

Ppm	No. of protons	Multiplicity	Assignment	
12.13	3 1H	Singlet broad	ОС-ОН	
9.54	1H	Singlet broad	С4-ОН	
9.13	1H	Singlet broad	С3-ОН	
7.43	1H	Doublet (J value:15.822 Hz)	С7-Н	
7.01	2H	Doublet(J value:22.793 Hz)	С2-Н	
6.97	2H	Doublet(J value:8.075 Hz)	С6-Н	
6.74	1H	Doublet(J value:8.091 Hz)	С5-Н	
6.19	1H	Doublet(J value:15.846 Hz)	С8-Н	

toluene: ethyl acetate: formic acid (36:12:5) in preparative TLC plates results in the number of bands that represent several compounds as in Figure 11.

Identification and characterization of the isolated constituents

The isolated constituents that result from preparative TLC were subjected to different

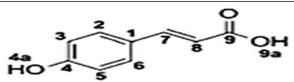
analysis identification techniques to determine their purity of them in addition to other several techniques that were applied to obtain the structural elucidation. These techniques include the following:

FTIR

The IR spectra of isolated caffeic and p-coumaric acids that gave similar results were

Table 8. Assignment of 1H-NMR spectral data of isolated p-coumaric acid

Ppm	No. of protons	Multiplicity	Assignment
12.15	1H	Singlet broad	ОС-ОН
9.96	1H	Singlet broad	С4-ОН
7.53	1H	Doublet(J value:8.466 Hz)	С3-Н, С5-Н
7.47	1H	Doublet(J value:15.911 Hz)	С7-Н
6.80	1H	Doublet(J value:8.477 Hz)	C2-H, C6-H
6.26	1H	Doublet(J value:15.917 Hz)	С8-Н



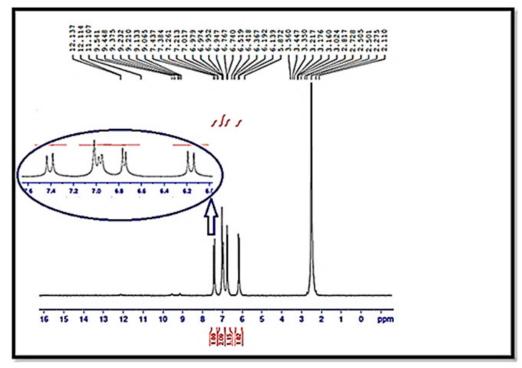


Fig. 14. 1H-NMR spectrum of isolated caffeic acid

matched with their standard; which indicated that isolated compounds are p-coumaric and caffeic acids as shown in Figures (12,13).

The isolated compounds' characteristic FTIR absorption bands with their standards (p-coumaric and caffeic acids) are listed in Tables 5 and 6, respectively^{27,28}.

Nuclear magnetic resonance (NMR) spectroscopy

Is a sophisticated and powerful analytical technique for the elucidation of organic compound

structures and identification of molecular interaction $^{\rm 28}$

For that reason, the identification of isolated, p-coumaric, and caffeic acids was further confirmed by NMR spectroscopy application. The isolated constituents' NMR spectra are in good agreement with works of literature on p-coumaric and caffeic acids as shown in Figures 14,15 and Tables 7 and 8 ^{26,29,30}.

Table 9. The specific ion of isolated caffeic acid and p-coumaric acid was monitored in the screening method

Isolated compounds	R _t (min)	Theoretical molecular weight (M)	Calculated molecular weight (M)	Fragmented ions (m/z)
Caffeic acid	5.6	180.16	180	179(M-H), 135(M-H-CO ₂), 161(M-H-H ₂ O), 134(M-H-HCO ₃)
p-coumaric acid	8.7	164.158	164	163(M-H), 119(M-H-CO ₂),113, and 101 (dissociation of the phenolic ring)

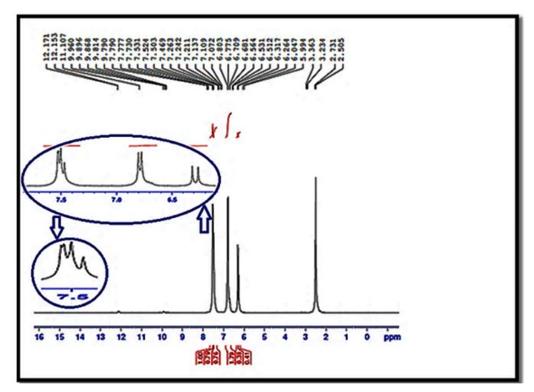


Fig. 15. 1H-NMR spectrum of isolated p-coumaric acid

LC/MS

The identification of isolated caffeic acid and p-coumaric acids were further confirmed by LC/MS. For structural scanning, full scan product ion was used instead of multiple reaction monitoring (MRM) since it was used for qualitative applications to obtain structural information.

According to the chromatographic conditions, Rt (retention times) of the analyzed constituents are 5.6 min for caffeic acid and 7.8 min for p-coumaric acid; in addition, the ions monitored by the MS method are illustrated in Table 9.

In the ionization conditions, most of the phenolic acids lost a molecule of COO (44 units) so ions detected by spectrometer are always in the form of [M-H-COO] . All these data are in good agreement with the works of literature on p-coumaric and caffeic acids³¹⁻³³.

These assignments were proved by full product ion scan liquid chromatography connected with a negative (-ve) ES ionization spectrum of caffeic acid (Figure 16) and p-coumaric acid (Figure 17).

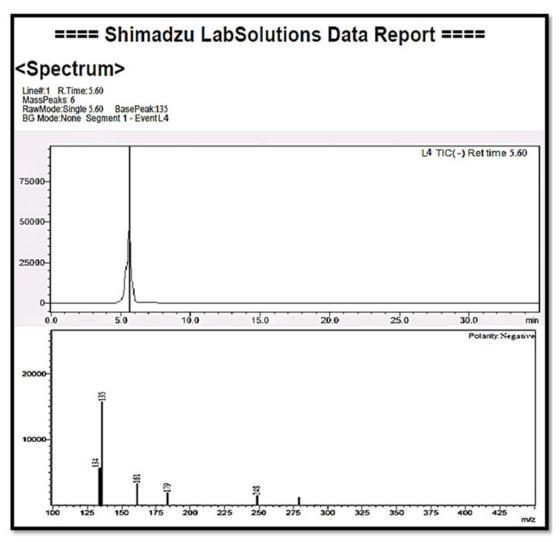


Fig. 16. Full product ion scan LC chromatogram and mass spectrum of isolated caffeic acid.

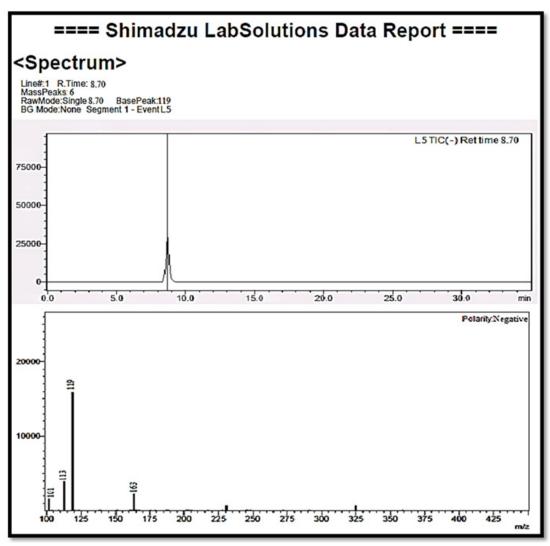


Fig. 17. Full product ion scan LC chromatogram and mass spectrum of isolated p-coumaric acid

CONCLUSION

Coriander (Coriandrum sativum L.) and celery (Apium graveolens L.) are excellent sources of caffeic and p-coumaric acids (phenolic acids). These compounds are considered a substantial class of natural components that are widely used in several countries as an ingredient in many herbal remedies and preparation. This work shows that the aerial parts extracts of the coriander and celery plant contain caffeic and p-coumaric acids, and quantitative estimation by HPLC shows that the concentrations of caffeic and p-coumaric acids

in the dry coriander are higher than in the fresh coriander and celery (dry and fresh). According to the HPLC result, the caffeic and p-coumaric acids were isolated from dry coriander by using the preparative TLC method. This study confirms that coriander showed significantly higher p-coumaric and caffeic acids contents than celery.

ACKNOWLEDGEMENT

None.

Conflict on Interest

There is no conflict of interest.

Funding Source

There are no funding sources.

REFERENCES

- Nhut PT, Quyen NT, Truc TT, Minh LV, An TN, Anh NH. Preliminary study on phytochemical, phenolic content, flavonoids, and antioxidant activity of *Coriandrum sativum* L. originating in Vietnam. *InIOP Conference Series: Materials* Science and Engineering 2020;991(1):012-022.
- Bhat S, Kaushal P, Kaur M, Sharma K. Coriander (Coriandrum sativum L.): Processing, nutritional and functional aspects. Afr J Plant Sci 2014; 8:5-33.
- 3. Pathak NL, Kasture SB, Bhatt NM. Phytochemical screening of *Coriander sativum* Linn. *Int J Pharm Sci Rev Res* 2011; 9:159-163.
- Ahmed EH, Abadi RS, Mohammed AM. Phytochemical screening, chemical composition and antioxidant activity of seeds essential oil of *Coriandrum sativum* L. from the Sudan. *International Journal of Herbal Medicine* 2018; 6(1): 01-04.
- Karmakar UK, Rahman MA, Roy DN, Sadhu SK, Ali ME. Chemical and biological investigations of *Coriandrum sativum* Linn. *IJPSR* 2011; 2(4): 999-1006.
- Ghani A. Medicinal plants of Bangladesh, 2nd Ed. The Asiatic Society of Bangladesh, Dhaka, 2003; pp 382.
- Rathore SS, Saxena SN, Balraj S. Potential health benefits of major seed spices. *Int J Seed Spices* 2013; 3:1-12.
- 8. Cortes-Eslava J, Gomez-Arroyo S, Villalobos-Pietrini R. Antimutagenicity of Coriander (*Coriandrum sativum*) juice on the mutagenesis produced by plant metabolites of aromatic amines. *J Toxicol Lett* 2004; 153:283-292.
- 9. Eidi M, Eidi A, Saeidi A, Molanaei S, Sadeghipour A, Bahar M et al. Effect of Coriander seed (*Coriandrum sativum* L) ethanol extract on insulin release from pancreatic beta cells in streptozotocin-induced diabetic rats. *J Phytother Res* 2012; 23:404-406.
- Alison MG, Peter RF. Insulin-releasing and insulin-like activity of the traditional antidiabetic plant *Coriander sativum* (coriander). *British J Nutr*1999; 81:203-209.
- 11. Sunil C, Agastian P, Kumarappan C, Ignacimuthu S. *In vitro* antioxidant, antidiabetic and antilipidemic activities of *Symplocos cochinchinensis* (Lour.) S. Moore bark. *J Food Chem Toxicol* 2012; 50:1547-1553.
- 12. Nair SN, Varghese A, Meenu B, Rejimon G, Neeraja E D. Comparative evaluation of *Coriandrum sativum Linn*. and *Apium graveolens* for Antimicrobial activity. *Research J. Pharm and Tech* 2017; 10(2):541-544.

- Sarmanovna TZ. Phytochemical Study of Odorous Celery Root (*Apium graveolens* L.) Grown in the North Caucasus. *Pharmacogn J* 2019; 11(3): 527-530.
- Uddin Z, Shad AA, Bakht J, Ullah I, Jan S. In vitro antimicrobial, antioxidant activity, and phytochemical screening of *Apium graveolens*. *Pak J Pharm Sci* 2015; 28 (5):1699-1704.
- Popovic M, Kaurinovic B, Triviv S, Mimica-Dukic N, Bursac M. Effect of celery (*Apium graeolens*) extracts on some biochemical parameters of oxidative stress in mice treated with carbon tetrachloride. *Phytother* 2006; 20: 531-536.
- Edziri H, Ammar S, Souad L, Mahjoub MA, Aouni M, Mighri Z, Verschaeve L. *In vitro* evaluation of antimicrobial and antioxidant activities of some Tunisian vegetables. *South Afri J Bot* 2012; 78: 252-256.
- Shad AA, Shah HU, Bakht J, Choudhary MI, Ullah J. Nutraceutical potential and bioassay of *Apium graveolens* L. grown in Khyber Pakhtunkhwa-Pakistan. *J Med Plant Res* 2011; 5: 5160-5166.
- Asgarpanah J and Kazemivash N. Phytochemistry, pharmacology and medicinal properties of Coriandrum sativum L. African Journal of Pharmacy and Pharmacology 2012; 6(31): 2340-2345.
- R.A. Mans D, Grant A, Pinas N. Plant-Based Ethnopharmacological Remedies for Hypertension in Suriname. *Herbal Medicine* 2017.
- 20. Iswantini D, Ramdhani TH, Darusman LK. *In vitro*, inhibition of celery (*Apium graveolens* l.) extract on the activity of xanthine oxidase and determination of its active compound. *Indo J Chem* 2012, 12 (3): 247-254.
- Medic-Saric M, Jasprica I, Smoicic-Bubalo A, Mornar A. Optimization of chromatographic conditions in thin layer. *Croatica Chemica Acta* 2004; 77(1-2):361-366.
- 22. Shah Sh M, Abdul-Jalil Th Z. Qualitative and Quantitative Estimation or Chemical Constituents from Leaves and Roots of Iraqi *Agave attenuata* by GC-MS and RP-HPLC. *Iraqi J Pharm Sci* 2022; 31(Sppl):75-85.
- Dubber MJ, Kanfer I. High-performance liquid chromatographic determination of selected flavonols in *Ginkgo biloba* solid oral dosage forms. *J Pharm Pharm Sci* 2004;7(3):303-309.
- Adams M, Weidenmann M, Tittel G, Bauer R. HPLC-MS trace analysis of atropine in *Lycium barbarum* berries. *Phyto Chem Anal* 2006; 17: 279-283.
- 25. MoCan A, Vlase L, Letitia AA, Vodnar D,

- Bischin C, Dumitrescu R.S., Crisan G. HPLC/MS analysis of Caffeic and chlorogenic acids from three Romanian Veronica species and their antioxidant and antimicrobial properties. *Farmacia* 2015; 63(6): 890-896.
- Kajdzanoska M, Gjamovski V, Stefova M. HPLC-DAD-ESI-MS Identification of phenolic compounds in cultivated Strawberry from Macedonia. *Maced J Chem Chem Eng* 2010; 29(2): 181-194.
- 27. Al-Hussaniy HA, Al-Kuraishy HM, Abdulameer AG. The Use of Panax Ginseng to Reduce the Cardiotoxicity of Doxorubicin and Study its Effect on Modulating Oxidative Stress, Inflammatory, and Apoptosis Pathways. *Macedonian Journal of Medical Sciences* 2022; 10:715-719.
- Viswannth A, Rao L and Babu P. Simple isolation and characterization of p-coumaric acid from *Cynodon dectylon* Linn (pers). *IJPAR* 2017; 6(2): 314-318.
- 29. Wong KC. Review of NMR spectroscopy: basic principle, concepts and application in chemistry. *J Chem Educ* 2014; 91: 1103-1104.

- 30. Lopez-Martinez LM, Ortega HS, Navarro RE, Mundo RR, Aguilar GC. A HNMR investigation of the interaction between phenolic acids found in Mango (*Manguifera indica* VC Ataulfo) and Papaya (*Carica papaya* CV Maradol) and 1,1-diphenyl-2,2 picrylhydrazal (DPPH) free radicals. *PLOS One* 2015; 11: 1-11.
- 31. Swislocka R, Sadowy MK, Kalinowska M, Lewandowski W. Spectroscopic (FTIR, FT-Raman, 1H and 13C NMR and theoretical studies) of p-coumaric acid and alkali metal p-coumarates. *Spectroscopy* 2012; 27: 35-48.
- Hossain MB, Rai DK, Brunton NP, Martin-Diana AB, Barry-Ryan C. Characterization of phenolic composition in Lamiaceae species by LC-ESI-MS/MS. J Agric Food Chem 2010; 58(19): 10576-10581.
- 33. Wv Z, Ma X, Fang D, Qi H, Ren W, Zhang G. Analysis of caffeic acid derivative from *Osmanthus xunnanensis* using electrospray ionization quadrupole time-of-flight mass spectrometry. *EJMS* 2009; 15:415-429.